

3-アミノフェノールのマウスを用いた  
経口投与によるがん原性試験（混水試験）報告書

試験番号：0712

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**TABLE A 1**

**SURVIVAL ANIMAL NUMBERS: MALE**

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
REPORT TYPE : A1 104  
SEX : MALE

# SURVIVAL ANIMAL NUMBERS

PAGE : 1

| Group Name | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|------------|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|            |                     | 0                      | 1              | 2              | 3              | 4              | 5              | 6              | 7              | 8              | 9              | 10             | 11             | 12             | 13             |
| Control    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 625 ppm    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 2500 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |

Number of survival/ Number of effective animals  
Survival rate(%)

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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ1j [Crj:BDF1]  
 REPORT TYPE : A1 104  
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 2

| Group Name   | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|--|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|  |                     | 14                     | 15             | 16             | 17             | 18             | 19             | 20             | 21             | 22             | 23             | 24             | 25             | 26             | 27             |
| Control  | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 625 ppm  | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 2500 ppm   | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| Number of survival/ Number of effective animals<br>Survival rate (%) |                     |                        |                |                |                |                |                |                |                |                |                |                |                |                |                |

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STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104  
SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 3

| Group Name | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|------------|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|            |                     | 28                     | 29             | 30             | 31             | 32             | 33             | 34             | 35             | 36             | 37             | 38             | 39             | 40             | 41             |
| Control    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  |
| 625 ppm    | 50                  | 49/50<br>98.0          | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 2500 ppm   | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |

Number of survival/ Number of effective animals  
Survival rate (%)

(HAN360)

BAIS4



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Cri:BDF1]  
 REPORT TYPE : A1 104  
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 4

| Group Name  | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |               |               |               |               |
|---|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|---------------|---------------|---------------|---------------|
|   |                     | 42                     | 43             | 44             | 45             | 46             | 47             | 48             | 49             | 50             | 51             | 52            | 53            | 54            | 55            |
| Control   | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0 | 49/50<br>98.0 | 49/50<br>98.0 | 48/50<br>96.0 |
| 625 ppm   | 50                  | 48/50<br>96.0          | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 48/50<br>96.0  | 47/50<br>94.0  | 47/50<br>94.0  | 47/50<br>94.0  | 47/50<br>94.0  | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 |
| 1250 ppm  | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0 | 49/50<br>98.0 | 49/50<br>98.0 | 49/50<br>98.0 |
| 2500 ppm  | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0 | 49/50<br>98.0 | 49/50<br>98.0 | 49/50<br>98.0 |
| Number of survival/ Number of effective animals<br>Survival rate(%) |                     |                        |                |                |                |                |                |                |                |                |                |               |               |               |               |

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BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 REPORT TYPE : A1 104  
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 5

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |       |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|            |                     | 56                     | 57    | 58    | 59    | 60    | 61    | 62    | 63    | 64    | 65    | 66    | 67    | 68    | 69    |
| Control    | 50                  | 48/50                  | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 |
|            |                     | 96.0                   | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 94.0  |
| 625 ppm    | 50                  | 47/50                  | 47/50 | 47/50 | 46/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 | 44/50 | 44/50 |
|            |                     | 94.0                   | 94.0  | 94.0  | 92.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 88.0  | 88.0  | 88.0  | 88.0  |
| 1250 ppm   | 50                  | 49/50                  | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 |
|            |                     | 98.0                   | 98.0  | 98.0  | 98.0  | 96.0  | 96.0  | 96.0  | 96.0  | 94.0  | 94.0  | 94.0  | 94.0  | 94.0  | 94.0  |
| 2500 ppm   | 50                  | 49/50                  | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 |
|            |                     | 98.0                   | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 96.0  | 96.0  |

Number of survival/ Number of effective animals  
 Survival rate(%)

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BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 REPORT TYPE : A1 104  
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 6

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |       |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|            |                     | 70                     | 71    | 72    | 73    | 74    | 75    | 76    | 77    | 78    | 79    | 80    | 81    | 82    | 83    |
| Control    | 50                  | 47/50                  | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 45/50 | 45/50 | 45/50 | 44/50 | 43/50 | 43/50 | 43/50 |
|            |                     | 94.0                   | 94.0  | 94.0  | 94.0  | 94.0  | 94.0  | 92.0  | 90.0  | 90.0  | 90.0  | 88.0  | 86.0  | 86.0  | 86.0  |
| 625 ppm    | 50                  | 44/50                  | 44/50 | 44/50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 43/50 | 43/50 | 42/50 | 42/50 | 42/50 | 42/50 |
|            |                     | 88.0                   | 88.0  | 88.0  | 88.0  | 88.0  | 86.0  | 86.0  | 86.0  | 86.0  | 86.0  | 84.0  | 84.0  | 84.0  | 84.0  |
| 1250 ppm   | 50                  | 47/50                  | 47/50 | 47/50 | 46/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 |
|            |                     | 94.0                   | 94.0  | 94.0  | 92.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 90.0  | 88.0  | 88.0  |
| 2500 ppm   | 50                  | 48/50                  | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 | 47/50 | 46/50 |
|            |                     | 96.0                   | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 94.0  | 94.0  | 92.0  |

Number of survival/ Number of effective animals  
 Survival rate(%)

(HAN360)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 REPORT TYPE : A1 104  
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 7

| Group Name                                      | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |       |       |       |       |       |       |       |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|   |                     | 84                     | 85    | 86    | 87    | 88    | 89    | 90    | 91    | 92    | 93    | 94    | 95    | 96    | 97    |
| Control   | 50                  | 42/50                  | 41/50 | 41/50 | 40/50 | 38/50 | 38/50 | 37/50 | 35/50 | 33/50 | 33/50 | 32/50 | 31/50 | 29/50 | 28/50 |
|   |                     | 84.0                   | 82.0  | 82.0  | 80.0  | 76.0  | 76.0  | 74.0  | 70.0  | 66.0  | 66.0  | 64.0  | 62.0  | 58.0  | 56.0  |
| 625 ppm   | 50                  | 42/50                  | 41/50 | 41/50 | 41/50 | 41/50 | 40/50 | 39/50 | 38/50 | 38/50 | 38/50 | 38/50 | 38/50 | 37/50 | 37/50 |
|   |                     | 84.0                   | 82.0  | 82.0  | 82.0  | 82.0  | 80.0  | 78.0  | 76.0  | 76.0  | 76.0  | 76.0  | 76.0  | 74.0  | 74.0  |
| 1250 ppm  | 50                  | 44/50                  | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 41/50 | 40/50 | 40/50 | 38/50 | 38/50 |
|   |                     | 88.0                   | 88.0  | 88.0  | 88.0  | 88.0  | 88.0  | 86.0  | 86.0  | 86.0  | 82.0  | 80.0  | 80.0  | 76.0  | 76.0  |
| 2500 ppm  | 50                  | 46/50                  | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 | 45/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 |
|   |                     | 92.0                   | 92.0  | 92.0  | 92.0  | 92.0  | 92.0  | 92.0  | 92.0  | 90.0  | 90.0  | 90.0  | 90.0  | 88.0  | 88.0  |
| Number of survival/ Number of effective animals |                     |                        |       |       |       |       |       |       |       |       |       |       |       |       |       |
| Survival rate (%)                               |                     |                        |       |       |       |       |       |       |       |       |       |       |       |       |       |

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BAIS4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1 104  
SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 8

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|
|            |                     | 98                     | 99    | 100   | 101   | 102   | 103   | 104   |
| Control    | 50                  | 28/50                  | 28/50 | 28/50 | 28/50 | 27/50 | 25/50 | 24/50 |
|            |                     | 56.0                   | 56.0  | 56.0  | 56.0  | 54.0  | 50.0  | 48.0  |
| 625 ppm    | 50                  | 36/50                  | 36/50 | 36/50 | 36/50 | 35/50 | 33/50 | 32/50 |
|            |                     | 72.0                   | 72.0  | 72.0  | 72.0  | 70.0  | 66.0  | 64.0  |
| 1250 ppm   | 50                  | 38/50                  | 38/50 | 38/50 | 36/50 | 34/50 | 33/50 | 31/50 |
|            |                     | 76.0                   | 76.0  | 76.0  | 72.0  | 68.0  | 66.0  | 62.0  |
| 2500 ppm   | 50                  | 44/50                  | 42/50 | 41/50 | 41/50 | 40/50 | 39/50 | 39/50 |
|            |                     | 88.0                   | 84.0  | 82.0  | 82.0  | 80.0  | 78.0  | 78.0  |

Number of survival/ Number of effective animals  
Survival rate (%)

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TABLE A 2

SURVIVAL ANIMAL NUMBERS: FEMALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 9

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |       |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|            |                     | 0                      | 1     | 2     | 3     | 4     | 5     | 6     | 7     | 8     | 9     | 10    | 11    | 12    | 13    |
| Control    | 50                  | 50/50                  | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 |
|            |                     | 100.0                  | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 625 ppm    | 50                  | 50/50                  | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 |
|            |                     | 100.0                  | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 1250 ppm   | 50                  | 50/50                  | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 |
|            |                     | 100.0                  | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 2500 ppm   | 50                  | 50/50                  | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 |
|            |                     | 100.0                  | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |

Number of survival/ Number of effective animals  
 Survival rate(%)

(HAN360)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 10

| Group Name | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|------------|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|            |                     | 14                     | 15             | 16             | 17             | 18             | 19             | 20             | 21             | 22             | 23             | 24             | 25             | 26             | 27             |
| Control    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 625 ppm    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 2500 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |

Number of survival/ Number of effective animals  
 Survival rate (%)

(HAN360)

BAIS4



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 11

| Group Name | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|------------|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|            |                     | 28                     | 29             | 30             | 31             | 32             | 33             | 34             | 35             | 36             | 37             | 38             | 39             | 40             | 41             |
| Control    | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 625 ppm    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |
| 2500 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |

Number of survival/ Number of effective animals  
 Survival rate(%)

(HAN360)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 12

| Group Name | Animals<br>At start | Administration (Weeks) |                |                |                |                |                |                |                |                |                |                |                |                |                |
|------------|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
|            |                     | 42                     | 43             | 44             | 45             | 46             | 47             | 48             | 49             | 50             | 51             | 52             | 53             | 54             | 55             |
| Control    | 50                  | 49/50<br>98.0          | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 625 ppm    | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  | 49/50<br>98.0  |
| 1250 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 49/50<br>98.0  |
| 2500 ppm   | 50                  | 50/50<br>100.0         | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 | 50/50<br>100.0 |

Number of survival/ Number of effective animals  
 Survival rate (%)

(HAN360)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 13

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |       |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|            |                     | 56                     | 57    | 58    | 59    | 60    | 61    | 62    | 63    | 64    | 65    | 66    | 67    | 68    | 69    |
| Control    | 50                  | 49/50                  | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 45/50 | 44/50 | 42/50 |
|            |                     | 98.0                   | 98.0  | 98.0  | 98.0  | 96.0  | 96.0  | 94.0  | 94.0  | 94.0  | 94.0  | 92.0  | 90.0  | 88.0  | 84.0  |
| 625 ppm    | 50                  | 49/50                  | 49/50 | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 |
|            |                     | 98.0                   | 98.0  | 98.0  | 98.0  | 98.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  |
| 1250 ppm   | 50                  | 49/50                  | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 | 47/50 |
|            |                     | 98.0                   | 98.0  | 98.0  | 98.0  | 98.0  | 98.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 96.0  | 94.0  | 94.0  |
| 2500 ppm   | 50                  | 50/50                  | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 |
|            |                     | 100.0                  | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |

Number of survival/ Number of effective animals  
 Survival rate (%)

(HAN360)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1 104  
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 14

| Group Name | Animals<br>At start | Administration (Weeks) |               |               |               |               |               |               |               |               |               |               |               |               |               |
|------------|---------------------|------------------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|
|            |                     | 70                     | 71            | 72            | 73            | 74            | 75            | 76            | 77            | 78            | 79            | 80            | 81            | 82            | 83            |
| Control    | 50                  | 41/50<br>82.0          | 40/50<br>80.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 38/50<br>76.0 | 38/50<br>76.0 | 38/50<br>76.0 | 38/50<br>76.0 | 38/50<br>76.0 |
| 625 ppm    | 50                  | 48/50<br>96.0          | 48/50<br>96.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 44/50<br>88.0 |
| 1250 ppm   | 50                  | 46/50<br>92.0          | 46/50<br>92.0 | 46/50<br>92.0 | 46/50<br>92.0 | 45/50<br>90.0 | 43/50<br>86.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 | 42/50<br>84.0 |
| 2500 ppm   | 50                  | 50/50<br>100.0         | 49/50<br>98.0 | 49/50<br>98.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 | 48/50<br>96.0 |

Number of survival/ Number of effective animals  
 Survival rate(%)

(HAN360)

BAIS4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
REPORT TYPE : A1 104  
SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 15

| Group Name | Animals<br>At start | Administration (Weeks) |               |               |               |               |               |               |               |               |               |               |               |               |               |
|------------|---------------------|------------------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|---------------|
|            |                     | 84                     | 85            | 86            | 87            | 88            | 89            | 90            | 91            | 92            | 93            | 94            | 95            | 96            | 97            |
| Control    | 50                  | 38/50<br>76.0          | 38/50<br>76.0 | 38/50<br>76.0 | 38/50<br>76.0 | 37/50<br>74.0 | 37/50<br>74.0 | 37/50<br>74.0 | 36/50<br>72.0 | 36/50<br>72.0 | 35/50<br>70.0 | 33/50<br>66.0 | 32/50<br>64.0 | 31/50<br>62.0 | 31/50<br>62.0 |
| 625 ppm    | 50                  | 43/50<br>86.0          | 43/50<br>86.0 | 43/50<br>86.0 | 42/50<br>84.0 | 41/50<br>82.0 | 40/50<br>80.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 39/50<br>78.0 | 38/50<br>76.0 | 36/50<br>72.0 | 35/50<br>70.0 | 34/50<br>68.0 |
| 1250 ppm   | 50                  | 42/50<br>84.0          | 41/50<br>82.0 | 41/50<br>82.0 | 41/50<br>82.0 | 41/50<br>82.0 | 41/50<br>82.0 | 40/50<br>80.0 | 40/50<br>80.0 | 40/50<br>80.0 | 39/50<br>78.0 | 38/50<br>76.0 | 38/50<br>76.0 | 38/50<br>76.0 | 35/50<br>70.0 |
| 2500 ppm   | 50                  | 47/50<br>94.0          | 47/50<br>94.0 | 47/50<br>94.0 | 47/50<br>94.0 | 46/50<br>92.0 | 46/50<br>92.0 | 45/50<br>90.0 | 45/50<br>90.0 | 43/50<br>86.0 | 43/50<br>86.0 | 43/50<br>86.0 | 43/50<br>86.0 | 43/50<br>86.0 | 43/50<br>86.0 |

Number of survival/ Number of effective animals  
Survival rate(%)

(HAN360)

BAIS4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
REPORT TYPE : A1 104  
SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 16

| Group Name | Animals<br>At start | Administration (Weeks) |       |       |       |       |       |       |
|------------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|
|            |                     | 98                     | 99    | 100   | 101   | 102   | 103   | 104   |
| Control    | 50                  | 31/50                  | 29/50 | 29/50 | 28/50 | 25/50 | 25/50 | 24/50 |
|            |                     | 62.0                   | 58.0  | 58.0  | 56.0  | 50.0  | 50.0  | 48.0  |
| 625 ppm    | 50                  | 33/50                  | 31/50 | 31/50 | 27/50 | 24/50 | 24/50 | 21/50 |
|            |                     | 66.0                   | 62.0  | 62.0  | 54.0  | 48.0  | 48.0  | 42.0  |
| 1250 ppm   | 50                  | 35/50                  | 35/50 | 34/50 | 33/50 | 33/50 | 33/50 | 33/50 |
|            |                     | 70.0                   | 70.0  | 68.0  | 66.0  | 66.0  | 66.0  | 66.0  |
| 2500 ppm   | 50                  | 42/50                  | 42/50 | 40/50 | 39/50 | 39/50 | 38/50 | 38/50 |
|            |                     | 84.0                   | 84.0  | 80.0  | 78.0  | 78.0  | 76.0  | 76.0  |

Number of survival/ Number of effective animals  
Survival rate(%)

(HAN360)

BAIS4

TABLE B 1

CLINICAL OBSERVATION: MALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : MALE

PAGE : 1

| Clinical sign         | Group Name | Administration Week-day |     |     |     |     |     |     |     |     |      |      |      |      |      |
|-----------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                       |            | 1-7                     | 2-7 | 3-7 | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
| DEATH                 | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 1   | 1   | 1   | 1   | 1   | 1   | 1   | 1    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 1   | 1    | 1    | 1    | 1    | 1    |
| MORIBUND SACRIFICE    | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION    | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT        | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| TREMOR                | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HINDLIMBS | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| SOILED                | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION          | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1 104

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| Clinical sign         | Group Name | Administration Week-day |      |      | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 15-7                    | 16-7 | 17-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| MORIBUND SACRIFICE    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HINDLIMBS | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
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| Clinical sign         | Group Name | Administration Week-day |      |      |   | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|-----------------------|------------|-------------------------|------|------|---|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 29-7                    | 30-7 | 31-7 |   |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                 | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
|                       | 625 ppm    | 2                       | 2    | 2    | 2 | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 1                       | 1    | 1    | 1 | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| MORIBUND SACRIFICE    | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION    | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT        | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HINDLIMBS | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT         | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION          | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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| Clinical sign          | Group Name | Administration Week-day |      |      | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
|------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                        |            | 43-7                    | 44-7 | 45-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                  | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    |
|                        | 625 ppm    | 2                       | 2    | 2    | 2    | 2    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                        | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| MORIBUND SACRIFICE     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HIND LIMBS | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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| Clinical sign          | Group Name | Administration Week-day |      |      | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                        |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                  | Control    | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 3    | 3    |
|                        | 625 ppm    | 3                       | 3    | 4    | 5    | 5    | 5    | 5    | 5    | 5    | 6    | 6    | 6    | 6    | 6    |
|                        | 1250 ppm   | 1                       | 1    | 1    | 2    | 2    | 2    | 2    | 3    | 3    | 3    | 3    | 3    | 3    | 3    |
|                        | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    |
| MORIBUND SACRIFICE     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HIND LIMBS | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTOR            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                        | 625 ppm    | 0                       | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                        | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    |

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| Clinical sign          | Group Name | Administration Week-day |      |      | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                        |            | 71-7                    | 72-7 | 73-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                  | Control    | 3                       | 3    | 3    | 3    | 3    | 4    | 5    | 5    | 5    | 5    | 6    | 6    | 6    | 7    |
|                        | 625 ppm    | 6                       | 6    | 6    | 6    | 7    | 7    | 7    | 7    | 7    | 8    | 8    | 8    | 8    | 8    |
|                        | 1250 ppm   | 3                       | 3    | 4    | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 6    | 6    | 6    |
|                        | 2500 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 3    | 3    | 4    | 4    |
| MORIBUND SACRIFICE     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| HUNCHBACK POSITION     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HIND LIMBS | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                 | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTOR            | Control    | 1                       | 1    | 0    | 1    | 2    | 1    | 1    | 1    | 1    | 2    | 1    | 1    | 1    | 0    |
|                        | 625 ppm    | 1                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|                        | 1250 ppm   | 1                       | 1    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 1    | 1    | 1    |
|                        | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |

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| Clinical sign         | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                 | Control    | 8                       | 8    | 9    | 11   | 11   | 11   | 13   | 15   | 15   | 16   | 17   | 19   | 20   | 20   |
|                       | 625 ppm    | 9                       | 9    | 9    | 9    | 10   | 10   | 11   | 11   | 11   | 11   | 11   | 12   | 12   | 13   |
|                       | 1250 ppm   | 6                       | 6    | 6    | 6    | 6    | 7    | 7    | 7    | 9    | 9    | 9    | 11   | 11   | 11   |
|                       | 2500 ppm   | 4                       | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 5    | 5    | 5    |
| MORIBUND SACRIFICE    | Control    | 1                       | 1    | 1    | 1    | 1    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| HUNCHBACK POSITION    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TREMOR                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| DRAGGING OF HINDLIMBS | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ABNORMAL GAIT         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    |
| PILOERECTION          | Control    | 0                       | 1    | 1    | 1    | 2    | 2    | 3    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 1                       | 1    | 1    | 1    | 2    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                       | 1250 ppm   | 1                       | 1    | 1    | 1    | 2    | 3    | 3    | 3    | 1    | 1    | 1    | 0    | 0    | 2    |
|                       | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    |

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| Clinical sign          | Group Name | Administration Week-day |       |       |       |       |       |
|------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                        |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| DEATH                  | Control    | 20                      | 20    | 20    | 21    | 23    | 24    |
|                        | 625 ppm    | 13                      | 13    | 13    | 14    | 15    | 16    |
|                        | 1250 ppm   | 11                      | 11    | 13    | 15    | 16    | 18    |
|                        | 2500 ppm   | 7                       | 8     | 8     | 9     | 10    | 10    |
| MORIBUND SACRIFICE     | Control    | 2                       | 2     | 2     | 2     | 2     | 2     |
|                        | 625 ppm    | 1                       | 1     | 1     | 1     | 2     | 2     |
|                        | 1250 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
|                        | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| HUNCHBACK POSITION     | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| PARALYTIC GAIT         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 1     | 1     | 1     |
|                        | 2500 ppm   | 0                       | 0     | 1     | 2     | 1     | 1     |
| TREMOR                 | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| DRAGGING OF HIND LIMBS | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| ABNORMAL GAIT          | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 0     | 1     | 0     |
|                        | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| SOILED                 | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                        | 2500 ppm   | 2                       | 2     | 2     | 1     | 0     | 0     |
| PILOERECTION           | Control    | 0                       | 2     | 2     | 1     | 0     | 0     |
|                        | 625 ppm    | 2                       | 2     | 2     | 1     | 1     | 0     |
|                        | 1250 ppm   | 2                       | 2     | 1     | 1     | 1     | 1     |
|                        | 2500 ppm   | 2                       | 2     | 2     | 1     | 0     | 1     |

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| Clinical sign         | Group Name | Administration Week-day |     |     | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
|-----------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                       |            | 1-7                     | 2-7 | 3-7 |     |     |     |     |     |     |      |      |      |      |      |
| FROG BELLY            | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 1   | 1   | 2   | 3   | 3   | 3   | 5   | 5   | 3    | 3    | 3    | 5    | 5    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 1   | 1   | 1   | 1   | 1   | 1   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 1   | 1   | 2   | 2   | 2    | 2    | 2    | 2    | 1    |
| M. EYE                | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |



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| Clinical sign         | Group Name | Administration Week-day |      |      |  | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|-----------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 15-7                    | 16-7 | 17-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| FROG BELLY            | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS         | Control    | 1                       | 2    | 3    |  | 3    | 3    | 3    | 3    | 4    | 4    | 3    | 3    | 3    | 4    | 4    |
|                       | 625 ppm    | 5                       | 6    | 7    |  | 7    | 7    | 7    | 8    | 8    | 8    | 7    | 7    | 7    | 8    | 9    |
|                       | 1250 ppm   | 0                       | 0    | 1    |  | 1    | 1    | 1    | 4    | 5    | 5    | 2    | 2    | 2    | 3    | 3    |
|                       | 2500 ppm   | 1                       | 2    | 2    |  | 3    | 3    | 3    | 3    | 4    | 4    | 3    | 3    | 3    | 3    | 4    |
| M. EYE                | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : MALE

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| Clinical sign         | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 29-7                    | 30-7 | 31-7 | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
| FROG BELLY            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS         | Control    | 4                       | 4    | 4    | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 4    | 4    |
|                       | 625 ppm    | 8                       | 6    | 6    | 6    | 6    | 7    | 7    | 7    | 7    | 6    | 6    | 6    | 6    | 6    |
|                       | 1250 ppm   | 3                       | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 4    | 5    | 5    | 5    | 5    |
|                       | 2500 ppm   | 4                       | 2    | 2    | 3    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    |
| M. EYE                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

SEX : MALE

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| Clinical sign         | Group Name | Administration Week-day |      |      | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 43-7                    | 44-7 | 45-7 |      |      |      |      |      |      |      |      |      |      |      |
| FROG BELLY            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
| CORNEAL OPACITY       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS         | Control    | 4                       | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 3    | 3    |
|                       | 625 ppm    | 6                       | 6    | 7    | 6    | 6    | 5    | 5    | 5    | 5    | 5    | 6    | 6    | 6    | 6    |
|                       | 1250 ppm   | 5                       | 5    | 5    | 5    | 5    | 5    | 5    | 5    | 6    | 5    | 5    | 4    | 4    | 4    |
|                       | 2500 ppm   | 4                       | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    |
| M. EYE                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

SEX : MALE

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| Clinical sign         | Group Name | Administration Week-day |      |      | 57-7 | 58-7 | 59-7 | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |      |      |      |
| FROG BELLY            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS         | Control    | 3                       | 3    | 3    | 4    | 4    | 4    | 4    | 4    | 6    | 6    | 5    | 5    | 6    | 5    | 5    | 5    | 5    |
|                       | 625 ppm    | 6                       | 6    | 6    | 6    | 6    | 6    | 6    | 6    | 6    | 7    | 6    | 6    | 6    | 6    | 6    | 8    | 8    |
|                       | 1250 ppm   | 4                       | 4    | 4    | 3    | 3    | 2    | 3    | 3    | 3    | 5    | 6    | 6    | 7    | 7    | 7    | 6    | 6    |
|                       | 2500 ppm   | 4                       | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    | 4    |
| M. EYE                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
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| Clinical sign         | Group Name | Administration Week-day |      |      | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 71-7                    | 72-7 | 73-7 |      |      |      |      |      |      |      |      |      |      |      |
| FROG BELLY            | Control    | 0                       | 0    | 0    | 0    | 2    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 0                       | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 1                       | 1    | 1    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    | 2    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
| INTERNAL MASS         | Control    | 5                       | 6    | 6    | 5    | 5    | 4    | 4    | 4    | 4    | 5    | 6    | 7    | 7    | 6    |
|                       | 625 ppm    | 8                       | 8    | 8    | 8    | 8    | 8    | 8    | 8    | 8    | 8    | 8    | 7    | 7    | 7    |
|                       | 1250 ppm   | 6                       | 6    | 6    | 7    | 7    | 7    | 7    | 7    | 7    | 7    | 7    | 6    | 6    | 7    |
|                       | 2500 ppm   | 4                       | 4    | 4    | 4    | 5    | 5    | 5    | 6    | 7    | 7    | 7    | 9    | 8    | 8    |
| M. EYE                | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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| Clinical sign         | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|-----------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                       |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| FROG BELLY            | Control    | 1                       | 1    | 1    | 1    | 1    | 2    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED PERI-GENITALIA | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS          | Control    | 1                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| GUM                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY       | Control    | 1                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS         | Control    | 2                       | 2    | 2    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    |
|                       | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 1    | 2    | 2    | 2    | 4    | 4    | 4    | 4    | 4    |
|                       | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 1    | 2    | 2    |
| INTERNAL MASS         | Control    | 6                       | 6    | 5    | 6    | 6    | 6    | 4    | 2    | 3    | 5    | 4    | 3    | 5    | 5    |
|                       | 625 ppm    | 6                       | 6    | 6    | 7    | 6    | 5    | 6    | 6    | 6    | 5    | 5    | 6    | 7    | 6    |
|                       | 1250 ppm   | 7                       | 7    | 8    | 11   | 11   | 10   | 10   | 10   | 9    | 6    | 6    | 4    | 7    | 9    |
|                       | 2500 ppm   | 9                       | 9    | 9    | 8    | 8    | 9    | 9    | 9    | 11   | 9    | 9    | 7    | 10   | 11   |
| M. EYE                | Control    | 1                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI EAR           | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                       | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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| Clinical sign         | Group Name | Administration Week-day |       |       |       |       |       |
|-----------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                       |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| FROG BELLY            | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 1                       | 1     | 1     | 1     | 1     | 1     |
|                       | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| SOILED PERI-GENITALIA | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| EXOPHTHALMOS          | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 1250 ppm   | 3                       | 3     | 3     | 3     | 3     | 2     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| GUM                   | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| CORNEAL OPACITY       | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 1250 ppm   | 2                       | 2     | 2     | 2     | 2     | 2     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| EXTERNAL MASS         | Control    | 0                       | 0     | 1     | 1     | 1     | 1     |
|                       | 625 ppm    | 2                       | 2     | 2     | 4     | 3     | 4     |
|                       | 1250 ppm   | 4                       | 5     | 5     | 4     | 4     | 3     |
|                       | 2500 ppm   | 2                       | 2     | 2     | 2     | 2     | 2     |
| INTERNAL MASS         | Control    | 7                       | 8     | 8     | 15    | 13    | 10    |
|                       | 625 ppm    | 7                       | 7     | 9     | 18    | 16    | 13    |
|                       | 1250 ppm   | 11                      | 10    | 8     | 13    | 12    | 10    |
|                       | 2500 ppm   | 9                       | 8     | 8     | 17    | 16    | 15    |
| M. EYE                | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 1250 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. PERI EAR           | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 625 ppm    | 0                       | 0     | 0     | 1     | 1     | 1     |
|                       | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                       | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |

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 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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| Clinical sign | Group Name | Administration Week-day |     |     |     |     |     |     |     |     |      |      |      |      |      |
|---------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|               |            | 1-7                     | 2-7 | 3-7 | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
| M. NECK       | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN    | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB  | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| CRUSTA        | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |



STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign | Group Name | Administration Week-day |      |      |  | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|---------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 15-7                    | 16-7 | 17-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CRUSTA        | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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| Clinical sign | Group Name | Administration Week-day |      |      |   | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|---------------|------------|-------------------------|------|------|---|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 29-7                    | 30-7 | 31-7 |   |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB  | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CRUSTA        | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign | Group Name | Administration Week-day |      |      |  | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
|---------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 43-7                    | 44-7 | 45-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 2    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CRUSTA        | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
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| Clinical sign | Group Name | Administration Week-day |      |      | 57-7 | 58-7 | 59-7 | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|---------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 2                       | 2    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CRUSTA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
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| Clinical sign | Group Name | Administration Week-day |      |      |   | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|---------------|------------|-------------------------|------|------|---|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 71-7                    | 72-7 | 73-7 |   |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB  | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 1                       | 1    | 1    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    | 1 | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    |
|               | 625 ppm    | 1                       | 1    | 1    | 1 | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 1250 ppm   | 1                       | 1    | 1    | 1 | 1    | 1    | 0    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
| CRUSTA        | Control    | 0                       | 0    | 1    | 1 | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 1    | 1    | 1    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0 | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0 | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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| Clinical sign | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|---------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|               |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| M. NECK       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| M. ABDOMEN    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
| M. GENITALIA  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    |
|               | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
| M. ANUS       | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL       | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    | 2    | 2    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ULCER         | Control    | 1                       | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 2    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EROSION       | Control    | 0                       | 1    | 1    | 1    | 1    | 2    | 2    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|               | 1250 ppm   | 0                       | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 0    | 1    | 2    | 0    | 0    | 0    |
|               | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| CRUSTA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|               | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
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| Clinical sign | Group Name | Administration Week-day |       |       |       |       |       |
|---------------|------------|-------------------------|-------|-------|-------|-------|-------|
|               |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| M. NECK       | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| M. ABDOMEN    | Control    | 0                       | 0     | 1     | 1     | 1     | 1     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 1                       | 1     | 1     | 1     | 1     | 0     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. HINDLIMB   | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| M. GENITALIA  | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 2                       | 2     | 2     | 3     | 2     | 3     |
|               | 1250 ppm   | 0                       | 1     | 1     | 1     | 1     | 1     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. ANUS       | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. TAIL       | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 2                       | 2     | 2     | 1     | 1     | 1     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| ULCER         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 1                       | 1     | 1     | 1     | 0     | 0     |
|               | 1250 ppm   | 2                       | 2     | 1     | 0     | 1     | 0     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| EROSION       | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 2                       | 2     | 2     | 1     | 2     | 1     |
|               | 1250 ppm   | 0                       | 0     | 0     | 2     | 2     | 2     |
|               | 2500 ppm   | 1                       | 1     | 1     | 1     | 0     | 0     |
| CRUSTA        | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|               | 1250 ppm   | 2                       | 1     | 1     | 1     | 0     | 0     |
|               | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |

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| Clinical sign       | Group Name | Administration Week-day |     |     |     |     |     |     |     |     |      |      |      |      |      |
|---------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                     |            | 1-7                     | 2-7 | 3-7 | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
| HEMORRHAGE          | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 49  | 49   | 49   | 49   | 49   | 49   |
| SMALL STOOL         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 1   | 1   | 0   | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0   | 3   | 2   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 3   | 1   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 0   | 4   | 1   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 1                       | 3   | 4   | 2   | 2   | 2   | 1   | 1   | 0   | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 50                      | 50  | 47  | 48  | 50  | 50  | 50  | 50  | 50  | 49   | 49   | 49   | 49   | 49   |
|                     | 625 ppm    | 50                      | 49  | 46  | 46  | 46  | 46  | 46  | 44  | 44  | 46   | 46   | 46   | 44   | 44   |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |



STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1 104

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ALL ANIMALS

SEX : MALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |    | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|---------------------|------------|-------------------------|------|------|----|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 15-7                    | 16-7 | 17-7 |    |      |      |      |      |      |      |      |      |      |      |      |
| HEMORRHAGE          | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50   | 50   | 50 | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 49                      | 49   | 49   | 49 | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 49                      | 48   | 47   | 47 | 47   | 47   | 47   | 47   | 46   | 46   | 47   | 47   | 47   | 46   | 46   |
|                     | 625 ppm    | 44                      | 43   | 42   | 42 | 42   | 42   | 42   | 41   | 41   | 41   | 42   | 42   | 42   | 41   | 40   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJj[Crj:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
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| Clinical sign       | Group Name | Administration |      | Week-day |    | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|---------------------|------------|----------------|------|----------|----|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 29-7           | 30-7 | 31-7     |    |      |      |      |      |      |      |      |      |      |      |      |
| HEMORRHAGE          | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50             | 50   | 50       | 50 | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 49             | 49   | 49       | 49 | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   |
| SMALL STOOL         | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 46             | 46   | 46       | 45 | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   |
|                     | 625 ppm    | 40             | 42   | 42       | 42 | 42   | 41   | 41   | 41   | 41   | 42   | 42   | 42   | 42   | 42   | 42   |
|                     | 1250 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0              | 0    | 0        | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : MALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 43-7                    | 44-7 | 45-7 | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
| HEMORRHAGE          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 49   | 49   | 49   | 49   | 49   |
|                     | 2500 ppm   | 49                      | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
| NON REMARKABLE      | Control    | 45                      | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 45   |
|                     | 625 ppm    | 42                      | 42   | 41   | 42   | 42   | 42   | 42   | 42   | 42   | 42   | 41   | 40   | 40   | 40   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : MALE

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| Clinical sign       | Group Name | Administration Week-day |      |      | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |
| HEMORRHAGE          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 49                      | 49   | 49   | 48   | 48   | 48   | 48   | 47   | 47   | 47   | 47   | 47   | 47   | 47   |
|                     | 2500 ppm   | 49                      | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 48   | 48   | 48   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 1    | 2    | 2    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 0    | 0    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
|                     | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 45                      | 45   | 45   | 44   | 43   | 42   | 43   | 41   | 41   | 42   | 41   | 40   | 40   | 40   |
|                     | 625 ppm    | 40                      | 40   | 40   | 39   | 39   | 38   | 38   | 38   | 37   | 37   | 37   | 37   | 37   | 35   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : MALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |    | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|---------------------|------------|-------------------------|------|------|----|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 71-7                    | 72-7 | 73-7 |    |      |      |      |      |      |      |      |      |      |      |      |
| HEMORRHAGE          | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 1                       | 1    | 1    | 1  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0  | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    | 1  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 47                      | 47   | 46   | 45 | 45   | 45   | 45   | 45   | 45   | 45   | 45   | 44   | 44   | 44   | 44   |
|                     | 2500 ppm   | 48                      | 48   | 48   | 48 | 48   | 48   | 48   | 48   | 48   | 48   | 47   | 47   | 46   | 46   | 46   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 1    | 2    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 1                       | 1    | 2    | 2  | 2    | 2    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    |
|                     | 625 ppm    | 1                       | 0    | 0    | 1  | 1    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 2                       | 0    | 2    | 1  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 40                      | 39   | 38   | 38 | 38   | 38   | 37   | 37   | 37   | 35   | 34   | 34   | 34   | 34   | 34   |
|                     | 625 ppm    | 35                      | 35   | 35   | 35 | 34   | 34   | 34   | 34   | 34   | 33   | 33   | 34   | 34   | 34   | 34   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
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SEX : MALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |  | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|---------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 85-7                    | 86-7 | 87-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| HEMORRHAGE          | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PROLAPSE OF PENIS   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| IRREGULAR BREATHING | Control    | 0                       | 1    | 1    |  | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 2    | 2    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| BROWN URINE         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 44                      | 44   | 44   |  | 44   | 44   | 43   | 43   | 43   | 41   | 39   | 40   | 38   | 38   | 38   |
|                     | 2500 ppm   | 46                      | 46   | 46   |  | 46   | 46   | 46   | 46   | 45   | 45   | 45   | 45   | 44   | 44   | 44   |
| SMALL STOOL         | Control    | 0                       | 0    | 1    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 1    | 0    | 1    | 1    | 1    | 0    | 0    | 1    |
| OLIGO-STOOL         | Control    | 1                       | 1    | 3    |  | 1    | 2    | 1    | 1    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 2    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 2    | 1    | 1    | 0    | 0    | 0    | 0    | 2    |
| NON REMARKABLE      | Control    | 33                      | 32   | 31   |  | 30   | 30   | 30   | 30   | 30   | 29   | 26   | 26   | 25   | 22   | 22   |
|                     | 625 ppm    | 34                      | 34   | 34   |  | 33   | 33   | 32   | 31   | 31   | 31   | 32   | 32   | 30   | 29   | 28   |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1 104

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| Clinical sign       | Group Name | Administration Week-day |       |       |       |       |       |
|---------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                     |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| HEMORRHAGE          | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 1                       | 1     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| TORTICOLLIS         | Control    | 1                       | 1     | 1     | 1     | 1     | 1     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 1                       | 1     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| PROLAPSE OF PENIS   | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 1                       | 1     | 1     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 1     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 0     | 0     |
| IRREGULAR BREATHING | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 1     | 0     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 0     | 0     |
| BROWN URINE         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 38                      | 38    | 36    | 34    | 33    | 31    |
|                     | 2500 ppm   | 42                      | 41    | 41    | 40    | 39    | 39    |
| SMALL STOOL         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 1     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 1     | 0     | 0     | 0     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| OLIGO-STOOL         | Control    | 1                       | 1     | 1     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 1     | 0     | 0     | 0     |
|                     | 1250 ppm   | 1                       | 1     | 1     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| NON REMARKABLE      | Control    | 20                      | 19    | 18    | 11    | 11    | 13    |
|                     | 625 ppm    | 28                      | 28    | 26    | 15    | 15    | 16    |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |

**TABLE B 2**

**CLINICAL OBSERVATION: FEMALE**



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
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| Clinical sign           | Group Name | Administration Week-day |     |     |     |     |     |     |     |     |      |      |      |      |      |
|-------------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                         |            | 1-7                     | 2-7 | 3-7 | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
| DEATH                   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| MORIBUND SACRIFICE      | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| PILOERECTOR             | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |

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ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
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| Clinical sign           | Group Name | Administration Week-day |      |      | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 15-7                    | 16-7 | 17-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| MORIBUND SACRIFICE      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION            | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
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SEX : FEMALE

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| Clinical sign           | Group Name | Administration Week-day |      |      | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 29-7                    | 30-7 | 31-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                   | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| MORIBUND SACRIFICE      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104

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SEX : FEMALE

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| Clinical sign           | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 43-7                    | 44-7 | 45-7 | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
| DEATH                   | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| MORIBUND SACRIFICE      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILORECTION             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
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SEX : FEMALE

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| Clinical sign           | Group Name | Administration Week-day |      |      |  | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|-------------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 57-7                    | 58-7 | 59-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                   | Control    | 1                       | 1    | 1    |  | 2    | 2    | 3    | 3    | 3    | 3    | 4    | 5    | 6    | 7    | 8    |
|                         | 625 ppm    | 1                       | 1    | 1    |  | 1    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 3    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| MORIBUND SACRIFICE      | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILORECTION             | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 3    | 2    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 2500 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104

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SEX : FEMALE

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| Clinical sign           | Group Name | Administration Week-day |      |      | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 71-7                    | 72-7 | 73-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                   | Control    | 9                       | 10   | 10   | 10   | 10   | 10   | 10   | 10   | 11   | 11   | 11   | 11   | 11   | 11   |
|                         | 625 ppm    | 2                       | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 5    | 6    |
|                         | 1250 ppm   | 3                       | 3    | 3    | 3    | 5    | 6    | 6    | 6    | 6    | 6    | 6    | 6    | 6    | 6    |
|                         | 2500 ppm   | 0                       | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    |
| MORIBUND SACRIFICE      | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
|                         | 1250 ppm   | 1                       | 1    | 1    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                         | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| SOILED                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILORECTION             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 1                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 2                       | 2    | 2    | 3    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 1                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    | 2    | 2    | 2    |
|                         | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign           | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                         |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| DEATH                   | Control    | 11                      | 11   | 11   | 12   | 12   | 12   | 13   | 13   | 14   | 16   | 17   | 18   | 18   | 18   |
|                         | 625 ppm    | 6                       | 6    | 7    | 8    | 9    | 10   | 10   | 10   | 10   | 11   | 12   | 12   | 13   | 14   |
|                         | 1250 ppm   | 7                       | 7    | 7    | 7    | 7    | 8    | 8    | 8    | 9    | 10   | 10   | 10   | 12   | 12   |
|                         | 2500 ppm   | 2                       | 2    | 2    | 3    | 3    | 4    | 4    | 6    | 6    | 6    | 6    | 6    | 6    | 7    |
| MORIBUND SACRIFICE      | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 3    | 3    | 3    |
|                         | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 3    | 3    |
|                         | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PARALYTIC GAIT          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
| SOILED                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| PILOERECTION            | Control    | 0                       | 0    | 1    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 2    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 1    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TRAUMA                  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| FROG BELLY              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 2    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 1    | 0    | 0    | 1    | 1    | 1    | 0    | 0    | 1    | 0    |
|                         | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 2500 ppm   | 0                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXOPHTHALMOS            | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                         | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                         | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                         | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 2    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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| Clinical sign           | Group Name | Administration Week-day |       |       |       |       |       |
|-------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                         |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| DEATH                   | Control    | 20                      | 20    | 21    | 23    | 23    | 24    |
|                         | 625 ppm    | 16                      | 16    | 20    | 22    | 22    | 25    |
|                         | 1250 ppm   | 12                      | 13    | 14    | 14    | 14    | 14    |
|                         | 2500 ppm   | 7                       | 7     | 8     | 8     | 9     | 9     |
| MORIBUND SACRIFICE      | Control    | 1                       | 1     | 1     | 2     | 2     | 2     |
|                         | 625 ppm    | 3                       | 3     | 3     | 4     | 4     | 4     |
|                         | 1250 ppm   | 3                       | 3     | 3     | 3     | 3     | 3     |
|                         | 2500 ppm   | 1                       | 3     | 3     | 3     | 3     | 3     |
| LOCOMOTOR MOVEMENT DECR | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| PARALYTIC GAIT          | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| SOILED                  | Control    | 1                       | 1     | 1     | 0     | 0     | 0     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| PILOERECTION            | Control    | 1                       | 2     | 1     | 1     | 1     | 1     |
|                         | 625 ppm    | 0                       | 1     | 2     | 1     | 2     | 1     |
|                         | 1250 ppm   | 0                       | 0     | 1     | 1     | 0     | 1     |
|                         | 2500 ppm   | 0                       | 0     | 0     | 1     | 1     | 1     |
| TRAUMA                  | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| FROG BELLY              | Control    | 2                       | 2     | 1     | 0     | 0     | 1     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 0                       | 0     | 1     | 2     | 2     | 2     |
|                         | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| EXOPHTHALMOS            | Control    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                         | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                         | 1250 ppm   | 2                       | 2     | 2     | 2     | 2     | 2     |
|                         | 2500 ppm   | 2                       | 2     | 2     | 2     | 2     | 2     |



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 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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| Clinical sign   | Group Name | Administration Week-day |     |     | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
|-----------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                 |            | 1-7                     | 2-7 | 3-7 |     |     |     |     |     |     |      |      |      |      |      |
| GUM             | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 1   | 1   | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. EYE          | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. PERI-MOUTH   | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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| Clinical sign   | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|-----------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 15-7                    | 16-7 | 17-7 | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
| GUM             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. EYE          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1 104

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| Clinical sign   | Group Name | Administration Week-day |      |      |  | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|-----------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 29-7                    | 30-7 | 31-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| GUM             | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| INTERNAL MASS   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
| M. EYE          | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign   | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|-----------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 43-7                    | 44-7 | 45-7 | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
| GUM             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EXTERNAL MASS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                 | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| INTERNAL MASS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 2    | 3    | 3    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 3    | 3    | 3    | 3    |
|                 | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| M. EYE          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
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| Clinical sign   | Group Name | Administration Week-day |      |      | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|-----------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |
| GUM             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EXTERNAL MASS   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 3    | 3    | 3    | 2    |
|                 | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 2    | 2    | 2    | 2    | 2    | 3    | 3    | 4    | 4    |
| INTERNAL MASS   | Control    | 3                       | 3    | 3    | 6    | 6    | 5    | 6    | 6    | 6    | 5    | 5    | 4    | 3    | 3    |
|                 | 625 ppm    | 1                       | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                 | 1250 ppm   | 3                       | 3    | 3    | 3    | 3    | 2    | 2    | 2    | 2    | 2    | 4    | 3    | 3    | 3    |
|                 | 2500 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
| M. EYE          | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1 104

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| Clinical sign   | Group Name | Administration Week-day |      |      |  | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|-----------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 71-7                    | 72-7 | 73-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| GUM             | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EXTERNAL MASS   | Control    | 0                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 625 ppm    | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 2                       | 2    | 2    |  | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                 | 2500 ppm   | 4                       | 4    | 3    |  | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 2    |
| INTERNAL MASS   | Control    | 1                       | 1    | 1    |  | 2    | 2    | 2    | 2    | 2    | 1    | 1    | 1    | 1    | 1    | 2    |
|                 | 625 ppm    | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 2    | 3    | 4    | 5    | 6    | 6    | 5    |
|                 | 1250 ppm   | 1                       | 1    | 1    |  | 2    | 1    | 0    | 0    | 0    | 0    | 1    | 1    | 2    | 2    | 2    |
|                 | 2500 ppm   | 2                       | 2    | 2    |  | 2    | 2    | 3    | 3    | 3    | 3    | 3    | 3    | 3    | 4    | 3    |
| M. EYE          | Control    | 0                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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| Clinical sign   | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|-----------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                 |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| GUM             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| CORNEAL OPACITY | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EXTERNAL MASS   | Control    | 2                       | 2    | 2    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 625 ppm    | 1                       | 1    | 1    | 3    | 4    | 3    | 3    | 3    | 3    | 3    | 2    | 2    | 2    | 2    |
|                 | 1250 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
|                 | 2500 ppm   | 2                       | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 3    | 3    | 4    | 4    | 4    | 4    |
| INTERNAL MASS   | Control    | 2                       | 2    | 2    | 2    | 3    | 4    | 3    | 5    | 4    | 4    | 3    | 3    | 4    | 4    |
|                 | 625 ppm    | 5                       | 5    | 4    | 3    | 3    | 2    | 2    | 3    | 4    | 4    | 6    | 7    | 7    | 6    |
|                 | 1250 ppm   | 1                       | 2    | 2    | 2    | 2    | 2    | 3    | 3    | 2    | 2    | 3    | 3    | 2    | 2    |
|                 | 2500 ppm   | 3                       | 3    | 4    | 4    | 4    | 4    | 4    | 3    | 3    | 4    | 8    | 8    | 7    | 7    |
| M. EYE          | Control    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| M. PERI-MOUTH   | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. NECK         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. FORELIMB     | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ABDOMEN      | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 625 ppm    | 0                       | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|                 | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                 | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

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| Clinical sign   | Group Name | Administration Week-day |       |       |       |       |       |
|-----------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                 |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| GUM             | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| CORNEAL OPACITY | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
|                 | 2500 ppm   | 2                       | 2     | 2     | 2     | 2     | 2     |
| EXTERNAL MASS   | Control    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                 | 625 ppm    | 2                       | 2     | 2     | 2     | 2     | 0     |
|                 | 1250 ppm   | 2                       | 3     | 4     | 4     | 4     | 5     |
|                 | 2500 ppm   | 5                       | 5     | 5     | 5     | 4     | 4     |
| INTERNAL MASS   | Control    | 3                       | 3     | 2     | 5     | 3     | 4     |
|                 | 625 ppm    | 4                       | 4     | 2     | 2     | 2     | 1     |
|                 | 1250 ppm   | 4                       | 5     | 6     | 10    | 9     | 9     |
|                 | 2500 ppm   | 8                       | 8     | 7     | 7     | 8     | 8     |
| M. EYE          | Control    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 1                       | 1     | 2     | 2     | 2     | 2     |
|                 | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| M. PERI-MOUTH   | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                 | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. NECK         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| M. FORELIMB     | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 0                       | 1     | 1     | 1     | 1     | 1     |
|                 | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. ABDOMEN      | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                 | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |



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| Clinical sign       | Group Name | Administration Week-day |     |     | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
|---------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                     |            | 1-7                     | 2-7 | 3-7 |     |     |     |     |     |     |      |      |      |      |      |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA        | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| EDEMA               | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |

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| Clinical sign       | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 15-7                    | 16-7 | 17-7 | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EDEMA               | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign       | Group Name | Administration Week-day |      |      | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 29-7                    | 30-7 | 31-7 |      |      |      |      |      |      |      |      |      |      |      |
| M. ANTERIOR. DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| EDEMA               | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104

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| Clinical sign       | Group Name | Administration Week-day |      |      | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 43-7                    | 44-7 | 45-7 |      |      |      |      |      |      |      |      |      |      |      |
| M. ANTERIOR. DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                     | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EDEMA               | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1 104

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SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |  | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|---------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 57-7                    | 58-7 | 59-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 1                       | 1    | 1    |  | 1    | 1    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    | 2    |
| EDEMA               | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1 104

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| Clinical sign       | Group Name | Administration Week-day |      |      | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 71-7                    | 72-7 | 73-7 |      |      |      |      |      |      |      |      |      |      |      |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HIND LIMB        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 1                       | 1    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 0    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. TAIL             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 2                       | 2    | 2    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EDEMA               | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
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SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |      |      |      |      |      |      |      |      |      |      |      |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 85-7                    | 86-7 | 87-7 | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. INTERSCAPULUM    | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. POSTERIOR DORSUM | Control    | 1                       | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. HINDLIMB         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
| M. GENITALIA        | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| M. ANUS             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
| M. TAIL             | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
| EDEMA               | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| ANEMIA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
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SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |       |       |       |       |       |
|---------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                     |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| M. ANTERIOR DORSUM  | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. INTERSCAPULUM    | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. POSTERIOR DORSUM | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 2     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. HINDLIMB         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| M. GENITALIA        | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| M. ANUS             | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 0     | 0     |
| M. TAIL             | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
|                     | 2500 ppm   | 1                       | 1     | 1     | 1     | 1     | 1     |
| EDEMA               | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 1                       | 0     | 0     | 0     | 0     | 0     |
| ANEMIA              | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |



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ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
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CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

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| Clinical sign       | Group Name | Administration Week-day |     |     |     |     |     |     |     |     |      |      |      |      |      |
|---------------------|------------|-------------------------|-----|-----|-----|-----|-----|-----|-----|-----|------|------|------|------|------|
|                     |            | 1-7                     | 2-7 | 3-7 | 4-7 | 5-7 | 6-7 | 7-7 | 8-7 | 9-7 | 10-7 | 11-7 | 12-7 | 13-7 | 14-7 |
| CRUSTA              | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 1    | 1    | 1    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50   | 50   | 50   | 50   | 50   |
| SMALL STOOL         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50  | 50   | 49   | 49   | 49   | 49   |
|                     | 625 ppm    | 50                      | 50  | 50  | 50  | 50  | 50  | 50  | 49  | 49  | 49   | 49   | 49   | 49   | 49   |
|                     | 1250 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0   | 0    | 0    | 0    | 0    | 0    |

(HAN190)

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STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
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SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |  | 18-7 | 19-7 | 20-7 | 21-7 | 22-7 | 23-7 | 24-7 | 25-7 | 26-7 | 27-7 | 28-7 |
|---------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 15-7                    | 16-7 | 17-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 1                       | 1    | 1    |  | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    |  | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50   | 50   |  | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 50                      | 50   | 50   |  | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 49                      | 49   | 49   |  | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   |
|                     | 625 ppm    | 50                      | 50   | 50   |  | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

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STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1 104

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SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      | 32-7 | 33-7 | 34-7 | 35-7 | 36-7 | 37-7 | 38-7 | 39-7 | 40-7 | 41-7 | 42-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 29-7                    | 30-7 | 31-7 |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 2500 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 49                      | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   |
|                     | 625 ppm    | 50                      | 50   | 50   | 50   | 50   | 49   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      | 46-7 | 47-7 | 48-7 | 49-7 | 50-7 | 51-7 | 52-7 | 53-7 | 54-7 | 55-7 | 56-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 43-7                    | 44-7 | 45-7 |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 49   | 49   |
|                     | 2500 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 1    | 0    | 1    | 1    | 0    | 0    | 0    | 1    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 1    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 1    | 0    | 0    | 1    | 2    | 1    | 1    | 1    | 1    | 2    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| NON REMARKABLE      | Control    | 49                      | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 47   | 46   | 46   |
|                     | 625 ppm    | 50                      | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 49   | 48   | 48   | 48   | 48   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      | 60-7 | 61-7 | 62-7 | 63-7 | 64-7 | 65-7 | 66-7 | 67-7 | 68-7 | 69-7 | 70-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 57-7                    | 58-7 | 59-7 |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 0    | 1    | 1    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 49                      | 49   | 49   | 49   | 49   | 48   | 48   | 48   | 48   | 48   | 48   | 47   | 47   | 46   |
|                     | 2500 ppm   | 50                      | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   | 50   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    | 1    | 1    | 0    | 0    | 1    | 1    | 0    | 0    | 1    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 1    | 2    | 2    | 1    | 0    | 0    | 0    | 0    | 0    | 2    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 1    | 1    | 2    | 2    | 3    | 2    | 2    | 0    | 1    | 0    | 1    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 2                       | 2    | 1    | 1    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 2    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 2    | 2    | 2    |
| NON REMARKABLE      | Control    | 46                      | 46   | 46   | 42   | 42   | 41   | 40   | 39   | 39   | 39   | 39   | 38   | 37   | 35   |
|                     | 625 ppm    | 48                      | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 46   | 46   | 46   | 46   | 45   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
REPORT TYPE : AT 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : FEMALE

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| Clinical sign       | Group Name | Administration Week-day |      |      |  | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
|---------------------|------------|-------------------------|------|------|--|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 71-7                    | 72-7 | 73-7 |  |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 1                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    |  | 3    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 0    |
| BROWN URINE         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 46                      | 46   | 46   |  | 45   | 43   | 42   | 42   | 42   | 41   | 41   | 42   | 42   | 42   | 42   |
|                     | 2500 ppm   | 49                      | 49   | 48   |  | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 48   | 47   |
| SMALL STOOL         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 1    | 1    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 2    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 1    |  | 3    | 2    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 0    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 1    | 1    | 0    | 0    | 1    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 3    | 0    | 0    | 0    |
|                     | 1250 ppm   | 1                       | 1    | 2    |  | 2    | 2    | 1    | 0    | 1    | 1    | 2    | 0    | 1    | 1    | 1    |
|                     | 2500 ppm   | 1                       | 1    | 2    |  | 3    | 1    | 1    | 1    | 1    | 1    | 1    | 0    | 1    | 1    | 1    |
| NON REMARKABLE      | Control    | 37                      | 37   | 37   |  | 36   | 36   | 36   | 35   | 36   | 36   | 36   | 35   | 36   | 36   | 35   |
|                     | 625 ppm    | 45                      | 45   | 45   |  | 45   | 45   | 45   | 45   | 43   | 43   | 42   | 41   | 39   | 37   | 37   |
|                     | 1250 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    |  | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ[Crl:BDF1]  
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

SEX : FEMALE

PAGE : 63

| Clinical sign       | Group Name | Administration Week-day |      |      | 88-7 | 89-7 | 90-7 | 91-7 | 92-7 | 93-7 | 94-7 | 95-7 | 96-7 | 97-7 | 98-7 |
|---------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
|                     |            | 85-7                    | 86-7 | 87-7 |      |      |      |      |      |      |      |      |      |      |      |
| CRUSTA              | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 1                       | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    | 1    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| TORTICOLLIS         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
| IRREGULAR BREATHING | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 1    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 2    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 0    | 1    |
| BROWN URINE         | Control    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 41                      | 41   | 41   | 41   | 41   | 40   | 40   | 40   | 39   | 38   | 38   | 38   | 35   | 35   |
|                     | 2500 ppm   | 47                      | 47   | 47   | 46   | 46   | 45   | 45   | 43   | 43   | 43   | 43   | 43   | 43   | 42   |
| SMALL STOOL         | Control    | 0                       | 1    | 1    | 0    | 0    | 0    | 0    | 1    | 1    | 3    | 1    | 0    | 1    | 1    |
|                     | 625 ppm    | 0                       | 1    | 2    | 1    | 1    | 0    | 0    | 0    | 0    | 1    | 0    | 0    | 1    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 2    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 1    | 1    | 1    |
| OLIGO-STOOL         | Control    | 0                       | 0    | 0    | 0    | 0    | 1    | 0    | 2    | 1    | 2    | 0    | 0    | 1    | 1    |
|                     | 625 ppm    | 0                       | 0    | 0    | 0    | 0    | 1    | 1    | 1    | 1    | 2    | 0    | 0    | 0    | 0    |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 2    | 0    | 0    |
|                     | 2500 ppm   | 1                       | 1    | 1    | 1    | 2    | 1    | 1    | 1    | 1    | 1    | 2    | 1    | 1    | 2    |
| NON REMARKABLE      | Control    | 34                      | 34   | 34   | 34   | 33   | 32   | 32   | 29   | 30   | 26   | 27   | 27   | 26   | 26   |
|                     | 625 ppm    | 37                      | 37   | 37   | 35   | 34   | 33   | 34   | 32   | 31   | 28   | 28   | 26   | 25   | 25   |
|                     | 1250 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |
|                     | 2500 ppm   | 0                       | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    | 0    |

(HAN190)

BA1S 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

SEX : FEMALE

PAGE : 64

| Clinical sign       | Group Name | Administration Week-day |       |       |       |       |       |
|---------------------|------------|-------------------------|-------|-------|-------|-------|-------|
|                     |            | 99-7                    | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| CRUSTA              | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 1                       | 1     | 1     | 1     | 1     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| TORTICOLLIS         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
| IRREGULAR BREATHING | Control    | 1                       | 1     | 1     | 0     | 0     | 1     |
|                     | 625 ppm    | 1                       | 2     | 3     | 2     | 3     | 1     |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 1                       | 0     | 0     | 0     | 0     | 0     |
| BROWN URINE         | Control    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 1250 ppm   | 35                      | 34    | 33    | 33    | 33    | 33    |
|                     | 2500 ppm   | 42                      | 40    | 39    | 39    | 38    | 38    |
| SMALL STOOL         | Control    | 0                       | 1     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 1     | 2     | 3     | 3     | 1     |
|                     | 1250 ppm   | 0                       | 1     | 0     | 0     | 0     | 2     |
|                     | 2500 ppm   | 1                       | 0     | 0     | 0     | 0     | 0     |
| OLIGO-STOOL         | Control    | 0                       | 1     | 0     | 0     | 0     | 0     |
|                     | 625 ppm    | 0                       | 3     | 2     | 3     | 3     | 0     |
|                     | 1250 ppm   | 0                       | 0     | 1     | 3     | 2     | 1     |
|                     | 2500 ppm   | 2                       | 1     | 0     | 0     | 0     | 0     |
| NON REMARKABLE      | Control    | 25                      | 25    | 25    | 19    | 21    | 19    |
|                     | 625 ppm    | 25                      | 23    | 21    | 19    | 19    | 19    |
|                     | 1250 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |
|                     | 2500 ppm   | 0                       | 0     | 0     | 0     | 0     | 0     |



TABLE C 1

BODY WEIGHT CHANGES AND  
SURVIVAL ANIMAL NUMBERS: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrlJ [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

MEAN BODY WEIGHTS AND SURVIVAL

PAGE : 1

| Week-Day<br>on Study | Control   |                           | 625 ppm   |                       |                   | 1250 ppm  |                       |                   | 2500 ppm  |                       |                   |
|----------------------|-----------|---------------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|
|                      | Av. Wt.   | No. of<br>Surviv.<br><50> | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 0-0                  | 24.2 (50) | 50/50                     | 24.2 (50) | 100                   | 50/50             | 24.2 (50) | 100                   | 50/50             | 24.2 (50) | 100                   | 50/50             |
| 1-7                  | 25.1 (50) | 50/50                     | 25.0 (50) | 100                   | 50/50             | 24.9 (50) | 99                    | 50/50             | 24.2 (50) | 96                    | 50/50             |
| 2-7                  | 25.9 (50) | 50/50                     | 25.7 (50) | 99                    | 50/50             | 25.7 (50) | 99                    | 50/50             | 24.8 (50) | 96                    | 50/50             |
| 3-7                  | 26.6 (50) | 50/50                     | 26.5 (49) | 100                   | 49/50             | 26.4 (50) | 99                    | 50/50             | 25.5 (50) | 96                    | 50/50             |
| 4-7                  | 27.3 (50) | 50/50                     | 27.5 (49) | 101                   | 49/50             | 27.4 (50) | 100                   | 50/50             | 26.4 (50) | 97                    | 50/50             |
| 5-7                  | 28.3 (50) | 50/50                     | 28.4 (49) | 100                   | 49/50             | 28.2 (50) | 100                   | 50/50             | 27.1 (50) | 96                    | 50/50             |
| 6-7                  | 29.0 (50) | 50/50                     | 29.2 (49) | 101                   | 49/50             | 28.9 (50) | 100                   | 50/50             | 27.4 (50) | 94                    | 50/50             |
| 7-7                  | 30.0 (50) | 50/50                     | 30.1 (49) | 100                   | 49/50             | 29.6 (50) | 99                    | 50/50             | 28.1 (50) | 94                    | 50/50             |
| 8-7                  | 30.8 (50) | 50/50                     | 30.8 (49) | 100                   | 49/50             | 30.3 (50) | 98                    | 50/50             | 28.7 (50) | 93                    | 50/50             |
| 9-7                  | 31.5 (50) | 50/50                     | 31.4 (49) | 100                   | 49/50             | 30.9 (50) | 98                    | 50/50             | 29.2 (49) | 93                    | 49/50             |
| 10-7                 | 32.0 (50) | 50/50                     | 32.0 (49) | 100                   | 49/50             | 31.6 (50) | 99                    | 50/50             | 30.1 (49) | 94                    | 49/50             |
| 11-7                 | 32.6 (50) | 50/50                     | 32.6 (49) | 100                   | 49/50             | 32.2 (50) | 99                    | 50/50             | 30.5 (49) | 94                    | 49/50             |
| 12-7                 | 33.7 (50) | 50/50                     | 33.6 (49) | 100                   | 49/50             | 33.3 (50) | 99                    | 50/50             | 31.3 (49) | 93                    | 49/50             |
| 13-7                 | 34.5 (50) | 50/50                     | 34.4 (49) | 100                   | 49/50             | 34.1 (50) | 99                    | 50/50             | 32.1 (49) | 93                    | 49/50             |
| 14-7                 | 35.3 (50) | 50/50                     | 34.9 (49) | 99                    | 49/50             | 34.8 (50) | 99                    | 50/50             | 32.8 (49) | 93                    | 49/50             |
| 18-7                 | 37.9 (50) | 50/50                     | 37.7 (49) | 99                    | 49/50             | 37.1 (50) | 98                    | 50/50             | 34.8 (49) | 92                    | 49/50             |
| 22-7                 | 40.1 (50) | 50/50                     | 39.7 (49) | 99                    | 49/50             | 39.0 (50) | 97                    | 50/50             | 36.4 (49) | 91                    | 49/50             |
| 26-7                 | 42.0 (50) | 50/50                     | 41.5 (49) | 99                    | 49/50             | 40.7 (50) | 97                    | 50/50             | 37.9 (49) | 90                    | 49/50             |
| 30-7                 | 44.0 (50) | 50/50                     | 43.5 (48) | 99                    | 48/50             | 42.5 (50) | 97                    | 50/50             | 39.3 (49) | 89                    | 49/50             |
| 34-7                 | 45.5 (50) | 50/50                     | 45.0 (48) | 99                    | 48/50             | 43.7 (50) | 96                    | 50/50             | 40.6 (49) | 89                    | 49/50             |
| 38-7                 | 47.2 (50) | 50/50                     | 46.7 (48) | 99                    | 48/50             | 45.3 (50) | 96                    | 50/50             | 42.3 (49) | 90                    | 49/50             |
| 42-7                 | 48.5 (49) | 49/50                     | 48.1 (48) | 99                    | 48/50             | 46.3 (50) | 95                    | 50/50             | 43.4 (49) | 89                    | 49/50             |
| 46-7                 | 49.6 (49) | 49/50                     | 48.8 (48) | 98                    | 48/50             | 47.0 (50) | 95                    | 50/50             | 44.2 (49) | 89                    | 49/50             |
| 50-7                 | 50.5 (49) | 49/50                     | 49.5 (47) | 98                    | 47/50             | 48.0 (50) | 95                    | 50/50             | 44.8 (49) | 89                    | 49/50             |
| 54-7                 | 51.0 (49) | 49/50                     | 49.9 (47) | 98                    | 47/50             | 48.8 (49) | 96                    | 49/50             | 45.4 (49) | 89                    | 49/50             |
| 58-7                 | 52.2 (48) | 48/50                     | 50.6 (47) | 97                    | 47/50             | 49.8 (49) | 95                    | 49/50             | 46.0 (49) | 88                    | 49/50             |
| 62-7                 | 52.0 (48) | 48/50                     | 51.7 (45) | 99                    | 45/50             | 49.7 (48) | 96                    | 48/50             | 46.2 (49) | 89                    | 49/50             |
| 66-7                 | 52.2 (48) | 48/50                     | 51.9 (44) | 99                    | 44/50             | 50.0 (47) | 96                    | 47/50             | 46.7 (49) | 89                    | 49/50             |
| 70-7                 | 52.2 (47) | 47/50                     | 51.6 (44) | 99                    | 44/50             | 50.2 (47) | 96                    | 47/50             | 46.9 (48) | 90                    | 48/50             |
| 74-7                 | 52.6 (47) | 47/50                     | 51.8 (44) | 98                    | 44/50             | 50.8 (45) | 97                    | 45/50             | 47.1 (48) | 90                    | 48/50             |
| 78-7                 | 52.4 (45) | 45/50                     | 52.0 (43) | 99                    | 43/50             | 51.8 (45) | 99                    | 45/50             | 47.8 (48) | 91                    | 48/50             |
| 82-7                 | 52.0 (43) | 43/50                     | 51.9 (42) | 100                   | 42/50             | 52.0 (44) | 100                   | 44/50             | 47.5 (47) | 91                    | 47/50             |
| 86-7                 | 50.8 (41) | 41/50                     | 52.4 (41) | 103                   | 41/50             | 51.3 (44) | 101                   | 44/50             | 47.8 (46) | 94                    | 46/50             |
| 90-7                 | 51.4 (37) | 37/50                     | 52.4 (39) | 102                   | 39/50             | 49.5 (43) | 96                    | 43/50             | 47.0 (46) | 91                    | 46/50             |
| 94-7                 | 51.6 (32) | 32/50                     | 51.0 (38) | 99                    | 38/50             | 49.5 (40) | 96                    | 40/50             | 45.7 (45) | 89                    | 45/50             |
| 98-7                 | 51.7 (28) | 28/50                     | 48.9 (36) | 95                    | 36/50             | 48.3 (38) | 93                    | 38/50             | 44.4 (44) | 86                    | 44/50             |
| 102-7                | 49.5 (27) | 27/50                     | 47.8 (35) | 97                    | 35/50             | 46.8 (34) | 95                    | 34/50             | 43.3 (40) | 87                    | 40/50             |
| 104-7                | 49.4 (24) | 24/50                     | 47.9 (32) | 97                    | 32/50             | 46.3 (31) | 94                    | 31/50             | 43.0 (39) | 87                    | 39/50             |

< >:No. of effective animals, ( ):No. of measured animals

Av. Wt. : g

**TABLE C 2**

**BODY WEIGHT CHANGES AND  
SURVIVAL ANIMAL NUMBERS: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BD01]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

MEAN BODY WEIGHTS AND SURVIVAL

PAGE : 2

| Week-Day<br>on Study | Control   |                           |  | 625 ppm   |                       |                   | 1250 ppm  |                       |                   | 2500 ppm  |                       |                   |
|----------------------|-----------|---------------------------|--|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|
|                      | Av. Wt.   | No. of<br>Surviv.<br><50> |  | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. Wt.   | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 0-0                  | 20.1 (50) | 50/50                     |  | 20.1 (50) | 100                   | 50/50             | 20.1 (50) | 100                   | 50/50             | 20.1 (50) | 100                   | 50/50             |
| 1-7                  | 20.5 (50) | 50/50                     |  | 20.2 (50) | 99                    | 50/50             | 20.1 (50) | 98                    | 50/50             | 19.9 (50) | 97                    | 50/50             |
| 2-7                  | 21.2 (50) | 50/50                     |  | 20.8 (50) | 98                    | 50/50             | 20.8 (50) | 98                    | 50/50             | 20.5 (50) | 97                    | 50/50             |
| 3-7                  | 21.7 (50) | 50/50                     |  | 21.6 (50) | 100                   | 50/50             | 21.2 (50) | 98                    | 50/50             | 21.0 (50) | 97                    | 50/50             |
| 4-7                  | 22.2 (50) | 50/50                     |  | 21.9 (50) | 99                    | 50/50             | 21.8 (50) | 98                    | 50/50             | 21.7 (50) | 98                    | 50/50             |
| 5-7                  | 22.7 (50) | 50/50                     |  | 22.5 (50) | 99                    | 50/50             | 22.3 (50) | 98                    | 50/50             | 22.1 (50) | 97                    | 50/50             |
| 6-7                  | 23.2 (50) | 50/50                     |  | 23.3 (50) | 100                   | 50/50             | 22.7 (50) | 98                    | 50/50             | 22.6 (50) | 97                    | 50/50             |
| 7-7                  | 23.8 (50) | 50/50                     |  | 23.6 (50) | 99                    | 50/50             | 23.2 (50) | 97                    | 50/50             | 23.1 (50) | 97                    | 50/50             |
| 8-7                  | 24.4 (50) | 50/50                     |  | 24.2 (50) | 99                    | 50/50             | 23.8 (50) | 98                    | 50/50             | 23.7 (50) | 97                    | 50/50             |
| 9-7                  | 24.6 (50) | 50/50                     |  | 24.4 (50) | 99                    | 50/50             | 24.1 (50) | 98                    | 50/50             | 23.9 (50) | 97                    | 50/50             |
| 10-7                 | 24.9 (50) | 50/50                     |  | 24.8 (50) | 100                   | 50/50             | 24.5 (50) | 98                    | 50/50             | 24.1 (50) | 97                    | 50/50             |
| 11-7                 | 24.8 (50) | 50/50                     |  | 24.7 (50) | 100                   | 50/50             | 24.2 (50) | 98                    | 50/50             | 24.2 (50) | 98                    | 50/50             |
| 12-7                 | 25.3 (50) | 50/50                     |  | 25.0 (50) | 99                    | 50/50             | 24.6 (50) | 97                    | 50/50             | 24.4 (50) | 96                    | 50/50             |
| 13-7                 | 25.8 (50) | 50/50                     |  | 25.4 (50) | 98                    | 50/50             | 25.1 (50) | 97                    | 50/50             | 25.0 (50) | 97                    | 50/50             |
| 14-7                 | 26.0 (50) | 50/50                     |  | 25.9 (50) | 100                   | 50/50             | 25.8 (50) | 99                    | 50/50             | 25.2 (50) | 97                    | 50/50             |
| 18-7                 | 26.6 (50) | 50/50                     |  | 27.2 (50) | 102                   | 50/50             | 26.3 (50) | 99                    | 50/50             | 26.0 (50) | 98                    | 50/50             |
| 22-7                 | 28.6 (49) | 49/50                     |  | 28.7 (50) | 100                   | 50/50             | 28.1 (50) | 98                    | 50/50             | 27.1 (50) | 95                    | 50/50             |
| 26-7                 | 29.3 (49) | 49/50                     |  | 29.6 (50) | 101                   | 50/50             | 28.8 (50) | 98                    | 50/50             | 27.5 (50) | 94                    | 50/50             |
| 30-7                 | 30.5 (49) | 49/50                     |  | 30.5 (50) | 100                   | 50/50             | 30.1 (50) | 99                    | 50/50             | 28.4 (50) | 93                    | 50/50             |
| 34-7                 | 32.1 (49) | 49/50                     |  | 31.7 (50) | 99                    | 50/50             | 31.0 (50) | 97                    | 50/50             | 29.3 (50) | 91                    | 50/50             |
| 38-7                 | 33.3 (49) | 49/50                     |  | 33.0 (50) | 99                    | 50/50             | 31.7 (50) | 95                    | 50/50             | 30.3 (50) | 91                    | 50/50             |
| 42-7                 | 34.0 (49) | 49/50                     |  | 33.9 (50) | 100                   | 50/50             | 32.5 (50) | 96                    | 50/50             | 30.9 (50) | 91                    | 50/50             |
| 46-7                 | 34.9 (49) | 49/50                     |  | 34.5 (50) | 99                    | 50/50             | 32.9 (50) | 94                    | 50/50             | 31.5 (50) | 90                    | 50/50             |
| 50-7                 | 35.4 (49) | 49/50                     |  | 35.5 (50) | 100                   | 50/50             | 33.6 (50) | 95                    | 50/50             | 32.1 (50) | 91                    | 50/50             |
| 54-7                 | 36.6 (49) | 49/50                     |  | 36.3 (49) | 99                    | 49/50             | 33.9 (50) | 93                    | 50/50             | 32.7 (50) | 89                    | 50/50             |
| 58-7                 | 37.3 (49) | 49/50                     |  | 37.1 (49) | 99                    | 49/50             | 34.7 (49) | 93                    | 49/50             | 33.0 (50) | 88                    | 50/50             |
| 62-7                 | 37.1 (47) | 47/50                     |  | 36.9 (48) | 99                    | 48/50             | 34.3 (48) | 92                    | 48/50             | 32.7 (50) | 88                    | 50/50             |
| 66-7                 | 36.8 (46) | 46/50                     |  | 36.6 (48) | 99                    | 48/50             | 34.5 (48) | 94                    | 48/50             | 32.4 (50) | 88                    | 50/50             |
| 70-7                 | 37.1 (41) | 41/50                     |  | 36.3 (48) | 98                    | 48/50             | 34.3 (46) | 92                    | 46/50             | 32.0 (50) | 86                    | 50/50             |
| 74-7                 | 37.6 (39) | 39/50                     |  | 36.8 (47) | 98                    | 47/50             | 35.0 (45) | 93                    | 45/50             | 32.1 (48) | 85                    | 48/50             |
| 78-7                 | 38.2 (39) | 39/50                     |  | 37.5 (47) | 98                    | 47/50             | 35.4 (42) | 93                    | 42/50             | 32.5 (48) | 85                    | 48/50             |
| 82-7                 | 38.0 (38) | 38/50                     |  | 37.0 (47) | 97                    | 47/50             | 35.0 (42) | 92                    | 42/50             | 32.4 (48) | 85                    | 48/50             |
| 86-7                 | 37.8 (38) | 38/50                     |  | 37.3 (43) | 99                    | 43/50             | 35.2 (41) | 93                    | 41/50             | 32.7 (47) | 87                    | 47/50             |
| 90-7                 | 37.8 (37) | 37/50                     |  | 36.6 (39) | 97                    | 39/50             | 34.7 (40) | 92                    | 40/50             | 32.1 (45) | 85                    | 45/50             |
| 94-7                 | 37.3 (33) | 33/50                     |  | 35.7 (38) | 96                    | 38/50             | 33.9 (38) | 91                    | 38/50             | 31.2 (43) | 84                    | 43/50             |
| 98-7                 | 37.1 (31) | 31/50                     |  | 35.5 (33) | 96                    | 33/50             | 33.9 (35) | 91                    | 35/50             | 30.4 (42) | 82                    | 42/50             |
| 102-7                | 36.3 (25) | 25/50                     |  | 34.8 (24) | 96                    | 24/50             | 33.7 (33) | 93                    | 33/50             | 30.8 (39) | 85                    | 39/50             |
| 104-7                | 37.0 (24) | 24/50                     |  | 34.5 (21) | 93                    | 21/50             | 34.4 (33) | 93                    | 33/50             | 30.5 (38) | 82                    | 38/50             |

< >:No. of effective animals, ( ):No. of measured animals

Av. Wt.: g

TABLE C 3

BODY WEIGHT CHANGES: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 1

| Group Name | Administration week-day |              |              |              |              |              |              |
|------------|-------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
|            | 0-0                     | 1-7          | 2-7          | 3-7          | 4-7          | 5-7          | 6-7          |
| Control    | 24.2 ± 0.7              | 25.1 ± 0.9   | 25.9 ± 0.9   | 26.6 ± 1.1   | 27.3 ± 1.3   | 28.3 ± 1.4   | 29.0 ± 1.5   |
| 625 ppm    | 24.2 ± 0.7              | 25.0 ± 0.7   | 25.7 ± 1.0   | 26.5 ± 1.1   | 27.5 ± 1.1   | 28.4 ± 1.2   | 29.2 ± 1.2   |
| 1250 ppm   | 24.2 ± 0.7              | 24.9 ± 0.9   | 25.7 ± 0.9   | 26.4 ± 1.0   | 27.4 ± 1.1   | 28.2 ± 1.2   | 28.9 ± 1.3   |
| 2500 ppm   | 24.2 ± 0.7              | 24.2 ± 1.0** | 24.8 ± 1.4** | 25.5 ± 1.3** | 26.4 ± 1.5** | 27.1 ± 1.5** | 27.4 ± 1.8** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 2

| Group Name  | Administration week-day |              |              |              |              |              |              |
|---|-------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
|   | 7-7                     | 8-7          | 9-7          | 10-7         | 11-7         | 12-7         | 13-7         |
| Control   | 30.0 ± 1.6              | 30.8 ± 1.7   | 31.5 ± 1.9   | 32.0 ± 2.0   | 32.6 ± 2.1   | 33.7 ± 2.2   | 34.5 ± 2.4   |
| 625 ppm   | 30.1 ± 1.4              | 30.8 ± 1.5   | 31.4 ± 1.7   | 32.0 ± 1.7   | 32.6 ± 1.7   | 33.6 ± 1.9   | 34.4 ± 2.0   |
| 1250 ppm  | 29.6 ± 1.5              | 30.3 ± 1.6   | 30.9 ± 1.8   | 31.6 ± 2.0   | 32.2 ± 2.1   | 33.3 ± 2.3   | 34.1 ± 2.4   |
| 2500 ppm  | 28.1 ± 2.1**            | 28.7 ± 2.2** | 29.2 ± 1.9** | 30.1 ± 2.1** | 30.5 ± 2.2** | 31.3 ± 2.3** | 32.1 ± 2.5** |
| Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett |                         |              |              |              |              |              |              |

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Cri:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 3

| Group Name | Administration week-day |       | 14-7  |       | 18-7  |       | 22-7  |       | 26-7  |       | 30-7  |       | 34-7  |       | 38-7 |  |
|------------|-------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|------|--|
|            |                         |       |       |       |       |       |       |       |       |       |       |       |       |       |      |  |
| Control    | 35.3±                   | 2.5   | 37.9± | 2.6   | 40.1± | 3.1   | 42.0± | 3.4   | 44.0± | 3.7   | 45.5± | 3.6   | 47.2± | 3.5   |      |  |
| 625 ppm    | 34.9±                   | 2.0   | 37.7± | 2.6   | 39.7± | 2.8   | 41.5± | 3.1   | 43.5± | 3.5   | 45.0± | 3.5   | 46.7± | 3.5   |      |  |
| 1250 ppm   | 34.8±                   | 2.4   | 37.1± | 2.8   | 39.0± | 3.4   | 40.7± | 3.8   | 42.5± | 4.1   | 43.7± | 4.2*  | 45.3± | 4.2*  |      |  |
| 2500 ppm   | 32.8±                   | 2.5** | 34.8± | 2.7** | 36.4± | 3.0** | 37.9± | 3.4** | 39.3± | 3.6** | 40.6± | 3.7** | 42.3± | 3.9** |      |  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 4

| Group Name | Administration |       | week-day |       |       |       |       |       |       |       |       |       |
|------------|----------------|-------|----------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
|            | 42-7           |       | 46-7     |       | 50-7  |       | 54-7  |       | 58-7  |       | 62-7  |       |
| Control    | 48.5±          | 3.5   | 49.6±    | 3.6   | 50.5± | 3.7   | 51.0± | 3.4   | 52.2± | 3.6   | 52.0± | 3.6   |
| 625 ppm    | 48.1±          | 3.6   | 48.8±    | 3.6   | 49.5± | 3.7   | 49.9± | 4.4   | 50.6± | 5.4   | 51.7± | 3.5   |
| 1250 ppm   | 46.3±          | 4.0*  | 47.0±    | 4.2** | 48.0± | 4.7** | 48.8± | 5.0*  | 49.8± | 5.1*  | 49.7± | 5.4*  |
| 2500 ppm   | 43.4±          | 4.1** | 44.2±    | 4.2** | 44.8± | 4.5** | 45.4± | 5.5** | 46.0± | 5.5** | 46.2± | 5.5** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 5

| Group Name | Administration week-day |       | 70-7   |       | 74-7   |       | 78-7   |       | 82-7   |     | 86-7   |      | 90-7   |       | 94-7 |  |
|------------|-------------------------|-------|--------|-------|--------|-------|--------|-------|--------|-----|--------|------|--------|-------|------|--|
|            |                         |       |        |       |        |       |        |       |        |     |        |      |        |       |      |  |
| Control    | 52.2 ±                  | 4.6   | 52.6 ± | 4.7   | 52.4 ± | 5.5   | 52.0 ± | 5.5   | 50.8 ± | 7.6 | 51.4 ± | 7.3  | 51.6 ± | 7.7   |      |  |
| 625 ppm    | 51.6 ±                  | 3.6   | 51.8 ± | 4.2   | 52.0 ± | 5.0   | 51.9 ± | 5.5   | 52.4 ± | 5.3 | 52.4 ± | 5.1  | 51.0 ± | 5.7   |      |  |
| 1250 ppm   | 50.2 ±                  | 5.5   | 50.8 ± | 6.1   | 51.8 ± | 6.0   | 52.0 ± | 5.9   | 51.3 ± | 6.4 | 49.5 ± | 7.9  | 49.5 ± | 7.2   |      |  |
| 2500 ppm   | 46.9 ±                  | 4.9** | 47.1 ± | 5.3** | 47.8 ± | 5.6** | 47.5 ± | 6.0** | 47.8 ± | 6.3 | 47.0 ± | 6.5* | 45.7 ± | 6.7** |      |  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 6

| Group Name | Administration week-day |             |             |
|------------|-------------------------|-------------|-------------|
|            | 98-7                    | 102-7       | 104-7       |
| Control    | 51.7± 6.4               | 49.5± 7.5   | 49.4± 7.6   |
| 625 ppm    | 48.9± 6.6               | 47.8± 6.4   | 47.9± 6.2   |
| 1250 ppm   | 48.3± 7.9               | 46.8± 8.2   | 46.3± 9.3   |
| 2500 ppm   | 44.4± 7.5**             | 43.3± 7.6** | 43.0± 7.8** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

**TABLE C 4**

**BODY WEIGHT CHANGES: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 7

| Group Name | Administration week-day |             |             |             |           |            |           |
|------------|-------------------------|-------------|-------------|-------------|-----------|------------|-----------|
|            | 0-0                     | 1-7         | 2-7         | 3-7         | 4-7       | 5-7        | 6-7       |
| Control    | 20.1± 0.8               | 20.5± 0.8   | 21.2± 1.1   | 21.7± 1.1   | 22.2± 1.3 | 22.7± 1.4  | 23.2± 1.3 |
| 625 ppm    | 20.1± 0.8               | 20.2± 0.8   | 20.8± 1.0   | 21.6± 1.1   | 21.9± 1.1 | 22.5± 1.1  | 23.3± 1.3 |
| 1250 ppm   | 20.1± 0.8               | 20.1± 0.9*  | 20.8± 0.9   | 21.2± 1.0   | 21.8± 1.0 | 22.3± 1.1  | 22.7± 1.1 |
| 2500 ppm   | 20.1± 0.8               | 19.9± 0.8** | 20.5± 0.8** | 21.0± 0.9** | 21.7± 1.0 | 22.1± 1.0* | 22.6± 1.2 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 8

| Group Name | Administration week-day |           |           |            |           |            |           |
|------------|-------------------------|-----------|-----------|------------|-----------|------------|-----------|
|            | 7-7                     | 8-7       | 9-7       | 10-7       | 11-7      | 12-7       | 13-7      |
| Control    | 23.8± 1.5               | 24.4± 1.7 | 24.6± 2.2 | 24.9± 2.0  | 24.8± 2.1 | 25.3± 2.3  | 25.8± 2.7 |
| 625 ppm    | 23.6± 1.4               | 24.2± 1.5 | 24.4± 1.5 | 24.8± 1.7  | 24.7± 1.6 | 25.0± 1.7  | 25.4± 2.0 |
| 1250 ppm   | 23.2± 1.2*              | 23.8± 1.1 | 24.1± 1.3 | 24.5± 1.3  | 24.2± 1.5 | 24.6± 1.5  | 25.1± 1.6 |
| 2500 ppm   | 23.1± 1.2*              | 23.7± 1.3 | 23.9± 1.4 | 24.1± 1.4* | 24.2± 1.5 | 24.4± 1.7* | 25.0± 1.7 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 9

| Group Name | Administration week-day |           |            |            |            |             |             |
|------------|-------------------------|-----------|------------|------------|------------|-------------|-------------|
|            | 14-7                    | 18-7      | 22-7       | 26-7       | 30-7       | 34-7        | 38-7        |
| Control    | 26.0± 2.7               | 26.6± 2.9 | 28.6± 3.8  | 29.3± 3.9  | 30.5± 4.7  | 32.1± 5.1   | 33.3± 4.9   |
| 625 ppm    | 25.9± 2.1               | 27.2± 2.1 | 28.7± 2.7  | 29.6± 3.4  | 30.5± 3.5  | 31.7± 3.9   | 33.0± 4.4   |
| 1250 ppm   | 25.8± 1.7               | 26.3± 2.0 | 28.1± 2.3  | 28.8± 2.8  | 30.1± 3.2  | 31.0± 3.5   | 31.7± 3.4   |
| 2500 ppm   | 25.2± 1.9               | 26.0± 2.1 | 27.1± 2.3* | 27.5± 2.6* | 28.4± 2.7* | 29.3± 2.8** | 30.3± 2.8** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 10

| Group Name | Administration week-day |  | 46-7        | 50-7        | 54-7        | 58-7        | 62-7        | 66-7        |
|------------|-------------------------|--|-------------|-------------|-------------|-------------|-------------|-------------|
|            | 42-7                    |  |             |             |             |             |             |             |
| Control    | 34.0± 5.9               |  | 34.9± 5.9   | 35.4± 6.3   | 36.6± 6.5   | 37.3± 5.9   | 37.1± 4.8   | 36.8± 4.9   |
| 625 ppm    | 33.9± 4.1               |  | 34.5± 4.3   | 35.5± 4.6   | 36.3± 4.6   | 37.1± 4.8   | 36.9± 4.8   | 36.6± 5.2   |
| 1250 ppm   | 32.5± 3.6               |  | 32.9± 3.7   | 33.6± 4.1   | 33.9± 4.2   | 34.7± 4.8   | 34.3± 4.5** | 34.5± 4.8   |
| 2500 ppm   | 30.9± 3.0**             |  | 31.5± 3.1** | 32.1± 3.5** | 32.7± 3.3** | 33.0± 3.4** | 32.7± 3.4** | 32.4± 3.2** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 11

| Group Name | Administration week-day |              |              |              |              |              |              |  |
|------------|-------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--|
|            | 70-7                    | 74-7         | 78-7         | 82-7         | 86-7         | 90-7         | 94-7         |  |
| Control    | 37.1 ± 5.2              | 37.6 ± 4.9   | 38.2 ± 4.9   | 38.0 ± 4.9   | 37.8 ± 4.6   | 37.8 ± 5.4   | 37.3 ± 6.2   |  |
| 625 ppm    | 36.3 ± 5.3              | 36.8 ± 5.0   | 37.5 ± 5.3   | 37.0 ± 5.8   | 37.3 ± 5.6   | 36.6 ± 6.3   | 35.7 ± 6.8   |  |
| 1250 ppm   | 34.3 ± 4.7*             | 35.0 ± 5.5*  | 35.4 ± 5.3*  | 35.0 ± 5.6*  | 35.2 ± 5.8   | 34.7 ± 5.8*  | 33.9 ± 6.0*  |  |
| 2500 ppm   | 32.0 ± 3.9**            | 32.1 ± 4.1** | 32.5 ± 4.7** | 32.4 ± 4.5** | 32.7 ± 5.1** | 32.1 ± 4.5** | 31.2 ± 4.7** |  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 12

| Group Name | Administration week-day |              |              |
|------------|-------------------------|--------------|--------------|
|            | 98-7                    | 102-7        | 104-7        |
| Control    | 37.1 ± 5.8              | 36.3 ± 6.3   | 37.0 ± 7.0   |
| 625 ppm    | 35.5 ± 6.4              | 34.8 ± 6.0   | 34.5 ± 5.7   |
| 1250 ppm   | 33.9 ± 5.9              | 33.7 ± 5.8   | 34.4 ± 7.3   |
| 2500 ppm   | 30.4 ± 4.6**            | 30.8 ± 4.3** | 30.5 ± 4.6** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

TABLE D 1

FOOD CONSUMPTION CHANGES AND  
SURVIVAL ANIMAL NUMBERS: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

MEAN FOOD CONSUMPTION (FC) AND SURVIVAL

PAGE : 1

| Week-Day<br>on Study | Control  |                           |  | 625 ppm  |                       |                   | 1250 ppm |                       |                   | 2500 ppm |                       |                   |
|----------------------|----------|---------------------------|--|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
|                      | Av. FC.  | No. of<br>Surviv.<br><50> |  | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 1-7                  | 4.0 (50) | 50/50                     |  | 3.9 (50) | 98                    | 50/50             | 3.9 (50) | 98                    | 50/50             | 3.5 (50) | 88                    | 50/50             |
| 2-7                  | 3.9 (50) | 50/50                     |  | 3.8 (50) | 97                    | 50/50             | 3.8 (50) | 97                    | 50/50             | 3.7 (50) | 95                    | 50/50             |
| 3-7                  | 3.8 (50) | 50/50                     |  | 3.7 (49) | 97                    | 49/50             | 3.7 (50) | 97                    | 50/50             | 3.6 (50) | 95                    | 50/50             |
| 4-7                  | 3.8 (50) | 50/50                     |  | 3.8 (49) | 100                   | 49/50             | 3.8 (50) | 100                   | 50/50             | 3.7 (50) | 97                    | 50/50             |
| 5-7                  | 3.9 (50) | 50/50                     |  | 3.8 (49) | 97                    | 49/50             | 3.8 (50) | 97                    | 50/50             | 3.7 (50) | 95                    | 50/50             |
| 6-7                  | 3.9 (50) | 50/50                     |  | 3.8 (49) | 97                    | 49/50             | 3.8 (50) | 97                    | 50/50             | 3.7 (50) | 95                    | 50/50             |
| 7-7                  | 4.0 (50) | 50/50                     |  | 3.9 (49) | 98                    | 49/50             | 3.9 (50) | 98                    | 50/50             | 3.7 (50) | 93                    | 50/50             |
| 8-7                  | 4.0 (50) | 50/50                     |  | 3.9 (49) | 98                    | 49/50             | 3.9 (50) | 98                    | 50/50             | 3.8 (50) | 95                    | 50/50             |
| 9-7                  | 4.2 (50) | 50/50                     |  | 4.0 (49) | 95                    | 49/50             | 4.0 (50) | 95                    | 50/50             | 3.9 (49) | 93                    | 49/50             |
| 10-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.1 (50) | 98                    | 50/50             | 3.9 (49) | 93                    | 49/50             |
| 11-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.1 (50) | 98                    | 50/50             | 4.0 (49) | 95                    | 49/50             |
| 12-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.1 (50) | 98                    | 50/50             | 4.0 (49) | 95                    | 49/50             |
| 13-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.1 (50) | 98                    | 50/50             | 4.0 (49) | 95                    | 49/50             |
| 14-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.2 (50) | 100                   | 50/50             | 4.1 (49) | 98                    | 49/50             |
| 18-7                 | 4.2 (50) | 50/50                     |  | 4.2 (49) | 100                   | 49/50             | 4.1 (50) | 98                    | 50/50             | 4.0 (49) | 95                    | 49/50             |
| 22-7                 | 4.3 (50) | 50/50                     |  | 4.1 (49) | 95                    | 49/50             | 4.1 (50) | 95                    | 50/50             | 4.0 (49) | 93                    | 49/50             |
| 26-7                 | 4.2 (50) | 50/50                     |  | 4.1 (49) | 98                    | 49/50             | 4.0 (50) | 95                    | 50/50             | 3.9 (49) | 93                    | 49/50             |
| 30-7                 | 4.4 (50) | 50/50                     |  | 4.2 (48) | 95                    | 48/50             | 4.1 (50) | 93                    | 50/50             | 4.0 (49) | 91                    | 49/50             |
| 34-7                 | 4.5 (50) | 50/50                     |  | 4.5 (48) | 100                   | 48/50             | 4.4 (50) | 98                    | 50/50             | 4.3 (49) | 96                    | 49/50             |
| 38-7                 | 4.5 (50) | 50/50                     |  | 4.5 (48) | 100                   | 48/50             | 4.4 (50) | 98                    | 50/50             | 4.4 (49) | 98                    | 49/50             |
| 42-7                 | 4.6 (49) | 49/50                     |  | 4.6 (48) | 100                   | 48/50             | 4.5 (50) | 98                    | 50/50             | 4.4 (49) | 96                    | 49/50             |
| 46-7                 | 4.7 (49) | 49/50                     |  | 4.6 (48) | 98                    | 48/50             | 4.5 (50) | 96                    | 50/50             | 4.4 (49) | 94                    | 49/50             |
| 50-7                 | 4.8 (49) | 49/50                     |  | 4.7 (47) | 98                    | 47/50             | 4.7 (50) | 98                    | 50/50             | 4.5 (49) | 94                    | 49/50             |
| 54-7                 | 4.6 (49) | 49/50                     |  | 4.7 (47) | 102                   | 47/50             | 4.6 (49) | 100                   | 49/50             | 4.6 (49) | 100                   | 49/50             |
| 58-7                 | 5.0 (48) | 48/50                     |  | 4.9 (47) | 98                    | 47/50             | 4.8 (49) | 96                    | 49/50             | 4.6 (49) | 92                    | 49/50             |
| 62-7                 | 4.7 (48) | 48/50                     |  | 4.7 (45) | 100                   | 45/50             | 4.6 (48) | 98                    | 48/50             | 4.5 (49) | 96                    | 49/50             |
| 66-7                 | 4.7 (48) | 48/50                     |  | 4.7 (44) | 100                   | 44/50             | 4.5 (47) | 96                    | 47/50             | 4.4 (49) | 94                    | 49/50             |
| 70-7                 | 4.9 (47) | 47/50                     |  | 4.8 (44) | 98                    | 44/50             | 4.7 (47) | 96                    | 47/50             | 4.5 (48) | 92                    | 48/50             |
| 74-7                 | 4.8 (47) | 47/50                     |  | 4.6 (44) | 96                    | 44/50             | 4.6 (45) | 96                    | 45/50             | 4.5 (48) | 94                    | 48/50             |
| 78-7                 | 4.8 (45) | 45/50                     |  | 4.7 (43) | 98                    | 43/50             | 4.7 (45) | 98                    | 45/50             | 4.4 (48) | 92                    | 48/50             |
| 82-7                 | 4.7 (43) | 43/50                     |  | 4.7 (42) | 100                   | 42/50             | 4.6 (44) | 98                    | 44/50             | 4.4 (47) | 94                    | 47/50             |
| 86-7                 | 4.6 (41) | 41/50                     |  | 4.9 (41) | 107                   | 41/50             | 4.7 (44) | 102                   | 44/50             | 4.6 (46) | 100                   | 46/50             |
| 90-7                 | 4.8 (37) | 37/50                     |  | 4.7 (39) | 98                    | 39/50             | 4.7 (43) | 98                    | 43/50             | 4.6 (46) | 96                    | 46/50             |
| 94-7                 | 4.7 (32) | 32/50                     |  | 4.6 (38) | 98                    | 38/50             | 4.7 (40) | 100                   | 40/50             | 4.4 (45) | 94                    | 45/50             |
| 98-7                 | 4.9 (28) | 28/50                     |  | 4.9 (36) | 100                   | 36/50             | 4.7 (38) | 96                    | 38/50             | 4.5 (44) | 92                    | 44/50             |
| 102-7                | 4.8 (27) | 27/50                     |  | 4.9 (35) | 102                   | 35/50             | 4.7 (34) | 98                    | 34/50             | 4.5 (40) | 94                    | 40/50             |
| 104-7                | 4.8 (24) | 24/50                     |  | 4.8 (32) | 100                   | 32/50             | 4.7 (31) | 98                    | 31/50             | 4.4 (39) | 92                    | 39/50             |

< >:No. of effective animals, ( ):No. of measured animals

Av. FC. : g

**TABLE D 2**

**FOOD CONSUMPTION CHANGES AND  
SURVIVAL ANIMAL NUMBERS: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

MEAN FOOD CONSUMPTION (FC) AND SURVIVAL

PAGE : 2

| Week-Day<br>on Study | Control  |                           |  | 625 ppm  |                       |                   | 1250 ppm |                       |                   | 2500 ppm |                       |                   |
|----------------------|----------|---------------------------|--|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
|                      | Av. FC.  | No. of<br>Surviv.<br><50> |  | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. FC.  | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 1-7                  | 3.6 (50) | 50/50                     |  | 3.5 (50) | 97                    | 50/50             | 3.5 (50) | 97                    | 50/50             | 3.3 (50) | 92                    | 50/50             |
| 2-7                  | 3.4 (50) | 50/50                     |  | 3.4 (50) | 100                   | 50/50             | 3.4 (50) | 100                   | 50/50             | 3.4 (50) | 100                   | 50/50             |
| 3-7                  | 3.4 (50) | 50/50                     |  | 3.4 (50) | 100                   | 50/50             | 3.3 (50) | 97                    | 50/50             | 3.3 (50) | 97                    | 50/50             |
| 4-7                  | 3.5 (50) | 50/50                     |  | 3.5 (50) | 100                   | 50/50             | 3.4 (50) | 97                    | 50/50             | 3.5 (50) | 100                   | 50/50             |
| 5-7                  | 3.5 (50) | 50/50                     |  | 3.5 (50) | 100                   | 50/50             | 3.5 (49) | 100                   | 50/50             | 3.5 (50) | 100                   | 50/50             |
| 6-7                  | 3.6 (50) | 50/50                     |  | 3.6 (50) | 100                   | 50/50             | 3.5 (50) | 97                    | 50/50             | 3.6 (50) | 100                   | 50/50             |
| 7-7                  | 3.7 (50) | 50/50                     |  | 3.6 (50) | 97                    | 50/50             | 3.6 (50) | 97                    | 50/50             | 3.7 (50) | 100                   | 50/50             |
| 8-7                  | 3.8 (50) | 50/50                     |  | 3.7 (50) | 97                    | 50/50             | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             |
| 9-7                  | 3.9 (50) | 50/50                     |  | 3.8 (50) | 97                    | 50/50             | 3.9 (50) | 100                   | 50/50             | 3.8 (50) | 97                    | 50/50             |
| 10-7                 | 3.8 (50) | 50/50                     |  | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             |
| 11-7                 | 3.9 (50) | 50/50                     |  | 3.8 (50) | 97                    | 50/50             | 3.8 (50) | 97                    | 50/50             | 3.9 (50) | 100                   | 50/50             |
| 12-7                 | 3.8 (50) | 50/50                     |  | 3.7 (50) | 97                    | 50/50             | 3.7 (50) | 97                    | 50/50             | 3.8 (50) | 100                   | 50/50             |
| 13-7                 | 3.9 (50) | 50/50                     |  | 3.8 (50) | 97                    | 50/50             | 3.8 (50) | 97                    | 50/50             | 3.9 (50) | 100                   | 50/50             |
| 14-7                 | 3.8 (50) | 50/50                     |  | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             |
| 18-7                 | 3.8 (50) | 50/50                     |  | 3.7 (50) | 97                    | 50/50             | 3.8 (50) | 100                   | 50/50             | 3.8 (50) | 100                   | 50/50             |
| 22-7                 | 3.9 (49) | 49/50                     |  | 3.8 (50) | 97                    | 50/50             | 3.8 (50) | 97                    | 50/50             | 3.8 (50) | 97                    | 50/50             |
| 26-7                 | 3.7 (49) | 49/50                     |  | 3.7 (50) | 100                   | 50/50             | 3.6 (50) | 97                    | 50/50             | 3.5 (50) | 95                    | 50/50             |
| 30-7                 | 3.8 (49) | 49/50                     |  | 3.6 (50) | 95                    | 50/50             | 3.7 (50) | 97                    | 50/50             | 3.6 (50) | 95                    | 50/50             |
| 34-7                 | 4.1 (49) | 49/50                     |  | 4.0 (50) | 98                    | 50/50             | 4.0 (50) | 98                    | 50/50             | 4.0 (50) | 98                    | 50/50             |
| 38-7                 | 4.3 (49) | 49/50                     |  | 4.1 (50) | 95                    | 50/50             | 4.1 (50) | 95                    | 50/50             | 4.0 (50) | 93                    | 50/50             |
| 42-7                 | 4.2 (49) | 49/50                     |  | 4.2 (50) | 100                   | 50/50             | 4.2 (50) | 100                   | 50/50             | 4.2 (50) | 100                   | 50/50             |
| 46-7                 | 4.2 (49) | 49/50                     |  | 4.2 (50) | 100                   | 50/50             | 4.1 (50) | 98                    | 50/50             | 4.2 (50) | 100                   | 50/50             |
| 50-7                 | 4.2 (49) | 49/50                     |  | 4.4 (50) | 105                   | 50/50             | 4.1 (50) | 98                    | 50/50             | 4.1 (50) | 98                    | 50/50             |
| 54-7                 | 4.2 (49) | 49/50                     |  | 4.2 (49) | 100                   | 49/50             | 4.1 (50) | 98                    | 50/50             | 4.1 (50) | 98                    | 50/50             |
| 58-7                 | 4.1 (49) | 49/50                     |  | 4.2 (49) | 102                   | 49/50             | 4.1 (49) | 100                   | 49/50             | 4.1 (50) | 100                   | 50/50             |
| 62-7                 | 4.2 (47) | 47/50                     |  | 4.1 (48) | 98                    | 48/50             | 3.9 (48) | 93                    | 48/50             | 4.0 (50) | 95                    | 50/50             |
| 66-7                 | 3.9 (46) | 46/50                     |  | 3.8 (48) | 97                    | 48/50             | 3.8 (48) | 97                    | 48/50             | 3.8 (50) | 97                    | 50/50             |
| 70-7                 | 4.0 (41) | 41/50                     |  | 3.9 (48) | 98                    | 48/50             | 3.8 (46) | 95                    | 46/50             | 3.8 (50) | 95                    | 50/50             |
| 74-7                 | 4.1 (39) | 39/50                     |  | 4.0 (47) | 98                    | 47/50             | 3.7 (45) | 90                    | 45/50             | 3.7 (48) | 90                    | 48/50             |
| 78-7                 | 4.2 (39) | 39/50                     |  | 4.1 (47) | 98                    | 47/50             | 4.0 (42) | 95                    | 42/50             | 3.8 (48) | 90                    | 48/50             |
| 82-7                 | 4.2 (38) | 38/50                     |  | 4.3 (47) | 102                   | 47/50             | 4.0 (42) | 95                    | 42/50             | 3.9 (48) | 93                    | 48/50             |
| 86-7                 | 4.3 (38) | 38/50                     |  | 4.2 (43) | 98                    | 43/50             | 3.9 (41) | 91                    | 41/50             | 3.8 (47) | 88                    | 47/50             |
| 90-7                 | 4.2 (37) | 37/50                     |  | 4.2 (39) | 100                   | 39/50             | 3.9 (40) | 93                    | 40/50             | 3.9 (45) | 93                    | 45/50             |
| 94-7                 | 4.0 (33) | 33/50                     |  | 4.0 (38) | 100                   | 38/50             | 3.9 (38) | 98                    | 38/50             | 3.9 (43) | 98                    | 43/50             |
| 98-7                 | 4.4 (31) | 31/50                     |  | 4.4 (33) | 100                   | 33/50             | 4.3 (35) | 98                    | 35/50             | 3.9 (42) | 89                    | 42/50             |
| 102-7                | 4.2 (25) | 25/50                     |  | 4.3 (24) | 102                   | 24/50             | 4.1 (33) | 98                    | 33/50             | 4.0 (39) | 95                    | 39/50             |
| 104-7                | 4.3 (24) | 24/50                     |  | 4.5 (21) | 105                   | 21/50             | 4.5 (33) | 105                   | 33/50             | 4.1 (38) | 95                    | 38/50             |

< >:No. of effective animals, ( ):No. of measured animals

Av. FC. : g

**TABLE D 3**

**FOOD CONSUMPTION CHANGES: MALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 1

| Group Name | Administration week-day (effective) |            |           |             |             |             |             |
|------------|-------------------------------------|------------|-----------|-------------|-------------|-------------|-------------|
|            | 1-7 (7)                             | 2-7 (7)    | 3-7 (7)   | 4-7 (7)     | 5-7 (7)     | 6-7 (7)     | 7-7 (7)     |
| Control    | 4.0 ± 0.3                           | 3.9 ± 0.2  | 3.8 ± 0.3 | 3.8 ± 0.3   | 3.9 ± 0.3   | 3.9 ± 0.2   | 4.0 ± 0.3   |
| 625 ppm    | 3.9 ± 0.2                           | 3.8 ± 0.4  | 3.7 ± 0.4 | 3.8 ± 0.2   | 3.8 ± 0.2   | 3.8 ± 0.3   | 3.9 ± 0.2   |
| 1250 ppm   | 3.9 ± 0.3*                          | 3.8 ± 0.3  | 3.7 ± 0.3 | 3.8 ± 0.4   | 3.8 ± 0.3   | 3.8 ± 0.2   | 3.9 ± 0.3*  |
| 2500 ppm   | 3.5 ± 0.2**                         | 3.7 ± 0.3* | 3.6 ± 0.3 | 3.7 ± 0.3** | 3.7 ± 0.3** | 3.7 ± 0.3** | 3.7 ± 0.3** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 2

| Group Name | Administration<br>8-7 (7) | week-day (effective)<br>9-7 (7) | 10-7 (7)    | 11-7 (7)    | 12-7 (7)    | 13-7 (7)    | 14-7 (7)  |
|------------|---------------------------|---------------------------------|-------------|-------------|-------------|-------------|-----------|
| Control    | 4.0 ± 0.3                 | 4.2 ± 0.3                       | 4.2 ± 0.3   | 4.2 ± 0.3   | 4.2 ± 0.3   | 4.2 ± 0.3   | 4.2 ± 0.3 |
| 625 ppm    | 3.9 ± 0.3                 | 4.0 ± 0.2                       | 4.1 ± 0.3   | 4.1 ± 0.3   | 4.1 ± 0.3   | 4.1 ± 0.3   | 4.1 ± 0.3 |
| 1250 ppm   | 3.9 ± 0.3                 | 4.0 ± 0.3                       | 4.1 ± 0.3   | 4.1 ± 0.3   | 4.1 ± 0.3*  | 4.1 ± 0.3   | 4.2 ± 0.3 |
| 2500 ppm   | 3.8 ± 0.3**               | 3.9 ± 0.2**                     | 3.9 ± 0.3** | 4.0 ± 0.3** | 4.0 ± 0.3** | 4.0 ± 0.3** | 4.1 ± 0.3 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crlj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 3

| Group Name | Administration week-day (effective) |             |             |             |             |            |             |
|------------|-------------------------------------|-------------|-------------|-------------|-------------|------------|-------------|
|            | 18-7 (7)                            | 22-7 (7)    | 26-7 (7)    | 30-7 (7)    | 34-7 (7)    | 38-7 (7)   | 42-7 (7)    |
| Control    | 4.2 ± 0.3                           | 4.3 ± 0.3   | 4.2 ± 0.3   | 4.4 ± 0.3   | 4.5 ± 0.3   | 4.5 ± 0.3  | 4.6 ± 0.3   |
| 625 ppm    | 4.2 ± 0.3                           | 4.1 ± 0.3*  | 4.1 ± 0.2*  | 4.2 ± 0.3   | 4.5 ± 0.3   | 4.5 ± 0.3  | 4.6 ± 0.3   |
| 1250 ppm   | 4.1 ± 0.3                           | 4.1 ± 0.3** | 4.0 ± 0.3** | 4.1 ± 0.4** | 4.4 ± 0.4   | 4.4 ± 0.3  | 4.5 ± 0.4*  |
| 2500 ppm   | 4.0 ± 0.2**                         | 4.0 ± 0.3** | 3.9 ± 0.3** | 4.0 ± 0.3** | 4.3 ± 0.3** | 4.4 ± 0.3* | 4.4 ± 0.3** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 4

| Group Name | Administration week-day(effective) |            |          |            |            |            |            |
|------------|------------------------------------|------------|----------|------------|------------|------------|------------|
|            | 46-7 (7)                           | 50-7 (7)   | 54-7 (7) | 58-7 (7)   | 62-7 (7)   | 66-7 (7)   | 70-7 (7)   |
| Control    | 4.7± 0.3                           | 4.8± 0.4   | 4.6± 0.4 | 5.0± 0.3   | 4.7± 0.3   | 4.7± 0.5   | 4.9± 0.4   |
| 625 ppm    | 4.6± 0.3                           | 4.7± 0.3   | 4.7± 0.3 | 4.9± 0.4   | 4.7± 0.3   | 4.7± 0.3   | 4.8± 0.3   |
| 1250 ppm   | 4.5± 0.4**                         | 4.7± 0.5   | 4.6± 0.4 | 4.8± 0.4   | 4.6± 0.4   | 4.5± 0.6   | 4.7± 0.4*  |
| 2500 ppm   | 4.4± 0.3**                         | 4.5± 0.4** | 4.6± 0.4 | 4.6± 0.4** | 4.5± 0.4** | 4.4± 0.4** | 4.5± 0.4** |

Significant difference.; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 5

| Group Name | Administration week-day (effective) |            |            |          |           |           |           |
|------------|-------------------------------------|------------|------------|----------|-----------|-----------|-----------|
|            | 74-7 (7)                            | 78-7 (7)   | 82-7 (7)   | 86-7 (7) | 90-7 (7)  | 94-7 (7)  | 98-7 (7)  |
| Control    | 4.8± 0.4                            | 4.8± 0.7   | 4.7± 0.5   | 4.6± 1.1 | 4.8± 0.5  | 4.7± 0.9  | 4.9± 0.5  |
| 625 ppm    | 4.6± 0.4                            | 4.7± 0.4   | 4.7± 0.4   | 4.9± 0.5 | 4.7± 0.7  | 4.6± 0.4  | 4.9± 0.5  |
| 1250 ppm   | 4.6± 0.4*                           | 4.7± 0.4   | 4.6± 0.4   | 4.7± 0.5 | 4.7± 0.5  | 4.7± 0.7  | 4.7± 0.7  |
| 2500 ppm   | 4.5± 0.3**                          | 4.4± 0.4** | 4.4± 0.5** | 4.6± 0.5 | 4.6± 0.5* | 4.4± 0.5* | 4.5± 0.6* |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 6

| Group Name | Administration week-day (effective) |            |
|------------|-------------------------------------|------------|
|            | 102-7 (7)                           | 104-7 (7)  |
| Control    | 4.8± 0.5                            | 4.8± 0.7   |
| 625 ppm    | 4.9± 0.8                            | 4.8± 0.6   |
| 1250 ppm   | 4.7± 0.8                            | 4.7± 0.7   |
| 2500 ppm   | 4.5± 0.6                            | 4.4± 0.4** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

**TABLE D 4**

**FOOD CONSUMPTION CHANGES: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 7

| Group Name | Administration week-day (effective) |          |          |          |          |          |          |
|------------|-------------------------------------|----------|----------|----------|----------|----------|----------|
|            | 1-7 (7)                             | 2-7 (7)  | 3-7 (7)  | 4-7 (7)  | 5-7 (7)  | 6-7 (7)  | 7-7 (7)  |
| Control    | 3.6± 0.3                            | 3.4± 0.3 | 3.4± 0.3 | 3.5± 0.4 | 3.5± 0.3 | 3.6± 0.3 | 3.7± 0.3 |
| 625 ppm    | 3.5± 0.2                            | 3.4± 0.2 | 3.4± 0.2 | 3.5± 0.2 | 3.5± 0.2 | 3.6± 0.3 | 3.6± 0.3 |
| 1250 ppm   | 3.5± 0.3                            | 3.4± 0.3 | 3.3± 0.2 | 3.4± 0.2 | 3.5± 0.2 | 3.5± 0.2 | 3.6± 0.3 |
| 2500 ppm   | 3.3± 0.3**                          | 3.4± 0.2 | 3.3± 0.2 | 3.5± 0.2 | 3.5± 0.2 | 3.6± 0.2 | 3.7± 0.2 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Crlj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 8

| Group Name | Administration<br>8-7 (7) | week-day (effective)<br>9-7 (7) | 10-7 (7) | 11-7 (7) | 12-7 (7) | 13-7 (7) | 14-7 (7) |
|------------|---------------------------|---------------------------------|----------|----------|----------|----------|----------|
| Control    | 3.8± 0.3                  | 3.9± 0.4                        | 3.8± 0.3 | 3.9± 0.3 | 3.8± 0.4 | 3.9± 0.4 | 3.8± 0.4 |
| 625 ppm    | 3.7± 0.3                  | 3.8± 0.3                        | 3.8± 0.3 | 3.8± 0.3 | 3.7± 0.3 | 3.8± 0.3 | 3.8± 0.3 |
| 1250 ppm   | 3.8± 0.2                  | 3.9± 0.2                        | 3.8± 0.3 | 3.8± 0.3 | 3.7± 0.3 | 3.8± 0.2 | 3.8± 0.3 |
| 2500 ppm   | 3.8± 0.3                  | 3.8± 0.3                        | 3.8± 0.3 | 3.9± 0.3 | 3.8± 0.3 | 3.9± 0.3 | 3.8± 0.3 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4



STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 9

| Group Name | Administration week-day (effective) |          |            |           |          |          |          |
|------------|-------------------------------------|----------|------------|-----------|----------|----------|----------|
|            | 18-7 (7)                            | 22-7 (7) | 26-7 (7)   | 30-7 (7)  | 34-7 (7) | 38-7 (7) | 42-7 (7) |
| Control    | 3.8± 0.3                            | 3.9± 0.4 | 3.7± 0.4   | 3.8± 0.4  | 4.1± 0.5 | 4.3± 0.4 | 4.2± 0.5 |
| 625 ppm    | 3.7± 0.4                            | 3.8± 0.4 | 3.7± 0.4   | 3.6± 0.5* | 4.0± 0.5 | 4.1± 0.5 | 4.2± 0.4 |
| 1250 ppm   | 3.8± 0.3                            | 3.8± 0.3 | 3.6± 0.3   | 3.7± 0.4  | 4.0± 0.4 | 4.1± 0.4 | 4.2± 0.6 |
| 2500 ppm   | 3.8± 0.3                            | 3.8± 0.3 | 3.5± 0.3** | 3.6± 0.4  | 4.0± 0.3 | 4.0± 0.4 | 4.2± 0.3 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 10

| Group Name | Administration week-day (effective) |          |          |          |            |          |          |
|------------|-------------------------------------|----------|----------|----------|------------|----------|----------|
|            | 46-7 (7)                            | 50-7 (7) | 54-7 (7) | 58-7 (7) | 62-7 (7)   | 66-7 (7) | 70-7 (7) |
| Control    | 4.2± 0.5                            | 4.2± 0.6 | 4.2± 0.5 | 4.1± 0.4 | 4.2± 0.6   | 3.9± 0.5 | 4.0± 0.6 |
| 625 ppm    | 4.2± 0.4                            | 4.4± 0.7 | 4.2± 0.6 | 4.2± 0.5 | 4.1± 0.5   | 3.8± 0.6 | 3.9± 0.6 |
| 1250 ppm   | 4.1± 0.4                            | 4.1± 0.5 | 4.1± 0.6 | 4.1± 0.6 | 3.9± 0.5** | 3.8± 0.5 | 3.8± 0.4 |
| 2500 ppm   | 4.2± 0.5                            | 4.1± 0.4 | 4.1± 0.4 | 4.1± 0.5 | 4.0± 0.4   | 3.8± 0.4 | 3.8± 0.4 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Cri:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 11

| Group Name | Administration week-day (effective) |            |           |            |          |          |            |
|------------|-------------------------------------|------------|-----------|------------|----------|----------|------------|
|            | 74-7 (7)                            | 78-7 (7)   | 82-7 (7)  | 86-7 (7)   | 90-7 (7) | 94-7 (7) | 98-7 (7)   |
| Control    | 4.1± 0.6                            | 4.2± 0.5   | 4.2± 0.5  | 4.3± 0.7   | 4.2± 0.6 | 4.0± 0.8 | 4.4± 1.0   |
| 625 ppm    | 4.0± 0.4                            | 4.1± 0.7   | 4.3± 1.1  | 4.2± 0.7   | 4.2± 0.7 | 4.0± 0.8 | 4.4± 0.9   |
| 1250 ppm   | 3.7± 0.7**                          | 4.0± 0.5   | 4.0± 0.6  | 3.9± 0.6*  | 3.9± 0.6 | 3.9± 0.6 | 4.3± 0.6   |
| 2500 ppm   | 3.7± 0.5*                           | 3.8± 0.5** | 3.9± 0.4* | 3.8± 0.5** | 3.9± 0.5 | 3.9± 0.7 | 3.9± 0.8** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Cri:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 12

| Group Name | Administration week-day(effective) |           |
|------------|------------------------------------|-----------|
|            | 102-7 (7)                          | 104-7 (7) |
| Control    | 4.2± 0.7                           | 4.3± 0.6  |
| 625 ppm    | 4.3± 1.2                           | 4.5± 0.7  |
| 1250 ppm   | 4.1± 0.6                           | 4.5± 0.8  |
| 2500 ppm   | 4.0± 0.5                           | 4.1± 0.6  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

TABLE E 1

WATER CONSUMPTION CHANGES AND  
SURVIVAL ANIMAL NUMBERS: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

MEAN WATER CONSUMPTION (WC) AND SURVIVAL

PAGE : 1

| Week-Day<br>on Study | Control  |                           | 625 ppm  |                       |                   | 1250 ppm |                       |                   | 2500 ppm |                       |                   |
|----------------------|----------|---------------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
|                      | Av. WC.  | No. of<br>Surviv.<br><50> | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 1-7                  | 4.7 (50) | 50/50                     | 4.1 (50) | 87                    | 50/50             | 3.4 (50) | 72                    | 50/50             | 2.7 (50) | 57                    | 50/50             |
| 2-7                  | 4.4 (49) | 50/50                     | 3.7 (50) | 84                    | 50/50             | 3.4 (50) | 77                    | 50/50             | 2.6 (50) | 59                    | 50/50             |
| 3-7                  | 4.3 (48) | 50/50                     | 3.9 (49) | 91                    | 49/50             | 3.4 (50) | 79                    | 50/50             | 2.7 (50) | 63                    | 50/50             |
| 4-7                  | 4.2 (48) | 50/50                     | 3.9 (49) | 93                    | 49/50             | 3.3 (50) | 79                    | 50/50             | 2.7 (50) | 64                    | 50/50             |
| 5-7                  | 4.4 (50) | 50/50                     | 3.9 (49) | 89                    | 49/50             | 3.4 (50) | 77                    | 50/50             | 2.8 (50) | 64                    | 50/50             |
| 6-7                  | 4.2 (50) | 50/50                     | 3.8 (49) | 90                    | 49/50             | 3.4 (50) | 81                    | 50/50             | 2.8 (50) | 67                    | 50/50             |
| 7-7                  | 4.1 (50) | 50/50                     | 3.7 (49) | 90                    | 49/50             | 3.5 (50) | 85                    | 50/50             | 2.9 (50) | 71                    | 50/50             |
| 8-7                  | 4.0 (50) | 50/50                     | 3.7 (49) | 93                    | 49/50             | 3.3 (50) | 83                    | 50/50             | 2.8 (50) | 70                    | 50/50             |
| 9-7                  | 4.1 (50) | 50/50                     | 3.7 (49) | 90                    | 49/50             | 3.5 (50) | 85                    | 50/50             | 2.9 (49) | 71                    | 49/50             |
| 10-7                 | 4.1 (50) | 50/50                     | 3.7 (49) | 90                    | 49/50             | 3.5 (50) | 85                    | 50/50             | 2.9 (49) | 71                    | 49/50             |
| 11-7                 | 4.0 (50) | 50/50                     | 3.8 (49) | 95                    | 49/50             | 3.4 (50) | 85                    | 50/50             | 3.0 (49) | 75                    | 49/50             |
| 12-7                 | 3.9 (50) | 50/50                     | 3.6 (49) | 92                    | 49/50             | 3.4 (50) | 87                    | 50/50             | 2.9 (49) | 74                    | 49/50             |
| 13-7                 | 3.9 (50) | 50/50                     | 3.6 (49) | 92                    | 49/50             | 3.4 (50) | 87                    | 50/50             | 3.0 (49) | 77                    | 49/50             |
| 14-7                 | 3.8 (50) | 50/50                     | 3.5 (49) | 92                    | 49/50             | 3.3 (50) | 87                    | 50/50             | 2.8 (49) | 74                    | 49/50             |
| 18-7                 | 3.5 (50) | 50/50                     | 3.4 (49) | 97                    | 49/50             | 3.2 (50) | 91                    | 50/50             | 2.7 (49) | 77                    | 49/50             |
| 22-7                 | 3.8 (50) | 50/50                     | 3.5 (49) | 92                    | 49/50             | 3.3 (50) | 87                    | 50/50             | 2.8 (49) | 74                    | 49/50             |
| 26-7                 | 3.6 (50) | 50/50                     | 3.4 (49) | 94                    | 49/50             | 3.4 (50) | 94                    | 50/50             | 2.8 (49) | 78                    | 49/50             |
| 30-7                 | 3.5 (50) | 50/50                     | 3.4 (48) | 97                    | 48/50             | 3.2 (50) | 91                    | 50/50             | 2.8 (49) | 80                    | 49/50             |
| 34-7                 | 3.7 (50) | 50/50                     | 3.6 (48) | 97                    | 48/50             | 3.4 (50) | 92                    | 50/50             | 3.0 (49) | 81                    | 49/50             |
| 38-7                 | 3.7 (50) | 50/50                     | 3.6 (48) | 97                    | 48/50             | 3.4 (50) | 92                    | 50/50             | 3.0 (49) | 81                    | 49/50             |
| 42-7                 | 3.8 (49) | 49/50                     | 3.7 (48) | 97                    | 48/50             | 3.5 (50) | 92                    | 50/50             | 3.2 (49) | 84                    | 49/50             |
| 46-7                 | 3.9 (49) | 49/50                     | 3.8 (48) | 97                    | 48/50             | 3.6 (50) | 92                    | 50/50             | 3.1 (49) | 79                    | 49/50             |
| 50-7                 | 3.8 (49) | 49/50                     | 3.7 (47) | 97                    | 47/50             | 3.5 (50) | 92                    | 50/50             | 3.1 (49) | 82                    | 49/50             |
| 54-7                 | 4.0 (49) | 49/50                     | 4.0 (47) | 100                   | 47/50             | 3.6 (49) | 90                    | 49/50             | 3.2 (49) | 80                    | 49/50             |
| 58-7                 | 4.1 (48) | 48/50                     | 4.0 (46) | 98                    | 47/50             | 3.7 (49) | 90                    | 49/50             | 3.2 (49) | 78                    | 49/50             |
| 62-7                 | 4.1 (48) | 48/50                     | 4.0 (45) | 98                    | 45/50             | 3.7 (48) | 90                    | 48/50             | 3.2 (49) | 78                    | 49/50             |
| 66-7                 | 4.3 (48) | 48/50                     | 4.3 (44) | 100                   | 44/50             | 4.0 (47) | 93                    | 47/50             | 3.4 (49) | 79                    | 49/50             |
| 70-7                 | 4.5 (47) | 47/50                     | 4.3 (44) | 96                    | 44/50             | 4.3 (47) | 96                    | 47/50             | 3.5 (48) | 78                    | 48/50             |
| 74-7                 | 4.6 (47) | 47/50                     | 4.2 (44) | 91                    | 44/50             | 4.1 (43) | 89                    | 45/50             | 3.6 (48) | 78                    | 48/50             |
| 78-7                 | 4.7 (42) | 45/50                     | 4.4 (42) | 94                    | 43/50             | 4.3 (43) | 91                    | 45/50             | 3.8 (48) | 81                    | 48/50             |
| 82-7                 | 4.5 (41) | 43/50                     | 4.3 (39) | 96                    | 42/50             | 4.2 (43) | 93                    | 44/50             | 3.8 (47) | 84                    | 47/50             |
| 86-7                 | 4.6 (40) | 41/50                     | 4.5 (39) | 98                    | 41/50             | 4.5 (43) | 98                    | 44/50             | 4.0 (46) | 87                    | 46/50             |
| 90-7                 | 4.9 (35) | 37/50                     | 4.7 (38) | 96                    | 39/50             | 4.5 (41) | 92                    | 43/50             | 4.0 (45) | 82                    | 46/50             |
| 94-7                 | 4.7 (31) | 32/50                     | 4.9 (38) | 104                   | 38/50             | 4.9 (39) | 104                   | 40/50             | 4.0 (44) | 85                    | 45/50             |
| 98-7                 | 5.0 (26) | 28/50                     | 5.1 (31) | 102                   | 36/50             | 4.8 (36) | 96                    | 38/50             | 4.2 (42) | 84                    | 44/50             |
| 102-7                | 5.1 (25) | 27/50                     | 4.9 (29) | 96                    | 35/50             | 4.4 (29) | 86                    | 34/50             | 4.1 (37) | 80                    | 40/50             |
| 104-7                | 5.4 (22) | 24/50                     | 5.1 (27) | 94                    | 32/50             | 4.7 (28) | 87                    | 31/50             | 4.2 (36) | 78                    | 39/50             |

< >:No. of effective animals, ( ) :No. of measured animals

Av. WC. : g

**TABLE E 2**

**WATER CONSUMPTION CHANGES AND  
SURVIVAL ANIMAL NUMBERS: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

MEAN WATER CONSUMPTION(WC) AND SURVIVAL

PAGE : 2

| Week-Day<br>on Study | Control  |                           |  | 625 ppm  |                       |                   | 1250 ppm |                       |                   | 2500 ppm |                       |                   |
|----------------------|----------|---------------------------|--|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
|                      | Av. WC.  | No. of<br>Surviv.<br><50> |  | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. | Av. WC.  | % of<br>cont.<br><50> | No. of<br>Surviv. |
| 1-7                  | 4.3 (50) | 50/50                     |  | 4.0 (50) | 93                    | 50/50             | 3.6 (50) | 84                    | 50/50             | 2.7 (50) | 63                    | 50/50             |
| 2-7                  | 4.1 (50) | 50/50                     |  | 3.6 (50) | 88                    | 50/50             | 3.4 (50) | 83                    | 50/50             | 2.7 (50) | 66                    | 50/50             |
| 3-7                  | 4.1 (50) | 50/50                     |  | 3.7 (50) | 90                    | 50/50             | 3.5 (50) | 85                    | 50/50             | 2.8 (50) | 68                    | 50/50             |
| 4-7                  | 4.0 (50) | 50/50                     |  | 3.6 (50) | 90                    | 50/50             | 3.6 (50) | 90                    | 50/50             | 2.9 (50) | 73                    | 50/50             |
| 5-7                  | 4.2 (50) | 50/50                     |  | 3.7 (50) | 88                    | 50/50             | 3.7 (50) | 88                    | 50/50             | 3.0 (50) | 71                    | 50/50             |
| 6-7                  | 4.0 (50) | 50/50                     |  | 3.7 (50) | 93                    | 50/50             | 3.6 (50) | 90                    | 50/50             | 3.0 (50) | 75                    | 50/50             |
| 7-7                  | 4.1 (50) | 50/50                     |  | 3.7 (50) | 90                    | 50/50             | 3.7 (50) | 90                    | 50/50             | 3.1 (50) | 76                    | 50/50             |
| 8-7                  | 4.1 (50) | 50/50                     |  | 3.6 (50) | 88                    | 50/50             | 3.6 (50) | 88                    | 50/50             | 3.1 (50) | 76                    | 50/50             |
| 9-7                  | 4.2 (50) | 50/50                     |  | 3.7 (50) | 88                    | 50/50             | 3.6 (50) | 86                    | 50/50             | 3.1 (50) | 74                    | 50/50             |
| 10-7                 | 4.1 (50) | 50/50                     |  | 3.7 (50) | 90                    | 50/50             | 3.7 (50) | 90                    | 50/50             | 3.1 (50) | 76                    | 50/50             |
| 11-7                 | 4.1 (50) | 50/50                     |  | 3.7 (50) | 90                    | 50/50             | 3.7 (50) | 90                    | 50/50             | 3.1 (50) | 76                    | 50/50             |
| 12-7                 | 4.2 (50) | 50/50                     |  | 3.6 (50) | 86                    | 50/50             | 3.5 (50) | 83                    | 50/50             | 3.1 (50) | 74                    | 50/50             |
| 13-7                 | 4.2 (50) | 50/50                     |  | 3.7 (50) | 88                    | 50/50             | 3.6 (50) | 86                    | 50/50             | 3.1 (50) | 74                    | 50/50             |
| 14-7                 | 4.0 (50) | 50/50                     |  | 3.6 (50) | 90                    | 50/50             | 3.6 (50) | 90                    | 50/50             | 3.0 (50) | 75                    | 50/50             |
| 18-7                 | 3.8 (50) | 50/50                     |  | 3.4 (50) | 89                    | 50/50             | 3.2 (50) | 84                    | 50/50             | 2.9 (50) | 76                    | 50/50             |
| 22-7                 | 4.1 (49) | 49/50                     |  | 3.5 (50) | 85                    | 50/50             | 3.3 (50) | 80                    | 50/50             | 3.0 (50) | 73                    | 50/50             |
| 26-7                 | 3.9 (49) | 49/50                     |  | 3.3 (50) | 85                    | 50/50             | 3.1 (50) | 79                    | 50/50             | 2.8 (50) | 72                    | 50/50             |
| 30-7                 | 3.9 (49) | 49/50                     |  | 3.3 (50) | 85                    | 50/50             | 3.2 (50) | 82                    | 50/50             | 2.8 (50) | 72                    | 50/50             |
| 34-7                 | 4.1 (49) | 49/50                     |  | 3.4 (50) | 83                    | 50/50             | 3.3 (50) | 80                    | 50/50             | 3.0 (50) | 73                    | 50/50             |
| 38-7                 | 4.0 (49) | 49/50                     |  | 3.4 (50) | 85                    | 50/50             | 3.3 (50) | 83                    | 50/50             | 2.9 (50) | 73                    | 50/50             |
| 42-7                 | 4.1 (49) | 49/50                     |  | 3.6 (50) | 88                    | 50/50             | 3.3 (50) | 80                    | 50/50             | 3.0 (50) | 73                    | 50/50             |
| 46-7                 | 4.1 (49) | 49/50                     |  | 3.6 (50) | 88                    | 50/50             | 3.4 (50) | 83                    | 50/50             | 3.0 (50) | 73                    | 50/50             |
| 50-7                 | 4.1 (49) | 49/50                     |  | 3.6 (49) | 88                    | 50/50             | 3.2 (50) | 78                    | 50/50             | 2.8 (50) | 68                    | 50/50             |
| 54-7                 | 4.0 (49) | 49/50                     |  | 3.4 (49) | 85                    | 49/50             | 3.3 (50) | 83                    | 50/50             | 2.9 (50) | 73                    | 50/50             |
| 58-7                 | 3.9 (49) | 49/50                     |  | 3.5 (49) | 90                    | 49/50             | 3.2 (49) | 82                    | 49/50             | 2.8 (50) | 72                    | 50/50             |
| 62-7                 | 3.9 (47) | 47/50                     |  | 3.5 (48) | 90                    | 48/50             | 3.2 (47) | 82                    | 48/50             | 2.8 (50) | 72                    | 50/50             |
| 66-7                 | 3.9 (46) | 46/50                     |  | 3.7 (48) | 95                    | 48/50             | 3.4 (47) | 87                    | 48/50             | 3.0 (50) | 77                    | 50/50             |
| 70-7                 | 4.0 (41) | 41/50                     |  | 3.5 (48) | 88                    | 48/50             | 3.4 (46) | 85                    | 46/50             | 3.1 (50) | 78                    | 50/50             |
| 74-7                 | 4.1 (39) | 39/50                     |  | 3.7 (45) | 90                    | 47/50             | 3.5 (44) | 85                    | 45/50             | 3.3 (48) | 80                    | 48/50             |
| 78-7                 | 4.2 (39) | 39/50                     |  | 3.9 (44) | 93                    | 47/50             | 3.8 (39) | 90                    | 42/50             | 3.2 (45) | 76                    | 48/50             |
| 82-7                 | 4.2 (37) | 38/50                     |  | 4.1 (43) | 98                    | 47/50             | 4.1 (40) | 98                    | 42/50             | 3.4 (46) | 81                    | 48/50             |
| 86-7                 | 4.2 (35) | 38/50                     |  | 4.2 (39) | 100                   | 43/50             | 4.2 (37) | 100                   | 41/50             | 3.5 (44) | 83                    | 47/50             |
| 90-7                 | 4.3 (35) | 37/50                     |  | 4.1 (34) | 95                    | 39/50             | 4.0 (33) | 93                    | 40/50             | 3.8 (43) | 88                    | 45/50             |
| 94-7                 | 4.1 (30) | 33/50                     |  | 4.1 (30) | 100                   | 38/50             | 4.6 (35) | 112                   | 38/50             | 3.6 (40) | 88                    | 43/50             |
| 98-7                 | 4.4 (26) | 31/50                     |  | 4.3 (27) | 98                    | 33/50             | 4.6 (30) | 105                   | 35/50             | 3.8 (40) | 86                    | 42/50             |
| 102-7                | 4.4 (21) | 25/50                     |  | 4.1 (19) | 93                    | 24/50             | 4.2 (23) | 95                    | 33/50             | 4.2 (36) | 95                    | 39/50             |
| 104-7                | 4.5 (20) | 24/50                     |  | 4.0 (15) | 89                    | 21/50             | 4.2 (21) | 93                    | 33/50             | 4.2 (36) | 93                    | 38/50             |

< >:No. of effective animals, ( ) :No. of measured animals Av. WC. : g



**TABLE E 3**

**WATER CONSUMPTION CHANGES: MALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 1

| Group Name | Administration |       | week-day (effective) |       |         |       |         |       |         |       |
|------------|----------------|-------|----------------------|-------|---------|-------|---------|-------|---------|-------|
|            | 1-7 (3)        |       | 2-7 (3)              |       | 3-7 (3) |       | 4-7 (3) |       | 5-7 (3) |       |
| Control    | 4.7±           | 1.1   | 4.4±                 | 0.7   | 4.3±    | 0.7   | 4.2±    | 0.7   | 4.4±    | 0.8   |
| 625 ppm    | 4.1±           | 0.4** | 3.7±                 | 0.5** | 3.9±    | 0.6*  | 3.9±    | 0.8** | 3.9±    | 0.6** |
| 1250 ppm   | 3.4±           | 0.4** | 3.4±                 | 0.5** | 3.4±    | 0.6** | 3.3±    | 0.6** | 3.4±    | 0.4** |
| 2500 ppm   | 2.7±           | 0.3** | 2.6±                 | 0.5** | 2.7±    | 0.5** | 2.7±    | 0.5** | 2.8±    | 0.5** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 2

| Group Name | Administration<br>8-7 (3) | week-day(effective)<br>9-7 (3) | 10-7 (3)    | 11-7 (3)    | 12-7 (3)    | 13-7 (3)    | 14-7 (3)    |
|------------|---------------------------|--------------------------------|-------------|-------------|-------------|-------------|-------------|
| Control    | 4.0 ± 0.6                 | 4.1 ± 0.7                      | 4.1 ± 0.6   | 4.0 ± 0.7   | 3.9 ± 0.6   | 3.9 ± 0.6   | 3.8 ± 0.5   |
| 625 ppm    | 3.7 ± 0.7**               | 3.7 ± 0.6**                    | 3.7 ± 0.7** | 3.8 ± 0.6   | 3.6 ± 0.6** | 3.6 ± 0.6*  | 3.5 ± 0.6** |
| 1250 ppm   | 3.3 ± 0.4**               | 3.5 ± 0.5**                    | 3.5 ± 0.4** | 3.4 ± 0.4** | 3.4 ± 0.4** | 3.4 ± 0.4** | 3.3 ± 0.3** |
| 2500 ppm   | 2.8 ± 0.4**               | 2.9 ± 0.5**                    | 2.9 ± 0.5** | 3.0 ± 0.4** | 2.9 ± 0.4** | 3.0 ± 0.5** | 2.8 ± 0.4** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 3

| Group Name | Administration<br>18-7 (3) | week-day(effective)<br>22-7 (3) | 26-7 (3)   | 30-7 (3)   | 34-7 (3)   | 38-7 (3)   | 42-7 (3)   |
|------------|----------------------------|---------------------------------|------------|------------|------------|------------|------------|
| Control    | 3.5± 0.4                   | 3.8± 0.4                        | 3.6± 0.4   | 3.5± 0.4   | 3.7± 0.4   | 3.7± 0.3   | 3.8± 0.3   |
| 625 ppm    | 3.4± 0.5                   | 3.5± 0.4**                      | 3.4± 0.5*  | 3.4± 0.5   | 3.6± 0.5   | 3.6± 0.4   | 3.7± 0.4   |
| 1250 ppm   | 3.2± 0.3**                 | 3.3± 0.3**                      | 3.4± 0.3*  | 3.2± 0.3** | 3.4± 0.3** | 3.4± 0.3** | 3.5± 0.4** |
| 2500 ppm   | 2.7± 0.4**                 | 2.8± 0.4**                      | 2.8± 0.4** | 2.8± 0.4** | 3.0± 0.5** | 3.0± 0.4** | 3.2± 0.5** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crlj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 4

| Group Name | Administration week-day (effective) |             |             |             |             |             |             |
|------------|-------------------------------------|-------------|-------------|-------------|-------------|-------------|-------------|
|            | 46-7 (3)                            | 50-7 (3)    | 54-7 (3)    | 58-7 (3)    | 62-7 (3)    | 66-7 (3)    | 70-7 (3)    |
| Control    | 3.9 ± 0.4                           | 3.8 ± 0.3   | 4.0 ± 0.4   | 4.1 ± 0.5   | 4.1 ± 0.6   | 4.3 ± 0.7   | 4.5 ± 0.6   |
| 625 ppm    | 3.8 ± 0.5                           | 3.7 ± 0.5   | 4.0 ± 0.8   | 4.0 ± 0.8   | 4.0 ± 0.6   | 4.3 ± 0.6   | 4.3 ± 0.6   |
| 1250 ppm   | 3.6 ± 0.4**                         | 3.5 ± 0.5** | 3.6 ± 0.4** | 3.7 ± 0.4** | 3.7 ± 0.5** | 4.0 ± 1.1** | 4.3 ± 1.2** |
| 2500 ppm   | 3.1 ± 0.4**                         | 3.1 ± 0.5** | 3.2 ± 0.5** | 3.2 ± 0.5** | 3.2 ± 0.4** | 3.4 ± 0.4** | 3.5 ± 0.5** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 5

| Group Name | Administration<br>74-7 (3) | week-day (effective)<br>78-7 (3) | 82-7 (3)    | 86-7 (3)    | 90-7 (3)    | 94-7 (3)    | 98-7 (3)    |
|------------|----------------------------|----------------------------------|-------------|-------------|-------------|-------------|-------------|
| Control    | 4.6 ± 0.9                  | 4.7 ± 1.0                        | 4.5 ± 0.9   | 4.6 ± 1.7   | 4.9 ± 1.0   | 4.7 ± 1.5   | 5.0 ± 0.8   |
| 625 ppm    | 4.2 ± 0.8                  | 4.4 ± 0.8                        | 4.3 ± 0.6   | 4.5 ± 0.7   | 4.7 ± 0.9   | 4.9 ± 1.5   | 5.1 ± 1.3   |
| 1250 ppm   | 4.1 ± 0.6**                | 4.3 ± 0.6                        | 4.2 ± 0.8   | 4.5 ± 1.0   | 4.5 ± 1.0   | 4.9 ± 1.5   | 4.8 ± 1.1   |
| 2500 ppm   | 3.6 ± 0.6**                | 3.8 ± 0.6**                      | 3.8 ± 0.9** | 4.0 ± 1.1** | 4.0 ± 1.0** | 4.0 ± 1.0** | 4.2 ± 1.0** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 6

| Group Name | Administration week-day(effective) |            |
|------------|------------------------------------|------------|
|            | 102-7 (3)                          | 104-7 (3)  |
| Control    | 5.1± 0.8                           | 5.4± 1.1   |
| 625 ppm    | 4.9± 1.0                           | 5.1± 1.0   |
| 1250 ppm   | 4.4± 1.2*                          | 4.7± 1.2   |
| 2500 ppm   | 4.1± 1.0**                         | 4.2± 1.0** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

**TABLE E 4**

**WATER CONSUMPTION CHANGES: FEMALE**



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 7

| Group Name | Administration<br>1-7 (3) | week-day (effective)<br>2-7 (3) | 3-7 (3)     | 4-7 (3)     | 5-7 (3)     | 6-7 (3)     | 7-7 (3)     |
|------------|---------------------------|---------------------------------|-------------|-------------|-------------|-------------|-------------|
| Control    | 4.3 ± 0.4                 | 4.1 ± 0.5                       | 4.1 ± 0.5   | 4.0 ± 0.5   | 4.2 ± 0.5   | 4.0 ± 0.4   | 4.1 ± 0.4   |
| 625 ppm    | 4.0 ± 0.4**               | 3.6 ± 0.3**                     | 3.7 ± 0.4** | 3.6 ± 0.3** | 3.7 ± 0.4** | 3.7 ± 0.3** | 3.7 ± 0.4** |
| 1250 ppm   | 3.6 ± 0.3**               | 3.4 ± 0.4**                     | 3.5 ± 0.5** | 3.6 ± 0.5** | 3.7 ± 0.4** | 3.6 ± 0.5** | 3.7 ± 0.6** |
| 2500 ppm   | 2.7 ± 0.3**               | 2.7 ± 0.3**                     | 2.8 ± 0.3** | 2.9 ± 0.3** | 3.0 ± 0.4** | 3.0 ± 0.4** | 3.1 ± 0.4** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 8

| Group Name | Administration<br>8-7 (3) | week-day(effective)<br>9-7 (3) | 10-7 (3)    | 11-7 (3)    | 12-7 (3)    | 13-7 (3)    | 14-7 (3)    |
|------------|---------------------------|--------------------------------|-------------|-------------|-------------|-------------|-------------|
| Control    | 4.1 ± 0.4                 | 4.2 ± 0.5                      | 4.1 ± 0.5   | 4.1 ± 0.4   | 4.2 ± 0.7   | 4.2 ± 0.6   | 4.0 ± 0.4   |
| 625 ppm    | 3.6 ± 0.4**               | 3.7 ± 0.4**                    | 3.7 ± 0.4** | 3.7 ± 0.4** | 3.6 ± 0.5** | 3.7 ± 0.4** | 3.6 ± 0.4** |
| 1250 ppm   | 3.6 ± 0.3**               | 3.6 ± 0.4**                    | 3.7 ± 0.4** | 3.7 ± 0.6** | 3.5 ± 0.4** | 3.6 ± 0.4** | 3.6 ± 0.6** |
| 2500 ppm   | 3.1 ± 0.4**               | 3.1 ± 0.4**                    | 3.1 ± 0.4** | 3.1 ± 0.4** | 3.1 ± 0.5** | 3.1 ± 0.5** | 3.0 ± 0.3** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 9

| Group Name | Administration week-day (effective) |            |            |            |            |            |            |
|------------|-------------------------------------|------------|------------|------------|------------|------------|------------|
|            | 18-7 (3)                            | 22-7 (3)   | 26-7 (3)   | 30-7 (3)   | 34-7 (3)   | 38-7 (3)   | 42-7 (3)   |
| Control    | 3.8± 0.4                            | 4.1± 0.6   | 3.9± 0.6   | 3.9± 0.7   | 4.1± 0.8   | 4.0± 0.7   | 4.1± 0.7   |
| 625 ppm    | 3.4± 0.3**                          | 3.5± 0.5** | 3.3± 0.4** | 3.3± 0.5** | 3.4± 0.6** | 3.4± 0.4** | 3.6± 0.4** |
| 1250 ppm   | 3.2± 0.3**                          | 3.3± 0.4** | 3.1± 0.3** | 3.2± 0.4** | 3.3± 0.3** | 3.3± 0.4** | 3.3± 0.4** |
| 2500 ppm   | 2.9± 0.4**                          | 3.0± 0.4** | 2.8± 0.4** | 2.8± 0.3** | 3.0± 0.4** | 2.9± 0.4** | 3.0± 0.3** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 10

| Group Name | Administration |  | week-day (effective) |  |             |  |             |  |
|------------|----------------|--|----------------------|--|-------------|--|-------------|--|
|            | 46-7 (3)       |  | 50-7 (3)             |  | 54-7 (3)    |  | 58-7 (3)    |  |
|            |                |  |                      |  |             |  |             |  |
| Control    | 4.1 ± 0.8      |  | 4.1 ± 0.8            |  | 4.0 ± 1.0   |  | 3.9 ± 0.6   |  |
| 625 ppm    | 3.6 ± 0.5**    |  | 3.6 ± 0.5**          |  | 3.4 ± 0.5** |  | 3.5 ± 0.5** |  |
| 1250 ppm   | 3.4 ± 0.7**    |  | 3.2 ± 0.4**          |  | 3.3 ± 0.7** |  | 3.2 ± 0.8** |  |
| 2500 ppm   | 3.0 ± 0.4**    |  | 2.8 ± 0.4**          |  | 2.9 ± 0.5** |  | 2.8 ± 0.4** |  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : g  
 REPORT TYPE : A1 104  
 SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 11

| Group Name | Administration week-day (effective) |             |             |             |             |            |           |
|------------|-------------------------------------|-------------|-------------|-------------|-------------|------------|-----------|
|            | 74-7 (3)                            | 78-7 (3)    | 82-7 (3)    | 86-7 (3)    | 90-7 (3)    | 94-7 (3)   | 98-7 (3)  |
| Control    | 4.1 ± 0.8                           | 4.2 ± 0.8   | 4.2 ± 1.0   | 4.2 ± 0.9   | 4.3 ± 0.9   | 4.1 ± 1.1  | 4.4 ± 1.1 |
| 625 ppm    | 3.7 ± 0.9                           | 3.9 ± 0.9   | 4.1 ± 1.2   | 4.2 ± 1.2   | 4.1 ± 1.4   | 4.1 ± 1.4  | 4.3 ± 1.5 |
| 1250 ppm   | 3.5 ± 1.0**                         | 3.8 ± 1.2** | 4.1 ± 1.5   | 4.2 ± 1.4   | 4.0 ± 1.3   | 4.6 ± 1.8  | 4.6 ± 1.6 |
| 2500 ppm   | 3.3 ± 1.4**                         | 3.2 ± 1.1** | 3.4 ± 1.5** | 3.5 ± 1.6** | 3.8 ± 1.7** | 3.6 ± 1.7* | 3.8 ± 1.8 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
UNIT : g  
REPORT TYPE : A1 104  
SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 12

| Group Name | Administration week-day (effective) |           |
|------------|-------------------------------------|-----------|
|            | 102-7 (3)                           | 104-7 (3) |
| Control    | 4.4± 1.2                            | 4.5± 1.1  |
| 625 ppm    | 4.1± 1.6                            | 4.0± 1.1  |
| 1250 ppm   | 4.2± 1.4                            | 4.2± 1.4  |
| 2500 ppm   | 4.2± 1.7                            | 4.2± 1.8  |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HAN260)

BAIS4

**TABLE F 1**

**CHEMICAL INTAKE CHANGES: MALE**

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
UNIT : mg/kg/day  
REPORT TYPE : A1 104  
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 1

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 1                      |    | 2    |    | 3    |    | 4    |    | 5    |    | 6    |    | 7    |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 103±                   | 10 | 90±  | 11 | 92±  | 14 | 88±  | 19 | 85±  | 15 | 81±  | 14 | 77±  | 12 |
| 1250 ppm   | 173±                   | 19 | 167± | 25 | 159± | 29 | 151± | 23 | 153± | 21 | 147± | 19 | 147± | 21 |
| 2500 ppm   | 283±                   | 32 | 258± | 47 | 263± | 50 | 256± | 47 | 259± | 45 | 258± | 51 | 262± | 50 |

(HAN300)

BAIS 5



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 2

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 8                      |    | 9    |    | 10   |    | 11   |    | 12   |    | 13   |    | 14   |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 74±                    | 15 | 74±  | 13 | 73±  | 14 | 73±  | 13 | 67±  | 13 | 66±  | 12 | 63±  | 12 |
| 1250 ppm   | 139±                   | 20 | 142± | 20 | 138± | 20 | 135± | 19 | 127± | 18 | 126± | 18 | 119± | 13 |
| 2500 ppm   | 245±                   | 43 | 250± | 46 | 242± | 40 | 244± | 41 | 232± | 41 | 232± | 43 | 218± | 37 |

(HAN300)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 3

| Group Name | Administration |    | (weeks) |    |      |    |      |    |      |    |      |    |      |    |
|------------|----------------|----|---------|----|------|----|------|----|------|----|------|----|------|----|
|            | 18             |    | 22      |    | 26   |    | 30   |    | 34   |    | 38   |    | 42   |    |
| Control    | 0±             | 0  | 0±      | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 56±            | 9  | 56±     | 8  | 52±  | 9  | 50±  | 8  | 50±  | 8  | 49±  | 7  | 49±  | 7  |
| 1250 ppm   | 110±           | 14 | 106±    | 13 | 104± | 12 | 96±  | 12 | 97±  | 13 | 96±  | 14 | 96±  | 13 |
| 2500 ppm   | 196±           | 36 | 195±    | 32 | 187± | 30 | 178± | 31 | 186± | 31 | 179± | 28 | 183± | 34 |

(HAN300)

BA1S5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Cri:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 4

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 46                     |    | 50   |    | 54   |    | 58   |    | 62   |    | 66   |    | 70   |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 50±                    | 8  | 47±  | 8  | 51±  | 18 | 50±  | 17 | 48±  | 7  | 52±  | 7  | 52±  | 8  |
| 1250 ppm   | 95±                    | 13 | 92±  | 13 | 93±  | 13 | 94±  | 12 | 95±  | 16 | 103± | 42 | 111± | 55 |
| 2500 ppm   | 178±                   | 27 | 176± | 30 | 177± | 28 | 172± | 26 | 177± | 25 | 183± | 26 | 189± | 35 |

(HAN300)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 5

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 74                     |    | 78   |    | 82   |    | 86   |    | 90   |    | 94   |    | 98   |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 52±                    | 13 | 53±  | 12 | 51±  | 10 | 53±  | 12 | 56±  | 15 | 62±  | 27 | 64±  | 22 |
| 1250 ppm   | 99±                    | 16 | 104± | 18 | 102± | 24 | 110± | 34 | 117± | 39 | 128± | 56 | 127± | 48 |
| 2500 ppm   | 196±                   | 42 | 202± | 48 | 203± | 74 | 215± | 87 | 219± | 76 | 227± | 83 | 236± | 72 |

(HAN300)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crlj:BDF1]  
UNIT : mg/kg/day  
REPORT TYPE : A1 104  
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 6

| Group Name | Administration (weeks) |    |      |    |
|------------|------------------------|----|------|----|
|            | 102                    |    | 104  |    |
| Control    | 0±                     | 0  | 0±   | 0  |
| 625 ppm    | 64±                    | 19 | 65±  | 19 |
| 1250 ppm   | 116±                   | 37 | 131± | 55 |
| 2500 ppm   | 242±                   | 81 | 250± | 86 |

(HAN300)

BAIS 5

**TABLE F 2**

**CHEMICAL INTAKE CHANGES: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 7

| Group Name | Administration (weeks) |         |         |         |         |         |         |  |  |  |  |  |  |  |
|------------|------------------------|---------|---------|---------|---------|---------|---------|--|--|--|--|--|--|--|
|            | 1                      | 2       | 3       | 4       | 5       | 6       | 7       |  |  |  |  |  |  |  |
| Control    | 0± 0                   | 0± 0    | 0± 0    | 0± 0    | 0± 0    | 0± 0    | 0± 0    |  |  |  |  |  |  |  |
| 625 ppm    | 123± 11                | 109± 9  | 106± 9  | 102± 10 | 103± 11 | 98± 9   | 99± 10  |  |  |  |  |  |  |  |
| 1250 ppm   | 225± 25                | 202± 21 | 210± 30 | 206± 33 | 208± 25 | 199± 28 | 201± 34 |  |  |  |  |  |  |  |
| 2500 ppm   | 340± 32                | 333± 33 | 335± 39 | 333± 39 | 339± 46 | 333± 46 | 336± 48 |  |  |  |  |  |  |  |

(HAN300)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 8

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 8                      |    | 9    |    | 10   |    | 11   |    | 12   |    | 13   |    | 14   |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 95±                    | 10 | 95±  | 10 | 94±  | 11 | 95±  | 9  | 91±  | 11 | 91±  | 10 | 86±  | 11 |
| 1250 ppm   | 192±                   | 20 | 186± | 23 | 187± | 22 | 191± | 33 | 180± | 23 | 180± | 25 | 176± | 32 |
| 2500 ppm   | 328±                   | 50 | 328± | 49 | 322± | 47 | 325± | 45 | 320± | 54 | 314± | 50 | 301± | 40 |

(HAN300)

BA1S5



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 9

| Group Name | Administration (weeks) |    |      |    |      |    |      |    |      |    |      |    |      |    |
|------------|------------------------|----|------|----|------|----|------|----|------|----|------|----|------|----|
|            | 18                     |    | 22   |    | 26   |    | 30   |    | 34   |    | 38   |    | 42   |    |
| Control    | 0±                     | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  |
| 625 ppm    | 79±                    | 10 | 78±  | 13 | 71±  | 10 | 68±  | 13 | 69±  | 15 | 66±  | 11 | 67±  | 12 |
| 1250 ppm   | 155±                   | 20 | 150± | 23 | 137± | 21 | 134± | 23 | 135± | 21 | 131± | 23 | 128± | 21 |
| 2500 ppm   | 279±                   | 43 | 274± | 43 | 259± | 44 | 247± | 39 | 255± | 43 | 244± | 45 | 242± | 33 |

(HAN300)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 10

| Group Name | Administration |    | (weeks) |    |      |    |      |    |      |    |      |    |      |     |
|------------|----------------|----|---------|----|------|----|------|----|------|----|------|----|------|-----|
|            | 46             |    | 50      |    | 54   |    | 58   |    | 62   |    | 66   |    | 70   |     |
| Control    | 0±             | 0  | 0±      | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0  | 0±   | 0   |
| 625 ppm    | 67±            | 14 | 63±     | 12 | 60±  | 13 | 60±  | 12 | 61±  | 15 | 65±  | 16 | 62±  | 17  |
| 1250 ppm   | 129±           | 30 | 119±    | 17 | 124± | 40 | 119± | 48 | 118± | 24 | 124± | 32 | 130± | 60  |
| 2500 ppm   | 239±           | 38 | 224±    | 39 | 224± | 42 | 216± | 34 | 220± | 47 | 237± | 69 | 247± | 109 |

(HAN300)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Crl:BDF1]  
 UNIT : mg/kg/day  
 REPORT TYPE : A1 104  
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
 ALL ANIMALS

PAGE : 11

| Group Name | Administration |     | (weeks) |     |      |     |      |     |      |     |      |     |      |     |
|------------|----------------|-----|---------|-----|------|-----|------|-----|------|-----|------|-----|------|-----|
|            | 74             |     | 78      |     | 82   |     | 86   |     | 90   |     | 94   |     | 98   |     |
| Control    | 0±             | 0   | 0±      | 0   | 0±   | 0   | 0±   | 0   | 0±   | 0   | 0±   | 0   | 0±   | 0   |
| 625 ppm    | 64±            | 22  | 66±     | 21  | 70±  | 27  | 72±  | 30  | 71±  | 38  | 72±  | 39  | 76±  | 35  |
| 1250 ppm   | 129±           | 52  | 137±    | 63  | 156± | 84  | 156± | 80  | 148± | 72  | 182± | 101 | 177± | 93  |
| 2500 ppm   | 267±           | 152 | 256±    | 143 | 280± | 178 | 281± | 169 | 309± | 188 | 307± | 191 | 337± | 209 |

(HAN300)

BAIS5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
UNIT : mg/kg/day  
REPORT TYPE : A1 104  
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)  
ALL ANIMALS

PAGE : 12

| Group Name | Administration (weeks) |     |      |     |
|------------|------------------------|-----|------|-----|
|            | 102                    |     | 104  |     |
| Control    | 0±                     | 0   | 0±   | 0   |
| 625 ppm    | 74±                    | 37  | 71±  | 21  |
| 1250 ppm   | 156±                   | 78  | 152± | 68  |
| 2500 ppm   | 356±                   | 192 | 360± | 200 |

(HAN300)

BAIS5

TABLE G 1

HEMATOLOGY: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Crl:BDF1]  
 MEASURE. TIME : 1  
 SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)  
 ALL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of<br>Animals | RED BLOOD CELL<br>10 <sup>6</sup> /μl |      | HEMOGLOBIN<br>g/dl |     | HEMATOCRIT<br>% |     | MCV<br>fl |       | MCH<br>pg |      | MCHC<br>g/dl |     | PLATELET<br>10 <sup>3</sup> /μl |     |
|------------|-------------------|---------------------------------------|------|--------------------|-----|-----------------|-----|-----------|-------|-----------|------|--------------|-----|---------------------------------|-----|
| Control    | 24                | 9.24±                                 | 1.47 | 13.1±              | 2.0 | 43.2±           | 6.5 | 46.9±     | 3.4   | 14.2±     | 0.9  | 30.2±        | 0.9 | 1921±                           | 524 |
| 625 ppm    | 32                | 8.83±                                 | 1.00 | 12.7±              | 1.5 | 42.4±           | 4.2 | 48.1±     | 1.7   | 14.4±     | 0.5  | 30.0±        | 0.9 | 1850±                           | 391 |
| 1250 ppm   | 31                | 8.63±                                 | 1.35 | 12.7±              | 1.8 | 42.1±           | 5.3 | 49.1±     | 2.9** | 14.7±     | 0.7  | 30.0±        | 1.0 | 1955±                           | 488 |
| 2500 ppm   | 38                | 8.42±                                 | 1.33 | 12.3±              | 1.9 | 41.2±           | 5.5 | 49.2±     | 2.8** | 14.6±     | 0.7* | 29.7±        | 1.0 | 1876±                           | 490 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
MEASURE TIME : 1  
SEX : MALE

HEMATOLOGY (SUMMARY)  
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of<br>Animals | RETICULOCYTE<br>% |       | METHEMOGLOBIN<br>% |       |
|------------|-------------------|-------------------|-------|--------------------|-------|
| Control    | 24                | 3.7±              | 3.7   | 0.4±               | 0.1   |
| 625 ppm    | 32                | 3.5±              | 1.6   | 0.5±               | 0.2   |
| 1250 ppm   | 31                | 4.3±              | 4.1   | 0.5±               | 0.3   |
| 2500 ppm   | 38                | 4.8±              | 3.3** | 0.7±               | 0.3** |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 MEASURE. TIME : 1  
 SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)  
 ALL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of<br>Animals | WBC<br>10 <sup>3</sup> /μl |       | Differential |    | WBC (%) |    | MONO |   | EOSINO |    | BASO |   | OTHER |   |
|------------|-------------------|----------------------------|-------|--------------|----|---------|----|------|---|--------|----|------|---|-------|---|
|            |                   |                            |       | NEUTRO       |    | LYMPHO  |    |      |   |        |    |      |   |       |   |
| Control    | 24                | 7.74±                      | 13.36 | 30±          | 14 | 62±     | 16 | 3±   | 2 | 3±     | 3  | 0±   | 0 | 3±    | 5 |
| 625 ppm    | 32                | 5.36±                      | 3.91  | 28±          | 16 | 63±     | 19 | 3±   | 2 | 4±     | 11 | 0±   | 0 | 3±    | 4 |
| 1250 ppm   | 31                | 4.37±                      | 2.40  | 34±          | 18 | 57±     | 20 | 3±   | 2 | 2±     | 2  | 0±   | 0 | 4±    | 6 |
| 2500 ppm   | 38                | 3.66±                      | 1.99* | 34±          | 18 | 59±     | 17 | 3±   | 2 | 1±     | 2  | 0±   | 0 | 3±    | 4 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS5



**TABLE G 2**

**HEMATOLOGY: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 MEASURE TIME : 1  
 SEX : FEMALE

HEMATOLOGY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of<br>Animals | RED BLOOD CELL<br>10 <sup>6</sup> /μl |       | HEMOGLOBIN<br>g/dl |      | HEMATOCRIT<br>% |     | MCV<br>fl |     | MCH<br>pg |     | MCHC<br>g/dl |       | PLATELET<br>10 <sup>3</sup> /μl |     |
|------------|-------------------|---------------------------------------|-------|--------------------|------|-----------------|-----|-----------|-----|-----------|-----|--------------|-------|---------------------------------|-----|
| Control    | 22                | 9.14±                                 | 1.10  | 13.3±              | 1.6  | 43.5±           | 4.1 | 47.9±     | 3.1 | 14.6±     | 0.6 | 30.4±        | 1.2   | 1081±                           | 353 |
| 625 ppm    | 21                | 9.21±                                 | 0.75  | 13.1±              | 1.2  | 44.2±           | 3.5 | 48.1±     | 1.9 | 14.3±     | 0.7 | 29.7±        | 1.0   | 1262±                           | 400 |
| 1250 ppm   | 28                | 8.64±                                 | 1.25  | 12.3±              | 1.9  | 41.8±           | 5.4 | 48.6±     | 3.0 | 14.3±     | 0.9 | 29.4±        | 1.2*  | 1150±                           | 413 |
| 2500 ppm   | 38                | 8.31±                                 | 1.43* | 11.8±              | 2.3* | 40.2±           | 6.2 | 48.7±     | 4.7 | 14.3±     | 1.3 | 29.3±        | 1.5** | 1232±                           | 485 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
MEASURE. TIME : 1  
SEX : FEMALE

HEMATOLOGY (SUMMARY)  
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 5

| Group Name | NO. of<br>Animals | RETICULOCYTE<br>% |       | METHEMOGLOBIN<br>% |       |
|------------|-------------------|-------------------|-------|--------------------|-------|
| Control    | 22                | 3.7±              | 4.1   | 0.3±               | 0.1   |
| 625 ppm    | 21                | 3.5±              | 1.2   | 0.4±               | 0.2   |
| 1250 ppm   | 28                | 4.7±              | 2.0** | 0.5±               | 0.2   |
| 2500 ppm   | 38                | 6.5±              | 5.3** | 0.6±               | 0.3** |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 MEASURE TIME : 1  
 SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)  
 ALL ANIMALS (105W)

PAGE : 6

| Group Name | NO. of<br>Animals | WBC<br>10 <sup>3</sup> /μl |       | Differential |    | WBC (%) |    | MONO |   | EOSINO |   | BASO |   | OTHER |    |
|------------|-------------------|----------------------------|-------|--------------|----|---------|----|------|---|--------|---|------|---|-------|----|
|            |                   |                            |       | NEUTRO       |    | LYMPHO  |    |      |   |        |   |      |   |       |    |
| Control    | 22                | 5.36 ±                     | 6.67  | 25 ±         | 15 | 61 ±    | 22 | 3 ±  | 2 | 2 ±    | 3 | 0 ±  | 0 | 10 ±  | 21 |
| 625 ppm    | 21                | 8.13 ±                     | 11.97 | 22 ±         | 12 | 57 ±    | 20 | 3 ±  | 3 | 1 ±    | 1 | 0 ±  | 0 | 17 ±  | 28 |
| 1250 ppm   | 28                | 3.61 ±                     | 2.06  | 32 ±         | 20 | 53 ±    | 25 | 2 ±  | 2 | 1 ±    | 1 | 0 ±  | 0 | 13 ±  | 11 |
| 2500 ppm   | 38                | 4.37 ±                     | 5.71  | 29 ±         | 18 | 53 ±    | 22 | 3 ±  | 2 | 2 ±    | 2 | 0 ±  | 0 | 14 ±  | 21 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS5

TABLE H 1

BIOCHEMISTRY: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 MEASURE. TIME : 1  
 SEX : MALE

BIOCHEMISTRY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of<br>Animals | TOTAL PROTEIN<br>g/dl |       | ALBUMIN<br>g/dl |     | A/G RATIO |     | T-BILIRUBIN<br>mg/dl |      | GLUCOSE<br>mg/dl |    | T-CHOLESTEROL<br>mg/dl |    | TRIGLYCERIDE<br>mg/dl |    |
|------------|-------------------|-----------------------|-------|-----------------|-----|-----------|-----|----------------------|------|------------------|----|------------------------|----|-----------------------|----|
| Control    | 24                | 5.4±                  | 0.7   | 2.6±            | 0.4 | 0.9±      | 0.2 | 0.11±                | 0.04 | 156±             | 37 | 149±                   | 71 | 47±                   | 21 |
| 625 ppm    | 32                | 5.1±                  | 0.5*  | 2.4±            | 0.3 | 0.9±      | 0.1 | 0.10±                | 0.03 | 158±             | 37 | 119±                   | 28 | 43±                   | 28 |
| 1250 ppm   | 31                | 5.1±                  | 0.5*  | 2.4±            | 0.3 | 1.0±      | 0.2 | 0.10±                | 0.02 | 157±             | 45 | 127±                   | 27 | 48±                   | 28 |
| 2500 ppm   | 38                | 5.0±                  | 0.5** | 2.4±            | 0.3 | 1.0±      | 0.1 | 0.12±                | 0.04 | 153±             | 37 | 129±                   | 31 | 49±                   | 35 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 MEASURE. TIME : 1  
 SEX : MALE

BIOCHEMISTRY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of<br>Animals | PHOSPHOLIPID<br>mg/dl |    | AST<br>U/L |     | ALT<br>U/L |     | LDH<br>U/L |      | ALP<br>U/L |     | G-GTP<br>U/L |   | CK<br>U/L |     |
|------------|-------------------|-----------------------|----|------------|-----|------------|-----|------------|------|------------|-----|--------------|---|-----------|-----|
| Control    | 24                | 247±                  | 86 | 85±        | 71  | 44±        | 49  | 268±       | 134  | 264±       | 276 | 1±           | 1 | 51±       | 21  |
| 625 ppm    | 32                | 206±                  | 48 | 100±       | 155 | 64±        | 164 | 314±       | 603  | 191±       | 66  | 1±           | 0 | 79±       | 191 |
| 1250 ppm   | 31                | 210±                  | 52 | 240±       | 833 | 136±       | 493 | 516±       | 1557 | 175±       | 47  | 1±           | 1 | 59±       | 38  |
| 2500 ppm   | 38                | 214±                  | 50 | 94±        | 126 | 43±        | 81  | 282±       | 344  | 178±       | 51  | 1±           | 1 | 73±       | 128 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 MEASURE TIME : 1  
 SEX : MALE

BIOCHEMISTRY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 3

| Group Name | NO. of<br>Animals | UREA NITROGEN<br>mg/dl |      | SODIUM<br>mEq/l |   | POTASSIUM<br>mEq/l |     | CHLORIDE<br>mEq/l |   | CALCIUM<br>mg/dl |     | INORGANIC PHOSPHORUS<br>mg/dl |     |
|------------|-------------------|------------------------|------|-----------------|---|--------------------|-----|-------------------|---|------------------|-----|-------------------------------|-----|
| Control    | 24                | 24.8±                  | 5.8  | 154±            | 2 | 4.3±               | 0.4 | 121±              | 3 | 9.2±             | 0.5 | 6.0±                          | 0.8 |
| 625 ppm    | 32                | 26.6±                  | 13.4 | 154±            | 2 | 4.3±               | 0.6 | 123±              | 3 | 8.9±             | 0.3 | 5.8±                          | 0.9 |
| 1250 ppm   | 31                | 30.2±                  | 16.9 | 154±            | 2 | 4.4±               | 0.8 | 123±              | 4 | 8.9±             | 0.3 | 6.0±                          | 1.2 |
| 2500 ppm   | 38                | 28.5±                  | 16.3 | 155±            | 4 | 4.3±               | 0.6 | 123±              | 5 | 8.9±             | 0.5 | 6.0±                          | 0.7 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5



**TABLE H 2**

**BIOCHEMISTRY: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 MEASURE. TIME : 1  
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of<br>Animals | TOTAL PROTEIN<br>g/dl |     | ALBUMIN<br>g/dl |     | A/G RATIO |     | T-BILIRUBIN<br>mg/dl |       | GLUCOSE<br>mg/dl |    | T-CHOLESTEROL<br>mg/dl |      | TRIGLYCERIDE<br>mg/dl |    |
|------------|-------------------|-----------------------|-----|-----------------|-----|-----------|-----|----------------------|-------|------------------|----|------------------------|------|-----------------------|----|
| Control    | 22                | 5.0±                  | 0.5 | 2.6±            | 0.2 | 1.1±      | 0.2 | 0.10±                | 0.02  | 118±             | 34 | 83±                    | 26   | 31±                   | 15 |
| 625 ppm    | 21                | 5.2±                  | 0.6 | 2.4±            | 0.3 | 0.9±      | 0.2 | 0.11±                | 0.05  | 106±             | 30 | 100±                   | 29   | 40±                   | 24 |
| 1250 ppm   | 28                | 5.1±                  | 0.6 | 2.6±            | 0.2 | 1.1±      | 0.2 | 0.11±                | 0.02  | 121±             | 28 | 110±                   | 48*  | 54±                   | 75 |
| 2500 ppm   | 38                | 5.2±                  | 0.8 | 2.6±            | 0.3 | 1.0±      | 0.2 | 0.13±                | 0.04* | 119±             | 24 | 112±                   | 39** | 49±                   | 55 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
MEASURE TIME : 1  
SEX : FEMALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS (105W)

PAGE : 5

| Group Name | NO. of<br>Animals | PHOSPHOLIPID<br>mg/dl |      | AST<br>U/L |     | ALT<br>U/L |    | LDH<br>U/L |      | ALP<br>U/L |     | G-GTP<br>U/L |   | CK<br>U/L |     |
|------------|-------------------|-----------------------|------|------------|-----|------------|----|------------|------|------------|-----|--------------|---|-----------|-----|
| Control    | 22                | 151±                  | 43   | 115±       | 79  | 44±        | 36 | 255±       | 145  | 412±       | 290 | 1±           | 1 | 84±       | 63  |
| 625 ppm    | 21                | 174±                  | 43   | 114±       | 61  | 42±        | 28 | 292±       | 187  | 330±       | 220 | 1±           | 1 | 211±      | 458 |
| 1250 ppm   | 28                | 197±                  | 81** | 115±       | 95  | 41±        | 35 | 394±       | 640  | 319±       | 176 | 1±           | 1 | 159±      | 222 |
| 2500 ppm   | 38                | 188±                  | 59*  | 148±       | 154 | 55±        | 67 | 631±       | 1182 | 251±       | 111 | 1±           | 1 | 97±       | 74  |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 MEASURE. TIME : 1  
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)  
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 6

| Group Name | NQ. of<br>Animals | UREA NITROGEN<br>mg/dl |      | SODIUM<br>mEq/l |   | POTASSIUM<br>mEq/l |     | CHLORIDE<br>mEq/l |   | CALCIUM<br>mg/dl |     | INORGANIC PHOSPHORUS<br>mg/dl |     |
|------------|-------------------|------------------------|------|-----------------|---|--------------------|-----|-------------------|---|------------------|-----|-------------------------------|-----|
| Control    | 22                | 23.6±                  | 16.2 | 152±            | 3 | 4.1±               | 0.6 | 121±              | 3 | 9.2±             | 0.5 | 6.1±                          | 1.3 |
| 625 ppm    | 21                | 27.9±                  | 25.2 | 153±            | 2 | 4.0±               | 0.4 | 123±              | 3 | 9.6±             | 1.4 | 7.4±                          | 2.9 |
| 1250 ppm   | 28                | 31.6±                  | 24.7 | 153±            | 4 | 4.1±               | 0.3 | 123±              | 4 | 9.5±             | 0.7 | 6.9±                          | 1.9 |
| 2500 ppm   | 38                | 28.8±                  | 17.6 | 153±            | 2 | 4.1±               | 0.5 | 122±              | 3 | 9.6±             | 0.9 | 7.3±                          | 2.5 |

Significant difference ; \* :  $P \leq 0.05$

\*\* :  $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 5

TABLE I 1

URINALYSIS: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 MEASURE. TIME : 1  
 SEX : MALE REPORT TYPE : A1

URINALYSIS

PAGE : 1

| Group Name | NO. of<br>Animals | pH_____ |     |     |     |     |     |     |   | CHI | Protein_____ |    |    |    |    |   |    |   | CHI | Glucose_____ |    |    |   |    |    |    |    | CHI | Ketone body_____ |   |    |   |    |    |   |  | CHI | Occult blood_____ |  |  |  |  | CHI |
|------------|-------------------|---------|-----|-----|-----|-----|-----|-----|---|-----|--------------|----|----|----|----|---|----|---|-----|--------------|----|----|---|----|----|----|----|-----|------------------|---|----|---|----|----|---|--|-----|-------------------|--|--|--|--|-----|
|            |                   | 5.0     | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | - |     | ±            | +  | 2+ | 3+ | 4+ | - | ±  | + |     | 2+           | 3+ | 4+ | - | ±  | +  | 2+ | 3+ |     | 4+               | - | ±  | + | 2+ | 3+ |   |  |     |                   |  |  |  |  |     |
| Control    | 25                | 0       | 5   | 8   | 3   | 1   | 8   | 0   |   | 0   | 5            | 16 | 2  | 2  | 0  |   | 25 | 0 | 0   | 0            | 0  | 0  |   | 10 | 14 | 0  | 1  | 0   | 0                |   | 25 | 0 | 0  | 0  | 0 |  |     |                   |  |  |  |  |     |
| 625 ppm    | 33                | 0       | 7   | 11  | 6   | 4   | 4   | 1   |   | 0   | 5            | 18 | 8  | 2  | 0  |   | 33 | 0 | 0   | 0            | 0  | 0  |   | 9  | 22 | 1  | 1  | 0   | 0                |   | 29 | 2 | 0  | 1  | 1 |  |     |                   |  |  |  |  |     |
| 1250 ppm   | 32                | 0       | 6   | 13  | 5   | 7   | 1   | 0   | * | 0   | 2            | 17 | 12 | 1  | 0  | * | 32 | 0 | 0   | 0            | 0  | 0  |   | 7  | 19 | 5  | 1  | 0   | 0                |   | 30 | 0 | 0  | 0  | 2 |  |     |                   |  |  |  |  |     |
| 2500 ppm   | 39                | 0       | 12  | 12  | 9   | 5   | 1   | 0   | * | 0   | 1            | 22 | 14 | 2  | 0  | * | 39 | 0 | 0   | 0            | 0  | 0  |   | 5  | 22 | 7  | 5  | 0   | 0                | * | 39 | 0 | 0  | 0  | 0 |  |     |                   |  |  |  |  |     |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
MEASURE. TIME : 1  
SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of<br>Animals | Urobilinogen |   |    |    |    | CHI |
|------------|-------------------|--------------|---|----|----|----|-----|
|            |                   | ±            | + | 2+ | 3+ | 4+ |     |
| Control    | 25                | 25           | 0 | 0  | 0  | 0  | 0   |
| 625 ppm    | 33                | 33           | 0 | 0  | 0  | 0  | 0   |
| 1250 ppm   | 32                | 32           | 0 | 0  | 0  | 0  | 0   |
| 2500 ppm   | 39                | 39           | 0 | 0  | 0  | 0  | 0   |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS 5

TABLE I 2

URINALYSIS: FEMALE



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 MEASURE. TIME : 1  
 SEX : FEMALE REPORT TYPE : A1

URINALYSIS

PAGE : 3

| Group Name | NO. of<br>Animals | pH_____ |     |     |     |     |     |     |    | CHI | Protein_____ |    |    |    |    |   | CHI | Glucose_____ |   |    |    |    |   | CHI | Ketone body |    |    |    |    |   | CHI | Occult blood |   |    |    |  | CHI |
|------------|-------------------|---------|-----|-----|-----|-----|-----|-----|----|-----|--------------|----|----|----|----|---|-----|--------------|---|----|----|----|---|-----|-------------|----|----|----|----|---|-----|--------------|---|----|----|--|-----|
|            |                   | 5.0     | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | -  |     | ±            | +  | 2+ | 3+ | 4+ | - |     | ±            | + | 2+ | 3+ | 4+ | - |     | ±           | +  | 2+ | 3+ | 4+ | - |     | ±            | + | 2+ | 3+ |  |     |
| Control    | 25                | 0       | 6   | 6   | 1   | 1   | 7   | 4   |    | 2   | 7            | 10 | 4  | 2  | 0  |   | 25  | 0            | 0 | 0  | 0  | 0  |   | 12  | 6           | 7  | 0  | 0  | 0  |   | 23  | 2            | 0 | 0  | 0  |  |     |
| 625 ppm    | 21                | 0       | 0   | 6   | 4   | 7   | 1   | 3   | ** | 2   | 7            | 6  | 6  | 0  | 0  |   | 21  | 0            | 0 | 0  | 0  | 0  |   | 9   | 6           | 6  | 0  | 0  | 0  |   | 21  | 0            | 0 | 0  | 0  |  |     |
| 1250 ppm   | 33                | 0       | 8   | 12  | 2   | 3   | 6   | 2   |    | 3   | 10           | 12 | 8  | 0  | 0  |   | 33  | 0            | 0 | 0  | 0  | 0  |   | 15  | 7           | 10 | 1  | 0  | 0  |   | 31  | 0            | 0 | 1  | 1  |  |     |
| 2500 ppm   | 38                | 0       | 9   | 9   | 7   | 7   | 3   | 3   |    | 1   | 10           | 11 | 9  | 6  | 1  |   | 38  | 0            | 0 | 0  | 0  | 0  |   | 12  | 9           | 11 | 3  | 3  | 0  |   | 37  | 0            | 0 | 0  | 1  |  |     |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj[Crlj:BDF1]  
MEASURE TIME : 1  
SEX : FEMALE REPORT TYPE : A1

URINALYSIS

PAGE : 4

| Group Name | NO. of<br>Animals | Urobilinogen |   |    |    |    | CHI |
|------------|-------------------|--------------|---|----|----|----|-----|
|            |                   | ±            | + | 2+ | 3+ | 4+ |     |
| Control    | 25                | 25           | 0 | 0  | 0  | 0  |     |
| 625 ppm    | 21                | 21           | 0 | 0  | 0  | 0  |     |
| 1250 ppm   | 33                | 33           | 0 | 0  | 0  | 0  |     |
| 2500 ppm   | 38                | 38           | 0 | 0  | 0  | 0  |     |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS 5

**TABLE J 1**

**GROSS FINDINGS: MALE: ALL ANIMALS**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 1

| Organ       | Findings                   | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------------|----------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| skin/app    | ulcer                      |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|             | erosion                    |                              | 1 ( 2)            | 6 ( 12)           | 8 ( 16)            | 1 ( 2)             |
|             | scab                       |                              | 2 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| subcutis    | edema                      |                              | 1 ( 2)            | 2 ( 4)            | 1 ( 2)             | 3 ( 6)             |
|             | dry                        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | mass                       |                              | 0 ( 0)            | 2 ( 4)            | 1 ( 2)             | 3 ( 6)             |
| nasal cavit | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| lung        | white zone                 |                              | 1 ( 2)            | 2 ( 4)            | 2 ( 4)             | 1 ( 2)             |
|             | nodule                     |                              | 11 ( 22)          | 10 ( 20)          | 8 ( 16)            | 9 ( 18)            |
|             | cyst                       |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
| lymph node  | enlarged                   |                              | 7 ( 14)           | 5 ( 10)           | 9 ( 18)            | 3 ( 6)             |
| thymus      | atrophic                   |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| spleen      | enlarged                   |                              | 6 ( 12)           | 2 ( 4)            | 6 ( 12)            | 2 ( 4)             |
|             | white zone                 |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 1 ( 2)             |
|             | nodule                     |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 1 ( 2)             |
|             | nodular                    |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
|             | accentuation of white pulp |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| heart       | white zone                 |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| oral cavity | nodule                     |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| stomach     | forestomach:erosion        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | glandular stomach:red zone |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
| duodenum    | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 2

| Organ       | Findings               | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------------|------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| small intes | red zone               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
|             | nodule                 |                              | 1 ( 2)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| liver       | enlarged               |                              | 2 ( 4)            | 2 ( 4)            | 4 ( 8)             | 0 ( 0)             |
|             | atrophic               |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|             | dark                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 2 ( 4)             |
|             | white zone             |                              | 9 ( 18)           | 0 ( 0)            | 5 ( 10)            | 3 ( 6)             |
|             | red zone               |                              | 2 ( 4)            | 1 ( 2)            | 2 ( 4)             | 2 ( 4)             |
|             | nodule                 |                              | 21 ( 42)          | 17 ( 34)          | 17 ( 34)           | 7 ( 14)            |
|             | cyst                   |                              | 2 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | nodular                |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | adhesion               |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|             | nodule                 |                              | 1 ( 2)            | 1 ( 2)            | 0 ( 0)             | 1 ( 2)             |
| pancreas    | enlarged               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| kidney      | small                  |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
|             | white zone             |                              | 1 ( 2)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|             | nodule                 |                              | 1 ( 2)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|             | deformed               |                              | 1 ( 2)            | 3 ( 6)            | 1 ( 2)             | 3 ( 6)             |
|             | granular               |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
|             | hydronephrosis         |                              | 9 ( 18)           | 11 ( 22)          | 6 ( 12)            | 7 ( 14)            |
|             | nodule                 |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| urin bladd  | thick                  |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | urine:marked retention |                              | 1 ( 2)            | 7 ( 14)           | 5 ( 10)            | 2 ( 4)             |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

GROSS FINDINGS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 3

| Organ       | Findings      | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------------|---------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| urin bladd  | urine:turbid  |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| thyroid     | dark          |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 3 ( 6)             |
|             | red zone      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
| testis      | enlarged      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
|             | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| epididymis  | nodule        |                              | 0 ( 0)            | 2 ( 4)            | 1 ( 2)             | 2 ( 4)             |
| prep/cli gl | nodule        |                              | 1 ( 2)            | 3 ( 6)            | 1 ( 2)             | 0 ( 0)             |
| brain       | red zone      |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
| eye         | turbid        |                              | 1 ( 2)            | 0 ( 0)            | 2 ( 4)             | 0 ( 0)             |
| Harder gl   | enlarged      |                              | 3 ( 6)            | 0 ( 0)            | 2 ( 4)             | 0 ( 0)             |
|             | nodule        |                              | 0 ( 0)            | 2 ( 4)            | 2 ( 4)             | 0 ( 0)             |
| pleura      | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| mediastinum | mass          |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| peritoneum  | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | mass          |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| abdominal c | hemorrhage    |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | ascites       |                              | 7 ( 14)           | 3 ( 6)            | 4 ( 8)             | 3 ( 6)             |
| mesenterium | nodule        |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| thoracic ca | hemorrhage    |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid |                              | 4 ( 8)            | 0 ( 0)            | 3 ( 6)             | 1 ( 2)             |
| other       | scab          |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

GROSS FINDINGS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 4

| Organ      | Findings        | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|------------|-----------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| other      | tail:nodule     |                              | 0 ( 0)            | 0 ( 0)            | 2 ( 4)             | 0 ( 0)             |
|            | hindlimb:nodule |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| whole body | anemic          |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

(HPT080)

BAIS 5

**TABLE J 2**

**GROSS FINDINGS: MALE: DEAD AND MORIBUND ANIMALS**



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 1

| Organ       | Findings                   | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>18 (%) | 1250 ppm<br>19 (%) | 2500 ppm<br>11 (%) |
|-------------|----------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| skin/app    | ulcer                      |                              | 0 ( 0)            | 1 ( 6)            | 1 ( 5)             | 0 ( 0)             |
|             | erosion                    |                              | 1 ( 4)            | 5 ( 28)           | 5 ( 26)            | 1 ( 9)             |
|             | scab                       |                              | 2 ( 8)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| subcutis    | edema                      |                              | 1 ( 4)            | 2 ( 11)           | 1 ( 5)             | 2 ( 18)            |
|             | dry                        |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | mass                       |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 2 ( 18)            |
| nasal cavit | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| lung        | white zone                 |                              | 1 ( 4)            | 1 ( 6)            | 1 ( 5)             | 0 ( 0)             |
|             | nodule                     |                              | 4 ( 15)           | 1 ( 6)            | 2 ( 11)            | 3 ( 27)            |
| lymph node  | enlarged                   |                              | 3 ( 12)           | 2 ( 11)           | 5 ( 26)            | 1 ( 9)             |
| thymus      | atrophic                   |                              | 0 ( 0)            | 1 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| spleen      | enlarged                   |                              | 5 ( 19)           | 1 ( 6)            | 6 ( 32)            | 1 ( 9)             |
|             | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| heart       | white zone                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| oral cavity | nodule                     |                              | 0 ( 0)            | 1 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| stomach     | forestomach:erosion        |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | glandular stomach:red zone |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| duodenum    | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 9)             |
| small intes | red zone                   |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| liver       | enlarged                   |                              | 2 ( 8)            | 2 ( 11)           | 4 ( 21)            | 0 ( 0)             |
|             | white zone                 |                              | 7 ( 27)           | 0 ( 0)            | 4 ( 21)            | 0 ( 0)             |
|             | red zone                   |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 2

| Organ       | Findings               | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>18 (%) | 1250 ppm<br>19 (%) | 2500 ppm<br>11 (%) |
|-------------|------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| liver       | nodule                 |                              | 8 ( 31)           | 4 ( 22)           | 5 ( 26)            | 3 ( 27)            |
|             | nodular                |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| pancreas    | nodule                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| kidney      | enlarged               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
|             | small                  |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
|             | white zone             |                              | 1 ( 4)            | 1 ( 6)            | 1 ( 5)             | 0 ( 0)             |
|             | nodule                 |                              | 1 ( 4)            | 1 ( 6)            | 1 ( 5)             | 0 ( 0)             |
|             | deformed               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
|             | granular               |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 9)             |
|             | hydronephrosis         |                              | 7 ( 27)           | 6 ( 33)           | 4 ( 21)            | 3 ( 27)            |
|             | urin bladd             |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | urine:marked retention |                              | 1 ( 4)            | 7 ( 39)           | 5 ( 26)            | 2 ( 18)            |
|             | urine:turbid           |                              | 0 ( 0)            | 1 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| thyroid     | red zone               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| testis      | enlarged               |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 9)             |
| epididymis  | nodule                 |                              | 0 ( 0)            | 2 ( 11)           | 1 ( 5)             | 1 ( 9)             |
| prep/cli gl | nodule                 |                              | 0 ( 0)            | 1 ( 6)            | 1 ( 5)             | 0 ( 0)             |
| brain       | red zone               |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 9)             |
| eye         | turbid                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| Harder gl   | enlarged               |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | nodule                 |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| pleura      | nodule                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 3

| Organ       | Findings        | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>18 (%) | 1250 ppm<br>19 (%) | 2500 ppm<br>11 (%) |
|-------------|-----------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| mediastinum | mass            |                              | 0 ( 0)            | 1 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| peritoneum  | nodule          |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | mass            |                              | 0 ( 0)            | 1 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| abdominal c | hemorrhage      |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid   |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | ascites         |                              | 7 ( 27)           | 3 ( 17)           | 3 ( 16)            | 3 ( 27)            |
| thoracic ca | hemorrhage      |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid   |                              | 3 ( 12)           | 0 ( 0)            | 2 ( 11)            | 0 ( 0)             |
| other       | scab            |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | tail:nodule     |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
|             | hindlimb:nodule |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 5)             | 0 ( 0)             |
| whole body  | anemic          |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

(HPT080)

BAIS 5

**TABLE J 3**

**GROSS FINDINGS: MALE: SACRIFICED ANIMALS**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 1

| Organ       | Findings                   | Group Name<br>NO. of Animals | Control<br>24 (%) | 625 ppm<br>32 (%) | 1250 ppm<br>31 (%) | 2500 ppm<br>39 (%) |
|-------------|----------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| skin/app    | erosion                    |                              | 0 ( 0)            | 1 ( 3)            | 3 ( 10)            | 0 ( 0)             |
| subcutis    | edema                      |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
|             | mass                       |                              | 0 ( 0)            | 2 ( 6)            | 0 ( 0)             | 1 ( 3)             |
| nasal cavit | nodule                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| lung        | white zone                 |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 3)             | 1 ( 3)             |
|             | nodule                     |                              | 7 ( 29)           | 9 ( 28)           | 6 ( 19)            | 6 ( 15)            |
|             | cyst                       |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| lymph node  | enlarged                   |                              | 4 ( 17)           | 3 ( 9)            | 4 ( 13)            | 2 ( 5)             |
| spleen      | enlarged                   |                              | 1 ( 4)            | 1 ( 3)            | 0 ( 0)             | 1 ( 3)             |
|             | white zone                 |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 1 ( 3)             |
|             | nodule                     |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 1 ( 3)             |
|             | nodular                    |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|             | accentuation of white pulp |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| small intes | nodule                     |                              | 1 ( 4)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| liver       | atrophic                   |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|             | dark                       |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 2 ( 5)             |
|             | white zone                 |                              | 2 ( 8)            | 0 ( 0)            | 1 ( 3)             | 3 ( 8)             |
|             | red zone                   |                              | 1 ( 4)            | 1 ( 3)            | 2 ( 6)             | 2 ( 5)             |
|             | nodule                     |                              | 13 ( 54)          | 13 ( 41)          | 12 ( 39)           | 4 ( 10)            |
|             | cyst                       |                              | 2 ( 8)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | adhesion                   |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| pancreas    | nodule                     |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 1 ( 3)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

GROSS FINDINGS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 2

| Organ       | Findings        | Group Name<br>NO. of Animals | Control<br>24 (%) | 625 ppm<br>32 (%) | 1250 ppm<br>31 (%) | 2500 ppm<br>39 (%) |
|-------------|-----------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| kidney      | enlarged        |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
|             | small           |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
|             | deformed        |                              | 1 ( 4)            | 3 ( 9)            | 0 ( 0)             | 3 ( 8)             |
|             | hydronephrosis  |                              | 2 ( 8)            | 5 ( 16)           | 2 ( 6)             | 4 ( 10)            |
| urin bladd  | nodule          |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| thyroid     | dark            |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 3 ( 8)             |
| testis      | enlarged        |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|             | nodule          |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| epididymis  | nodule          |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| prep/cli gl | nodule          |                              | 1 ( 4)            | 2 ( 6)            | 0 ( 0)             | 0 ( 0)             |
| eye         | turbid          |                              | 0 ( 0)            | 0 ( 0)            | 2 ( 6)             | 0 ( 0)             |
| Harder gl   | enlarged        |                              | 2 ( 8)            | 0 ( 0)            | 2 ( 6)             | 0 ( 0)             |
|             | nodule          |                              | 0 ( 0)            | 2 ( 6)            | 1 ( 3)             | 0 ( 0)             |
| abdominal c | ascites         |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
| mesenterium | nodule          |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| thoracic ca | pleural fluid   |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 3)             | 1 ( 3)             |
| other       | tail:nodule     |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|             | hindlimb:nodule |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |

**TABLE J 4**

**GROSS FINDINGS: FEMALE: ALL ANIMALS**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 5

| Organ       | Findings                 | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------------|--------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| skin/app    | scab                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
| subcutis    | edema                    |                              | 3 ( 6)            | 3 ( 6)            | 4 ( 8)             | 2 ( 4)             |
|             | mass                     |                              | 1 ( 2)            | 5 ( 10)           | 2 ( 4)             | 5 ( 10)            |
| lung        | white zone               |                              | 1 ( 2)            | 1 ( 2)            | 2 ( 4)             | 0 ( 0)             |
|             | red zone                 |                              | 2 ( 4)            | 2 ( 4)            | 0 ( 0)             | 2 ( 4)             |
|             | nodule                   |                              | 1 ( 2)            | 6 ( 12)           | 2 ( 4)             | 2 ( 4)             |
| lymph node  | enlarged                 |                              | 8 ( 16)           | 6 ( 12)           | 10 ( 20)           | 9 ( 18)            |
| spleen      | enlarged                 |                              | 12 ( 24)          | 12 ( 24)          | 8 ( 16)            | 7 ( 14)            |
|             | nodule                   |                              | 1 ( 2)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| heart       | white zone               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| stomach     | forestomach:nodule       |                              | 1 ( 2)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|             | glandular stomach:nodule |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|             | glandular stomach:thick  |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 1 ( 2)             |
| small intes | nodule                   |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|             | adhesion                 |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| liver       | enlarged                 |                              | 10 ( 20)          | 7 ( 14)           | 3 ( 6)             | 1 ( 2)             |
|             | dark                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 4 ( 8)             |
|             | white zone               |                              | 9 ( 18)           | 8 ( 16)           | 4 ( 8)             | 2 ( 4)             |
|             | red zone                 |                              | 3 ( 6)            | 0 ( 0)            | 4 ( 8)             | 4 ( 8)             |
|             | black zone               |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | nodule                   |                              | 7 ( 14)           | 10 ( 20)          | 6 ( 12)            | 4 ( 8)             |
|             | cyst                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 6

| Organ      | Findings               | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|------------|------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| liver      | deformed               |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|            | rough                  |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
| pancreas   | nodule                 |                              | 2 ( 4)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
| kidney     | enlarged               |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|            | small                  |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
|            | pale                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
|            | white zone             |                              | 1 ( 2)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
|            | nodule                 |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|            | cyst                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 2 ( 4)             |
|            | deformed               |                              | 3 ( 6)            | 8 ( 16)           | 7 ( 14)            | 11 ( 22)           |
|            | hydronephrosis         |                              | 4 ( 8)            | 4 ( 8)            | 6 ( 12)            | 5 ( 10)            |
| ureter     | dilated                |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| urin bladd | nodule                 |                              | 1 ( 2)            | 1 ( 2)            | 1 ( 2)             | 0 ( 0)             |
|            | urine:marked retention |                              | 1 ( 2)            | 2 ( 4)            | 0 ( 0)             | 0 ( 0)             |
| pituitary  | enlarged               |                              | 1 ( 2)            | 4 ( 8)            | 3 ( 6)             | 4 ( 8)             |
|            | red zone               |                              | 2 ( 4)            | 3 ( 6)            | 3 ( 6)             | 0 ( 0)             |
|            | nodule                 |                              | 1 ( 2)            | 3 ( 6)            | 2 ( 4)             | 2 ( 4)             |
|            | cyst                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 2)             |
| thyroid    | dark                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 5 ( 10)            |
| ovary      | enlarged               |                              | 7 ( 14)           | 8 ( 16)           | 4 ( 8)             | 0 ( 0)             |
|            | nodule                 |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
|            | cyst                   |                              | 3 ( 6)            | 7 ( 14)           | 8 ( 16)            | 7 ( 14)            |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 7

| Organ       | Findings      | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------------|---------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| uterus      | nodule        |                              | 19 ( 38)          | 12 ( 24)          | 9 ( 18)            | 10 ( 20)           |
|             | dilated lumen |                              | 1 ( 2)            | 0 ( 0)            | 1 ( 2)             | 2 ( 4)             |
| vagina      | nodule        |                              | 0 ( 0)            | 2 ( 4)            | 0 ( 0)             | 0 ( 0)             |
| brain       | red zone      |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 1 ( 2)             |
|             | black zone    |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 1 ( 2)             |
|             | deformed      |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| eye         | turbid        |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 2 ( 4)             |
| Harder gl   | enlarged      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 2 ( 4)             |
|             | nodule        |                              | 0 ( 0)            | 0 ( 0)            | 2 ( 4)             | 1 ( 2)             |
| muscle      | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |
| bone        | nodule        |                              | 0 ( 0)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| pleura      | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| mediastinum | mass          |                              | 4 ( 8)            | 4 ( 8)            | 1 ( 2)             | 0 ( 0)             |
| peritoneum  | nodule        |                              | 1 ( 2)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
|             | adhesion      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 0 ( 0)             |
|             | thick         |                              | 3 ( 6)            | 1 ( 2)            | 0 ( 0)             | 0 ( 0)             |
| retroperit  | mass          |                              | 0 ( 0)            | 1 ( 2)            | 1 ( 2)             | 1 ( 2)             |
| abdominal c | hemorrhage    |                              | 1 ( 2)            | 2 ( 4)            | 1 ( 2)             | 0 ( 0)             |
|             | ascites       |                              | 10 ( 20)          | 10 ( 20)          | 10 ( 20)           | 4 ( 8)             |
| thoracic ca | hemorrhage    |                              | 3 ( 6)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid |                              | 14 ( 28)          | 12 ( 24)          | 9 ( 18)            | 6 ( 12)            |
| other       | tail:nodule   |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 2)             | 1 ( 2)             |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 8

| Organ | Findings    | Group Name<br>NO. of Animals | Control<br>50 (%) | 625 ppm<br>50 (%) | 1250 ppm<br>50 (%) | 2500 ppm<br>50 (%) |
|-------|-------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| other | nose:nodule |                              | 1 ( 2)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

(HPT080)

BAIS 5

**TABLE J 5**

**GROSS FINDINGS: FEMALE: DEAD AND MORIBUND ANIMALS**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 4

| Organ       | Findings                 | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>29 (%) | 1250 ppm<br>17 (%) | 2500 ppm<br>12 (%) |
|-------------|--------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| skin/app    | scab                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 8)             |
| subcutis    | edema                    |                              | 3 ( 12)           | 3 ( 10)           | 3 ( 18)            | 2 ( 17)            |
|             | mass                     |                              | 1 ( 4)            | 4 ( 14)           | 0 ( 0)             | 3 ( 25)            |
| lung        | white zone               |                              | 1 ( 4)            | 1 ( 3)            | 2 ( 12)            | 0 ( 0)             |
|             | red zone                 |                              | 2 ( 8)            | 2 ( 7)            | 0 ( 0)             | 2 ( 17)            |
|             | nodule                   |                              | 1 ( 4)            | 1 ( 3)            | 0 ( 0)             | 1 ( 8)             |
| lymph node  | enlarged                 |                              | 6 ( 23)           | 6 ( 21)           | 5 ( 29)            | 3 ( 25)            |
| spleen      | enlarged                 |                              | 8 ( 31)           | 10 ( 34)          | 4 ( 24)            | 4 ( 33)            |
|             | nodule                   |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 6)             | 0 ( 0)             |
| heart       | white zone               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 6)             | 1 ( 8)             |
| stomach     | forestomach:nodule       |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|             | glandular stomach:nodule |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|             | glandular stomach:thick  |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| small intes | nodule                   |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|             | adhesion                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| liver       | enlarged                 |                              | 9 ( 35)           | 7 ( 24)           | 2 ( 12)            | 0 ( 0)             |
|             | white zone               |                              | 7 ( 27)           | 8 ( 28)           | 3 ( 18)            | 2 ( 17)            |
|             | red zone                 |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 6)             | 0 ( 0)             |
|             | black zone               |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | nodule                   |                              | 4 ( 15)           | 6 ( 21)           | 2 ( 12)            | 0 ( 0)             |
|             | deformed                 |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|             | rough                    |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 8)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 5

| Organ      | Findings               | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>29 (%) | 1250 ppm<br>17 (%) | 2500 ppm<br>12 (%) |
|------------|------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| pancreas   | nodule                 |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 6)             | 0 ( 0)             |
| kidney     | enlarged               |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|            | small                  |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 6)             | 0 ( 0)             |
|            | pale                   |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 8)             |
|            | white zone             |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 1 ( 8)             |
|            | nodule                 |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|            | deformed               |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 6)             | 1 ( 8)             |
|            | hydronephrosis         |                              | 3 ( 12)           | 2 ( 7)            | 1 ( 6)             | 1 ( 8)             |
| ureter     | dilated                |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| urin bladd | nodule                 |                              | 1 ( 4)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
|            | urine:marked retention |                              | 1 ( 4)            | 2 ( 7)            | 0 ( 0)             | 0 ( 0)             |
| pituitary  | enlarged               |                              | 0 ( 0)            | 3 ( 10)           | 0 ( 0)             | 2 ( 17)            |
|            | red zone               |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 6)             | 0 ( 0)             |
| ovary      | enlarged               |                              | 6 ( 23)           | 7 ( 24)           | 2 ( 12)            | 0 ( 0)             |
|            | cyst                   |                              | 0 ( 0)            | 3 ( 10)           | 1 ( 6)             | 0 ( 0)             |
| uterus     | nodule                 |                              | 11 ( 42)          | 10 ( 34)          | 4 ( 24)            | 2 ( 17)            |
| vagina     | nodule                 |                              | 0 ( 0)            | 2 ( 7)            | 0 ( 0)             | 0 ( 0)             |
| brain      | red zone               |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 6)             | 1 ( 8)             |
|            | black zone             |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 6)             | 1 ( 8)             |
|            | deformed               |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| Harder gl  | nodule                 |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 8)             |
| muscle     | nodule                 |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 6)             | 1 ( 8)             |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 6

| Organ       | Findings      | Group Name<br>NO. of Animals | Control<br>26 (%) | 625 ppm<br>29 (%) | 1250 ppm<br>17 (%) | 2500 ppm<br>12 (%) |
|-------------|---------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| bone        | nodule        |                              | 0 ( 0)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| pleura      | nodule        |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| mediastinum | mass          |                              | 4 ( 15)           | 4 ( 14)           | 1 ( 6)             | 0 ( 0)             |
| peritoneum  | nodule        |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | adhesion      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 6)             | 0 ( 0)             |
|             | thick         |                              | 2 ( 8)            | 1 ( 3)            | 0 ( 0)             | 0 ( 0)             |
| retroperit  | mass          |                              | 0 ( 0)            | 1 ( 3)            | 1 ( 6)             | 1 ( 8)             |
| abdominal c | hemorrhage    |                              | 1 ( 4)            | 2 ( 7)            | 1 ( 6)             | 0 ( 0)             |
|             | ascites       |                              | 7 ( 27)           | 10 ( 34)          | 5 ( 29)            | 2 ( 17)            |
| thoracic ca | hemorrhage    |                              | 3 ( 12)           | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|             | pleural fluid |                              | 11 ( 42)          | 12 ( 41)          | 4 ( 24)            | 3 ( 25)            |
| other       | nose:nodule   |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |

(HPT080)

BAIS 5

**TABLE J 6**

**GROSS FINDINGS: FEMALE: SACRIFICED ANIMALS**



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 3

| Organ      | Findings                 | Group Name<br>NO. of Animals | Control<br>24 (%) | 625 ppm<br>21 (%) | 1250 ppm<br>33 (%) | 2500 ppm<br>38 (%) |
|------------|--------------------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| subcutis   | edema                    |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|            | mass                     |                              | 0 ( 0)            | 1 ( 5)            | 2 ( 6)             | 2 ( 5)             |
| lung       | nodule                   |                              | 0 ( 0)            | 5 ( 24)           | 2 ( 6)             | 1 ( 3)             |
| lymph node | enlarged                 |                              | 2 ( 8)            | 0 ( 0)            | 5 ( 15)            | 6 ( 16)            |
| spleen     | enlarged                 |                              | 4 ( 17)           | 2 ( 10)           | 4 ( 12)            | 3 ( 8)             |
|            | nodule                   |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| stomach    | forestomach:nodule       |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
|            | glandular stomach:nodule |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|            | glandular stomach:thick  |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| liver      | enlarged                 |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 3)             | 1 ( 3)             |
|            | dark                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 4 ( 11)            |
|            | white zone               |                              | 2 ( 8)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|            | red zone                 |                              | 2 ( 8)            | 0 ( 0)            | 3 ( 9)             | 4 ( 11)            |
|            | nodule                   |                              | 3 ( 13)           | 4 ( 19)           | 4 ( 12)            | 4 ( 11)            |
|            | cyst                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| pancreas   | nodule                   |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| kidney     | enlarged                 |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|            | white zone               |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|            | cyst                     |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 2 ( 5)             |
|            | deformed                 |                              | 3 ( 13)           | 7 ( 33)           | 6 ( 18)            | 10 ( 26)           |
|            | hydronephrosis           |                              | 1 ( 4)            | 2 ( 10)           | 5 ( 15)            | 4 ( 11)            |
| urin bladd | nodule                   |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
SACRIFICED ANIMALS (105W)

PAGE : 4

| Organ       | Findings      | Group Name<br>NO. of Animals | Control<br>24 (%) | 625 ppm<br>21 (%) | 1250 ppm<br>33 (%) | 2500 ppm<br>38 (%) |
|-------------|---------------|------------------------------|-------------------|-------------------|--------------------|--------------------|
| pituitary   | enlarged      |                              | 1 ( 4)            | 1 ( 5)            | 3 ( 9)             | 2 ( 5)             |
|             | red zone      |                              | 2 ( 8)            | 2 ( 10)           | 2 ( 6)             | 0 ( 0)             |
|             | nodule        |                              | 1 ( 4)            | 3 ( 14)           | 2 ( 6)             | 2 ( 5)             |
|             | cyst          |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 1 ( 3)             |
| thyroid     | dark          |                              | 0 ( 0)            | 0 ( 0)            | 0 ( 0)             | 5 ( 13)            |
| ovary       | enlarged      |                              | 1 ( 4)            | 1 ( 5)            | 2 ( 6)             | 0 ( 0)             |
|             | nodule        |                              | 0 ( 0)            | 1 ( 5)            | 0 ( 0)             | 0 ( 0)             |
|             | cyst          |                              | 3 ( 13)           | 4 ( 19)           | 7 ( 21)            | 7 ( 18)            |
| uterus      | nodule        |                              | 8 ( 33)           | 2 ( 10)           | 5 ( 15)            | 8 ( 21)            |
|             | dilated lumen |                              | 1 ( 4)            | 0 ( 0)            | 1 ( 3)             | 2 ( 5)             |
| eye         | turbid        |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 2 ( 5)             |
| Harder gl   | enlarged      |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 2 ( 5)             |
|             | nodule        |                              | 0 ( 0)            | 0 ( 0)            | 2 ( 6)             | 0 ( 0)             |
| peritoneum  | nodule        |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 0 ( 0)             |
|             | thick         |                              | 1 ( 4)            | 0 ( 0)            | 0 ( 0)             | 0 ( 0)             |
| abdominal c | ascites       |                              | 3 ( 13)           | 0 ( 0)            | 5 ( 15)            | 2 ( 5)             |
| thoracic ca | pleural fluid |                              | 3 ( 13)           | 0 ( 0)            | 5 ( 15)            | 3 ( 8)             |
| other       | tail:nodule   |                              | 0 ( 0)            | 0 ( 0)            | 1 ( 3)             | 1 ( 3)             |

TABLE K 1

ORGAN WEIGHT, ABSOLUTE: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJj[Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE  
 UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
 SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of<br>Animals | Body Weight | ADRENALS |       | TESTES  |       | HEART   |       | LUNGS   |       | KIDNEYS |       |
|------------|-------------------|-------------|----------|-------|---------|-------|---------|-------|---------|-------|---------|-------|
| Control    | 24                | 44.9 ± 7.9  | 0.010 ±  | 0.003 | 0.235 ± | 0.047 | 0.215 ± | 0.022 | 0.259 ± | 0.167 | 0.662 ± | 0.081 |
| 625 ppm    | 32                | 44.0 ± 6.2  | 0.010 ±  | 0.002 | 0.239 ± | 0.049 | 0.207 ± | 0.021 | 0.224 ± | 0.075 | 1.087 ± | 1.970 |
| 1250 ppm   | 31                | 42.3 ± 9.6  | 0.009 ±  | 0.002 | 0.248 ± | 0.074 | 0.216 ± | 0.027 | 0.208 ± | 0.043 | 0.696 ± | 0.252 |
| 2500 ppm   | 38                | 39.3 ± 7.8* | 0.010 ±  | 0.002 | 0.232 ± | 0.034 | 0.205 ± | 0.033 | 0.214 ± | 0.048 | 0.945 ± | 1.196 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE  
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of<br>Animals | SPLEEN |       | LIVER  |       | BRAIN  |       |
|------------|-------------------|--------|-------|--------|-------|--------|-------|
| Control    | 24                | 0.139± | 0.142 | 1.853± | 0.470 | 0.454± | 0.013 |
| 625 ppm    | 32                | 0.145± | 0.114 | 1.679± | 0.283 | 0.455± | 0.015 |
| 1250 ppm   | 31                | 0.147± | 0.231 | 1.777± | 0.442 | 0.454± | 0.013 |
| 2500 ppm   | 38                | 0.137± | 0.149 | 1.584± | 0.219 | 0.455± | 0.012 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS 5

TABLE K 2

ORGAN WEIGHT, ABSOLUTE: FEMALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | Body Weight  | ADRENALS |       | OVARIES |       | HEART   |       | LUNGS   |       | KIDNEYS |       |
|------------|----------------|--------------|----------|-------|---------|-------|---------|-------|---------|-------|---------|-------|
| Control    | 22             | 33.1 ± 6.3   | 0.014 ±  | 0.002 | 0.044 ± | 0.056 | 0.173 ± | 0.025 | 0.206 ± | 0.076 | 0.455 ± | 0.110 |
| 625 ppm    | 21             | 30.9 ± 5.7   | 0.013 ±  | 0.002 | 0.045 ± | 0.032 | 0.175 ± | 0.026 | 0.247 ± | 0.139 | 0.535 ± | 0.263 |
| 1250 ppm   | 30             | 30.9 ± 5.9   | 0.013 ±  | 0.002 | 0.044 ± | 0.033 | 0.171 ± | 0.022 | 0.210 ± | 0.041 | 0.504 ± | 0.190 |
| 2500 ppm   | 38             | 27.7 ± 3.9** | 0.013 ±  | 0.002 | 0.088 ± | 0.221 | 0.166 ± | 0.022 | 0.203 ± | 0.038 | 0.515 ± | 0.239 |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of<br>Animals | SPLEEN |       | LIVER  |       | BRAIN  |        |
|------------|-------------------|--------|-------|--------|-------|--------|--------|
| Control    | 22                | 0.214± | 0.150 | 1.534± | 0.427 | 0.476± | 0.018  |
| 625 ppm    | 21                | 0.217± | 0.160 | 1.449± | 0.199 | 0.481± | 0.017  |
| 1250 ppm   | 30                | 0.311± | 0.385 | 1.568± | 0.313 | 0.474± | 0.016  |
| 2500 ppm   | 38                | 0.340± | 0.573 | 1.550± | 0.612 | 0.463± | 0.016* |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS5



TABLE L 1

ORGAN WEIGHT, RELATIVE: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE  
 UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)  
 SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of<br>Animals | Body Weight<br>(g) | ADRENALS      | TESTES        | HEART         | LUNGS         | KIDNEYS        |
|------------|-------------------|--------------------|---------------|---------------|---------------|---------------|----------------|
| Control    | 24                | 44.9 ± 7.9         | 0.023 ± 0.006 | 0.534 ± 0.124 | 0.490 ± 0.080 | 0.604 ± 0.476 | 1.516 ± 0.323  |
| 625 ppm    | 32                | 44.0 ± 6.2         | 0.023 ± 0.006 | 0.553 ± 0.126 | 0.478 ± 0.062 | 0.529 ± 0.226 | 2.526 ± 4.711  |
| 1250 ppm   | 31                | 42.3 ± 9.6         | 0.024 ± 0.010 | 0.607 ± 0.184 | 0.537 ± 0.159 | 0.530 ± 0.220 | 1.707 ± 0.584  |
| 2500 ppm   | 38                | 39.3 ± 7.8*        | 0.026 ± 0.007 | 0.612 ± 0.146 | 0.533 ± 0.100 | 0.571 ± 0.187 | 2.587 ± 3.600* |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj[Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE  
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | SPLEEN       | LIVER        | BRAIN         |
|------------|----------------|--------------|--------------|---------------|
| Control    | 24             | 0.332± 0.386 | 4.318± 1.678 | 1.043± 0.189  |
| 625 ppm    | 32             | 0.341± 0.290 | 3.885± 0.864 | 1.056± 0.167  |
| 1250 ppm   | 31             | 0.353± 0.430 | 4.331± 1.161 | 1.136± 0.301  |
| 2500 ppm   | 38             | 0.358± 0.362 | 4.163± 1.012 | 1.204± 0.240* |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 5

TABLE L 2

ORGAN WEIGHT, RELATIVE: FEMALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj[Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of<br>Animals | Body Weight<br>(g) | ADRENALS       | OVARIES       | HEART         | LUNGS         | KIDNEYS         |
|------------|-------------------|--------------------|----------------|---------------|---------------|---------------|-----------------|
| Control    | 22                | 33.1 ± 6.3         | 0.042 ± 0.008  | 0.135 ± 0.164 | 0.537 ± 0.105 | 0.650 ± 0.318 | 1.399 ± 0.343   |
| 625 ppm    | 21                | 30.9 ± 5.7         | 0.042 ± 0.008  | 0.150 ± 0.105 | 0.582 ± 0.129 | 0.837 ± 0.515 | 1.841 ± 1.190   |
| 1250 ppm   | 30                | 30.9 ± 5.9         | 0.043 ± 0.008  | 0.150 ± 0.115 | 0.570 ± 0.113 | 0.708 ± 0.202 | 1.715 ± 0.890   |
| 2500 ppm   | 38                | 27.7 ± 3.9**       | 0.047 ± 0.009* | 0.300 ± 0.678 | 0.608 ± 0.087 | 0.757 ± 0.213 | 1.903 ± 0.971** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ[Crl:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | SPLEEN       | LIVER        | BRAIN          |
|------------|----------------|--------------|--------------|----------------|
| Control    | 22             | 0.673± 0.523 | 4.684± 1.046 | 1.484± 0.260   |
| 625 ppm    | 21             | 0.713± 0.584 | 4.772± 0.735 | 1.606± 0.302   |
| 1250 ppm   | 30             | 0.971± 1.038 | 5.201± 1.219 | 1.587± 0.290   |
| 2500 ppm   | 38             | 1.257± 2.205 | 5.600± 1.938 | 1.701± 0.224** |

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 5

TABLE M 1

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

MALE: ALL ANIMALS

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 1

| Organ                            | Findings                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |           |           |           | 625 ppm<br>50 |           |           |           | 1250 ppm<br>50 |           |           |           | 2500 ppm<br>50 |           |           |           |
|----------------------------------|---------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                                  |                           |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Integumentary system/appandage) |                           |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| skin/app                         | ulcer                     |  | <50>          |           |           |           | <50>          |           |           |           | <50>           |           |           |           | <50>           |           |           |           |
|                                  |                           | 1<br>( 2)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 2<br>( 4)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                                  | erosion                   |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) |
|                                  |                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | inflammation              |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  |                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| subcutis                         | squamous cell hyperplasia |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                                  |                           |  | 3<br>( 6)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                                  | epidermal cyst            |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  |                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | hematoma                  |  | <50>          |           |           |           | <50>          |           |           |           | <50>           |           |           |           | <50>           |           |           |           |
|                                  |                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| subcutis                         | inflammation              |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                                  |                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
< a > a : Number of animals examined at the site  
b : Number of animals with lesion  
( c ) c : b / a \* 100  
Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 2

| Organ                            | Findings                                    | Group Name<br>No. of Animals on Study<br>Grade |       |       |       | Control<br>50 |       |       |       | 625 ppm<br>50 |       |       |       | 1250 ppm<br>50 |       |       |       | 2500 ppm<br>50 |       |       |       |
|----------------------------------|---|--|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|----------------|-------|-------|-------|----------------|-------|-------|-------|
|                                  |   | 1+   | 2+    | 3+    | 4+    | 1+            | 2+    | 3+    | 4+    | 1+            | 2+    | 3+    | 4+    | 1+             | 2+    | 3+    | 4+    | 1+             | 2+    | 3+    | 4+    |
|                                  |   | (%)  | (%)   | (%)   | (%)   | (%)           | (%)   | (%)   | (%)   | (%)           | (%)   | (%)   | (%)   | (%)            | (%)   | (%)   | (%)   | (%)            | (%)   | (%)   | (%)   |
| (Integumentary system/appandage) |   |  |       |       |       |               |       |       |       |               |       |       |       |                |       |       |       |                |       |       |       |
| subcutis                         |   | <50>   |       |       |       | <50>          |       |       |       | <50>          |       |       |       | <50>           |       |       |       | <50>           |       |       |       |
|                                  | xanthogranuloma                             | 0  | 0     | 0     | 0     | 0             | 1     | 0     | 0     | 1             | 0     | 0     | 0     | 0              | 0     | 0     | 0     | 0              | 0     | 0     | 0     |
|                                  |   | ( 0 )  | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )         | ( 2 ) | ( 0 ) | ( 0 ) | ( 2 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) |
| (Respiratory system)             |   |  |       |       |       |               |       |       |       |               |       |       |       |                |       |       |       |                |       |       |       |
| nasal cavit                      |   | <50>   |       |       |       | <50>          |       |       |       | <50>          |       |       |       | <50>           |       |       |       | <50>           |       |       |       |
|                                  | eosinophilic change:olfactory epithelium    | 16   | 0     | 0     | 0     | 13            | 0     | 0     | 0     | 13            | 0     | 0     | 0     | 14             | 0     | 0     | 0     | 14             | 0     | 0     | 0     |
|                                  |   | ( 32 )   | ( 0 ) | ( 0 ) | ( 0 ) | ( 26 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 26 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 28 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 28 )         | ( 0 ) | ( 0 ) | ( 0 ) |
|                                  | eosinophilic change:respiratory epithelium  | 23   | 0     | 0     | 0     | 21            | 0     | 0     | 0     | 19            | 0     | 0     | 0     | 18             | 0     | 0     | 0     | 18             | 0     | 0     | 0     |
|                                  |   | ( 46 )   | ( 0 ) | ( 0 ) | ( 0 ) | ( 42 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 38 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 36 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 36 )         | ( 0 ) | ( 0 ) | ( 0 ) |
|                                  | inflammation:foreign body                   | 1  | 0     | 0     | 0     | 0             | 0     | 0     | 0     | 0             | 0     | 0     | 0     | 0              | 0     | 0     | 0     | 0              | 0     | 0     | 0     |
|                                  |   | ( 2 )  | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) |
|                                  | respiratory metaplasia:olfactory epithelium | 9  | 1     | 0     | 0     | 5             | 0     | 0     | 0     | 7             | 0     | 0     | 0     | 5              | 0     | 0     | 0     | 5              | 0     | 0     | 0     |
|                                  |   | ( 18 )   | ( 2 ) | ( 0 ) | ( 0 ) | ( 10 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 14 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 10 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 10 )         | ( 0 ) | ( 0 ) | ( 0 ) |
|                                  | respiratory metaplasia:gland                | 9  | 1     | 0     | 0     | 8             | 0     | 0     | 0     | 7             | 1     | 0     | 0     | 10             | 0     | 0     | 0     | 10             | 0     | 0     | 0     |
|                                  |   | ( 18 )   | ( 2 ) | ( 0 ) | ( 0 ) | ( 16 )        | ( 0 ) | ( 0 ) | ( 0 ) | ( 14 )        | ( 2 ) | ( 0 ) | ( 0 ) | ( 20 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 20 )         | ( 0 ) | ( 0 ) | ( 0 ) |
| nasopharynx                      |   | <50>   |       |       |       | <50>          |       |       |       | <50>          |       |       |       | <50>           |       |       |       | <50>           |       |       |       |
|                                  | eosinophilic change                         | 0  | 0     | 0     | 0     | 1             | 0     | 0     | 0     | 0             | 0     | 0     | 0     | 0              | 0     | 0     | 0     | 0              | 0     | 0     | 0     |
|                                  |   | ( 0 )  | ( 0 ) | ( 0 ) | ( 0 ) | ( 2 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )         | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )          | ( 0 ) | ( 0 ) | ( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 3

| Organ_____           | Findings_____                            | Group Name              | Control    |            |            |            | 625 ppm     |            |            |            | 1250 ppm   |            |            |            | 2500 ppm    |            |            |            |            |
|----------------------|--|-------------------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|
|                      |  | No. of Animals on Study | 50         |            |            |            | 50          |            |            |            | 50         |            |            |            | 50          |            |            |            |            |
|                      |  | Grade                   | 1+         | 2+         | 3+         | 4+         | 1+          | 2+         | 3+         | 4+         | 1+         | 2+         | 3+         | 4+         | 1+          | 2+         | 3+         | 4+         |            |
|                      |  |                         | (%)        | (%)        | (%)        | (%)        | (%)         | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)         | (%)        | (%)        | (%)        |            |
|                      |  |                         |            |            |            |            |             |            |            |            |            |            |            |            |             |            |            |            |            |
| (Respiratory system) |  |                         |            |            |            |            |             |            |            |            |            |            |            |            |             |            |            |            |            |
| lung                 |  |                         | <50>       |            |            |            | <50>        |            |            |            | <50>       |            |            |            | <50>        |            |            |            |            |
|                      | congestion                               |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                      | hemorrhage                               |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                      | deposit of amyloid                       |                         | 4<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 6<br>( 12 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 16 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                      | inflammatory infiltration                |                         | 2<br>( 4 ) | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                      | bronchiolar-alveolar cell hyperplasia    |                         | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                      | eosinophilic change:bronchial epithelium |                         | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

{Hematopoietic system}

|             |            |  |            |            |            |            |            |            |            |            |            |            |            |            |            |            |            |            |
|-------------|------------|--|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| bone marrow | congestion |  | <50>       |            |            |            | <50>       |            |            |            | <50>       |            |            |            | <50>       |            |            |            |
|             |            |  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe

< a > a : Number of animals examined at the site

b b : Number of animals with lesion

( c ) c : b / a \* 100

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 4

| Organ_____             | Findings_____            | Group Name              | Control   |           |           |             | 625 ppm   |           |           |             | 1250 ppm  |           |              |             | 2500 ppm    |           |              |  |
|------------------------|--------------------------|-------------------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|--------------|-------------|-------------|-----------|--------------|--|
|                        |                          | No. of Animals on Study | 50        |           |           |             | 50        |           |           |             | 50        |           |              |             | 50          |           |              |  |
| Grade                  |                          | 1+                      | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+           | 1+          | 2+          | 3+        | 4+           |  |
|                        |                          | (%)                     | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)          | (%)         | (%)         | (%)       | (%)          |  |
| (Hematopoietic system) |                          |                         |           |           |           |             |           |           |           |             |           |           |              |             |             |           |              |  |
| bone marrow            |                          | <50>                    |           |           |           | <50>        |           |           |           | <50>        |           |           |              | <50>        |             |           |              |  |
|                        | increased hematopoiesis  | 12<br>( 24)             | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 11<br>( 22) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 10<br>( 20) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 9<br>( 18)  | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |
|                        | granulopoiesis:increased | 2<br>( 4)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 12)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 2<br>( 4)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |
| lymph node             |                          | <50>                    |           |           |           | <50>        |           |           |           | <50>        |           |           |              | <50>        |             |           |              |  |
|                        | lymphadenitis            | 1<br>( 2)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |
| thymus                 |                          | <50>                    |           |           |           | <50>        |           |           |           | <50>        |           |           |              | <50>        |             |           |              |  |
|                        | atrophy                  | 0<br>( 0)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |
| spleen                 |                          | <50>                    |           |           |           | <50>        |           |           |           | <50>        |           |           |              | <50>        |             |           |              |  |
|                        | deposit of hemosiderin   | 20<br>( 40)             | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 26<br>( 52) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 30<br>( 60) | 4<br>( 8) | 0<br>( 0) | 0<br>( 0) ** | 32<br>( 64) | 14<br>( 28) | 2<br>( 4) | 0<br>( 0) ** |  |
|                        | deposit of melanin       | 0<br>( 0)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |
|                        | fibrosis:focal           | 0<br>( 0)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)    |  |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 5

| Organ                  | Findings                     | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |             |           |           | 625 ppm<br>50 |            |           |           | 1250 ppm<br>50 |             |           |           | 2500 ppm<br>50 |             |           |           |
|------------------------|------------------------------|--|---------------|-------------|-----------|-----------|---------------|------------|-----------|-----------|----------------|-------------|-----------|-----------|----------------|-------------|-----------|-----------|
|                        |                              |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%) |
| (Hematopoietic system) |                              |  |               |             |           |           |               |            |           |           |                |             |           |           |                |             |           |           |
| spleen                 |                              |  | <50>          |             |           |           | <50>          |            |           |           | <50>           |             |           |           | <50>           |             |           |           |
|                        | extramedullary hematopoiesis |  | 11<br>( 22)   | 11<br>( 22) | 0<br>( 0) | 0<br>( 0) | 14<br>( 28)   | 8<br>( 16) | 0<br>( 0) | 0<br>( 0) | 13<br>( 26)    | 10<br>( 20) | 1<br>( 2) | 0<br>( 0) | 20<br>( 40)    | 12<br>( 24) | 1<br>( 2) | 0<br>( 0) |
|                        | follicular hyperplasia       |  | 2<br>( 4)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 3<br>( 6)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |
| (Circulatory system)   |                              |  |               |             |           |           |               |            |           |           |                |             |           |           |                |             |           |           |
| heart                  |                              |  | <50>          |             |           |           | <50>          |            |           |           | <50>           |             |           |           | <50>           |             |           |           |
|                        | deposit of amyloid           |  | 2<br>( 4)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 7<br>( 14)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |
|                        | mineralization               |  | 2<br>( 4)     | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |
|                        | myocardial fibrosis          |  | 17<br>( 34)   | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 10<br>( 20)   | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 19<br>( 38)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 13<br>( 26)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |
|                        | arteritis                    |  | 1<br>( 2)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) |
| artery/aort            |                              |  | <50>          |             |           |           | <50>          |            |           |           | <50>           |             |           |           | <50>           |             |           |           |
|                        | mineralization               |  | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 6

| Organ              | Findings                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|--------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                    |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Digestive system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| oral cavity        | hematoma                  |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                    |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| tooth              | dysplasia                 |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                    |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| tongue             | squamous cell hyperplasia |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                    |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | epidermal cyst            |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    |                           | arteritis                                      | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| stomach            | ulcer:forestomach         |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                    |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | hyperplasia:forestomach   | 0<br>( 0 )                                     | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 7

| Organ              | Findings                      | Group Name              | Control     |           |           |           | 625 ppm     |           |           |           | 1250 ppm   |           |           |           | 2500 ppm    |           |           |           |
|--------------------|-------------------------------|-------------------------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|
|                    |                               | No. of Animals on Study | 50          |           |           |           | 50          |           |           |           | 50         |           |           |           | 50          |           |           |           |
|                    |                               | Grade                   | 1+          | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        | 1+         | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        |
|                    |                               |                         | (%)         | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       | (%)        | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       |
| (Digestive system) |                               |                         |             |           |           |           |             |           |           |           |            |           |           |           |             |           |           |           |
| stomach            | erosion:glandular stomach     |                         | <50>        |           |           |           | <50>        |           |           |           | <50>       |           |           |           | <50>        |           |           |           |
|                    |                               | 1<br>( 2)               | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 3<br>( 6) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 4<br>( 8) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |           |
|                    | hyperplasia:glandular stomach |                         | 12<br>( 24) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 12<br>( 24) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 9<br>( 18) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 14<br>( 28) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| small intes        | ulcer                         |                         | <50>        |           |           |           | <50>        |           |           |           | <50>       |           |           |           | <50>        |           |           |           |
|                    |                               | 0<br>( 0)               | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) |           |
|                    | hemorrhage                    |                         | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| liver              | angiectasis                   |                         | <50>        |           |           |           | <50>        |           |           |           | <50>       |           |           |           | <50>        |           |           |           |
|                    |                               | 1<br>( 2)               | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) |           |
|                    | necrosis:central              |                         | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | necrosis:focal                |                         | 2<br>( 4)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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 Significant difference : \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 8

| Organ_____         | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|--------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                    |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Digestive system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| liver              | necrosis:single cell      |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                    |                           |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                    | collapse                  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                    | inflammatory infiltration |  | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                    | inflammatory cell nest    |  | 11<br>( 22 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 9<br>( 18 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 16 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 13<br>( 26 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | clear cell focus          |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | acidophilic cell focus    |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | basophilic cell focus     |  | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 6<br>( 12 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| mixed cell focus   |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe

< a > a : Number of animals examined at the site

b b : Number of animals with lesion

( c ) c : b / a \* 100

Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

(HPT150)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j[Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 9

| Organ              | Findings                        | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |           |           |           | 625 ppm<br>50 |           |           |           | 1250 ppm<br>50 |           |           |           | 2500 ppm<br>50 |           |           |              |
|--------------------|---------------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|--------------|
|                    |                                 |  | 1+            | 2+        | 3+        | 4+        | 1+            | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+           |
|                    |                                 |  | (%)           | (%)       | (%)       | (%)       | (%)           | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)          |
| (Digestive system) |                                 |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |              |
| liver              | biliary cyst                    |  | 2<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    |
|                    | intestinal metaplasia:bile duct |  | 0<br>( 0)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    |
|                    | deposit of brown pigment        |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 10<br>( 20)    | 0<br>( 0) | 0<br>( 0) | 0 **<br>( 0) |
| pancreas           | cyst                            |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    |
| (Urinary system)   |                                 |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |              |
| kidney             | atrophy                         |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 2<br>( 4) | 0<br>( 0) | 0<br>( 0)    |
|                    | cyst                            |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)    |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

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| Organ_____       | Findings_____             | Group Name              | Control   |            |           |           | 1250 ppm  |            |           |           | 2500 ppm   |           |           |           |            |           |           |           |
|------------------|---------------------------|-------------------------|-----------|------------|-----------|-----------|-----------|------------|-----------|-----------|------------|-----------|-----------|-----------|------------|-----------|-----------|-----------|
|                  |                           | No. of Animals on Study | 50        |            |           |           | 50        |            |           |           | 50         |           |           |           |            |           |           |           |
|                  |                           | Grade                   | 1+        | 2+         | 3+        | 4+        | 1+        | 2+         | 3+        | 4+        | 1+         | 2+        | 3+        | 4+        | 1+         | 2+        | 3+        | 4+        |
|                  |                           |                         | (%)       | (%)        | (%)       | (%)       | (%)       | (%)        | (%)       | (%)       | (%)        | (%)       | (%)       | (%)       | (%)        | (%)       | (%)       | (%)       |
| <hr/>            |                           |                         |           |            |           |           |           |            |           |           |            |           |           |           |            |           |           |           |
| (Urinary system) |                           |                         |           |            |           |           |           |            |           |           |            |           |           |           |            |           |           |           |
| kidney           |                           |                         | <50>      |            |           |           | <50>      |            |           |           | <50>       |           |           |           | <50>       |           |           |           |
|                  | hyaline droplet           |                         | 1<br>( 2) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 3<br>( 6) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 5<br>( 10) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                  | deposit of amyloid        |                         | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                  | hyaline cast              |                         | 1<br>( 2) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                  | inflammatory infiltration |                         | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                  | lymphocytic infiltration  |                         | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 2<br>( 4) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                  | scar                      |                         | 1<br>( 2) | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 2<br>( 4) | 3<br>( 6)  | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 3<br>( 6) | 0<br>( 0) | 0<br>( 0) |
|                  | inflammatory polyp        |                         | 0<br>( 0) | 2<br>( 4)  | 4<br>( 8) | 0<br>( 0) | 1<br>( 2) | 3<br>( 6)  | 2<br>( 4) | 0<br>( 0) | 1<br>( 2)  | 4<br>( 8) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)  | 1<br>( 2) | 1<br>( 2) | 0<br>( 0) |
| hydronephrosis   |                           | 1<br>( 2)               | 2<br>( 4) | 6<br>( 12) | 1<br>( 2) | 0<br>( 0) | 3<br>( 6) | 8<br>( 16) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)  | 3<br>( 6) | 0<br>( 0) | 0<br>( 0) | 5<br>( 10) | 2<br>( 4) | 0<br>( 0) |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 11

| Organ            | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |             |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|------------------|-------------------------------|--|---------------|------------|------------|------------|---------------|-------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                  |                               |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system) |                               |  |               |            |            |            |               |             |            |            |                |            |            |            |                |            |            |            |
| kidney           |                               |  | <50>          |            |            |            | <50>          |             |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                  | pyelonephritis                |  | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )  | 1<br>( 2 ) | 0<br>( 0 ) | 1<br>( 2 )     | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | mineralization:papilla        |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | mineralization:cortex         |  | 2<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | regeneration:proximal tubule  |  | 19<br>( 38 )  | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 17<br>( 34 )  | 2<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 ) | 17<br>( 34 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 21<br>( 42 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | urothelial hyperplasia:pelvis |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
| urin bladd       |                               |  | <50>          |            |            |            | <50>          |             |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                  | dilatation                    |  | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 7<br>( 14 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 4<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | ulcer                         |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | simple tubule hyperplasia     |  | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 12

| Organ                 | Findings                 | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |               |
|-----------------------|--------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|---------------|
|                       |                          |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)     |
| (Urinary system)      |                          |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |               |
| urin bladd            | xanthogranuloma          |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |               |
|                       |                          | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |               |
| (Endocrine system)    |                          |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |               |
| pituitary             | Rathke pouch             |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <49>           |            |            |               |
|                       |                          | 2<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |               |
| thyroid               | cystic thyroid follicle  |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |               |
|                       |                          | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |               |
|                       | deposit of brown pigment |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 10<br>( 20 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0 **<br>( 0 ) |
| adrenal               | hyperplasia:medulla      |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |               |
|                       |                          | 0<br>( 0 )                                     | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |               |
| (Reproductive system) |                          |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |               |
| testis                | mineralization           |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |               |
|                       |                          | 2<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) |               |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 13

| Organ  | Findings                | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|--|-------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|  |                         |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Reproductive system)  |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| testis   | xanthogranuloma         |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|  |                         | 0<br>( 0 )                                     | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
| epididymis   | spermatogenic granuloma |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|  |                         | 4<br>( 8 )                                     | 2<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
| prostate   | inflammation            |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|  |                         | 0<br>( 0 )                                     | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
| prep/cli gl  | duct ectasia            |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|  |                         | 1<br>( 2 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
| (Nervous system)   |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| brain  | hemorrhage              |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|  |                         | 1<br>( 2 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|  | mineralization          |  | 18<br>( 36 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 16<br>( 32 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe                |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| < a > a : Number of animals examined at the site                       |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| b : Number of animals with lesion                                      |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| ( c ) c : b / a * 100  |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |                         |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 14

| Organ                            | Findings                 | Group Name              | Control   |           |           |           | 625 ppm   |           |           |           | 1250 ppm  |           |           |           | 2500 ppm  |           |           |           |
|----------------------------------|--------------------------|-------------------------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|
|                                  |                          | No. of Animals on Study | 50        |           |           |           | 50        |           |           |           | 50        |           |           |           | 50        |           |           |           |
|                                  |                          | Grade                   | 1+        | 2+        | 3+        | 4+        | 1+        | 2+        | 3+        | 4+        | 1+        | 2+        | 3+        | 4+        | 1+        | 2+        | 3+        | 4+        |
|                                  |                          |                         | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       | (%)       |
| (Special sense organs/appendage) |                          |                         |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |
| eye                              | cataract                 |                         | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           |
|                                  |                          | 2<br>( 4)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
|                                  | keratitis                |                         | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| Harder gl                        | lymphocytic infiltration |                         | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           |
|                                  |                          | 0<br>( 0)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
|                                  | hyperplasia              |                         | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Musculoskeletal system)         |                          |                         |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |
| muscle                           | mineralization           |                         | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           |
|                                  |                          | 1<br>( 2)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
| bone                             | osteosclerosis           |                         | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           |
|                                  |                          | 1<br>( 2)               | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

TABLE M 2

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

MALE: DEAD AND MORIBUND ANIMALS

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 1

| Organ                            | Findings                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |           |           |           | 625 ppm<br>18 |            |           |           | 1250 ppm<br>19 |            |           |           | 2500 ppm<br>11 |           |           |           |
|----------------------------------|---------------------------|--|---------------|-----------|-----------|-----------|---------------|------------|-----------|-----------|----------------|------------|-----------|-----------|----------------|-----------|-----------|-----------|
|                                  |                           |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| {Integumentary system/appandage} |                           |  |               |           |           |           |               |            |           |           |                |            |           |           |                |           |           |           |
| skin/app                         |                           |  | <26>          |           |           |           | <18>          |            |           |           | <19>           |            |           |           | <11>           |           |           |           |
|                                  | ulcer                     |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 2<br>( 11) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 2<br>( 11) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | erosion                   |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)     | 1<br>( 6)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 2<br>( 11) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 9) | 0<br>( 0) | 0<br>( 0) |
|                                  | inflammation              |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 5)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | squamous cell hyperplasia |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | scab                      |  | 3<br>( 12)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | epidermal cyst            |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 2<br>( 11) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| subcutis                         |                           |  | <26>          |           |           |           | <18>          |            |           |           | <19>           |            |           |           | <11>           |           |           |           |
|                                  | hematoma                  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 5)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | xanthogranuloma           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 6)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 2

| Organ                  | Findings                                    | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |           |           |           | 625 ppm<br>18 |           |           |           | 1250 ppm<br>19 |           |           |           | 2500 ppm<br>11 |           |           |           |
|------------------------|---|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                        |   |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Respiratory system)   |   |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| nasal cavit            |   |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |
|                        | eosinophilic change:olfactory epithelium    |  | 7<br>( 27)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 22)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 26)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 9)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | eosinophilic change:respiratory epithelium  |  | 10<br>( 38)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 7<br>( 39)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 32)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 18)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | inflammation:foreign body                   |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | respiratory metaplasia:olfactory epithelium |  | 6<br>( 23)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 16)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | respiratory metaplasia:gland                |  | 4<br>( 15)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 11)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 11)     | 1<br>( 5) | 0<br>( 0) | 0<br>( 0) | 2<br>( 18)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| lung                   | congestion                                  |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |
|                        |   |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 6) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | inflammatory infiltration                   |  | 1<br>( 4)     | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 9)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Hematopoietic system) |   |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| bone marrow            |   |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |
|                        | congestion                                  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 6) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 { c } c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 3

| Organ                  | Findings                     | Group Name<br>No. of Animals on Study<br>Grade |       |      |      | Control<br>26 |       |      |      | 625 ppm<br>18 |       |      |      | 1250 ppm<br>19 |       |      |      | 2500 ppm<br>11 |       |      |      |
|------------------------|------------------------------|--|-------|------|------|---------------|-------|------|------|---------------|-------|------|------|----------------|-------|------|------|----------------|-------|------|------|
|                        |                              | 1+   | 2+    | 3+   | 4+   | 1+            | 2+    | 3+   | 4+   | 1+            | 2+    | 3+   | 4+   | 1+             | 2+    | 3+   | 4+   | 1+             | 2+    | 3+   | 4+   |
|                        |                              | (%)  | (%)   | (%)  | (%)  | (%)           | (%)   | (%)  | (%)  | (%)           | (%)   | (%)  | (%)  | (%)            | (%)   | (%)  | (%)  | (%)            | (%)   | (%)  | (%)  |
| (Hematopoietic system) |                              |  |       |      |      |               |       |      |      |               |       |      |      |                |       |      |      |                |       |      |      |
| bone marrow            | increased hematopoiesis      | <26>   |       |      |      | 12            | 0     | 0    | 0    | <18>          |       |      |      | <19>           |       |      |      | <11>           |       |      |      |
|                        |                              | ( 46)  | ( 0)  | ( 0) | ( 0) | ( 61)         | ( 0)  | ( 0) | ( 0) | ( 37)         | ( 0)  | ( 0) | ( 0) | ( 73)          | ( 0)  | ( 0) | ( 0) | ( 73)          | ( 0)  | ( 0) | ( 0) |
|                        | granulopoiesis:increased     | 1  | 0     | 0    | 0    | 4             | 0     | 0    | 0    | 3             | 0     | 0    | 0    | 1              | 0     | 0    | 0    | 1              | 0     | 0    | 0    |
|                        |                              | ( 4)   | ( 0)  | ( 0) | ( 0) | ( 22)         | ( 0)  | ( 0) | ( 0) | ( 16)         | ( 0)  | ( 0) | ( 0) | ( 9)           | ( 0)  | ( 0) | ( 0) | ( 9)           | ( 0)  | ( 0) | ( 0) |
| lymph node             | lymphadenitis                | <26>   |       |      |      | 1             | 0     | 0    | 0    | <18>          |       |      |      | <19>           |       |      |      | <11>           |       |      |      |
|                        |                              | ( 4)   | ( 0)  | ( 0) | ( 0) | ( 0)          | ( 0)  | ( 6) | ( 0) | ( 0)          | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) |
| thymus                 | atrophy                      | <26>   |       |      |      | 0             | 0     | 0    | 0    | <18>          |       |      |      | <19>           |       |      |      | <11>           |       |      |      |
|                        |                              | ( 0)   | ( 0)  | ( 0) | ( 0) | ( 0)          | ( 0)  | ( 6) | ( 0) | ( 0)          | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) |
| spleen                 | deposit of hemosiderin       | <26>   |       |      |      | 11            | 0     | 0    | 0    | <18>          |       |      |      | <19>           |       |      |      | <11>           |       |      |      |
|                        |                              | ( 42)  | ( 0)  | ( 0) | ( 0) | ( 50)         | ( 0)  | ( 0) | ( 0) | ( 47)         | ( 0)  | ( 0) | ( 0) | ( 73)          | ( 18) | ( 9) | ( 0) | ( 73)          | ( 18) | ( 9) | ( 0) |
|                        | fibrosis:focal               | 0  | 0     | 0    | 0    | 0             | 0     | 0    | 0    | 1             | 0     | 0    | 0    | 0              | 0     | 0    | 0    | 0              | 0     | 0    | 0    |
|                        |                              | ( 0)   | ( 0)  | ( 0) | ( 0) | ( 0)          | ( 0)  | ( 0) | ( 0) | ( 5)          | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) | ( 0)           | ( 0)  | ( 0) | ( 0) |
|                        | extramedullary hematopoiesis | 4  | 10    | 0    | 0    | 4             | 6     | 0    | 0    | 0             | 6     | 1    | 0    | 2              | 3     | 0    | 0    | 2              | 3     | 0    | 0    |
|                        |                              | ( 15)  | ( 38) | ( 0) | ( 0) | ( 22)         | ( 33) | ( 0) | ( 0) | ( 0)          | ( 32) | ( 5) | ( 0) | ( 18)          | ( 27) | ( 0) | ( 0) | ( 18)          | ( 27) | ( 0) | ( 0) |

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 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 4

| Organ_____           | Findings_____       | Group Name              | Control    |           |           |           | 625 ppm    |           |           |           | 1250 ppm    |           |           |           | 2500 ppm  |            |           |           |           |
|----------------------|---------------------|-------------------------|------------|-----------|-----------|-----------|------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-----------|------------|-----------|-----------|-----------|
|                      |                     | No. of Animals on Study | 26         |           |           |           | 18         |           |           |           | 19          |           |           |           | 11        |            |           |           |           |
|                      |                     | Grade                   | 1+         | 2+        | 3+        | 4+        | 1+         | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        | 1+        | 2+         | 3+        | 4+        |           |
|                      |                     |                         | (%)        | (%)       | (%)       | (%)       | (%)        | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       | (%)       | (%)        | (%)       | (%)       |           |
| (Circulatory system) |                     |                         |            |           |           |           |            |           |           |           |             |           |           |           |           |            |           |           |           |
| heart                |                     |                         | <26>       |           |           |           | <18>       |           |           |           | <19>        |           |           |           | <11>      |            |           |           |           |
|                      | deposit of amyloid  |                         | 1<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |           |
|                      | mineralization      |                         | 2<br>( 8)  | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |           |
|                      | myocardial fibrosis |                         | 6<br>( 23) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 28) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 11<br>( 58) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0 *       | 3<br>( 27) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | arteritis           |                         | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 9) | 0<br>( 0) | 0<br>( 0) |
| artery/aort          |                     |                         | <26>       |           |           |           | <18>       |           |           |           | <19>        |           |           |           | <11>      |            |           |           |           |
|                      | mineralization      |                         | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |           |
| (Digestive system)   |                     |                         |            |           |           |           |            |           |           |           |             |           |           |           |           |            |           |           |           |
| oral cavity          |                     |                         | <26>       |           |           |           | <18>       |           |           |           | <19>        |           |           |           | <11>      |            |           |           |           |
|                      | hematoma            |                         | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 6) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

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 ( c ) c : b / a \* 100  
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(HPT150)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ_____         | Findings_____                 | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |           |           |           | 625 ppm<br>18 |           |           |           | 1250 ppm<br>19 |           |           |           | 2500 ppm<br>11 |           |           |           |           |
|--------------------|-------------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|-----------|
|                    |                               |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |           |
| (Digestive system) |                               |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |           |
| tooth              | dysplasia                     |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |           |
|                    |                               | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 6) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
| tongue             | epidermal cyst                |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |           |
|                    |                               | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 6) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
|                    | arteritis                     |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| stomach            | ulcer:forestomach             |  | <26>          |           |           |           | <18>          |           |           |           | <19>           |           |           |           | <11>           |           |           |           |           |
|                    |                               | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 5) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |           |
|                    | hyperplasia:forestomach       |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | erosion:glandular stomach     |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | hyperplasia:glandular stomach |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 11)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ              | Findings                  | Group Name              | Control |      |      |      | 625 ppm |      |      |      | 1250 ppm |      |      |      | 2500 ppm |      |      |      |      |
|--------------------|---------------------------|-------------------------|---------|------|------|------|---------|------|------|------|----------|------|------|------|----------|------|------|------|------|
|                    |                           | No. of Animals on Study | 26      |      |      |      | 18      |      |      |      | 19       |      |      |      | 11       |      |      |      |      |
|                    |                           | Grade                   | 1+      | 2+   | 3+   | 4+   | 1+      | 2+   | 3+   | 4+   | 1+       | 2+   | 3+   | 4+   | 1+       | 2+   | 3+   | 4+   |      |
|                    |                           |                         | (%)     | (%)  | (%)  | (%)  | (%)     | (%)  | (%)  | (%)  | (%)      | (%)  | (%)  | (%)  | (%)      | (%)  | (%)  | (%)  |      |
| (Digestive system) |                           |                         |         |      |      |      |         |      |      |      |          |      |      |      |          |      |      |      |      |
| small intes        | hemorrhage                |                         | <26>    |      |      |      | <18>    |      |      |      | <19>     |      |      |      | <11>     |      |      |      |      |
|                    |                           | 0                       | 0       | 0    | 0    | 0    | 0       | 0    | 0    | 0    | 1        | 0    | 0    | 0    | 0        | 0    | 0    | 0    |      |
|                    |                           |                         | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)    | ( 0) | ( 0) | ( 0) | ( 5)     | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) |      |
| liver              | necrosis:focal            |                         | <26>    |      |      |      | <18>    |      |      |      | <19>     |      |      |      | <11>     |      |      |      |      |
|                    |                           | 0                       | 0       | 0    | 0    | 1    | 1       | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0        | 0    | 0    | 0    |      |
|                    |                           | ( 0)                    | ( 0)    | ( 0) | ( 0) | ( 6) | ( 6)    | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) |      |
|                    | inflammatory infiltration |                         | 0       | 1    | 0    | 0    | 0       | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0    |
|                    |                           |                         | ( 0)    | ( 4) | ( 0) | ( 0) | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 0) |
|                    | inflammatory cell nest    |                         | 2       | 0    | 0    | 0    | 0       | 0    | 0    | 0    | 1        | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0    |
|                    |                           | ( 8)                    | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)    | ( 0) | ( 0) | ( 5) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) |      |
|                    | basophilic cell focus     |                         | 0       | 0    | 0    | 0    | 1       | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0        | 0    | 0    | 0    |      |
|                    |                           |                         | ( 0)    | ( 0) | ( 0) | ( 0) | ( 6)    | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) |      |
|                    | deposit of brown pigment  |                         | 0       | 0    | 0    | 0    | 0       | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 2        | 0    | 0    | 0    |      |
|                    |                           |                         | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 18)    | ( 0) | ( 0) | ( 0) |      |
| (Urinary system)   |                           |                         |         |      |      |      |         |      |      |      |          |      |      |      |          |      |      |      |      |
| kidney             | atrophy                   |                         | <26>    |      |      |      | <18>    |      |      |      | <19>     |      |      |      | <11>     |      |      |      |      |
|                    |                           | 0                       | 0       | 0    | 0    | 0    | 0       | 0    | 0    | 0    | 0        | 0    | 0    | 0    | 0        | 1    | 0    | 0    |      |
|                    | ( 0)                      | ( 0)                    | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)    | ( 0) | ( 0) | ( 0) | ( 0)     | ( 0) | ( 0) | ( 0) | ( 9)     | ( 0) | ( 0) |      |      |

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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 7

| Organ_____       | Findings_____                | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |           |            |           | 625 ppm<br>18 |            |            |           | 1250 ppm<br>19 |            |            |           | 2500 ppm<br>11 |            |           |           |
|------------------|------------------------------|--|---------------|-----------|------------|-----------|---------------|------------|------------|-----------|----------------|------------|------------|-----------|----------------|------------|-----------|-----------|
|                  |                              |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%)  | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) |
| (Urinary system) |                              |  |               |           |            |           |               |            |            |           |                |            |            |           |                |            |           |           |
| kidney           |                              |  | <26>          |           |            |           | <18>          |            |            |           | <19>           |            |            |           | <11>           |            |           |           |
|                  | hyaline droplet              |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 2<br>( 11)    | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 5<br>( 26)     | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 1<br>( 9)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                  | deposit of amyloid           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 1<br>( 9)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                  | inflammatory polyp           |  | 0<br>( 0)     | 1<br>( 4) | 4<br>( 15) | 0<br>( 0) | 0<br>( 0)     | 3<br>( 17) | 2<br>( 11) | 0<br>( 0) | 1<br>( 5)      | 3<br>( 16) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 1<br>( 9) | 0<br>( 0) |
|                  | hydronephrosis               |  | 1<br>( 4)     | 0<br>( 0) | 6<br>( 23) | 1<br>( 4) | 0<br>( 0)     | 0<br>( 0)  | 6<br>( 33) | 0<br>( 0) | 0<br>( 0)      | 2<br>( 11) | 3<br>( 16) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 9)  | 1<br>( 9) | 0<br>( 0) |
|                  | pyelonephritis               |  | 0<br>( 0)     | 1<br>( 4) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 1<br>( 6)  | 0<br>( 0) | 1<br>( 5)      | 2<br>( 11) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                  | mineralization:cortex        |  | 2<br>( 8)     | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
| urin bladd       | regeneration:proximal tubule |  | 4<br>( 15)    | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0) | 1<br>( 6)     | 1<br>( 6)  | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)  | 0<br>( 0) | 1<br>( 9)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                  | dilatation                   |  | <26>          |           |            |           | <18>          |            |            |           | <19>           |            |            |           | <11>           |            |           |           |
|                  |                              |  | 0<br>( 0)     | 1<br>( 4) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)     | 7<br>( 39) | 0<br>( 0)  | 0<br>( 0) | 1<br>( 5)      | 4<br>( 21) | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)      | 2<br>( 18) | 0<br>( 0) | 0<br>( 0) |

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STUDY NO. : 0712  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ_____            | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |            |            | 625 ppm<br>18 |            |            |            | 1250 ppm<br>19 |            |            |            | 2500 ppm<br>11 |            |            |            |
|-----------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                       |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system)      |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| urin bladd            | ulcer                     |  | <26>          |            |            |            | <18>          |            |            |            | <19>           |            |            |            | <11>           |            |            |            |
|                       |                           |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 5 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                       | simple tubule hyperplasia |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 9 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Endocrine system)    |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| thyroid               | deposit of brown pigment  |  | <26>          |            |            |            | <18>          |            |            |            | <19>           |            |            |            | <11>           |            |            |            |
|                       |                           |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 5 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 9 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Reproductive system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| testis                | mineralization            |  | <26>          |            |            |            | <18>          |            |            |            | <19>           |            |            |            | <11>           |            |            |            |
|                       |                           |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| epididymis            | spermatogenic granuloma   |  | <26>          |            |            |            | <18>          |            |            |            | <19>           |            |            |            | <11>           |            |            |            |
|                       |                           |  | 2<br>( 8 )    | 2<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 6 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

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 < a > a : Number of animals examined at the site  
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(HPT150)

BA1S5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 9

| Organ                            | Findings       | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |            |             | 625 ppm<br>18 |            |            |             | 1250 ppm<br>19 |            |            |             | 2500 ppm<br>11 |            |            |            |
|----------------------------------|----------------|--|---------------|------------|------------|-------------|---------------|------------|------------|-------------|----------------|------------|------------|-------------|----------------|------------|------------|------------|
|                                  |                |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Reproductive system)            |                |  |               |            |            |             |               |            |            |             |                |            |            |             |                |            |            |            |
| prostate                         | inflammation   |  | <26>          |            |            |             | <18>          |            |            |             | <19>           |            |            |             | <11>           |            |            |            |
|                                  |                | 0<br>( 0 )                                     | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 2<br>( 11 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| prep/cli gl                      | duct ectasia   |  | <26>          |            |            |             | <18>          |            |            |             | <19>           |            |            |             | <11>           |            |            |            |
|                                  |                | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 5 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 9 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Nervous system)                 |                |  |               |            |            |             |               |            |            |             |                |            |            |             |                |            |            |            |
| brain                            | hemorrhage     |  | <26>          |            |            |             | <18>          |            |            |             | <19>           |            |            |             | <11>           |            |            |            |
|                                  |                | 1<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                                  | mineralization |  | <26>          |            |            |             | <18>          |            |            |             | <19>           |            |            |             | <11>           |            |            |            |
|                                  |                | 7<br>( 27 )                                    | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 11 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 37 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 36 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Special sense organs/appendage) |                |  |               |            |            |             |               |            |            |             |                |            |            |             |                |            |            |            |
| Harder gl                        | hyperplasia    |  | <26>          |            |            |             | <18>          |            |            |             | <19>           |            |            |             | <11>           |            |            |            |
|                                  |                | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

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PAGE : 10

| Organ                    | Findings       | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |      |      |      | 625 ppm<br>18 |      |      |      | 1250 ppm<br>19 |      |      |      | 2500 ppm<br>11 |      |      |      |
|--------------------------|----------------|--|---------------|------|------|------|---------------|------|------|------|----------------|------|------|------|----------------|------|------|------|
|                          |                |  | 1+            | 2+   | 3+   | 4+   | 1+            | 2+   | 3+   | 4+   | 1+             | 2+   | 3+   | 4+   | 1+             | 2+   | 3+   | 4+   |
|                          |                |  | (%)           | (%)  | (%)  | (%)  | (%)           | (%)  | (%)  | (%)  | (%)            | (%)  | (%)  | (%)  | (%)            | (%)  | (%)  | (%)  |
| (Musculoskeletal system) |                |  |               |      |      |      |               |      |      |      |                |      |      |      |                |      |      |      |
| muscle                   | mineralization |  | <26>          |      |      |      | <18>          |      |      |      | <19>           |      |      |      | <11>           |      |      |      |
|                          |                |  | 1             | 0    | 0    | 0    | 0             | 0    | 0    | 0    | 0              | 0    | 0    | 0    | 0              | 0    | 0    | 0    |
|                          |                |  | ( 4)          | ( 0) | ( 0) | ( 0) | ( 0)          | ( 0) | ( 0) | ( 0) | ( 0)           | ( 0) | ( 0) | ( 0) | ( 0)           | ( 0) | ( 0) | ( 0) |
| bone                     | osteosclerosis |  | <26>          |      |      |      | <18>          |      |      |      | <19>           |      |      |      | <11>           |      |      |      |
|                          |                |  | 1             | 0    | 0    | 0    | 0             | 0    | 0    | 0    | 0              | 0    | 0    | 0    | 0              | 0    | 0    | 0    |
|                          |                |  | ( 4)          | ( 0) | ( 0) | ( 0) | ( 0)          | ( 0) | ( 0) | ( 0) | ( 0)           | ( 0) | ( 0) | ( 0) | ( 0)           | ( 0) | ( 0) | ( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

(HPT150)

BAIS5



TABLE M 3

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

MALE: SACRIFICED ANIMALS

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 1

| Organ_____                       | Findings_____                            | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |           | 625 ppm<br>32 |           |           |           | 1250 ppm<br>31 |           |           |           | 2500 ppm<br>39 |           |           |           |
|----------------------------------|--|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                                  |  |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| [Integumentary system/appandage] |  |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| skin/app                         |  |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                                  | erosion                                  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | squamous cell hyperplasia                |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 10)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | epidermal cyst                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| subcutis                         |  |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                                  | inflammation                             |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                                  | xanthogranuloma                          |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| [Respiratory system]             |  |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| nasal cavit                      |  |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                                  | eosinophilic change:olfactory epithelium |  | 9<br>( 38)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 9<br>( 28)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 8<br>( 26)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 13<br>( 33)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
SACRIFICED ANIMALS (105W)

PAGE : 2

| Organ                | Findings                                    | Group Name<br>No. of Animals on Study<br>Grade |           |           |           | Control<br>24 |           |           |           | 625 ppm<br>32 |           |           |           | 1250 ppm<br>31 |           |           |           | 2500 ppm<br>39 |           |           |           |
|----------------------|---|--|-----------|-----------|-----------|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                      |   | 1+   | 2+        | 3+        | 4+        | 1+            | 2+        | 3+        | 4+        | 1+            | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+        |
|                      |   | (%)  | (%)       | (%)       | (%)       | (%)           | (%)       | (%)       | (%)       | (%)           | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)       |
| (Respiratory system) |   |  |           |           |           |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| nasal cavit          | eosinophilic change:respiratory epithelium  | 13<br>( 54)                                    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 14<br>( 44)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 13<br>( 42)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 16<br>( 41)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | respiratory metaplasia:olfactory epithelium | 3<br>( 13)                                     | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 4<br>( 13)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 13)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 13)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | respiratory metaplasia:gland                | 5<br>( 21)                                     | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 6<br>( 19)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 16)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 8<br>( 21)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| nasopharynx          | eosinophilic change                         | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      |   |  |           |           |           |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| lung                 | hemorrhage                                  | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | deposit of amyloid                          | 4<br>( 17)                                     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 19)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 13)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 8<br>( 21)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | inflammatory infiltration                   | 1<br>( 4)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
< a > a : Number of animals examined at the site  
b : Number of animals with lesion  
( c ) c : b / a \* 100  
Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 3

| Organ                  | Findings                                 | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |             | 625 ppm<br>32 |           |           |             | 1250 ppm<br>31 |           |           |             | 2500 ppm<br>39 |           |           |           |
|------------------------|--|--|---------------|-----------|-----------|-------------|---------------|-----------|-----------|-------------|----------------|-----------|-----------|-------------|----------------|-----------|-----------|-----------|
|                        |  |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Respiratory system)   |  |  |               |           |           |             |               |           |           |             |                |           |           |             |                |           |           |           |
| lung                   | bronchiolar-alveolar cell hyperplasia    |  | <24>          |           |           |             | <32>          |           |           |             | <31>           |           |           |             | <39>           |           |           |           |
|                        |  | 2<br>( 8)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)   | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                        | eosinophilic change:bronchial epithelium |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| (Hematopoietic system) |  |  |               |           |           |             |               |           |           |             |                |           |           |             |                |           |           |           |
| bone marrow            | increased hematopoiesis                  |  | <24>          |           |           |             | <32>          |           |           |             | <31>           |           |           |             | <39>           |           |           |           |
|                        |  | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 3<br>( 10)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                        | granulopoiesis:increased                 |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 2<br>( 6)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) |           |
| spleen                 | deposit of hemosiderin                   |  | <24>          |           |           |             | <32>          |           |           |             | <31>           |           |           |             | <39>           |           |           |           |
|                        |  | 9<br>( 38)                                     | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 17<br>( 53) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 21<br>( 68) | 4<br>( 13)     | 0<br>( 0) | 0<br>( 0) | 24<br>( 62) | 12<br>( 31)    | 1<br>( 3) | 0<br>( 0) |           |
|                        | deposit of melanin                       |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

(HPT150)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 4

| Organ                  | Findings                     | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |           | 625 ppm<br>32 |           |           |             | 1250 ppm<br>31 |            |           |           | 2500 ppm<br>39 |            |           |             |
|------------------------|------------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-------------|----------------|------------|-----------|-----------|----------------|------------|-----------|-------------|
|                        |                              |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)   |
| (Hematopoietic system) |                              |  |               |           |           |           |               |           |           |             |                |            |           |           |                |            |           |             |
| spleen                 |                              |  | <24>          |           |           |           | <32>          |           |           |             | <31>           |            |           |           | <39>           |            |           |             |
|                        | extramedullary hematopoiesis |  | 7<br>( 29)    | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 10<br>( 31)   | 2<br>( 6) | 0<br>( 0) | 0<br>( 0)   | 13<br>( 42)    | 4<br>( 13) | 0<br>( 0) | 0<br>( 0) | 18<br>( 46)    | 9<br>( 23) | 1<br>( 3) | 0 *<br>( 0) |
|                        | follicular hyperplasia       |  | 2<br>( 8)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 9)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
|                        |                              |  |               |           |           |           |               |           |           |             |                |            |           |           |                |            |           |             |
| (Circulatory system)   |                              |  |               |           |           |           |               |           |           |             |                |            |           |           |                |            |           |             |
| heart                  |                              |  | <24>          |           |           |           | <32>          |           |           |             | <31>           |            |           |           | <39>           |            |           |             |
|                        | deposit of amyloid           |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 16)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 5<br>( 16)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 7<br>( 18)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
|                        | myocardial fibrosis          |  | 11<br>( 46)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 16)    | 0<br>( 0) | 0<br>( 0) | 0 *<br>( 0) | 8<br>( 26)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 10<br>( 26)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
|                        | arteritis                    |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
| (Digestive system)     |                              |  |               |           |           |           |               |           |           |             |                |            |           |           |                |            |           |             |
| tooth                  |                              |  | <24>          |           |           |           | <32>          |           |           |             | <31>           |            |           |           | <39>           |            |           |             |
|                        | dysplasia                    |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
 Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 5

| Organ_____         | Findings_____                 | Group Name              | Control      |            |            |            | 625 ppm      |            |            |            | 1250 ppm    |            |            |            | 2500 ppm     |            |            |            |
|--------------------|-------------------------------|-------------------------|--------------|------------|------------|------------|--------------|------------|------------|------------|-------------|------------|------------|------------|--------------|------------|------------|------------|
|                    |                               | No. of Animals on Study | 24           |            |            |            | 32           |            |            |            | 31          |            |            |            | 39           |            |            |            |
|                    |                               | Grade                   | 1+           | 2+         | 3+         | 4+         | 1+           | 2+         | 3+         | 4+         | 1+          | 2+         | 3+         | 4+         | 1+           | 2+         | 3+         | 4+         |
|                    |                               |                         | (%)          | (%)        | (%)        | (%)        | (%)          | (%)        | (%)        | (%)        | (%)         | (%)        | (%)        | (%)        | (%)          | (%)        | (%)        | (%)        |
| (Digestive system) |                               |                         |              |            |            |            |              |            |            |            |             |            |            |            |              |            |            |            |
| tongue             |                               |                         | <24>         |            |            |            | <32>         |            |            |            | <31>        |            |            |            | <39>         |            |            |            |
|                    | squamous cell hyperplasia     |                         | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | arteritis                     |                         | 1<br>( 4 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| stomach            |                               |                         | <24>         |            |            |            | <32>         |            |            |            | <31>        |            |            |            | <39>         |            |            |            |
|                    | hyperplasia:forestomach       |                         | 0<br>( 0 )   | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | erosion:glandular stomach     |                         | 1<br>( 4 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 9 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 10 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | hyperplasia:glandular stomach |                         | 11<br>( 46 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 10<br>( 31 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 9<br>( 29 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 14<br>( 36 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| small intes        |                               |                         | <24>         |            |            |            | <32>         |            |            |            | <31>        |            |            |            | <39>         |            |            |            |
|                    | ulcer                         |                         | 0<br>( 0 )   | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| liver              |                               |                         | <24>         |            |            |            | <32>         |            |            |            | <31>        |            |            |            | <39>         |            |            |            |
|                    | angiectasis                   |                         | 1<br>( 4 )   | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
SACRIFICED ANIMALS (105W)

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| Organ              | Findings               | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |            |            | 625 ppm<br>32 |            |            |            | 1250 ppm<br>31 |            |            |            | 2500 ppm<br>39 |            |            |            |
|--------------------|------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                    |                        |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Digestive system) |                        |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| liver              |                        |  | <24>          |            |            |            | <32>          |            |            |            | <31>           |            |            |            | <39>           |            |            |            |
|                    | necrosis:central       |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | necrosis:focal         |  | 2<br>( 8 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | necrosis:single cell   |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | collapse               |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | inflammatory cell nest |  | 9<br>( 38 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 9<br>( 28 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 23 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 13<br>( 33 )   | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | clear cell focus       |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | acidophilic cell focus |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | basophilic cell focus  |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 5<br>( 16 )   | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 10 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
< a > a : Number of animals examined at the site  
b : Number of animals with lesion  
( c ) c : b / a \* 100  
Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ_____         | Findings_____                   | Group Name              | Control    |            |            |            | 625 ppm    |            |            |            | 1250 ppm   |            |            |            | 2500 ppm    |            |            |              |
|--------------------|---------------------------------|-------------------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|--------------|
|                    |                                 | No. of Animals on Study | 1+         | 2+         | 3+         | 4+         | 1+         | 2+         | 3+         | 4+         | 1+         | 2+         | 3+         | 4+         | 1+          | 2+         | 3+         | 4+           |
| Grade              |                                 |                         | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)        | (%)         | (%)        | (%)        | (%)          |
| <hr/>              |                                 |                         |            |            |            |            |            |            |            |            |            |            |            |            |             |            |            |              |
| {Digestive system} |                                 |                         |            |            |            |            |            |            |            |            |            |            |            |            |             |            |            |              |
| liver              |                                 |                         | <24>       |            |            |            | <32>       |            |            |            | <31>       |            |            |            | <39>        |            |            |              |
|                    | mixed cell focus                |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                    | biliary cyst                    |                         | 2<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                    | intestinal metaplasia:bile duct |                         | 0<br>( 0 ) | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                    | deposit of brown pigment        |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 21 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0 *<br>( 0 ) |
| pancreas           |                                 |                         | <24>       |            |            |            | <32>       |            |            |            | <31>       |            |            |            | <39>        |            |            |              |
|                    | cyst                            |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
| {Urinary system}   |                                 |                         |            |            |            |            |            |            |            |            |            |            |            |            |             |            |            |              |
| kidney             |                                 |                         | <24>       |            |            |            | <32>       |            |            |            | <31>       |            |            |            | <39>        |            |            |              |
|                    | atrophy                         |                         | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 )   |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ_____       | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |            |            | 625 ppm<br>32 |            |            |            | 1250 ppm<br>31 |            |            |            | 2500 ppm<br>39 |             |            |            |
|------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|-------------|------------|------------|
|                  |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |             |            |            |
| kidney           |                           |  | <24>          |            |            |            | <32>          |            |            |            | <31>           |            |            |            | <39>           |             |            |            |
|                  | cyst                      |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | hyaline droplet           |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | hyaline cast              |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | inflammatory infiltration |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | lymphocytic infiltration  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | scar                      |  | 1<br>( 4 )    | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 )    | 3<br>( 9 ) | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 13 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 3<br>( 8 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | inflammatory polyp        |  | 0<br>( 0 )    | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 1<br>( 3 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | hydronephrosis            |  | 0<br>( 0 )    | 2<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 3<br>( 9 ) | 2<br>( 6 ) | 0<br>( 0 ) | 0<br>( 0 )     | 2<br>( 6 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 4<br>( 10 ) | 1<br>( 3 ) | 0<br>( 0 ) |

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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ              | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |           | 625 ppm<br>32 |           |           |           | 1250 ppm<br>31 |           |           |           | 2500 ppm<br>39 |           |           |           |
|--------------------|-------------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                    |                               |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Urinary system)   |                               |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| kidney             |                               |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                    | mineralization:papilla        |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | regeneration:proximal tubule  |  | 15<br>( 63)   | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 16<br>( 50)   | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 17<br>( 55)    | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 20<br>( 51)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | urothelial hyperplasia:pelvis |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| urin bladd         |                               |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                    | xanthogranuloma               |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Endocrine system) |                               |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| pituitary          |                               |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                    | Rathke pouch                  |  | 2<br>( 8)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| thyroid            |                               |  | <24>          |           |           |           | <32>          |           |           |           | <31>           |           |           |           | <39>           |           |           |           |
|                    | cystic thyroid follicle       |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ_____            | Findings_____            | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |            |            | 625 ppm<br>32 |            |            |             | 1250 ppm<br>31 |            |            |             | 2500 ppm<br>39 |            |            |            |
|-----------------------|--------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|-------------|----------------|------------|------------|-------------|----------------|------------|------------|------------|
|                       |                          |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Endocrine system)    |                          |  |               |            |            |            |               |            |            |             |                |            |            |             |                |            |            |            |
| thyroid               | deposit of brown pigment |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 9<br>( 23 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| adrenal               | hyperplasia:medulla      |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 0<br>( 0 )                                     | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Reproductive system) |                          |  |               |            |            |            |               |            |            |             |                |            |            |             |                |            |            |            |
| testis                | mineralization           |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 1<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )  | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                       | xanthogranuloma          |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 0<br>( 0 )                                     | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| epididymis            | spermatogenic granuloma  |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 2<br>( 8 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 10 ) | 2<br>( 6 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| prep/cli gl           | duct ectasia             |  | <24>          |            |            |            | <32>          |            |            |             | <31>           |            |            |             | <39>           |            |            |            |
|                       |                          | 1<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 11

| Organ                            | Findings  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |             |           |            | 625 ppm<br>32 |           |           |            | 1250 ppm<br>31 |           |           |            | 2500 ppm<br>39 |           |           |           |  |
|----------------------------------|---|--|---------------|-------------|-----------|------------|---------------|-----------|-----------|------------|----------------|-----------|-----------|------------|----------------|-----------|-----------|-----------|--|
|                                  |   |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |  |
| (Nervous system)                 |   |  |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
| brain                            | mineralization  |  | <24>          |             |           |            | <32>          |           |           |            | <31>           |           |           |            | <39>           |           |           |           |  |
|                                  |   | 11<br>( 46)                                    | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 9<br>( 28) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 9<br>( 29) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 7<br>( 18) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0 *       |  |
| (Special sense organs/appendage) |   |  |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
| eye                              | cataract  |  | <24>          |             |           |            | <32>          |           |           |            | <31>           |           |           |            | <39>           |           |           |           |  |
|                                  |   | 2<br>( 8)                                      | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 2<br>( 6)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0         |  |
|                                  |   | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0         |  |
| Harder gl                        | lymphocytic infiltration  |  | <24>          |             |           |            | <32>          |           |           |            | <31>           |           |           |            | <39>           |           |           |           |  |
|                                  |   | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0         |  |
|                                  |   | 1<br>( 4)                                      | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0         |  |
| Grade                            | 1+ : Slight   | 2+ : Moderate                                  | 3+ : Marked   | 4+ : Severe |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
|                                  | < a >   | a : Number of animals examined at the site     |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
|                                  | b   | b : Number of animals with lesion              |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
|                                  | ( c )   | c : b / a * 100                                |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |
|                                  | Significant difference ;    * : P ≤ 0.05    ** : P ≤ 0.01    Test of Chi Square |  |               |             |           |            |               |           |           |            |                |           |           |            |                |           |           |           |  |

(HPT150)

BAIS5

TABLE M 4

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

FEMALE: ALL ANIMALS

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 15

| Organ                            | Findings                                    | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |             |            |            | 625 ppm<br>50 |              |            |            | 1250 ppm<br>50 |              |            |             | 2500 ppm<br>50 |             |            |            |
|----------------------------------|---|--|---------------|-------------|------------|------------|---------------|--------------|------------|------------|----------------|--------------|------------|-------------|----------------|-------------|------------|------------|
|                                  |   |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)    | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)    | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  |
| (Integumentary system/appandage) |   |  |               |             |            |            |               |              |            |            |                |              |            |             |                |             |            |            |
| skin/app                         | epidermal cyst                              |  | <50>          |             |            |            | <50>          |              |            |            | <50>           |              |            |             | <50>           |             |            |            |
|                                  |   | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 )   | 0<br>( 0 ) | 1<br>( 2 )  | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
| (Respiratory system)             |   |  |               |             |            |            |               |              |            |            |                |              |            |             |                |             |            |            |
| nasal cavit                      | eosinophilic change:olfactory epithelium    |  | <50>          |             |            |            | <50>          |              |            |            | <50>           |              |            |             | <50>           |             |            |            |
|                                  |   | 4<br>( 8 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 3<br>( 6 ) | 1<br>( 2 )    | 0<br>( 0 )   | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )     | 0<br>( 0 )   | 0<br>( 0 ) | 6<br>( 12 ) | 1<br>( 2 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
|                                  | eosinophilic change:respiratory epithelium  |  | 28<br>( 56 )  | 8<br>( 16 ) | 0<br>( 0 ) | 0<br>( 0 ) | 26<br>( 52 )  | 2<br>( 4 )   | 0<br>( 0 ) | 0<br>( 0 ) | 28<br>( 56 )   | 3<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 )  | 33<br>( 66 )   | 5<br>( 10 ) | 1<br>( 2 ) | 0<br>( 0 ) |
|                                  |   | inflammation:respiratory epithelium            |               | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 2 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                                  | respiratory metaplasia:olfactory epithelium |  |               | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 2<br>( 4 )   | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 3<br>( 6 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                                  |   | respiratory metaplasia:gland                   |               | 9<br>( 18 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 11<br>( 22 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 10<br>( 20 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )     | 9<br>( 18 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| nasopharynx                      | eosinophilic change                         |  | <50>          |             |            |            | <50>          |              |            |            | <50>           |              |            |             | <50>           |             |            |            |
|                                  |   | 3<br>( 6 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )    | 0<br>( 0 )   | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 )   | 0<br>( 0 ) | 2<br>( 4 )  | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 16

| Organ_____             | Findings_____                         | Group Name              | Control      |            |            |            | 625 ppm      |            |            |            | 1250 ppm     |            |            |            | 2500 ppm    |            |            |            |
|------------------------|---------------------------------------|-------------------------|--------------|------------|------------|------------|--------------|------------|------------|------------|--------------|------------|------------|------------|-------------|------------|------------|------------|
|                        |                                       | No. of Animals on Study | 50           |            |            |            | 50           |            |            |            | 50           |            |            |            | 50          |            |            |            |
|                        |                                       | Grade                   | 1+           | 2+         | 3+         | 4+         | 1+           | 2+         | 3+         | 4+         | 1+           | 2+         | 3+         | 4+         | 1+          | 2+         | 3+         | 4+         |
|                        |                                       |                         | (%)          | (%)        | (%)        | (%)        | (%)          | (%)        | (%)        | (%)        | (%)          | (%)        | (%)        | (%)        | (%)         | (%)        | (%)        | (%)        |
| (Respiratory system)   |                                       |                         |              |            |            |            |              |            |            |            |              |            |            |            |             |            |            |            |
| lung                   |                                       |                         | <50>         |            |            |            | <50>         |            |            |            | <50>         |            |            |            | <50>        |            |            |            |
|                        | congestion                            |                         | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | deposit of amyloid                    |                         | 3<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 14 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 5<br>( 10 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | inflammatory infiltration             |                         | 3<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | lymphocytic infiltration              |                         | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | bronchiolar-alveolar cell hyperplasia |                         | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Hematopoietic system) |                                       |                         |              |            |            |            |              |            |            |            |              |            |            |            |             |            |            |            |
| bone marrow            |                                       |                         | <50>         |            |            |            | <50>         |            |            |            | <50>         |            |            |            | <50>        |            |            |            |
|                        | granulation                           |                         | 0<br>( 0 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | increased hematopoiesis               |                         | 10<br>( 20 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 13<br>( 26 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 16 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 17

| Organ                  | Findings                 | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |             |             | 625 ppm<br>50 |             |             |             | 1250 ppm<br>50 |             |              |             | 2500 ppm<br>50 |             |              |           |
|------------------------|--------------------------|--|---------------|------------|-------------|-------------|---------------|-------------|-------------|-------------|----------------|-------------|--------------|-------------|----------------|-------------|--------------|-----------|
|                        |                          |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)   | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)   | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)    | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)    | 4+<br>(%) |
| (Hematopoietic system) |                          |  |               |            |             |             |               |             |             |             |                |             |              |             |                |             |              |           |
| bone marrow            | granulopoiesis:increased |  | <50>          |            |             |             | <50>          |             |             |             | <50>           |             |              |             | <50>           |             |              |           |
|                        |                          | 5<br>( 10)                                     | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 3<br>( 6)   | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0)   | 4<br>( 8)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    | 5<br>( 10)  | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    |           |
| lymph node             | lymphadenitis            |  | <50>          |            |             |             | <50>          |             |             |             | <50>           |             |              |             | <50>           |             |              |           |
|                        |                          | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0)   | 1<br>( 2)   | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0)    | 0<br>( 0)   | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0)    |           |
| spleen                 | deposit of hemosiderin   |  | <50>          |            |             |             | <50>          |             |             |             | <50>           |             |              |             | <50>           |             |              |           |
|                        |                          | 30<br>( 60)                                    | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 34<br>( 68) | 3<br>( 6)     | 0<br>( 0)   | 0<br>( 0)   | 38<br>( 76) | 6<br>( 12)     | 0<br>( 0)   | 0<br>( 0) ** | 30<br>( 60) | 12<br>( 24)    | 5<br>( 10)  | 0<br>( 0) ** |           |
|                        | deposit of melanin       |  | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)   | 1<br>( 2)     | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    |           |
|                        |                          | extramedullary hematopoiesis                   |               | 7<br>( 14) | 11<br>( 22) | 1<br>( 2)   | 0<br>( 0)     | 13<br>( 26) | 13<br>( 26) | 2<br>( 4)   | 0<br>( 0)      | 13<br>( 26) | 10<br>( 20)  | 0<br>( 0)   | 0<br>( 0)      | 19<br>( 38) | 12<br>( 24)  | 0<br>( 0) |
|                        | follicular hyperplasia   |  | 1<br>( 2)     | 1<br>( 2)  | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)   | 1<br>( 2)      | 2<br>( 4)   | 0<br>( 0)    | 0<br>( 0)   | 1<br>( 2)      | 0<br>( 0)   | 0<br>( 0)    | 0<br>( 0) |
| (Circulatory system)   |                          |  |               |            |             |             |               |             |             |             |                |             |              |             |                |             |              |           |
| heart                  | thrombus                 |  | <50>          |            |             |             | <50>          |             |             |             | <50>           |             |              |             | <50>           |             |              |           |
|                        |                          | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 3<br>( 6)   | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 18

| Organ                | Findings                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |           |           |           | 625 ppm<br>50 |           |           |           | 1250 ppm<br>50 |           |           |           | 2500 ppm<br>50 |           |           |           |
|----------------------|---------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                      |                           |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Circulatory system) |                           |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| heart                | deposit of amyloid        |  | <50>          |           |           |           | <50>          |           |           |           | <50>           |           |           |           | <50>           |           |           |           |
|                      |                           |  | 2<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 12)     | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | mineralization            |  | 2<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | inflammatory infiltration |  | 1<br>( 2)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | myocardial fibrosis       |  | 13<br>( 26)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 13<br>( 26)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 20<br>( 40)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 17<br>( 34)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | arteritis                 |  | 1<br>( 2)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

(Digestive system)

|        |                           |  |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |
|--------|---------------------------|--|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|
| tongue | inflammatory infiltration |  | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           | <50>      |           |           |           |
|        |                           |  | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|        | arteritis                 |  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 19

| Organ_____         | Findings_____                 | Group Name              | Control    |           |           |           | 625 ppm     |           |           |           | 1250 ppm    |           |           |           | 2500 ppm    |           |           |           |
|--------------------|-------------------------------|-------------------------|------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|
|                    |                               | No. of Animals on Study | 50         |           |           |           | 50          |           |           |           | 50          |           |           |           | 50          |           |           |           |
|                    |                               | Grade                   | 1+         | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        | 1+          | 2+        | 3+        | 4+        |
|                    |                               |                         | (%)        | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       | (%)         | (%)       | (%)       | (%)       |
| <hr/>              |                               |                         |            |           |           |           |             |           |           |           |             |           |           |           |             |           |           |           |
| (Digestive system) |                               |                         |            |           |           |           |             |           |           |           |             |           |           |           |             |           |           |           |
| stomach            |                               |                         | <50>       |           |           |           | <50>        |           |           |           | <50>        |           |           |           | <50>        |           |           |           |
|                    | hyperplasia:forestomach       |                         | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 6)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | erosion:glandular stomach     |                         | 2<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 6)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | hyperplasia:glandular stomach |                         | 8<br>( 16) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 11<br>( 22) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 12<br>( 24) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 12<br>( 24) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| small intes        |                               |                         | <50>       |           |           |           | <50>        |           |           |           | <50>        |           |           |           | <50>        |           |           |           |
|                    | adhesion                      |                         | 0<br>( 0)  | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| liver              |                               |                         | <50>       |           |           |           | <50>        |           |           |           | <50>        |           |           |           | <50>        |           |           |           |
|                    | angiectasis                   |                         | 1<br>( 2)  | 3<br>( 6) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 3<br>( 6) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 2<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 3<br>( 6) | 0<br>( 0) | 0<br>( 0) |
|                    | necrosis:central              |                         | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 2) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | necrosis:focal                |                         | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 20

| Organ_____         | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |               |
|--------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|---------------|
|                    |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)     |
| (Digestive system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |               |
| liver              |                           |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |               |
|                    | inflammatory infiltration |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    |
|                    | lymphocytic infiltration  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    |
|                    | inflammatory cell nest    |  | 14<br>( 28 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 14<br>( 28 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 12<br>( 24 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 17<br>( 34 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 )    |
|                    | acidophilic cell focus    |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    |
|                    | biliary cyst              |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    |
|                    | deposit of brown pigment  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 10<br>( 20 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0 **<br>( 0 ) |

(Urinary system)

|        |      |  |            |            |            |            |            |            |            |            |            |            |            |            |            |            |            |            |
|--------|------|--|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| kidney | cyst |  | <50>       |            |            |            | <50>       |            |            |            | <50>       |            |            |            | <50>       |            |            |            |
|        |      |  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 21

| Organ            | Findings                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |           |           | 625 ppm<br>50 |            |           |           | 1250 ppm<br>50 |            |           |           | 2500 ppm<br>50 |            |           |             |
|------------------|---------------------------|--|---------------|------------|-----------|-----------|---------------|------------|-----------|-----------|----------------|------------|-----------|-----------|----------------|------------|-----------|-------------|
|                  |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)   |
| (Urinary system) |                           |  |               |            |           |           |               |            |           |           |                |            |           |           |                |            |           |             |
| kidney           |                           |  | <50>          |            |           |           | <50>          |            |           |           | <50>           |            |           |           | <50>           |            |           |             |
|                  | hyaline droplet           |  | 14<br>( 28)   | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 9<br>( 18)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 10<br>( 20)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)     | 0<br>( 0)  | 0<br>( 0) | 0 *<br>( 0) |
|                  | hyaline cast              |  | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 3<br>( 6) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) |             |
|                  | inflammatory infiltration |  | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) |             |
|                  | lymphocytic infiltration  |  | 4<br>( 8)     | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 3<br>( 6)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
|                  | osseous metaplasia        |  | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 2)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
|                  | scar                      |  | 3<br>( 6)     | 5<br>( 10) | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)     | 5<br>( 10) | 0<br>( 0) | 0<br>( 0) | 2<br>( 4)      | 9<br>( 18) | 1<br>( 2) | 0<br>( 0) | 6<br>( 12)     | 5<br>( 10) | 0<br>( 0) | 0<br>( 0)   |
|                  | inflammatory polyp        |  | 1<br>( 2)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 3<br>( 6)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 4<br>( 8)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   |
| hydronephrosis   |                           | 0<br>( 0)                                      | 3<br>( 6)     | 1<br>( 2)  | 0<br>( 0) | 0<br>( 0) | 5<br>( 10)    | 1<br>( 2)  | 0<br>( 0) | 1<br>( 2) | 4<br>( 8)      | 2<br>( 4)  | 0<br>( 0) | 1<br>( 2) | 6<br>( 12)     | 1<br>( 2)  | 0<br>( 0) |             |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
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STUDY NO. : 0712  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 22

| Organ            | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|------------------|-------------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                  |                               |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system) |                               |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| kidney           | tubular necrosis              |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                  |                               |  | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | papillary necrosis            |  | 3<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 14 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 6<br>( 12 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | mineralization:papilla        |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | glomerulosclerosis            |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | regeneration:proximal tubule  |  | 2<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 8 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | desquamation:pelvis           |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| ureter           | urothelial hyperplasia:pelvis |  | 2<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                  | dilatation                    |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                  |                               |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

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 < a > a : Number of animals examined at the site  
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 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 23

| Organ   | Findings   | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|---|--|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|   |  |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system)  |  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| urin bladd  |  |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|   | dilatation   |  | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|   | lymphocytic infiltration   |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|   | xanthogranuloma  |  | 0<br>( 0 )    | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Endocrine system)  |  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| pituitary   |  |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|   | angiectasis  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|   | cyst   |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|   | hyperplasia  |  | 3<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 )    | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 8 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 8 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|   | Rathke pouch   |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| Grade   | 1+ : Slight      2+ : Moderate      3+ : Marked      4+ : Severe |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| < a >   | a : Number of animals examined at the site                       |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| b   | b : Number of animals with lesion                                |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| ( c )   | c : b / a * 100  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| Significant difference ;    * : P ≤ 0.05    ** : P ≤ 0.01    Test of Chi Square |  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

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| Organ                 | Findings                               | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |              |
|-----------------------|--|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|--------------|
|                       |  |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)    |
| (Endocrine system)    |  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |              |
| thyroid               | arteritis                              |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |              |
|                       |  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                       | deposit of brown pigment               |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 14 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0 *<br>( 0 ) |
| adrenal               | spindle-cell hyperplasia               |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |              |
|                       |  |  | 4<br>( 8 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                       | fatty change:corticomedullary junction |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |              |
| (Reproductive system) |  |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |              |
| ovary                 | thrombus                               |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |              |
|                       |  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |
|                       | cyst                                   |  | 3<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 16 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 5<br>( 10 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 7<br>( 14 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )   |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

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| Organ_____                       | Findings_____                  | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |            |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|----------------------------------|--------------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                                  |                                |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| [Reproductive system]            |                                |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| uterus                           | dilatation                     |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                                  |                                |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                                  | cystic endometrial hyperplasia |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                                  |                                |  | 9<br>( 18 )   | 4<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 8<br>( 16 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 )   | 1<br>( 2 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 )   | 2<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| [Nervous system]                 |                                |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| brain                            | mineralization                 |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                                  |                                |  | 12<br>( 24 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 11<br>( 22 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 13<br>( 26 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 10<br>( 20 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| spinal cord                      | gliosis                        |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                                  |                                |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| [Special sense organs/appendage] |                                |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| eye                              | keratitis                      |  | <50>          |            |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |            |
|                                  |                                |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

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| Organ  | Findings                                   | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |             |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |            |
|--|--|--|---------------|-------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|  |  |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Special sense organs/appendage)                                       |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| Harder gl  | lymphocytic infiltration                   |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|  | hyperplasia                                |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Musculoskeletal system)   |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| muscle   | hematoma                                   |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|  | mineralization                             |  | 1<br>( 2 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 6 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| bone   | osteosclerosis                             |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 4 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|  |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| (Body cavities)  |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| pleura   | inflammation                               |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|  |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| Grade  | 1+ : Slight                                | 2+ : Moderate                                  | 3+ : Marked   | 4+ : Severe |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| < a >  | a : Number of animals examined at the site |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| b  | b : Number of animals with lesion          |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| ( c )  | c : b / a * 100                            |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |            |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

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| Organ_____   | Findings_____                              | Group Name<br>No. of Animals on Study<br>Grade | Control<br>50 |             |            |            | 625 ppm<br>50 |            |            |            | 1250 ppm<br>50 |            |            |            | 2500 ppm<br>50 |            |            |           |
|--|--|--|---------------|-------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|-----------|
|  |  |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%) |
| (Body cavities)  |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| mediastinum  | inflammatory infiltration                  |  | <50>          |             |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |           |
|  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
| peritoneum   | inflammation                               |  | <50>          |             |            |            | <50>          |            |            |            | <50>           |            |            |            | <50>           |            |            |           |
|  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 1<br>( 2 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 2 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
| Grade  | 1+ : Slight                                | 2+ : Moderate                                  | 3+ : Marked   | 4+ : Severe |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| < a >  | a : Number of animals examined at the site |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| b  | b : Number of animals with lesion          |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| ( c )  | c : b / a * 100                            |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |

(HPT150)

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TABLE M 5

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

FEMALE: DEAD AND MORIBUND ANIMALS

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ_____   | Findings_____                               | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |             |           |           | 625 ppm<br>29 |           |           |           | 1250 ppm<br>17 |           |           |           | 2500 ppm<br>12 |           |           |           |
|--|---|--|---------------|-------------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|  |   |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Respiratory system)   |   |  |               |             |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| nasal cavit  |   |  | <26>          |             |           |           | <29>          |           |           |           | <17>           |           |           |           | <12>           |           |           |           |
|  | eosinophilic change:olfactory epithelium    |  | 3<br>( 12)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 17)     | 1<br>( 8) | 0<br>( 0) | 0<br>( 0) |
|  | eosinophilic change:respiratory epithelium  |  | 14<br>( 54)   | 2<br>( 8)   | 0<br>( 0) | 0<br>( 0) | 13<br>( 45)   | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 7<br>( 41)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 42)     | 1<br>( 8) | 0<br>( 0) | 0<br>( 0) |
|  | inflammation:respiratory epithelium         |  | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|  | respiratory metaplasia:olfactory epithelium |  | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|  | respiratory metaplasia:gland                |  | 5<br>( 19)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 5<br>( 17)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 4<br>( 24)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 25)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| nasopharynx  |   |  | <26>          |             |           |           | <29>          |           |           |           | <17>           |           |           |           | <12>           |           |           |           |
|  | eosinophilic change                         |  | 1<br>( 4)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| lung   |   |  | <26>          |             |           |           | <29>          |           |           |           | <17>           |           |           |           | <12>           |           |           |           |
|  | congestion                                  |  | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 8)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| Grade  | 1+ : Slight                                 | 2+ : Moderate                                  | 3+ : Marked   | 4+ : Severe |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| < a >  | a : Number of animals examined at the site  |  |               |             |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| b  | b : Number of animals with lesion           |  |               |             |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| ( c )  | c : b / a * 100                             |  |               |             |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |   |  |               |             |           |           |               |           |           |           |                |           |           |           |                |           |           |           |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 12

| Organ                  | Findings                     | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |             |           |             | 625 ppm<br>29 |             |           |             | 1250 ppm<br>17 |            |             |            | 2500 ppm<br>12 |            |             |           |
|------------------------|------------------------------|--|---------------|-------------|-----------|-------------|---------------|-------------|-----------|-------------|----------------|------------|-------------|------------|----------------|------------|-------------|-----------|
|                        |                              |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)   | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)   | 4+<br>(%) |
| [Respiratory system]   |                              |  |               |             |           |             |               |             |           |             |                |            |             |            |                |            |             |           |
| lung                   | deposit of amyloid           |  | <26>          |             |           |             | <29>          |             |           |             | <17>           |            |             |            | <12>           |            |             |           |
|                        |                              | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 1<br>( 6)   | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)   |           |
|                        | inflammatory infiltration    |  | 3<br>( 12)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 1<br>( 3)   | 0<br>( 0) | 0<br>( 0)   | 2<br>( 12)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)  | 2<br>( 17)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0) |
| [Hematopoietic system] |                              |  |               |             |           |             |               |             |           |             |                |            |             |            |                |            |             |           |
| bone marrow            | increased hematopoiesis      |  | <26>          |             |           |             | <29>          |             |           |             | <17>           |            |             |            | <12>           |            |             |           |
|                        |                              | 10<br>( 38)                                    | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 12<br>( 41) | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 8<br>( 47)  | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)   | 5<br>( 42) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)   |           |
|                        | granulopoiesis:increased     |  | 4<br>( 15)    | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 2<br>( 7)     | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 2<br>( 12)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0)   |           |
| spleen                 | deposit of hemosiderin       |  | <26>          |             |           |             | <29>          |             |           |             | <17>           |            |             |            | <12>           |            |             |           |
|                        |                              | 12<br>( 46)                                    | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 16<br>( 55) | 0<br>( 0)     | 0<br>( 0)   | 0<br>( 0) | 12<br>( 71) | 2<br>( 12)     | 0<br>( 0)  | 0<br>( 0) * | 8<br>( 67) | 2<br>( 17)     | 0<br>( 0)  | 0<br>( 0) * |           |
|                        | extramedullary hematopoiesis |  | 2<br>( 8)     | 11<br>( 42) | 1<br>( 4) | 0<br>( 0)   | 4<br>( 14)    | 13<br>( 45) | 2<br>( 7) | 0<br>( 0)   | 2<br>( 12)     | 7<br>( 41) | 0<br>( 0)   | 0<br>( 0)  | 2<br>( 17)     | 3<br>( 25) | 0<br>( 0)   | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
< a > a : Number of animals examined at the site  
b : Number of animals with lesion  
( c ) c : b / a \* 100  
Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$  Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [CrJ:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ_____             | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |             |            |             | 625 ppm<br>29 |             |            |            | 1250 ppm<br>17 |             |            |            | 2500 ppm<br>12 |             |            |            |
|------------------------|---------------------------|--|---------------|-------------|------------|-------------|---------------|-------------|------------|------------|----------------|-------------|------------|------------|----------------|-------------|------------|------------|
|                        |                           |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  |
| (Hematopoietic system) |                           |  |               |             |            |             |               |             |            |            |                |             |            |            |                |             |            |            |
| spleen                 | follicular hyperplasia    |  | <26>          |             |            |             | <29>          |             |            |            | <17>           |             |            |            | <12>           |             |            |            |
|                        |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 1<br>( 8 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
| (Circulatory system)   |                           |  |               |             |            |             |               |             |            |            |                |             |            |            |                |             |            |            |
| heart                  | thrombus                  |  | <26>          |             |            |             | <29>          |             |            |            | <17>           |             |            |            | <12>           |             |            |            |
|                        |                           | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 3<br>( 10 ) | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
|                        | deposit of amyloid        |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 )  | 1<br>( 3 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
|                        |                           | mineralization                                 |               | 2<br>( 8 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 4<br>( 14 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 6 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 8 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | inflammatory infiltration |  |               | 1<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                        |                           | myocardial fibrosis                            |               | 7<br>( 27 ) | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 9<br>( 31 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 8<br>( 47 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 5<br>( 42 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                        | arteritis                 |  |               | 1<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 2<br>( 12 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ              | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |           |           |           | 625 ppm<br>29 |            |           |           | 1250 ppm<br>17 |           |           |           | 2500 ppm<br>12 |           |           |           |
|--------------------|-------------------------------|--|---------------|-----------|-----------|-----------|---------------|------------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                    |                               |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| [Digestive system] |                               |  |               |           |           |           |               |            |           |           |                |           |           |           |                |           |           |           |
| tongue             |                               |  | <26>          |           |           |           | <29>          |            |           |           | <17>           |           |           |           | <12>           |           |           |           |
|                    | inflammatory infiltration     |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | arteritis                     |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| stomach            |                               |  | <26>          |           |           |           | <29>          |            |           |           | <17>           |           |           |           | <12>           |           |           |           |
|                    | hyperplasia:forestomach       |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | erosion:glandular stomach     |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                    | hyperplasia:glandular stomach |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 7)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| small intes        |                               |  | <26>          |           |           |           | <29>          |            |           |           | <17>           |           |           |           | <12>           |           |           |           |
|                    | adhesion                      |  | 0<br>( 0)     | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| liver              |                               |  | <26>          |           |           |           | <29>          |            |           |           | <17>           |           |           |           | <12>           |           |           |           |
|                    | angiectasis                   |  | 0<br>( 0)     | 1<br>( 4) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 3<br>( 10) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ_____         | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |            |            | 625 ppm<br>29 |            |            |            | 1250 ppm<br>17 |            |            |            | 2500 ppm<br>12 |            |            |            |
|--------------------|---------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                    |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Digestive system) |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| liver              | necrosis:central          |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | necrosis:focal            |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | inflammatory infiltration |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | inflammatory cell nest    |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 7 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | deposit of brown pigment  |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Urinary system)   |                           |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| kidney             | hyaline droplet           |  | 11<br>( 42 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 9<br>( 31 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 6<br>( 35 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | hyaline cast              |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 17 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 16

| Organ_____         | Findings_____             | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |            |            | 625 ppm<br>29 |             |            |            | 1250 ppm<br>17 |             |            |            | 2500 ppm<br>12 |            |            |            |
|--------------------|---------------------------|--|---------------|------------|------------|------------|---------------|-------------|------------|------------|----------------|-------------|------------|------------|----------------|------------|------------|------------|
|                    |                           |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system)   |                           |  |               |            |            |            |               |             |            |            |                |             |            |            |                |            |            |            |
| kidney             |                           |  | <26>          |            |            |            | <29>          |             |            |            | <17>           |             |            |            | <12>           |            |            |            |
|                    | inflammatory infiltration |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | lymphocytic infiltration  |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | osseous metaplasia        |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | scar                      |  | 2<br>( 8 )    | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 2<br>( 7 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 2<br>( 12 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | inflammatory polyp        |  | 1<br>( 4 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                    | hydronephrosis            |  | 0<br>( 0 )    | 2<br>( 8 ) | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 )    | 3<br>( 10 ) | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 1<br>( 6 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 1<br>( 8 ) | 0<br>( 0 ) |
|                    | tubular necrosis          |  | 0<br>( 0 )    | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| papillary necrosis |                           | 1<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 1<br>( 6 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ            | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26   |            |            |            | 625 ppm<br>29 |            |            |            | 1250 ppm<br>17 |            |            |            | 2500 ppm<br>12 |            |            |            |
|------------------|-------------------------------|--|-----------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                  |                               |  | 1+<br>(%)       | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Urinary system) |                               |  |                 |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| kidney           | glomerulosclerosis            |  | <26>            |            |            |            | <29>          |            |            |            | <17>           |            |            |            | <12>           |            |            |            |
|                  |                               | 0<br>( 0 )                                     | 0<br>( 0 )      | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                  | regeneration:proximal tubule  |  | 0<br>( 0 )      | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 1<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  |                               | desquamation:pelvis                            |                 | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| ureter           | urothelial hyperplasia:pelvis |  | 0<br>( 0 )      | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  |                               | dilatation                                     |                 | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  | urin bladd                    | dilatation                                     |                 | 1<br>( 4 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  |                               |  | xanthogranuloma |            | 0<br>( 0 ) | 1<br>( 4 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

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| Organ                 | Findings                 | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |            |            | 625 ppm<br>29 |            |            |            | 1250 ppm<br>17 |            |            |            | 2500 ppm<br>12 |            |            |            |
|-----------------------|--------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                       |                          |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| (Endocrine system)    |                          |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| pituitary             | hyperplasia              |  | <26>          |            |            |            | <29>          |            |            |            | <17>           |            |            |            | <12>           |            |            |            |
|                       |                          |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                       | Rathke pouch             |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| thyroid               | arteritis                |  | <26>          |            |            |            | <29>          |            |            |            | <17>           |            |            |            | <12>           |            |            |            |
|                       |                          |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 6 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                       | deposit of brown pigment |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| (Reproductive system) |                          |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| ovary                 | thrombus                 |  | <26>          |            |            |            | <29>          |            |            |            | <17>           |            |            |            | <12>           |            |            |            |
|                       |                          |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                       | cyst                     |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 10 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 19

| Organ                    | Findings                       | Group Name<br>No. of Animals on Study<br>Grade | Control<br>26 |            |           |            | 625 ppm<br>29 |           |           |            | 1250 ppm<br>17 |           |           |            | 2500 ppm<br>12 |           |           |           |
|--------------------------|--------------------------------|--|---------------|------------|-----------|------------|---------------|-----------|-----------|------------|----------------|-----------|-----------|------------|----------------|-----------|-----------|-----------|
|                          |                                |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Reproductive system)    |                                |  |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| uterus                   | cystic endometrial hyperplasia |  | <26>          |            |           |            | <29>          |           |           |            | <17>           |           |           |            | <12>           |           |           |           |
|                          |                                | 4<br>( 15)                                     | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 3<br>( 10) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 1<br>( 8)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| (Nervous system)         |                                |  |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| brain                    | mineralization                 |  | <26>          |            |           |            | <29>          |           |           |            | <17>           |           |           |            | <12>           |           |           |           |
|                          |                                | 4<br>( 15)                                     | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 6<br>( 21) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 4<br>( 24) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 2<br>( 17) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| spinal cord              | gliosis                        |  | <26>          |            |           |            | <29>          |           |           |            | <17>           |           |           |            | <12>           |           |           |           |
|                          |                                | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 6)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| (Musculoskeletal system) |                                |  |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| muscle                   | hematoma                       |  | <26>          |            |           |            | <29>          |           |           |            | <17>           |           |           |            | <12>           |           |           |           |
|                          |                                | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
|                          | mineralization                 |  | <26>          |            |           |            | <29>          |           |           |            | <17>           |           |           |            | <12>           |           |           |           |
| 1<br>( 4)                | 0<br>( 0)                      | 0<br>( 0)                                      | 0<br>( 0)     | 3<br>( 10) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 1<br>( 6) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 1<br>( 8) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      |           |           |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 20

| Organ_____   | Findings_____                              | Group Name              | Control     |             |       |       | 625 ppm |       |       |       | 1250 ppm |       |       |       | 2500 ppm |       |       |       |
|--|--|-------------------------|-------------|-------------|-------|-------|---------|-------|-------|-------|----------|-------|-------|-------|----------|-------|-------|-------|
|  |  | No. of Animals on Study | 26          |             |       |       | 29      |       |       |       | 17       |       |       |       | 12       |       |       |       |
|  |  | Grade                   | 1+          | 2+          | 3+    | 4+    | 1+      | 2+    | 3+    | 4+    | 1+       | 2+    | 3+    | 4+    | 1+       | 2+    | 3+    | 4+    |
|  |  |                         | (%)         | (%)         | (%)   | (%)   | (%)     | (%)   | (%)   | (%)   | (%)      | (%)   | (%)   | (%)   | (%)      | (%)   | (%)   | (%)   |
| [Body cavities]  |  |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| pleura   | inflammation                               |                         | <26>        |             |       |       | <29>    |       |       |       | <17>     |       |       |       | <12>     |       |       |       |
|  |  | 0                       | 0           | 0           | 0     | 0     | 0       | 0     | 0     | 0     | 0        | 1     | 0     | 0     | 0        | 0     | 0     | 0     |
|  |  |                         | ( 0 )       | ( 0 )       | ( 0 ) | ( 0 ) | ( 0 )   | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )    | ( 6 ) | ( 0 ) | ( 0 ) | ( 0 )    | ( 0 ) | ( 0 ) | ( 0 ) |
| peritoneum   | inflammation                               |                         | <26>        |             |       |       | <29>    |       |       |       | <17>     |       |       |       | <12>     |       |       |       |
|  |  | 0                       | 0           | 1           | 0     | 0     | 0       | 0     | 0     | 1     | 0        | 0     | 0     | 0     | 0        | 0     | 0     | 0     |
|  |  |                         | ( 0 )       | ( 0 )       | ( 4 ) | ( 0 ) | ( 0 )   | ( 0 ) | ( 0 ) | ( 0 ) | ( 6 )    | ( 0 ) | ( 0 ) | ( 0 ) | ( 0 )    | ( 0 ) | ( 0 ) | ( 0 ) |
| Grade  | 1+ : Slight                                | 2+ : Moderate           | 3+ : Marked | 4+ : Severe |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| < a >  | a : Number of animals examined at the site |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| b  | b : Number of animals with lesion          |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| ( c )  | c : b / a * 100                            |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |  |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| (HPT150)   |  |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |
| RAIS   |  |                         |             |             |       |       |         |       |       |       |          |       |       |       |          |       |       |       |

BAIS5

TABLE M 6

HISTOPATHOLOGICAL FINDINGS:

NON-NEOPLASTIC LESIONS:

FEMALE: SACRIFICED ANIMALS

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 12

| Organ                            | Findings                                   | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |             |            |             | 625 ppm<br>21 |            |            |            | 1250 ppm<br>33 |            |            |             | 2500 ppm<br>38 |             |            |            |
|----------------------------------|--|--|---------------|-------------|------------|-------------|---------------|------------|------------|------------|----------------|------------|------------|-------------|----------------|-------------|------------|------------|
|                                  |  |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  |
| (Integumentary system/appandage) |  |  |               |             |            |             |               |            |            |            |                |            |            |             |                |             |            |            |
| skin/app                         | epidermal cyst                             |  | <24>          |             |            |             | <21>          |            |            |            | <33>           |            |            |             | <38>           |             |            |            |
|                                  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )  | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
| (Respiratory system)             |  |  |               |             |            |             |               |            |            |            |                |            |            |             |                |             |            |            |
| nasal cavit                      | eosinophilic change:olfactory epithelium   |  | <24>          |             |            |             | <21>          |            |            |            | <33>           |            |            |             | <38>           |             |            |            |
|                                  |  | 1<br>( 4 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 3<br>( 14 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 4<br>( 11 ) | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |
|                                  | eosinophilic change:respiratory epithelium |  | 14<br>( 58 )  | 6<br>( 25 ) | 0<br>( 0 ) | 0<br>( 0 )  | 13<br>( 62 )  | 1<br>( 5 ) | 0<br>( 0 ) | 0<br>( 0 ) | 21<br>( 64 )   | 3<br>( 9 ) | 0<br>( 0 ) | 0<br>( 0 )  | 28<br>( 74 )   | 4<br>( 11 ) | 1<br>( 3 ) | 0<br>( 0 ) |
|                                  |  | respiratory metaplasia:olfactory epithelium    |               | 1<br>( 4 )  | 0<br>( 0 ) | 0<br>( 0 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 3<br>( 8 )     | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
|                                  | respiratory metaplasia:gland               |  | 4<br>( 17 )   | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 )  | 6<br>( 29 )   | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 6<br>( 18 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )  | 6<br>( 16 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) |
| nasopharynx                      | eosinophilic change                        |  | <24>          |             |            |             | <21>          |            |            |            | <33>           |            |            |             | <38>           |             |            |            |
|                                  |  | 2<br>( 8 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 1<br>( 5 )  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 )  | 0<br>( 0 )     | 0<br>( 0 )  | 0<br>( 0 ) |            |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 13

| Organ_____             | Findings_____                         | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |           | 625 ppm<br>21 |           |           |           | 1250 ppm<br>33 |           |           |           | 2500 ppm<br>38 |           |           |           |
|------------------------|---------------------------------------|--|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                        |                                       |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Respiratory system)   |                                       |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| lung                   |                                       |  | <24>          |           |           |           | <21>          |           |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                        | deposit of amyloid                    |  | 3<br>( 13)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 14)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 18)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 13)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | inflammatory infiltration             |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | lymphocytic infiltration              |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | bronchiolar-alveolar cell hyperplasia |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Hematopoietic system) |                                       |  |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |           |
| bone marrow            |                                       |  | <24>          |           |           |           | <21>          |           |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                        | granulation                           |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 5) | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) |
|                        | increased hematopoiesis               |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 9)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 3<br>( 8)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                        | granulopoiesis:increased              |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 5<br>( 13)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ                  | Findings               | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |           |             | 625 ppm<br>21 |            |             |             | 1250 ppm<br>33 |             |           |             | 2500 ppm<br>38 |             |              |           |
|------------------------|------------------------|--|---------------|------------|-----------|-------------|---------------|------------|-------------|-------------|----------------|-------------|-----------|-------------|----------------|-------------|--------------|-----------|
|                        |                        |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)   | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%)   | 3+<br>(%)    | 4+<br>(%) |
| (Hematopoietic system) |                        |  |               |            |           |             |               |            |             |             |                |             |           |             |                |             |              |           |
| lymph node             | lymphadenitis          |  | <24>          |            |           |             | <21>          |            |             |             | <33>           |             |           |             | <38>           |             |              |           |
|                        |                        | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 1<br>( 3)   | 1<br>( 3)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)      | 0<br>( 0)   | 0<br>( 0)    |           |
| spleen                 | deposit of hemosiderin |  | <24>          |            |           |             | <21>          |            |             |             | <33>           |             |           |             | <38>           |             |              |           |
|                        |                        | 18<br>( 75)                                    | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 18<br>( 86) | 3<br>( 14)    | 0<br>( 0)  | 0 *<br>( 0) | 26<br>( 79) | 4<br>( 12)     | 0<br>( 0)   | 0<br>( 0) | 22<br>( 58) | 10<br>( 26)    | 5<br>( 13)  | 0 **<br>( 0) |           |
|                        | deposit of melanin     |  | 1<br>( 4)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)   | 1<br>( 5)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    |           |
|                        |                        | extramedullary hematopoiesis                   |               | 5<br>( 21) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)     | 9<br>( 43) | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)      | 11<br>( 33) | 3<br>( 9) | 0<br>( 0)   | 0<br>( 0)      | 17<br>( 45) | 9<br>( 24)   | 0<br>( 0) |
|                        | follicular hyperplasia |  |               | 1<br>( 4)  | 1<br>( 4) | 0<br>( 0)   | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 0<br>( 0)   | 0<br>( 0)      | 1<br>( 3)   | 2<br>( 6) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    | 0<br>( 0) |
|                        |                        | (Circulatory system)                           |               |            |           |             |               |            |             |             |                |             |           |             |                |             |              |           |
| heart                  | deposit of amyloid     |  | <24>          |            |           |             | <21>          |            |             |             | <33>           |             |           |             | <38>           |             |              |           |
|                        |                        | 2<br>( 8)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 3<br>( 14)  | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0)   | 6<br>( 18)  | 1<br>( 3)      | 0<br>( 0)   | 0<br>( 0) | 5<br>( 13)  | 0<br>( 0)      | 0<br>( 0)   | 0<br>( 0)    |           |

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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrJ [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ                | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |            | 625 ppm<br>21 |           |           |             | 1250 ppm<br>33 |           |           |             | 2500 ppm<br>38 |           |           |           |
|----------------------|-------------------------------|--|---------------|-----------|-----------|------------|---------------|-----------|-----------|-------------|----------------|-----------|-----------|-------------|----------------|-----------|-----------|-----------|
|                      |                               |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)   | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Circulatory system) |                               |  |               |           |           |            |               |           |           |             |                |           |           |             |                |           |           |           |
| heart                | myocardial fibrosis           |  | <24>          |           |           |            | <21>          |           |           |             | <33>           |           |           |             | <38>           |           |           |           |
|                      |                               | 6<br>( 25)                                     | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 4<br>( 19) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 12<br>( 36) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 12<br>( 32) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | arteritis                     |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 5)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Digestive system)   |                               |  |               |           |           |            |               |           |           |             |                |           |           |             |                |           |           |           |
| stomach              | hyperplasia:forestomach       |  | <24>          |           |           |            | <21>          |           |           |             | <33>           |           |           |             | <38>           |           |           |           |
|                      |                               | 1<br>( 4)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 2<br>( 10) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | erosion:glandular stomach     |  | 1<br>( 4)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 5)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 3<br>( 8)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                      | hyperplasia:glandular stomach |  | 7<br>( 29)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 9<br>( 43)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 11<br>( 33)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 12<br>( 32)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| small intes          | adhesion                      |  | <24>          |           |           |            | <21>          |           |           |             | <33>           |           |           |             | <38>           |           |           |           |
|                      |                               | 0<br>( 0)                                      | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)   | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ              | Findings                 | Group Name<br>No. of Animals on Study<br>Grade |           |           |           | Control<br>24 |           |           |           | 625 ppm<br>21 |           |           |           | 1250 ppm<br>33 |           |           |           | 2500 ppm<br>38 |           |           |             |
|--------------------|--------------------------|--|-----------|-----------|-----------|---------------|-----------|-----------|-----------|---------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-------------|
|                    |                          | 1+   | 2+        | 3+        | 4+        | 1+            | 2+        | 3+        | 4+        | 1+            | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+        | 1+             | 2+        | 3+        | 4+          |
|                    |                          | (%)  | (%)       | (%)       | (%)       | (%)           | (%)       | (%)       | (%)       | (%)           | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)       | (%)            | (%)       | (%)       | (%)         |
| (Digestive system) |                          |  |           |           |           |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |             |
| liver              |                          | <24>   |           |           |           | <21>          |           |           |           | <33>          |           |           |           | <38>           |           |           |           |                |           |           |             |
|                    | angiectasis              | 1<br>( 4)                                      | 0<br>( 8) | 0<br>( 0) | 0<br>( 0) | 1<br>( 5)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)     | 0<br>( 6) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 3<br>( 8) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | necrosis:focal           | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | lymphocytic infiltration | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | inflammatory cell nest   | 13<br>( 54)                                    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 12<br>( 57)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 12<br>( 36)   | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 16<br>( 42)    | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | acidophilic cell focus   | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | biliary cyst             | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |
|                    | deposit of brown pigment | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 9<br>( 24)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0 *<br>( 0) |
| (Urinary system)   |                          |  |           |           |           |               |           |           |           |               |           |           |           |                |           |           |           |                |           |           |             |
| kidney             |                          | <24>   |           |           |           | <21>          |           |           |           | <33>          |           |           |           | <38>           |           |           |           |                |           |           |             |
|                    | cyst                     | 0<br>( 0)                                      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 5)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)   |

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 < a > a : Number of animals examined at the site  
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 ( c ) c : b / a \* 100  
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STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
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HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ_____         | Findings_____            | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |           |           | 625 ppm<br>21 |            |           |            | 1250 ppm<br>33 |            |           |            | 2500 ppm<br>38 |            |           |           |
|--------------------|--------------------------|--|---------------|------------|-----------|-----------|---------------|------------|-----------|------------|----------------|------------|-----------|------------|----------------|------------|-----------|-----------|
|                    |                          |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) |
| (Urinary system)   |                          |  |               |            |           |           |               |            |           |            |                |            |           |            |                |            |           |           |
| kidney             |                          |  | <24>          |            |           |           | <21>          |            |           |            | <33>           |            |           |            | <38>           |            |           |           |
|                    | hyaline droplet          |  | 3<br>( 13)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 4<br>( 12)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 4<br>( 11)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                    | hyaline cast             |  | 1<br>( 4)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                    | lymphocytic infiltration |  | 3<br>( 13)    | 1<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 3<br>( 14)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 2<br>( 5)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                    | osseous metaplasia       |  | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                    | scar                     |  | 1<br>( 4)     | 4<br>( 17) | 0<br>( 0) | 0<br>( 0) | 3<br>( 14)    | 3<br>( 14) | 0<br>( 0) | 0<br>( 0)  | 2<br>( 6)      | 7<br>( 21) | 1<br>( 3) | 0<br>( 0)  | 5<br>( 13)     | 5<br>( 13) | 0<br>( 0) | 0<br>( 0) |
|                    | inflammatory polyp       |  | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 2<br>( 10)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 3<br>( 9)      | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 5<br>( 13)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) |
|                    | hydronephrosis           |  | 0<br>( 0)     | 1<br>( 4)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 2<br>( 10) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 4<br>( 12) | 1<br>( 3) | 0<br>( 0)  | 1<br>( 3)      | 6<br>( 16) | 0<br>( 0) | 0<br>( 0) |
| papillary necrosis |                          | 2<br>( 8)                                      | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 6<br>( 18) | 1<br>( 3)      | 0<br>( 0)  | 0<br>( 0) | 6<br>( 16) | 0<br>( 0)      | 0<br>( 0)  | 0<br>( 0) |           |

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 SACRIFICED ANIMALS (105W)

PAGE : 18

| Organ            | Findings                      | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |            |            |            | 625 ppm<br>21 |            |            |            | 1250 ppm<br>33 |            |            |            | 2500 ppm<br>38 |            |            |            |
|------------------|-------------------------------|--|---------------|------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|------------|
|                  |                               |  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  |
| {Urinary system} |                               |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| kidney           | mineralization:papilla        |  | <24>          |            |            |            | <21>          |            |            |            | <33>           |            |            |            | <38>           |            |            |            |
|                  |                               | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                  | glomerulosclerosis            |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  |                               | regeneration:proximal tubule                   |               | 2<br>( 8 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 1<br>( 5 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 3<br>( 8 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
| urin bladd       | urothelial hyperplasia:pelvis |  | 2<br>( 8 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) |
|                  |                               | lymphocytic infiltration                       |               | <24>       |            |            |               | <21>       |            |            |                | <33>       |            |            |                | <38>       |            |            |
|                  | 0<br>( 0 )                    |  | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |
|                  | {Endocrine system}            |  |               |            |            |            |               |            |            |            |                |            |            |            |                |            |            |            |
| pituitary        | angiectasis                   |  | <24>          |            |            |            | <21>          |            |            |            | <33>           |            |            |            | <38>           |            |            |            |
|                  |                               | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |            |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

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| Organ_____            | Findings_____                          | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |           |           |           | 625 ppm<br>21 |            |           |           | 1250 ppm<br>33 |           |           |           | 2500 ppm<br>38 |           |           |           |
|-----------------------|--|--|---------------|-----------|-----------|-----------|---------------|------------|-----------|-----------|----------------|-----------|-----------|-----------|----------------|-----------|-----------|-----------|
|                       |  |  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| (Endocrine system)    |  |  |               |           |           |           |               |            |           |           |                |           |           |           |                |           |           |           |
| pituitary             | cyst                                   |  | <24>          |           |           |           | <21>          |            |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                       |  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                       | hyperplasia                            |  | 3<br>( 13)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 10)    | 2<br>( 10) | 0<br>( 0) | 0<br>( 0) | 4<br>( 12)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 4<br>( 11)     | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) |
| thyroid               | deposit of brown pigment               |  | <24>          |           |           |           | <21>          |            |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                       |  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 6<br>( 16)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| adrenal               | spindle-cell hyperplasia               |  | <24>          |           |           |           | <21>          |            |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                       |  |  | 4<br>( 17)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 2<br>( 10)    | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 2<br>( 6)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
|                       | fatty change:corticomedullary junction |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 1<br>( 3) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |
| (Reproductive system) |  |  |               |           |           |           |               |            |           |           |                |           |           |           |                |           |           |           |
| ovary                 | thrombus                               |  | <24>          |           |           |           | <21>          |            |           |           | <33>           |           |           |           | <38>           |           |           |           |
|                       |  |  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)     | 1<br>( 5)  | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0) |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105W)

PAGE : 20

|                                  |                                | Group Name<br>No. of Animals on Study | Control<br>24 |            |           |            | 625 ppm<br>21 |           |           |            | 1250 ppm<br>33 |           |           |            | 2500 ppm<br>38 |           |           |           |
|----------------------------------|--------------------------------|---------------------------------------|---------------|------------|-----------|------------|---------------|-----------|-----------|------------|----------------|-----------|-----------|------------|----------------|-----------|-----------|-----------|
| Organ                            | Findings                       | Grade                                 | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%) | 3+<br>(%) | 4+<br>(%) |
| [Reproductive system]            |                                |                                       |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| ovary                            | cyst                           |                                       | <24>          |            |           |            | <21>          |           |           |            | <33>           |           |           |            | <38>           |           |           |           |
|                                  |                                | 3<br>( 13)                            | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 5<br>( 24) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 5<br>( 15) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 7<br>( 18) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| uterus                           | dilatation                     |                                       | <24>          |            |           |            | <21>          |           |           |            | <33>           |           |           |            | <38>           |           |           |           |
|                                  |                                | 0<br>( 0)                             | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 1<br>( 3)      | 0<br>( 0) | 0<br>( 0) |           |
|                                  | cystic endometrial hyperplasia |                                       | 5<br>( 21)    | 4<br>( 17) | 0<br>( 0) | 0<br>( 0)  | 5<br>( 24)    | 0<br>( 0) | 0<br>( 0) | 0<br>( 0)  | 10<br>( 30)    | 1<br>( 3) | 0<br>( 0) | 0<br>( 0)  | 10<br>( 26)    | 2<br>( 5) | 0<br>( 0) | 0<br>( 0) |
| [Nervous system]                 |                                |                                       |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| brain                            | mineralization                 |                                       | <24>          |            |           |            | <21>          |           |           |            | <33>           |           |           |            | <38>           |           |           |           |
|                                  |                                | 8<br>( 33)                            | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 5<br>( 24) | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 9<br>( 27) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 8<br>( 21) | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |
| [Special sense organs/appendage] |                                |                                       |               |            |           |            |               |           |           |            |                |           |           |            |                |           |           |           |
| eye                              | keratitis                      |                                       | <24>          |            |           |            | <21>          |           |           |            | <33>           |           |           |            | <38>           |           |           |           |
|                                  |                                | 0<br>( 0)                             | 0<br>( 0)     | 0<br>( 0)  | 0<br>( 0) | 0<br>( 0)  | 0<br>( 0)     | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) | 1<br>( 3)  | 0<br>( 0)      | 0<br>( 0) | 0<br>( 0) |           |

Grade 1+ : Slight 2+ : Moderate 3+ : Marked 4+ : Severe  
 < a > a : Number of animals examined at the site  
 b : Number of animals with lesion  
 ( c ) c : b / a \* 100  
 Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 SACRIFICED ANIMALS (105%)

PAGE : 21

| Organ  | Findings                                   | Group Name<br>No. of Animals on Study<br>Grade | Control<br>24 |             |            |            | 625 ppm<br>21 |            |            |            | 1250 ppm<br>33 |            |            |            | 2500 ppm<br>38 |            |            |           |
|--|--|--|---------------|-------------|------------|------------|---------------|------------|------------|------------|----------------|------------|------------|------------|----------------|------------|------------|-----------|
|  |  |  | 1+<br>(%)     | 2+<br>(%)   | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)     | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%)  | 1+<br>(%)      | 2+<br>(%)  | 3+<br>(%)  | 4+<br>(%) |
| (Special sense organs/appendage)                                       |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| Harder gl  | lymphocytic infiltration                   |  | <24>          |             |            |            | <21>          |            |            |            | <33>           |            |            |            | <38>           |            |            |           |
|  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 1<br>( 5 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 5 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
|  | hyperplasia                                |  | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 1<br>( 3 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
| (Musculoskeletal system)   |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| bone   | osteosclerosis                             |  | <24>          |             |            |            | <21>          |            |            |            | <33>           |            |            |            | <38>           |            |            |           |
|  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 2<br>( 6 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
| (Body cavities)  |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| mediastinum  | inflammatory infiltration                  |  | <24>          |             |            |            | <21>          |            |            |            | <33>           |            |            |            | <38>           |            |            |           |
|  |  | 0<br>( 0 )                                     | 0<br>( 0 )    | 0<br>( 0 )  | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )    | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) | 1<br>( 3 ) | 0<br>( 0 )     | 0<br>( 0 ) | 0<br>( 0 ) |           |
| Grade  | 1+ : Slight                                | 2+ : Moderate                                  | 3+ : Marked   | 4+ : Severe |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| < a >  | a : Number of animals examined at the site |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| b  | b : Number of animals with lesion          |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| ( c )  | c : b / a * 100                            |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |
| Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square |  |  |               |             |            |            |               |            |            |            |                |            |            |            |                |            |            |           |



TABLE N 1

NUMBER OF ANIMALS WITH TUMORS AND  
NUMBER OF TUMORS-TIME RELATED: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

PAGE : 1

| Time-related Weeks | Items                               | Group Name | Control | 625 ppm | 1250 ppm | 2500 ppm |
|--------------------|-------------------------------------|------------|---------|---------|----------|----------|
| 0 - 52             | NO. OF EXAMINED ANIMALS             |            | 1       | 3       | 1        | 1        |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 1       | 0       | 0        | 0        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 1       | 0       | 0        | 0        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 0       | 0       | 0        | 0        |
|                    | NO. OF BENIGN TUMORS                |            | 0       | 0       | 0        | 0        |
|                    | NO. OF MALIGNANT TUMORS             |            | 1       | 0       | 0        | 0        |
|                    | NO. OF TOTAL TUMORS                 |            | 1       | 0       | 0        | 0        |
| 53 - 78            | NO. OF EXAMINED ANIMALS             |            | 4       | 4       | 4        | 1        |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 2       | 2       | 1        | 0        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 2       | 2       | 1        | 0        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 0       | 0       | 0        | 0        |
|                    | NO. OF BENIGN TUMORS                |            | 0       | 1       | 0        | 0        |
|                    | NO. OF MALIGNANT TUMORS             |            | 2       | 1       | 1        | 0        |
|                    | NO. OF TOTAL TUMORS                 |            | 2       | 2       | 1        | 0        |
| 79 - 104           | NO. OF EXAMINED ANIMALS             |            | 21      | 11      | 14       | 9        |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 18      | 6       | 11       | 7        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 13      | 6       | 10       | 6        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 5       | 0       | 1        | 1        |
|                    | NO. OF BENIGN TUMORS                |            | 8       | 1       | 1        | 3        |
|                    | NO. OF MALIGNANT TUMORS             |            | 18      | 5       | 11       | 5        |
|                    | NO. OF TOTAL TUMORS                 |            | 26      | 6       | 12       | 8        |
| 105 - 105          | NO. OF EXAMINED ANIMALS             |            | 24      | 32      | 31       | 39       |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 19      | 27      | 25       | 17       |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 7       | 15      | 18       | 15       |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 12      | 12      | 7        | 2        |
|                    | NO. OF BENIGN TUMORS                |            | 20      | 18      | 20       | 6        |
|                    | NO. OF MALIGNANT TUMORS             |            | 14      | 21      | 13       | 13       |
|                    | NO. OF TOTAL TUMORS                 |            | 34      | 39      | 33       | 19       |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

PAGE : 2

| Time-related<br>Weeks | Items                               | Group Name | Control | 625 ppm | 1250 ppm | 2500 ppm |
|-----------------------|-------------------------------------|------------|---------|---------|----------|----------|
| 0 - 105               | NO. OF EXAMINED ANIMALS             |            | 50      | 50      | 50       | 50       |
|                       | NO. OF ANIMALS WITH TUMORS          |            | 40      | 35      | 37       | 24       |
|                       | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 23      | 23      | 29       | 21       |
|                       | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 17      | 12      | 8        | 3        |
|                       | NO. OF BENIGN TUMORS                |            | 28      | 20      | 21       | 9        |
|                       | NO. OF MALIGNANT TUMORS             |            | 35      | 27      | 25       | 18       |
|                       | NO. OF TOTAL TUMORS                 |            | 63      | 47      | 46       | 27       |

(HPT070)

BA1S5

TABLE N 2

NUMBER OF ANIMALS WITH TUMORS AND  
NUMBER OF TUMORS-TIME RELATED: FEMALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

PAGE : 3

| Time-related Weeks | Items                               | Group Name | Control | 625 ppm | 1250 ppm | 2500 ppm |
|--------------------|-------------------------------------|------------|---------|---------|----------|----------|
| 0 - 52             | NO. OF EXAMINED ANIMALS             |            | 1       | 1       | 0        | 0        |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 0       | 1       | 0        | 0        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 0       | 1       | 0        | 0        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 0       | 0       | 0        | 0        |
|                    | NO. OF BENIGN TUMORS                |            | 0       | 0       | 0        | 0        |
|                    | NO. OF MALIGNANT TUMORS             |            | 0       | 1       | 0        | 0        |
|                    | NO. OF TOTAL TUMORS                 |            | 0       | 1       | 0        | 0        |
| 53 - 78            | NO. OF EXAMINED ANIMALS             |            | 10      | 2       | 8        | 2        |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 10      | 2       | 5        | 1        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 10      | 2       | 5        | 0        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 0       | 0       | 0        | 1        |
|                    | NO. OF BENIGN TUMORS                |            | 0       | 0       | 0        | 1        |
|                    | NO. OF MALIGNANT TUMORS             |            | 10      | 2       | 5        | 1        |
|                    | NO. OF TOTAL TUMORS                 |            | 10      | 2       | 5        | 2        |
| 79 - 104           | NO. OF EXAMINED ANIMALS             |            | 15      | 26      | 9        | 10       |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 14      | 25      | 9        | 9        |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 10      | 14      | 7        | 4        |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 4       | 11      | 2        | 5        |
|                    | NO. OF BENIGN TUMORS                |            | 5       | 16      | 3        | 4        |
|                    | NO. OF MALIGNANT TUMORS             |            | 15      | 26      | 8        | 13       |
|                    | NO. OF TOTAL TUMORS                 |            | 20      | 42      | 11       | 17       |
| 105 - 105          | NO. OF EXAMINED ANIMALS             |            | 24      | 21      | 33       | 38       |
|                    | NO. OF ANIMALS WITH TUMORS          |            | 19      | 15      | 28       | 24       |
|                    | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 13      | 9       | 15       | 16       |
|                    | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 6       | 6       | 13       | 8        |
|                    | NO. OF BENIGN TUMORS                |            | 13      | 11      | 23       | 20       |
|                    | NO. OF MALIGNANT TUMORS             |            | 15      | 15      | 24       | 18       |
|                    | NO. OF TOTAL TUMORS                 |            | 28      | 26      | 47       | 38       |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

PAGE : 4

| Time-related<br>Weeks | Items                               | Group Name | Control | 625 ppm | 1250 ppm | 2500 ppm |
|-----------------------|-------------------------------------|------------|---------|---------|----------|----------|
| 0 - 105               | NO. OF EXAMINED ANIMALS             |            | 50      | 50      | 50       | 50       |
|                       | NO. OF ANIMALS WITH TUMORS          |            | 43      | 43      | 42       | 34       |
|                       | NO. OF ANIMALS WITH SINGLE TUMORS   |            | 33      | 26      | 27       | 20       |
|                       | NO. OF ANIMALS WITH MULTIPLE TUMORS |            | 10      | 17      | 15       | 14       |
|                       | NO. OF BENIGN TUMORS                |            | 18      | 27      | 26       | 25       |
|                       | NO. OF MALIGNANT TUMORS             |            | 40      | 44      | 37       | 32       |
|                       | NO. OF TOTAL TUMORS                 |            | 58      | 71      | 63       | 57       |

(HPT070)

BAIS5

TABLE O 1

HISTOPATHOLOGICAL FINDINGS:

NEOPLASTIC LESIONS: MALE

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 1

| Organ                            | Findings                       | Group Name<br>No. of animals on Study | Control<br>50    | 625 ppm<br>50   | 1250 ppm<br>50   | 2500 ppm<br>50   |
|----------------------------------|--------------------------------|---------------------------------------|------------------|-----------------|------------------|------------------|
| (Integumentary system/appandage) |                                |                                       |                  |                 |                  |                  |
| subcutis                         | histiocytic sarcoma            |                                       | <50><br>0 ( 0%)  | <50><br>1 ( 2%) | <50><br>1 ( 2%)  | <50><br>1 ( 2%)  |
| (Respiratory system)             |                                |                                       |                  |                 |                  |                  |
| lung                             | bronchiolar-alveolar adenoma   |                                       | <50><br>6 ( 12%) | <50><br>2 ( 4%) | <50><br>3 ( 6%)  | <50><br>6 ( 12%) |
|                                  | bronchiolar-alveolar carcinoma |                                       | 10 ( 20%)        | 9 ( 18%)        | 2 ( 4%)          | 3 ( 6%)          |
| (Hematopoietic system)           |                                |                                       |                  |                 |                  |                  |
| lymph node                       | malignant lymphoma             |                                       | <50><br>7 ( 14%) | <50><br>4 ( 8%) | <50><br>9 ( 18%) | <50><br>6 ( 12%) |
|                                  | mastcytoma:malignant           |                                       | 1 ( 2%)          | 1 ( 2%)         | 0 ( 0%)          | 0 ( 0%)          |
| spleen                           | mastcytoma:benign              |                                       | <50><br>0 ( 0%)  | <50><br>1 ( 2%) | <50><br>1 ( 2%)  | <50><br>0 ( 0%)  |
|                                  | hemangioma                     |                                       | 1 ( 2%)          | 3 ( 6%)         | 1 ( 2%)          | 1 ( 2%)          |
|                                  | malignant lymphoma             |                                       | 0 ( 0%)          | 0 ( 0%)         | 1 ( 2%)          | 1 ( 2%)          |
|                                  | hemangiosarcoma                |                                       | 0 ( 0%)          | 1 ( 2%)         | 0 ( 0%)          | 0 ( 0%)          |
| (Digestive system)               |                                |                                       |                  |                 |                  |                  |
| liver                            | hemangioma                     |                                       | <50><br>2 ( 4%)  | <50><br>3 ( 6%) | <50><br>3 ( 6%)  | <50><br>1 ( 2%)  |

< a > a : Number of animals examined at the site  
 b ( c ) b : Number of animals with neoplasm c : b / a \* 100



STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crj[Crl:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 2

| Organ                            | Findings                 | Group Name<br>No. of animals on Study | Control<br>50     | 625 ppm<br>50    | 1250 ppm<br>50   | 2500 ppm<br>50  |
|----------------------------------|--------------------------|---------------------------------------|-------------------|------------------|------------------|-----------------|
| (Digestive system)               |                          |                                       |                   |                  |                  |                 |
| liver                            | hepatocellular adenoma   |                                       | <50><br>16 ( 32%) | <50><br>8 ( 16%) | <50><br>9 ( 18%) | <50><br>1 ( 2%) |
|                                  | histiocytic sarcoma      |                                       | 6 ( 12%)          | 2 ( 4%)          | 4 ( 8%)          | 1 ( 2%)         |
|                                  | hemangiosarcoma          |                                       | 0 ( 0%)           | 1 ( 2%)          | 0 ( 0%)          | 1 ( 2%)         |
|                                  | hepatocellular carcinoma |                                       | 7 ( 14%)          | 6 ( 12%)         | 5 ( 10%)         | 2 ( 4%)         |
|                                  | hepatoblastoma           |                                       | 1 ( 2%)           | 0 ( 0%)          | 0 ( 0%)          | 0 ( 0%)         |
| (Urinary system)                 |                          |                                       |                   |                  |                  |                 |
| urin bladd                       | histiocytic sarcoma      |                                       | <50><br>1 ( 2%)   | <50><br>0 ( 0%)  | <50><br>0 ( 0%)  | <50><br>0 ( 0%) |
| (Endocrine system)               |                          |                                       |                   |                  |                  |                 |
| adrenal                          | pheochromocytoma         |                                       | <50><br>1 ( 2%)   | <50><br>0 ( 0%)  | <50><br>0 ( 0%)  | <50><br>0 ( 0%) |
| (Reproductive system)            |                          |                                       |                   |                  |                  |                 |
| epididymis                       | histiocytic sarcoma      |                                       | <50><br>0 ( 0%)   | <50><br>2 ( 4%)  | <50><br>1 ( 2%)  | <50><br>2 ( 4%) |
| (Special sense organs/appendage) |                          |                                       |                   |                  |                  |                 |
| Harder gl                        | adenoma                  |                                       | <50><br>2 ( 4%)   | <50><br>3 ( 6%)  | <50><br>4 ( 8%)  | <50><br>0 ( 0%) |

< a > a : Number of animals examined at the site  
b ( c ) b : Number of animals with neoplasm c : b / a \* 100

(HPT085)

BA1S5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 3

| Organ                            | Findings           | Group Name<br>No. of animals on Study | Control<br>50   | 625 ppm<br>50   | 1250 ppm<br>50  | 2500 ppm<br>50  |
|----------------------------------|--------------------|---------------------------------------|-----------------|-----------------|-----------------|-----------------|
| (Special sense organs/appendage) |                    |                                       |                 |                 |                 |                 |
| Harder gl                        | adenocarcinoma     |                                       | <50><br>1 ( 2%) | <50><br>0 ( 0%) | <50><br>0 ( 0%) | <50><br>0 ( 0%) |
| (Musculoskeletal system)         |                    |                                       |                 |                 |                 |                 |
| muscle                           | leiomyosarcoma     |                                       | <50><br>0 ( 0%) | <50><br>0 ( 0%) | <50><br>1 ( 2%) | <50><br>0 ( 0%) |
|                                  | hemangiosarcoma    |                                       | 0 ( 0%)         | 0 ( 0%)         | 0 ( 0%)         | 1 ( 2%)         |
| vertebra                         | chordoma:malignant |                                       | <50><br>0 ( 0%) | <50><br>0 ( 0%) | <50><br>1 ( 2%) | <50><br>0 ( 0%) |
| (Body cavities)                  |                    |                                       |                 |                 |                 |                 |
| peritoneum                       | leiomyosarcoma     |                                       | <50><br>1 ( 2%) | <50><br>0 ( 0%) | <50><br>0 ( 0%) | <50><br>0 ( 0%) |

< a > a : Number of animals examined at the site  
 b ( c ) b : Number of animals with neoplasm c : b / a \* 100

(HPT085)

BAIS5

**TABLE O 2**

**HISTOPATHOLOGICAL FINDINGS:**

**NEOPLASTIC LESIONS: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 4

| Organ                            | Findings                       | Group Name<br>No. of animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|----------------------------------|--------------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Integumentary system/appandage) |                                |                                       |               |               |                |                |
| skin/app                         |                                |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | trichoepithelioma:malignant    |                                       | 0 ( 0%)       | 0 ( 0%)       | 0 ( 0%)        | 1 ( 2%)        |
| subcutis                         |                                |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | fibroma                        |                                       | 0 ( 0%)       | 0 ( 0%)       | 1 ( 2%)        | 0 ( 0%)        |
|                                  | hemangioma                     |                                       | 0 ( 0%)       | 1 ( 2%)       | 0 ( 0%)        | 0 ( 0%)        |
|                                  | fibrosarcoma                   |                                       | 0 ( 0%)       | 2 ( 4%)       | 0 ( 0%)        | 1 ( 2%)        |
|                                  | mastcytoma:malignant           |                                       | 0 ( 0%)       | 1 ( 2%)       | 0 ( 0%)        | 0 ( 0%)        |
| (Respiratory system)             |                                |                                       |               |               |                |                |
| lung                             |                                |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | bronchiolar-alveolar adenoma   |                                       | 2 ( 4%)       | 3 ( 6%)       | 1 ( 2%)        | 1 ( 2%)        |
|                                  | hemangioma                     |                                       | 0 ( 0%)       | 1 ( 2%)       | 0 ( 0%)        | 0 ( 0%)        |
|                                  | bronchiolar-alveolar carcinoma |                                       | 1 ( 2%)       | 4 ( 8%)       | 1 ( 2%)        | 2 ( 4%)        |
| (Hematopoietic system)           |                                |                                       |               |               |                |                |
| lymph node                       |                                |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | malignant lymphoma             |                                       | 18 ( 36%)     | 17 ( 34%)     | 21 ( 42%)      | 10 ( 20%)      |
| spleen                           |                                |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | hemangioma                     |                                       | 0 ( 0%)       | 0 ( 0%)       | 1 ( 2%)        | 0 ( 0%)        |
|                                  | mastcytoma:malignant           |                                       | 0 ( 0%)       | 0 ( 0%)       | 0 ( 0%)        | 1 ( 2%)        |

< a > a : Number of animals examined at the site  
 b ( c ) b : Number of animals with neoplasm c : b / a \* 100

(HPT085)

BAIS5

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 5

| Organ                 | Findings                 | Group Name<br>No. of animals on Study | Control<br>50   | 625 ppm<br>50    | 1250 ppm<br>50   | 2500 ppm<br>50   |
|-----------------------|--------------------------|---------------------------------------|-----------------|------------------|------------------|------------------|
| (Digestive system)    |                          |                                       |                 |                  |                  |                  |
| stomach               | squamous cell papilloma  |                                       | <50><br>1 ( 2%) | <50><br>1 ( 2%)  | <50><br>0 ( 0%)  | <50><br>0 ( 0%)  |
| liver                 | hemangioma               |                                       | <50><br>1 ( 2%) | <50><br>1 ( 2%)  | <50><br>1 ( 2%)  | <50><br>3 ( 6%)  |
|                       | hepatocellular adenoma   |                                       | 4 ( 8%)         | 6 ( 12%)         | 4 ( 8%)          | 5 ( 10%)         |
|                       | liposarcoma              |                                       | 1 ( 2%)         | 0 ( 0%)          | 0 ( 0%)          | 0 ( 0%)          |
|                       | histiocytic sarcoma      |                                       | 2 ( 4%)         | 3 ( 6%)          | 0 ( 0%)          | 1 ( 2%)          |
|                       | hemangiosarcoma          |                                       | 0 ( 0%)         | 1 ( 2%)          | 0 ( 0%)          | 0 ( 0%)          |
|                       | hepatocellular carcinoma |                                       | 0 ( 0%)         | 0 ( 0%)          | 1 ( 2%)          | 0 ( 0%)          |
| gall bladd            | papillary adenoma        |                                       | <50><br>0 ( 0%) | <50><br>0 ( 0%)  | <50><br>0 ( 0%)  | <50><br>1 ( 2%)  |
| (Endocrine system)    |                          |                                       |                 |                  |                  |                  |
| pituitary             | adenoma                  |                                       | <50><br>4 ( 8%) | <50><br>9 ( 18%) | <50><br>6 ( 12%) | <50><br>7 ( 14%) |
| (Reproductive system) |                          |                                       |                 |                  |                  |                  |
| ovary                 | hemangioma               |                                       | <50><br>0 ( 0%) | <50><br>2 ( 4%)  | <50><br>2 ( 4%)  | <50><br>1 ( 2%)  |
|                       | papillary adenoma        |                                       | 1 ( 2%)         | 0 ( 0%)          | 4 ( 8%)          | 1 ( 2%)          |

< a > a : Number of animals examined at the site  
b ( c ) b : Number of animals with neoplasm c : b / a \* 100

(HPT085)

BAIS5

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 6

| Organ                            | Findings                  | Group Name<br>No. of animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|----------------------------------|---------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Reproductive system)            |                           |                                       |               |               |                |                |
| uterus                           |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leiomyoma                 |                                       | 1 ( 2%)       | 1 ( 2%)       | 0 ( 0%)        | 0 ( 0%)        |
|                                  | endometrial stromal polyp |                                       | 3 ( 6%)       | 2 ( 4%)       | 2 ( 4%)        | 2 ( 4%)        |
|                                  | histiocytic sarcoma       |                                       | 16 ( 32%)     | 14 ( 28%)     | 12 ( 24%)      | 12 ( 24%)      |
| mammary gl                       |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | adenocarcinoma            |                                       | 0 ( 0%)       | 1 ( 2%)       | 0 ( 0%)        | 3 ( 6%)        |
|                                  | adenosquamous carcinoma   |                                       | 0 ( 0%)       | 1 ( 2%)       | 0 ( 0%)        | 1 ( 2%)        |
| (Special sense organs/appendage) |                           |                                       |               |               |                |                |
| Harder gl                        |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | adenoma                   |                                       | 1 ( 2%)       | 0 ( 0%)       | 4 ( 8%)        | 3 ( 6%)        |
|                                  | adenocarcinoma            |                                       | 1 ( 2%)       | 0 ( 0%)       | 0 ( 0%)        | 0 ( 0%)        |
| (Musculoskeletal system)         |                           |                                       |               |               |                |                |
| muscle                           |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leiomyosarcoma            |                                       | 1 ( 2%)       | 0 ( 0%)       | 0 ( 0%)        | 0 ( 0%)        |
| (Body cavities)                  |                           |                                       |               |               |                |                |
| mediastinum                      |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | histiocytic sarcoma       |                                       | 0 ( 0%)       | 0 ( 0%)       | 1 ( 2%)        | 0 ( 0%)        |
| retroperit                       |                           |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | fibroma                   |                                       | 0 ( 0%)       | 0 ( 0%)       | 0 ( 0%)        | 1 ( 2%)        |

< a > a : Number of animals examined at the site  
 b ( c ) b : Number of animals with neoplasm c : b / a \* 100

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 7

| Organ_____      | Findings_____  | Group Name              | Control | 625 ppm | 1250 ppm | 2500 ppm |
|-----------------|----------------|-------------------------|---------|---------|----------|----------|
|                 |                | No. of animals on Study | 50      | 50      | 50       | 50       |
| <hr/>           |                |                         |         |         |          |          |
| {Body cavities} |                |                         |         |         |          |          |
| retroperit      |                |                         | <50>    | <50>    | <50>     | <50>     |
|                 | leiomyosarcoma |                         | 0 ( 0%) | 0 ( 0%) | 1 ( 2%)  | 0 ( 0%)  |

< a > a : Number of animals examined at the site  
b ( c ) b : Number of animals with neoplasm c : b / a \* 100

(HPT085)

BAIS5

TABLE P 1

NEOPLASTIC LESIONS-INCIDENCE AND  
STATISTICAL ANALYSIS: MALE



STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 SEX : MALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 1

| Group Name  | Control       | 625 ppm       | 1250 ppm     | 2500 ppm     |
|---|---------------|---------------|--------------|--------------|
| SITE : lung<br>TUMOR : bronchiolar-alveolar adenoma                                 |               |               |              |              |
| Tumor rate  |               |               |              |              |
| Overall rates (a)   | 6/50 ( 12.0)  | 2/50 ( 4.0)   | 3/50 ( 6.0)  | 6/50 ( 12.0) |
| Adjusted rates (b)  | 19.23         | 6.25          | 7.32         | 13.04        |
| Terminal rates (c)  | 4/24 ( 16.7)  | 2/32 ( 6.3)   | 2/31 ( 6.5)  | 4/39 ( 10.3) |
| Statistical analysis  |               |               |              |              |
| Peto test   |               |               |              |              |
| Standard method (d)   | P = -----     |               |              |              |
| Prevalence method (d)   | P = 0.4977    |               |              |              |
| Combined analysis (d)   | P = -----     |               |              |              |
| Cochran-Armitage test (e)   | P = 0.6997    |               |              |              |
| Fisher Exact test (e)   |               | P = 0.1343    | P = 0.2435   | P = 0.6202   |
| SITE : lung<br>TUMOR : bronchiolar-alveolar carcinoma                               |               |               |              |              |
| Tumor rate  |               |               |              |              |
| Overall rates (a)   | 10/50 ( 20.0) | 9/50 ( 18.0)  | 2/50 ( 4.0)  | 3/50 ( 6.0)  |
| Adjusted rates (b)  | 20.83         | 25.71         | 6.45         | 5.13         |
| Terminal rates (c)  | 5/24 ( 20.8)  | 8/32 ( 25.0)  | 2/31 ( 6.5)  | 2/39 ( 5.1)  |
| Statistical analysis  |               |               |              |              |
| Peto test   |               |               |              |              |
| Standard method (d)   | P = 0.8551    |               |              |              |
| Prevalence method (d)   | P = 0.9978    |               |              |              |
| Combined analysis (d)   | P = 0.9989    |               |              |              |
| Cochran-Armitage test (e)   | P = 0.0124*   |               |              |              |
| Fisher Exact test (e)   |               | P = 0.5000    | P = 0.0139*  | P = 0.0357*  |
| SITE : lung<br>TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar carcinoma |               |               |              |              |
| Tumor rate  |               |               |              |              |
| Overall rates (a)   | 16/50 ( 32.0) | 10/50 ( 20.0) | 5/50 ( 10.0) | 9/50 ( 18.0) |
| Adjusted rates (b)  | 38.46         | 28.57         | 12.90        | 17.39        |
| Terminal rates (c)  | 9/24 ( 37.5)  | 9/32 ( 28.1)  | 4/31 ( 12.9) | 6/39 ( 15.4) |
| Statistical analysis  |               |               |              |              |
| Peto test   |               |               |              |              |
| Standard method (d)   | P = 0.8551    |               |              |              |
| Prevalence method (d)   | P = 0.9753    |               |              |              |
| Combined analysis (d)   | P = 0.9872    |               |              |              |
| Cochran-Armitage test (e)   | P = 0.0943    |               |              |              |
| Fisher Exact test (e)   |               | P = 0.1271    | P = 0.0064** | P = 0.0826   |

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 2

| Group Name   | Control      | 625 ppm      | 1250 ppm     | 2500 ppm     |
|--|--------------|--------------|--------------|--------------|
| SITE : lymph node<br>TUMOR : malignant lymphoma      |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                    | 7/50 ( 14.0) | 4/50 ( 8.0)  | 9/50 ( 18.0) | 6/50 ( 12.0) |
| Adjusted rates (b)                                   | 16.67        | 9.38         | 16.13        | 12.82        |
| Terminal rates (c)                                   | 4/24 ( 16.7) | 3/32 ( 9.4)  | 5/31 ( 16.1) | 5/39 ( 12.8) |
| Statistical analysis                                 |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                  | P = 0.8199   |              |              |              |
| Prevalence method (d)                                | P = 0.5467   |              |              |              |
| Combined analysis (d)                                | P = 0.7459   |              |              |              |
| Cochran-Armitage test (e)                            | P = 0.9433   |              |              |              |
| Fisher Exact test (e)                                |              | P = 0.2623   | P = 0.3929   | P = 0.5000   |
| SITE : spleen<br>TUMOR : hemangioma                  |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                    | 1/50 ( 2.0)  | 3/50 ( 6.0)  | 1/50 ( 2.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)                                   | 3.33         | 9.38         | 3.23         | 2.56         |
| Terminal rates (c)                                   | 0/24 ( 0.0)  | 3/32 ( 9.4)  | 1/31 ( 3.2)  | 1/39 ( 2.6)  |
| Statistical analysis                                 |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                  | P = -----    |              |              |              |
| Prevalence method (d)                                | P = 0.7576   |              |              |              |
| Combined analysis (d)                                | P = -----    |              |              |              |
| Cochran-Armitage test (e)                            | P = 0.6742   |              |              |              |
| Fisher Exact test (e)                                |              | P = 0.3087   | P = 0.7525   | P = 0.7525   |
| SITE : spleen<br>TUMOR : hemangioma, hemangiosarcoma |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                    | 1/50 ( 2.0)  | 4/50 ( 8.0)  | 1/50 ( 2.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)                                   | 3.33         | 12.50        | 3.23         | 2.56         |
| Terminal rates (c)                                   | 0/24 ( 0.0)  | 4/32 ( 12.5) | 1/31 ( 3.2)  | 1/39 ( 2.6)  |
| Statistical analysis                                 |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                  | P = -----    |              |              |              |
| Prevalence method (d)                                | P = 0.8184   |              |              |              |
| Combined analysis (d)                                | P = -----    |              |              |              |
| Cochran-Armitage test (e)                            | P = 0.5583   |              |              |              |
| Fisher Exact test (e)                                |              | P = 0.1811   | P = 0.7525   | P = 0.7525   |

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 3

| Group Name                                     | Control       | 625 ppm      | 1250 ppm     | 2500 ppm     |
|--|---------------|--------------|--------------|--------------|
| SITE : liver<br>TUMOR : hemangioma             |               |              |              |              |
| Tumor rate                                     |               |              |              |              |
| Overall rates (a)                              | 2/50 ( 4.0)   | 3/50 ( 6.0)  | 3/50 ( 6.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)                             | 4.17          | 9.38         | 9.68         | 2.56         |
| Terminal rates (c)                             | 1/24 ( 4.2)   | 3/32 ( 9.4)  | 3/31 ( 9.7)  | 1/39 ( 2.6)  |
| Statistical analysis                           |               |              |              |              |
| Peto test                                      |               |              |              |              |
| Standard method (d)                            | P = 0.9629 ?  |              |              |              |
| Prevalence method (d)                          | P = 0.7503    |              |              |              |
| Combined analysis (d)                          | P = 0.8614    |              |              |              |
| Cochran-Armitage test (e)                      | P = 0.5259    |              |              |              |
| Fisher Exact test (e)                          |               | P = 0.5000   | P = 0.5000   | P = 0.5000   |
| SITE : liver<br>TUMOR : hepatocellular adenoma |               |              |              |              |
| Tumor rate                                     |               |              |              |              |
| Overall rates (a)                              | 16/50 ( 32.0) | 8/50 ( 16.0) | 9/50 ( 18.0) | 1/50 ( 2.0)  |
| Adjusted rates (b)                             | 50.00         | 21.21        | 29.03        | 2.13         |
| Terminal rates (c)                             | 12/24 ( 50.0) | 6/32 ( 18.8) | 9/31 ( 29.0) | 0/39 ( 0.0)  |
| Statistical analysis                           |               |              |              |              |
| Peto test                                      |               |              |              |              |
| Standard method (d)                            | P = -----     |              |              |              |
| Prevalence method (d)                          | P = 1.0000    |              |              |              |
| Combined analysis (d)                          | P = -----     |              |              |              |
| Cochran-Armitage test (e)                      | P = 0.0002**  |              |              |              |
| Fisher Exact test (e)                          |               | P = 0.0500   | P = 0.0826   | P < 0.0001** |
| SITE : liver<br>TUMOR : histiocytic sarcoma    |               |              |              |              |
| Tumor rate                                     |               |              |              |              |
| Overall rates (a)                              | 6/50 ( 12.0)  | 2/50 ( 4.0)  | 4/50 ( 8.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)                             | 0.0           | 0.0          | 0.0          | 2.56         |
| Terminal rates (c)                             | 0/24 ( 0.0)   | 0/32 ( 0.0)  | 0/31 ( 0.0)  | 1/39 ( 2.6)  |
| Statistical analysis                           |               |              |              |              |
| Peto test                                      |               |              |              |              |
| Standard method (d)                            | P = 0.9956    |              |              |              |
| Prevalence method (d)                          | P = 0.1404    |              |              |              |
| Combined analysis (d)                          | P = 0.9804    |              |              |              |
| Cochran-Armitage test (e)                      | P = 0.0897    |              |              |              |
| Fisher Exact test (e)                          |               | P = 0.1343   | P = 0.3703   | P = 0.0559   |

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 4

| Group Name   | Control       | 625 ppm       | 1250 ppm      | 2500 ppm     |
|--|---------------|---------------|---------------|--------------|
| SITE : liver<br>TUMOR : hepatocellular carcinoma                         |               |               |               |              |
| Tumor rate   |               |               |               |              |
| Overall rates (a)  | 7/50 ( 14.0)  | 6/50 ( 12.0)  | 5/50 ( 10.0)  | 2/50 ( 4.0)  |
| Adjusted rates (b)   | 16.67         | 18.75         | 12.90         | 2.56         |
| Terminal rates (c)   | 4/24 ( 16.7)  | 6/32 ( 18.8)  | 4/31 ( 12.9)  | 1/39 ( 2.6)  |
| Statistical analysis   |               |               |               |              |
| Peto test  |               |               |               |              |
| Standard method (d)  | P = 0.6956    |               |               |              |
| Prevalence method (d)  | P = 0.9914    |               |               |              |
| Combined analysis (d)  | P = 0.9910    |               |               |              |
| Cochran-Armitage test (e)  | P = 0.0796    |               |               |              |
| Fisher Exact test (e)  |               | P = 0.5000    | P = 0.3798    | P = 0.0798   |
| SITE : liver<br>TUMOR : hemangioma, hemangiosarcoma                      |               |               |               |              |
| Tumor rate   |               |               |               |              |
| Overall rates (a)  | 2/50 ( 4.0)   | 4/50 ( 8.0)   | 3/50 ( 6.0)   | 2/50 ( 4.0)  |
| Adjusted rates (b)   | 4.17          | 12.50         | 9.68          | 2.56         |
| Terminal rates (c)   | 1/24 ( 4.2)   | 4/32 ( 12.5)  | 3/31 ( 9.7)   | 1/39 ( 2.6)  |
| Statistical analysis   |               |               |               |              |
| Peto test  |               |               |               |              |
| Standard method (d)  | P = 0.4556    |               |               |              |
| Prevalence method (d)  | P = 0.8076    |               |               |              |
| Combined analysis (d)  | P = 0.7729    |               |               |              |
| Cochran-Armitage test (e)  | P = 0.7932    |               |               |              |
| Fisher Exact test (e)  |               | P = 0.3389    | P = 0.5000    | P = 0.6913   |
| SITE : liver<br>TUMOR : hepatocellular adenoma, hepatocellular carcinoma |               |               |               |              |
| Tumor rate   |               |               |               |              |
| Overall rates (a)  | 21/50 ( 42.0) | 13/50 ( 26.0) | 12/50 ( 24.0) | 3/50 ( 6.0)  |
| Adjusted rates (b)   | 58.33         | 36.36         | 35.48         | 4.35         |
| Terminal rates (c)   | 14/24 ( 58.3) | 11/32 ( 34.4) | 11/31 ( 35.5) | 1/39 ( 2.6)  |
| Statistical analysis   |               |               |               |              |
| Peto test  |               |               |               |              |
| Standard method (d)  | P = 0.6956    |               |               |              |
| Prevalence method (d)  | P = 1.0000    |               |               |              |
| Combined analysis (d)  | P = 1.0000    |               |               |              |
| Cochran-Armitage test (e)  | P < 0.0001**  |               |               |              |
| Fisher Exact test (e)  |               | P = 0.0695    | P = 0.0441*   | P < 0.0001** |

STUDY No. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
SEX : MALE

# NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 5

| Group Name  | Control     | 625 ppm     | 1250 ppm     | 2500 ppm    |
|---|-------------|-------------|--------------|-------------|
| SITE : Harderian gland<br>TUMOR : adenoma                 |             |             |              |             |
| Tumor rate  |             |             |              |             |
| Overall rates (a)   | 2/50 ( 4.0) | 3/50 ( 6.0) | 4/50 ( 8.0)  | 0/50 ( 0.0) |
| Adjusted rates (b)  | 8.33        | 9.38        | 12.90        | 0.0         |
| Terminal rates (c)  | 2/24 ( 8.3) | 3/32 ( 9.4) | 4/31 ( 12.9) | 0/39 ( 0.0) |
| Statistical analysis                                      |             |             |              |             |
| Peto test   |             |             |              |             |
| Standard method (d)                                       | P = -----   |             |              |             |
| Prevalence method (d)                                     | P = 0.9440  |             |              |             |
| Combined analysis (d)                                     | P = -----   |             |              |             |
| Cochran-Armitage test (e)                                 | P = 0.2733  |             |              |             |
| Fisher Exact test (e)                                     |             | P = 0.5000  | P = 0.3389   | P = 0.2475  |
| SITE : Harderian gland<br>TUMOR : adenoma, adenocarcinoma |             |             |              |             |
| Tumor rate  |             |             |              |             |
| Overall rates (a)   | 3/50 ( 6.0) | 3/50 ( 6.0) | 4/50 ( 8.0)  | 0/50 ( 0.0) |
| Adjusted rates (b)  | 8.33        | 9.38        | 12.90        | 0.0         |
| Terminal rates (c)  | 2/24 ( 8.3) | 3/32 ( 9.4) | 4/31 ( 12.9) | 0/39 ( 0.0) |
| Statistical analysis                                      |             |             |              |             |
| Peto test   |             |             |              |             |
| Standard method (d)                                       | P = -----   |             |              |             |
| Prevalence method (d)                                     | P = 0.9682  |             |              |             |
| Combined analysis (d)                                     | P = -----   |             |              |             |
| Cochran-Armitage test (e)                                 | P = 0.1539  |             |              |             |
| Fisher Exact test (e)                                     |             | P = 0.6611  | P = 0.5000   | P = 0.1212  |

(HPT360A)

BA1S5

- (a) : Number of tumor-bearing animals/number of animals examined at the site.  
(b) : Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.  
(c) : Observed tumor incidence at terminal kill.  
(d) : Beneath the control incidence are the P-values associated with the trend test.  
Standard method : Death analysis  
Prevalence method : Incidental tumor test  
Combined analysis : Death analysis + Incidental tumor test  
(e) : The Cochran-Armitage and Fisher exact test compare directly the overall incidence rates.  
? : The conditional probabilities of the largest and smallest possible outcomes can not be estimated or this P-value is beyond the estimated P-value.  
----- : There is no data which should be statistical analysis.  
Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$   
N.C. : Statistical value cannot be calculated and was not significant.

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 SEX : MALE

# NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 1

| Group Name                                     | Control      | 625 ppm      | 1250 ppm     | 2500 ppm    |
|--|--------------|--------------|--------------|-------------|
| SITE : ALL SITE<br>TUMOR : hemangioma          |              |              |              |             |
| Tumor rate                                     |              |              |              |             |
| Overall rates (a)                              | 3/50 ( 6.0)  | 5/50 ( 10.0) | 3/50 ( 6.0)  | 2/50 ( 4.0) |
| Adjusted rates (b)                             | 6.90         | 15.63        | 9.68         | 5.13        |
| Terminal rates (c)                             | 1/24 ( 4.2)  | 5/32 ( 15.6) | 3/31 ( 9.7)  | 2/39 ( 5.1) |
| Statistical analysis                           |              |              |              |             |
| Peto test                                      |              |              |              |             |
| Standard method (d)                            | P = 0.9629 ? |              |              |             |
| Prevalence method (d)                          | P = 0.8186   |              |              |             |
| Combined analysis (d)                          | P = 0.8953   |              |              |             |
| Cochran-Armitage test (e)                      | P = 0.4671   |              |              |             |
| Fisher Exact test (e)                          |              | P = 0.3575   | P = 0.6611   | P = 0.5000  |
| SITE : ALL SITE<br>TUMOR : histiocytic sarcoma |              |              |              |             |
| Tumor rate                                     |              |              |              |             |
| Overall rates (a)                              | 7/50 ( 14.0) | 5/50 ( 10.0) | 6/50 ( 12.0) | 4/50 ( 8.0) |
| Adjusted rates (b)                             | 0.0          | 4.88         | 0.0          | 7.69        |
| Terminal rates (c)                             | 0/24 ( 0.0)  | 1/32 ( 3.1)  | 0/31 ( 0.0)  | 3/39 ( 7.7) |
| Statistical analysis                           |              |              |              |             |
| Peto test                                      |              |              |              |             |
| Standard method (d)                            | P = 0.9879   |              |              |             |
| Prevalence method (d)                          | P = 0.0861   |              |              |             |
| Combined analysis (d)                          | P = 0.8952   |              |              |             |
| Cochran-Armitage test (e)                      | P = 0.4007   |              |              |             |
| Fisher Exact test (e)                          |              | P = 0.3798   | P = 0.5000   | P = 0.2623  |

(HPT360A)

BA155

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 2

| Group Name                                    | Control      | 625 ppm     | 1250 ppm      | 2500 ppm     |
|---|--------------|-------------|---------------|--------------|
| SITE : ALL SITE<br>TUMOR : malignant lymphoma |              |             |               |              |
| Tumor rate                                    |              |             |               |              |
| Overall rates (a)                             | 7/50 ( 14.0) | 4/50 ( 8.0) | 10/50 ( 20.0) | 7/50 ( 14.0) |
| Adjusted rates (b)                            | 16.67        | 9.38        | 19.35         | 15.38        |
| Terminal rates (c)                            | 4/24 ( 16.7) | 3/32 ( 9.4) | 6/31 ( 19.4)  | 6/39 ( 15.4) |
| Statistical analysis                          |              |             |               |              |
| Peto test                                     |              |             |               |              |
| Standard method (d)                           | P = 0.8199   |             |               |              |
| Prevalence method (d)                         | P = 0.4130   |             |               |              |
| Combined analysis (d)                         | P = 0.6450   |             |               |              |
| Cochran-Armitage test (e)                     | P = 0.6793   |             |               |              |
| Fisher Exact test (e)                         |              | P = 0.2623  | P = 0.2977    | P = 0.6129   |

(HPT360A)

BAIS5

- (a) : Number of tumor-bearing animals/number of animals examined at the site.  
 (b) : Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.  
 (c) : Observed tumor incidence at terminal kill.  
 (d) : Beneath the control incidence are the P-values associated with the trend test.  
     Standard method : Death analysis  
     Prevalence method : Incidental tumor test  
     Combined analysis : Death analysis + Incidental tumor test  
 (e) : The Cochran-Armitage and Fisher exact test compare directly the overall incidence rates.  
 ? : The conditional probabilities of the largest and smallest possible outcomes can not be estimated or this P-value is beyond the estimated P-value.  
 — : There is no data which should be statistical analysis.  
 Significant difference ; \* :  $P \leq 0.05$  \*\* :  $P \leq 0.01$   
 N.C. : Statistical value cannot be calculated and was not significant.

**TABLE P 2**

**NEOPLASTIC LESIONS-INCIDENCE AND  
STATISTICAL ANALYSIS: FEMALE**



STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crlj:BDF1]  
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 6

| Group Name  | Control     | 625 ppm      | 1250 ppm    | 2500 ppm    |
|---|-------------|--------------|-------------|-------------|
| SITE : lung<br>TUMOR : bronchiolar-alveolar adenoma                                 |             |              |             |             |
| Tumor rate  |             |              |             |             |
| Overall rates (a)   | 2/50 ( 4.0) | 3/50 ( 6.0)  | 1/50 ( 2.0) | 1/50 ( 2.0) |
| Adjusted rates (b)  | 8.33        | 9.52         | 3.03        | 2.63        |
| Terminal rates (c)  | 2/24 ( 8.3) | 2/21 ( 9.5)  | 1/33 ( 3.0) | 1/38 ( 2.6) |
| Statistical analysis  |             |              |             |             |
| Peto test   |             |              |             |             |
| Standard method (d)   | P = -----   |              |             |             |
| Prevalence method (d)   | P = 0.8884  |              |             |             |
| Combined analysis (d)   | P = -----   |              |             |             |
| Cochran-Armitage test (e)   | P = 0.3979  |              |             |             |
| Fisher Exact test (e)   |             | P = 0.5000   | P = 0.5000  | P = 0.5000  |
| SITE : lung<br>TUMOR : bronchiolar-alveolar carcinoma                               |             |              |             |             |
| Tumor rate  |             |              |             |             |
| Overall rates (a)   | 1/50 ( 2.0) | 4/50 ( 8.0)  | 1/50 ( 2.0) | 2/50 ( 4.0) |
| Adjusted rates (b)  | 0.0         | 14.29        | 3.03        | 2.63        |
| Terminal rates (c)  | 0/24 ( 0.0) | 3/21 ( 14.3) | 1/33 ( 3.0) | 1/38 ( 2.6) |
| Statistical analysis  |             |              |             |             |
| Peto test   |             |              |             |             |
| Standard method (d)   | P = 0.4490  |              |             |             |
| Prevalence method (d)   | P = 0.6716  |              |             |             |
| Combined analysis (d)   | P = 0.6326  |              |             |             |
| Cochran-Armitage test (e)   | P = 1.0000  |              |             |             |
| Fisher Exact test (e)   |             | P = 0.1811   | P = 0.7525  | P = 0.5000  |
| SITE : lung<br>TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar carcinoma |             |              |             |             |
| Tumor rate  |             |              |             |             |
| Overall rates (a)   | 3/50 ( 6.0) | 7/50 ( 14.0) | 2/50 ( 4.0) | 3/50 ( 6.0) |
| Adjusted rates (b)  | 8.33        | 23.81        | 6.06        | 5.26        |
| Terminal rates (c)  | 2/24 ( 8.3) | 5/21 ( 23.8) | 2/33 ( 6.1) | 2/38 ( 5.3) |
| Statistical analysis  |             |              |             |             |
| Peto test   |             |              |             |             |
| Standard method (d)   | P = 0.4490  |              |             |             |
| Prevalence method (d)   | P = 0.8831  |              |             |             |
| Combined analysis (d)   | P = 0.8556  |              |             |             |
| Cochran-Armitage test (e)   | P = 0.5552  |              |             |             |
| Fisher Exact test (e)   |             | P = 0.1589   | P = 0.5000  | P = 0.6611  |

STUDY No. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 7

| Group Name                                      | Control       | 625 ppm       | 1250 ppm      | 2500 ppm      |
|---|---------------|---------------|---------------|---------------|
| SITE : lymph node<br>TUMOR : malignant lymphoma |               |               |               |               |
| Tumor rate                                      |               |               |               |               |
| Overall rates (a)                               | 18/50 ( 36.0) | 17/50 ( 34.0) | 21/50 ( 42.0) | 10/50 ( 20.0) |
| Adjusted rates (b)                              | 41.67         | 33.33         | 45.45         | 15.79         |
| Terminal rates (c)                              | 10/24 ( 41.7) | 7/21 ( 33.3)  | 15/33 ( 45.5) | 6/38 ( 15.8)  |
| Statistical analysis                            |               |               |               |               |
| Peto test                                       |               |               |               |               |
| Standard method (d)                             | P = 0.9682    |               |               |               |
| Prevalence method (d)                           | P = 0.9890    |               |               |               |
| Combined analysis (d)                           | P = 0.9984    |               |               |               |
| Cochran-Armitage test (e)                       | P = 0.0934    |               |               |               |
| Fisher Exact test (e)                           |               | P = 0.5000    | P = 0.3410    | P = 0.0591    |
| SITE : liver<br>TUMOR : hemangioma              |               |               |               |               |
| Tumor rate                                      |               |               |               |               |
| Overall rates (a)                               | 1/50 ( 2.0)   | 1/50 ( 2.0)   | 1/50 ( 2.0)   | 3/50 ( 6.0)   |
| Adjusted rates (b)                              | 4.17          | 2.27          | 2.94          | 7.89          |
| Terminal rates (c)                              | 1/24 ( 4.2)   | 0/21 ( 0.0)   | 0/33 ( 0.0)   | 3/38 ( 7.9)   |
| Statistical analysis                            |               |               |               |               |
| Peto test                                       |               |               |               |               |
| Standard method (d)                             | P = -----     |               |               |               |
| Prevalence method (d)                           | P = 0.1578    |               |               |               |
| Combined analysis (d)                           | P = -----     |               |               |               |
| Cochran-Armitage test (e)                       | P = 0.2072    |               |               |               |
| Fisher Exact test (e)                           |               | P = 0.7525    | P = 0.7525    | P = 0.3087    |
| SITE : liver<br>TUMOR : hepatocellular adenoma  |               |               |               |               |
| Tumor rate                                      |               |               |               |               |
| Overall rates (a)                               | 4/50 ( 8.0)   | 6/50 ( 12.0)  | 4/50 ( 8.0)   | 5/50 ( 10.0)  |
| Adjusted rates (b)                              | 12.00         | 17.86         | 12.12         | 13.16         |
| Terminal rates (c)                              | 2/24 ( 8.3)   | 2/21 ( 9.5)   | 4/33 ( 12.1)  | 5/38 ( 13.2)  |
| Statistical analysis                            |               |               |               |               |
| Peto test                                       |               |               |               |               |
| Standard method (d)                             | P = -----     |               |               |               |
| Prevalence method (d)                           | P = 0.6464    |               |               |               |
| Combined analysis (d)                           | P = -----     |               |               |               |
| Cochran-Armitage test (e)                       | P = 0.9027    |               |               |               |
| Fisher Exact test (e)                           |               | P = 0.3703    | P = 0.6425    | P = 0.5000    |

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 SEX : FEMALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 8

| Group Name   | Control     | 625 ppm      | 1250 ppm     | 2500 ppm     |
|--|-------------|--------------|--------------|--------------|
| SITE : liver<br>TUMOR : histiocytic sarcoma                              |             |              |              |              |
| Tumor rate   |             |              |              |              |
| Overall rates (a)  | 2/50 ( 4.0) | 3/50 ( 6.0)  | 0/50 ( 0.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)   | 0.0         | 3.13         | 0.0          | 0.0          |
| Terminal rates (c)   | 0/24 ( 0.0) | 0/21 ( 0.0)  | 0/33 ( 0.0)  | 0/38 ( 0.0)  |
| Statistical analysis   |             |              |              |              |
| Peto test  |             |              |              |              |
| Standard method (d)  | P = 0.8579  |              |              |              |
| Prevalence method (d)  | P = 0.6376  |              |              |              |
| Combined analysis (d)  | P = 0.8942  |              |              |              |
| Cochran-Armitage test (e)  | P = 0.3266  |              |              |              |
| Fisher Exact test (e)  |             | P = 0.5000   | P = 0.2475   | P = 0.5000   |
| SITE : liver<br>TUMOR : hemangioma, hemangiosarcoma                      |             |              |              |              |
| Tumor rate   |             |              |              |              |
| Overall rates (a)  | 1/50 ( 2.0) | 2/50 ( 4.0)  | 1/50 ( 2.0)  | 3/50 ( 6.0)  |
| Adjusted rates (b)   | 4.17        | 4.76         | 2.94         | 7.89         |
| Terminal rates (c)   | 1/24 ( 4.2) | 1/21 ( 4.8)  | 0/33 ( 0.0)  | 3/38 ( 7.9)  |
| Statistical analysis   |             |              |              |              |
| Peto test  |             |              |              |              |
| Standard method (d)  | P = -----   |              |              |              |
| Prevalence method (d)  | P = 0.2414  |              |              |              |
| Combined analysis (d)  | P = -----   |              |              |              |
| Cochran-Armitage test (e)  | P = 0.3293  |              |              |              |
| Fisher Exact test (e)  |             | P = 0.5000   | P = 0.7525   | P = 0.3087   |
| SITE : liver<br>TUMOR : hepatocellular adenoma, hepatocellular carcinoma |             |              |              |              |
| Tumor rate   |             |              |              |              |
| Overall rates (a)  | 4/50 ( 8.0) | 6/50 ( 12.0) | 5/50 ( 10.0) | 5/50 ( 10.0) |
| Adjusted rates (b)   | 12.00       | 17.86        | 15.15        | 13.16        |
| Terminal rates (c)   | 2/24 ( 8.3) | 2/21 ( 9.5)  | 5/33 ( 15.2) | 5/38 ( 13.2) |
| Statistical analysis   |             |              |              |              |
| Peto test  |             |              |              |              |
| Standard method (d)  | P = -----   |              |              |              |
| Prevalence method (d)  | P = 0.6407  |              |              |              |
| Combined analysis (d)  | P = -----   |              |              |              |
| Cochran-Armitage test (e)  | P = 0.8734  |              |              |              |
| Fisher Exact test (e)  |             | P = 0.3703   | P = 0.5000   | P = 0.5000   |

STUDY No. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
SEX : FEMALE

# NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 9

| Group Name   | Control      | 625 ppm      | 1250 ppm     | 2500 ppm     |
|--|--------------|--------------|--------------|--------------|
| SITE : pituitary gland<br>TUMOR : adenoma          |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                  | 4/50 ( 8.0)  | 9/50 ( 18.0) | 6/50 ( 12.0) | 7/50 ( 14.0) |
| Adjusted rates (b)                                 | 16.67        | 23.81        | 15.38        | 13.16        |
| Terminal rates (c)                                 | 4/24 ( 16.7) | 5/21 ( 23.8) | 5/33 ( 15.2) | 5/38 ( 13.2) |
| Statistical analysis                               |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                | P = 0.2302   |              |              |              |
| Prevalence method (d)                              | P = 0.7216   |              |              |              |
| Combined analysis (d)                              | P = 0.5825   |              |              |              |
| Cochran-Armitage test (e)                          | P = 0.6188   |              |              |              |
| Fisher Exact test (e)                              |              | P = 0.1168   | P = 0.3703   | P = 0.2623   |
| SITE : ovary<br>TUMOR : papillary adenoma          |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                  | 1/50 ( 2.0)  | 0/50 ( 0.0)  | 4/50 ( 8.0)  | 1/50 ( 2.0)  |
| Adjusted rates (b)                                 | 3.23         | 0.0          | 12.12        | 2.27         |
| Terminal rates (c)                                 | 0/24 ( 0.0)  | 0/21 ( 0.0)  | 4/33 ( 12.1) | 0/38 ( 0.0)  |
| Statistical analysis                               |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                | P = -----    |              |              |              |
| Prevalence method (d)                              | P = 0.3991   |              |              |              |
| Combined analysis (d)                              | P = -----    |              |              |              |
| Cochran-Armitage test (e)                          | P = 0.6742   |              |              |              |
| Fisher Exact test (e)                              |              | P = 0.5000   | P = 0.1811   | P = 0.7525   |
| SITE : uterus<br>TUMOR : endometrial stromal polyp |              |              |              |              |
| Tumor rate   |              |              |              |              |
| Overall rates (a)                                  | 3/50 ( 6.0)  | 2/50 ( 4.0)  | 2/50 ( 4.0)  | 2/50 ( 4.0)  |
| Adjusted rates (b)                                 | 11.11        | 6.25         | 5.13         | 5.26         |
| Terminal rates (c)                                 | 2/24 ( 8.3)  | 0/21 ( 0.0)  | 1/33 ( 3.0)  | 2/38 ( 5.3)  |
| Statistical analysis                               |              |              |              |              |
| Peto test  |              |              |              |              |
| Standard method (d)                                | P = -----    |              |              |              |
| Prevalence method (d)                              | P = 0.7312   |              |              |              |
| Combined analysis (d)                              | P = -----    |              |              |              |
| Cochran-Armitage test (e)                          | P = 0.6865   |              |              |              |
| Fisher Exact test (e)                              |              | P = 0.5000   | P = 0.5000   | P = 0.5000   |

STUDY No. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 10

| Group Name  | Control       | 625 ppm       | 1250 ppm      | 2500 ppm      |
|---|---------------|---------------|---------------|---------------|
| SITE : uterus<br>TUMOR : histiocytic sarcoma                            |               |               |               |               |
| Tumor rate  |               |               |               |               |
| Overall rates (a)   | 16/50 ( 32.0) | 14/50 ( 28.0) | 12/50 ( 24.0) | 12/50 ( 24.0) |
| Adjusted rates (b)  | 18.52         | 13.04         | 21.21         | 27.27         |
| Terminal rates (c)  | 4/24 ( 16.7)  | 2/21 ( 9.5)   | 7/33 ( 21.2)  | 9/38 ( 23.7)  |
| Statistical analysis  |               |               |               |               |
| Peto test   |               |               |               |               |
| Standard method (d)   | P = 1.0000    |               |               |               |
| Prevalence method (d)   | P = 0.0444*   |               |               |               |
| Combined analysis (d)   | P = 0.9459    |               |               |               |
| Cochran-Armitage test (e)   | P = 0.3600    |               |               |               |
| Fisher Exact test (e)   |               | P = 0.4138    | P = 0.2522    | P = 0.2522    |
| SITE : mammary gland<br>TUMOR : adenocarcinoma                          |               |               |               |               |
| Tumor rate  |               |               |               |               |
| Overall rates (a)   | 0/50 ( 0.0)   | 1/50 ( 2.0)   | 0/50 ( 0.0)   | 3/50 ( 6.0)   |
| Adjusted rates (b)  | 0.0           | 4.76          | 0.0           | 4.88          |
| Terminal rates (c)  | 0/24 ( 0.0)   | 1/21 ( 4.8)   | 0/33 ( 0.0)   | 1/38 ( 2.6)   |
| Statistical analysis  |               |               |               |               |
| Peto test   |               |               |               |               |
| Standard method (d)   | P = 0.1467    |               |               |               |
| Prevalence method (d)   | P = 0.1237    |               |               |               |
| Combined analysis (d)   | P = 0.0483*   |               |               |               |
| Cochran-Armitage test (e)   | P = 0.0405*   |               |               |               |
| Fisher Exact test (e)   |               | P = 0.5000    | P = N. C.     | P = 0.1212    |
| SITE : mammary gland<br>TUMOR : adenocarcinoma, adenosquamous carcinoma |               |               |               |               |
| Tumor rate  |               |               |               |               |
| Overall rates (a)   | 0/50 ( 0.0)   | 2/50 ( 4.0)   | 0/50 ( 0.0)   | 4/50 ( 8.0)   |
| Adjusted rates (b)  | 0.0           | 4.76          | 0.0           | 7.32          |
| Terminal rates (c)  | 0/24 ( 0.0)   | 1/21 ( 4.8)   | 0/33 ( 0.0)   | 1/38 ( 2.6)   |
| Statistical analysis  |               |               |               |               |
| Peto test   |               |               |               |               |
| Standard method (d)   | P = 0.3193    |               |               |               |
| Prevalence method (d)   | P = 0.0446*   |               |               |               |
| Combined analysis (d)   | P = 0.0469*   |               |               |               |
| Cochran-Armitage test (e)   | P = 0.0356*   |               |               |               |
| Fisher Exact test (e)   |               | P = 0.2475    | P = N. C.     | P = 0.0587    |

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 SEX : FEMALE

# NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 11

| Group Name  | Control      | 625 ppm     | 1250 ppm     | 2500 ppm    |
|---|--------------|-------------|--------------|-------------|
| SITE : Harderian gland<br>TUMOR : adenoma                 |              |             |              |             |
| Tumor rate  |              |             |              |             |
| Overall rates (a)   | 1/50 ( 2.0)  | 0/50 ( 0.0) | 4/50 ( 8.0)  | 3/50 ( 6.0) |
| Adjusted rates (b)  | 3.70         | 0.0         | 12.12        | 7.32        |
| Terminal rates (c)  | 0/24 ( 0.0)  | 0/21 ( 0.0) | 4/33 ( 12.1) | 2/38 ( 5.3) |
| Statistical analysis                                      |              |             |              |             |
| Peto test   |              |             |              |             |
| Standard method (d)                                       | P = -----    |             |              |             |
| Prevalence method (d)                                     | P = 0.1492   |             |              |             |
| Combined analysis (d)                                     | P = -----    |             |              |             |
| Cochran-Armitage test (e)                                 | P = 0.1432   |             |              |             |
| Fisher Exact test (e)                                     |              | P = 0.5000  | P = 0.1811   | P = 0.3087  |
| SITE : Harderian gland<br>TUMOR : adenoma, adenocarcinoma |              |             |              |             |
| Tumor rate  |              |             |              |             |
| Overall rates (a)   | 2/50 ( 4.0)  | 0/50 ( 0.0) | 4/50 ( 8.0)  | 3/50 ( 6.0) |
| Adjusted rates (b)  | 3.85         | 0.0         | 12.12        | 7.32        |
| Terminal rates (c)  | 0/24 ( 0.0)  | 0/21 ( 0.0) | 4/33 ( 12.1) | 2/38 ( 5.3) |
| Statistical analysis                                      |              |             |              |             |
| Peto test   |              |             |              |             |
| Standard method (d)                                       | P = 0.9209 ? |             |              |             |
| Prevalence method (d)                                     | P = 0.1549   |             |              |             |
| Combined analysis (d)                                     | P = 0.3059   |             |              |             |
| Cochran-Armitage test (e)                                 | P = 0.3270   |             |              |             |
| Fisher Exact test (e)                                     |              | P = 0.2475  | P = 0.3389   | P = 0.5000  |

(HPT360A)

BAIS5

- (a) : Number of tumor-bearing animals/number of animals examined at the site.  
 (b) : Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.  
 (c) : Observed tumor incidence at terminal kill.  
 (d) : Beneath the control incidence are the P-values associated with the trend test.  
     Standard method : Death analysis  
     Prevalence method : Incidental tumor test  
     Combined analysis : Death analysis + Incidental tumor test  
 (e) : The Cochran-Armitage and Fisher exact test compare directly the overall incidence rates.  
 ? : The conditional probabilities of the largest and smallest possible outcomes cannot be estimated or this P-value is beyond the estimated P-value.  
 ----- : There is no data which should be statistical analysis.  
 Significant difference ; \* :  $P \leq 0.05$     \*\* :  $P \leq 0.01$   
 N.C. : Statistical value cannot be calculated and was not significant.

STUDY No. : 0712  
ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 3

| Group Name                                     | Control       | 625 ppm       | 1250 ppm      | 2500 ppm      |
|--|---------------|---------------|---------------|---------------|
| SITE : ALL SITE<br>TUMOR : hemangioma          |               |               |               |               |
| Tumor rate                                     |               |               |               |               |
| Overall rates (a)                              | 1/50 ( 2.0)   | 4/50 ( 8.0)   | 4/50 ( 8.0)   | 4/50 ( 8.0)   |
| Adjusted rates (b)                             | 4.17          | 8.70          | 11.76         | 10.53         |
| Terminal rates (c)                             | 1/24 ( 4.2)   | 1/21 ( 4.8)   | 3/33 ( 9.1)   | 4/38 ( 10.5)  |
| Statistical analysis                           |               |               |               |               |
| Peto test                                      |               |               |               |               |
| Standard method (d)                            | P = 0.6336    |               |               |               |
| Prevalence method (d)                          | P = 0.2193    |               |               |               |
| Combined analysis (d)                          | P = 0.2863    |               |               |               |
| Cochran-Armitage test (e)                      | P = 0.3086    |               |               |               |
| Fisher Exact test (e)                          |               | P = 0.1811    | P = 0.1811    | P = 0.1811    |
| SITE : ALL SITE<br>TUMOR : histiocytic sarcoma |               |               |               |               |
| Tumor rate                                     |               |               |               |               |
| Overall rates (a)                              | 18/50 ( 36.0) | 17/50 ( 34.0) | 13/50 ( 26.0) | 13/50 ( 26.0) |
| Adjusted rates (b)                             | 18.52         | 13.79         | 21.21         | 27.91         |
| Terminal rates (c)                             | 4/24 ( 16.7)  | 2/21 ( 9.5)   | 7/33 ( 21.2)  | 9/38 ( 23.7)  |
| Statistical analysis                           |               |               |               |               |
| Peto test                                      |               |               |               |               |
| Standard method (d)                            | P = 1.0000    |               |               |               |
| Prevalence method (d)                          | P = 0.0586    |               |               |               |
| Combined analysis (d)                          | P = 0.9717    |               |               |               |
| Cochran-Armitage test (e)                      | P = 0.2224    |               |               |               |
| Fisher Exact test (e)                          |               | P = 0.5000    | P = 0.1937    | P = 0.1937    |
| (HPT360A)                                      |               |               |               |               |

BA1S5

STUDY No. : 0712  
 ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
 SEX : FEMALE

# NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 4

| Group Name                 | Control        | 625 ppm        | 1250 ppm       | 2500 ppm       |
|----------------------------|----------------|----------------|----------------|----------------|
| SITE : ALL SITE            |                |                |                |                |
| TUMOR : malignant lymphoma |                |                |                |                |
| Tumor rate                 |                |                |                |                |
| Overall rates (a)          | 18/50 ( 36. 0) | 17/50 ( 34. 0) | 21/50 ( 42. 0) | 10/50 ( 20. 0) |
| Adjusted rates (b)         | 41. 67         | 33. 33         | 45. 45         | 15. 79         |
| Terminal rates (c)         | 10/24 ( 41. 7) | 7/21 ( 33. 3)  | 15/33 ( 45. 5) | 6/38 ( 15. 8)  |
| Statistical analysis       |                |                |                |                |
| Peto test                  |                |                |                |                |
| Standard method (d)        | P = 0. 9682    |                |                |                |
| Prevalence method (d)      | P = 0. 9890    |                |                |                |
| Combined analysis (d)      | P = 0. 9984    |                |                |                |
| Cochran-Armitage test (e)  | P = 0. 0934    |                |                |                |
| Fisher Exact test (e)      |                | P = 0. 5000    | P = 0. 3410    | P = 0. 0591    |

(HPT360A)

BAIS5

- (a) : Number of tumor-bearing animals/number of animals examined at the site.  
 (b) : Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.  
 (c) : Observed tumor incidence at terminal kill.  
 (d) : Beneath the control incidence are the P-values associated with the trend test.  
     Standard method : Death analysis  
     Prevalence method : Incidental tumor test  
     Combined analysis : Death analysis + Incidental tumor test  
 (e) : The Cochran-Armitage and Fisher exact test compare directly the overall incidence rates.  
 ? : The conditional probabilities of the largest and smallest possible outcomes can not be estimated or this P-value is beyond the estimated P-value.  
 ----- : There is no data which should be statistical analysis.  
 Significant difference ; \* :  $P \leq 0. 05$     \*\* :  $P \leq 0. 01$   
 N.C. : Statistical value cannot be calculated and was not significant.



TABLE Q 1

HISTOPATHOLOGICAL FINDINGS:

METASTASIS OF TUMOR: MALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 1

| Organ                            | Findings  | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|----------------------------------|---|---------------------------------------|---------------|---------------|----------------|----------------|
| (Integumentary system/appandage) |   |                                       |               |               |                |                |
| skin/app                         | metastasis:liver tumor  |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>0      |
| subcutis                         | metastasis:epididymis tumor   |                                       | <50><br>0     | <50><br>0     | <50><br>0      | <50><br>1      |
| (Respiratory system)             |   |                                       |               |               |                |                |
| nasal cavit                      | leukemic cell infiltration  |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>1      |
|                                  | metastasis:liver tumor  |                                       | 0             | 0             | 1              | 0              |
|                                  | metastasis:epididymis tumor   |                                       | 0             | 0             | 0              | 2              |
| lung                             | leukemic cell infiltration  |                                       | <50><br>3     | <50><br>1     | <50><br>7      | <50><br>1      |
|                                  | metastasis:liver tumor  |                                       | 2             | 1             | 3              | 1              |
|                                  | metastasis:subcutis tumor   |                                       | 0             | 1             | 0              | 1              |
|                                  | metastasis:muscle tumor   |                                       | 0             | 0             | 1              | 0              |
|                                  | metastasis:vertebra tumor   |                                       | 0             | 0             | 1              | 0              |
|                                  | metastasis:epididymis tumor   |                                       | 0             | 1             | 0              | 1              |
| (Hematopoietic system)           |   |                                       |               |               |                |                |
| bone marrow                      | leukemic cell infiltration  |                                       | <50><br>1     | <50><br>2     | <50><br>4      | <50><br>1      |
|                                  | metastasis:liver tumor  |                                       | 2             | 1             | 3              | 0              |
| < a ><br>b                       | a : Number of animals examined at the site<br>b : Number of animals with lesion |                                       |               |               |                |                |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 2

| Organ                  | Findings  | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|------------------------|---|---------------------------------------|---------------|---------------|----------------|----------------|
| (Hematopoietic system) |   |                                       |               |               |                |                |
| bone marrow            | metastasis:subcutis tumor   |                                       | <50><br>0     | <50><br>0     | <50><br>0      | <50><br>1      |
|                        | metastasis:lymph node tumor   |                                       | 1             | 0             | 0              | 0              |
| lymph node             | metastasis:liver tumor  |                                       | <50><br>3     | <50><br>0     | <50><br>1      | <50><br>0      |
|                        | metastasis:lymph node tumor   |                                       | 1             | 1             | 0              | 0              |
| spleen                 | leukemic cell infiltration  |                                       | <50><br>7     | <50><br>3     | <50><br>6      | <50><br>6      |
|                        | metastasis:liver tumor  |                                       | 2             | 1             | 3              | 0              |
|                        | metastasis:lymph node tumor   |                                       | 1             | 1             | 0              | 0              |
| (Circulatory system)   |   |                                       |               |               |                |                |
| heart                  | leukemic cell infiltration  |                                       | <50><br>0     | <50><br>0     | <50><br>3      | <50><br>2      |
|                        | metastasis:liver tumor  |                                       | 1             | 0             | 0              | 0              |
|                        | metastasis:lung tumor   |                                       | 1             | 0             | 0              | 0              |
|                        | metastasis:epididymis tumor   |                                       | 0             | 1             | 0              | 0              |
| (Digestive system)     |   |                                       |               |               |                |                |
| tongue                 | leukemic cell infiltration  |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>0      |
| salivary gl            | leukemic cell infiltration  |                                       | <50><br>0     | <50><br>0     | <50><br>0      | <50><br>1      |
| < a ><br>b             | a : Number of animals examined at the site<br>b : Number of animals with lesion |                                       |               |               |                |                |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 3

| Organ              | Findings                                   | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|--------------------|--|---------------------------------------|---------------|---------------|----------------|----------------|
| (Digestive system) |  |                                       |               |               |                |                |
| salivary gl        | metastasis:lymph node tumor                |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>0      |
| small intes        | metastasis:epididymis tumor                |                                       | <50><br>0     | <50><br>0     | <50><br>0      | <50><br>1      |
|                    | metastasis:lymph node tumor                |                                       | 0             | 1             | 0              | 0              |
| liver              | leukemic cell infiltration                 |                                       | <50><br>3     | <50><br>0     | <50><br>2      | <50><br>1      |
|                    | metastasis:peritoneum tumor                |                                       | 1             | 0             | 0              | 0              |
|                    | metastasis:subcutis tumor                  |                                       | 0             | 1             | 1              | 1              |
|                    | metastasis:muscle tumor                    |                                       | 0             | 0             | 1              | 0              |
|                    | metastasis:epididymis tumor                |                                       | 0             | 0             | 1              | 1              |
|                    | metastasis:lymph node tumor                |                                       | 0             | 1             | 0              | 0              |
| pancreas           | leukemic cell infiltration                 |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>0      |
|                    | metastasis:peritoneum tumor                |                                       | 1             | 0             | 0              | 0              |
| (Urinary system)   |  |                                       |               |               |                |                |
| kidney             | leukemic cell infiltration                 |                                       | <50><br>2     | <50><br>0     | <50><br>0      | <50><br>1      |
|                    | metastasis:liver tumor                     |                                       | 1             | 0             | 1              | 0              |
|                    | metastasis:subcutis tumor                  |                                       | 0             | 0             | 0              | 1              |
| < a >              | a : Number of animals examined at the site |                                       |               |               |                |                |
| b                  | b : Number of animals with lesion          |                                       |               |               |                |                |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Crl:BDF1]  
 REPORT TYPE : A1  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 4

| Group Name                       |                             | Control   | 625 ppm   | 1250 ppm  | 2500 ppm  |
|----------------------------------|-----------------------------|---|-----------|-----------|-----------|
| No. of Animals on Study          |                             | 50  | 50        | 50        | 50        |
| Organ                            | Findings                    |   |           |           |           |
| (Urinary system)                 |                             |   |           |           |           |
| kidney                           | metastasis:lung tumor       | <50><br>1   | <50><br>0 | <50><br>0 | <50><br>0 |
|                                  | metastasis:epididymis tumor | 0   | 0         | 0         | 1         |
| urin bladd                       | metastasis:liver tumor      | <50><br>0   | <50><br>0 | <50><br>1 | <50><br>0 |
| (Reproductive system)            |                             |   |           |           |           |
| testis                           | metastasis:epididymis tumor | <50><br>0   | <50><br>1 | <50><br>0 | <50><br>1 |
|                                  | leukemic cell infiltration  | <50><br>0   | <50><br>0 | <50><br>0 | <50><br>1 |
| (Nervous system)                 |                             |   |           |           |           |
| brain                            | leukemic cell infiltration  | <50><br>0   | <50><br>1 | <50><br>0 | <50><br>1 |
| (Special sense organs/appendage) |                             |   |           |           |           |
| Harder gl                        | leukemic cell infiltration  | <50><br>1   | <50><br>0 | <50><br>1 | <50><br>1 |
|                                  | metastasis:liver tumor      | 1   | 0         | 0         | 0         |
| (Musculoskeletal system)         |                             |   |           |           |           |
| muscle                           | leukemic cell infiltration  | <50><br>1   | <50><br>0 | <50><br>0 | <50><br>1 |
| < a ><br>b                       |                             | a : Number of animals examined at the site<br>b : Number of animals with lesion |           |           |           |

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj[Crl:BDF1]  
REPORT TYPE : A1  
SEX : MALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 5

|                 |  | Group Name              | Control | 625 ppm | 1250 ppm | 2500 ppm |
|-----------------|--|-------------------------|---------|---------|----------|----------|
|                 |  | No. of Animals on Study | 50      | 50      | 50       | 50       |
| Organ           | Findings                                   |                         |         |         |          |          |
| (Body cavities) |  |                         |         |         |          |          |
| pleura          |  |                         | <50>    | <50>    | <50>     | <50>     |
|                 | metastasis:lung tumor                      |                         | 1       | 0       | 0        | 0        |
| mediastinum     |  |                         | <50>    | <50>    | <50>     | <50>     |
|                 | leukemic cell infiltration                 |                         | 0       | 0       | 1        | 0        |
|                 | metastasis:lung tumor                      |                         | 1       | 0       | 0        | 0        |
|                 | metastasis:epididymis tumor                |                         | 0       | 1       | 0        | 0        |
| peritoneum      |  |                         | <50>    | <50>    | <50>     | <50>     |
|                 | leukemic cell infiltration                 |                         | 0       | 0       | 1        | 0        |
|                 | metastasis:liver tumor                     |                         | 2       | 0       | 1        | 0        |
|                 | metastasis:epididymis tumor                |                         | 0       | 2       | 0        | 0        |
| < a >           | a : Number of animals examined at the site |                         |         |         |          |          |
| b               | b : Number of animals with lesion          |                         |         |         |          |          |

(JPT150)

BA1S5

**TABLE Q 2**

**HISTOPATHOLOGICAL FINDINGS:**

**METASTASIS OF TUMOR: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/CrIj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 6

| Organ                            | Findings                         | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|----------------------------------|----------------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Integumentary system/appandage) |                                  |                                       |               |               |                |                |
| skin/app                         |                                  |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leukemic cell infiltration       |                                       | 3             | 0             | 1              | 0              |
| subcutis                         |                                  |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | metastasis:uterus tumor          |                                       | 1             | 0             | 0              | 0              |
|                                  | metastasis:lung tumor            |                                       | 0             | 0             | 0              | 1              |
| (Respiratory system)             |                                  |                                       |               |               |                |                |
| nasal cavit                      |                                  |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leukemic cell infiltration       |                                       | 1             | 0             | 1              | 0              |
|                                  | metastasis:uterus tumor          |                                       | 1             | 2             | 0              | 0              |
|                                  | metastasis:subcutis tumor        |                                       | 0             | 1             | 0              | 0              |
|                                  | metastasis:lung tumor            |                                       | 1             | 0             | 0              | 0              |
|                                  | metastasis:Harderian gland tumor |                                       | 1             | 0             | 0              | 0              |
| larynx                           |                                  |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leukemic cell infiltration       |                                       | 0             | 1             | 0              | 0              |
| lung                             |                                  |                                       | <50>          | <50>          | <50>           | <50>           |
|                                  | leukemic cell infiltration       |                                       | 12            | 9             | 13             | 6              |
|                                  | metastasis:liver tumor           |                                       | 1             | 1             | 0              | 1              |
|                                  | metastasis:uterus tumor          |                                       | 7             | 4             | 4              | 0              |
|                                  | metastasis:subcutis tumor        |                                       | 0             | 1             | 0              | 0              |
|                                  | metastasis:muscle tumor          |                                       | 1             | 0             | 0              | 0              |

< a > a : Number of animals examined at the site  
 b b : Number of animals with lesion



STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crj[Cri:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 7

|                        |                                  | Group Name                                 | Control    | 625 ppm    | 1250 ppm   | 2500 ppm  |
|------------------------|----------------------------------|--|------------|------------|------------|-----------|
|                        |                                  | No. of Animals on Study                    | 50         | 50         | 50         | 50        |
| Organ                  | Findings                         |  |            |            |            |           |
| (Respiratory system)   |                                  |  |            |            |            |           |
| lung                   | metastasis:Harderian gland tumor |  | <50><br>1  | <50><br>0  | <50><br>0  | <50><br>0 |
|                        | metastasis:mediastinum tumor     |  | 0          | 0          | 1          | 0         |
| (Hematopoietic system) |                                  |  |            |            |            |           |
| bone marrow            | leukemic cell infiltration       |  | <50><br>7  | <50><br>5  | <50><br>4  | <50><br>2 |
|                        | metastasis:liver tumor           |  | 0          | 0          | 0          | 1         |
|                        | metastasis:uterus tumor          |  | 5          | 4          | 2          | 0         |
|                        | metastasis:subcutis tumor        |  | 0          | 1          | 0          | 0         |
| lymph node             | metastasis:liver tumor           |  | <50><br>0  | <50><br>1  | <50><br>0  | <50><br>0 |
|                        | metastasis:uterus tumor          |  | 2          | 0          | 0          | 0         |
|                        | metastasis:subcutis tumor        |  | 0          | 1          | 0          | 0         |
|                        | metastasis:lung tumor            |  | 0          | 0          | 0          | 1         |
|                        | metastasis:Harderian gland tumor |  | 1          | 0          | 0          | 0         |
| spleen                 | leukemic cell infiltration       |  | <50><br>15 | <50><br>10 | <50><br>19 | <50><br>9 |
|                        | metastasis:liver tumor           |  | 0          | 1          | 0          | 0         |
|                        | metastasis:uterus tumor          |  | 1          | 0          | 0          | 0         |
|                        | metastasis:subcutis tumor        |  | 0          | 1          | 0          | 0         |
| < a >                  |                                  | a : Number of animals examined at the site |            |            |            |           |
| b                      |                                  | b : Number of animals with lesion          |            |            |            |           |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 8

| Group Name              |  | Control   | 625 ppm   | 1250 ppm  | 2500 ppm  |
|-------------------------|--|-----------|-----------|-----------|-----------|
| No. of Animals on Study |  | 50        | 50        | 50        | 50        |
| Organ                   | Findings                                   |           |           |           |           |
| (Circulatory system)    |  |           |           |           |           |
| heart                   | leukemic cell infiltration                 | <50><br>7 | <50><br>2 | <50><br>6 | <50><br>2 |
|                         | metastasis:uterus tumor                    | 0         | 1         | 0         | 1         |
|                         | metastasis:subcutis tumor                  | 0         | 1         | 0         | 0         |
|                         | metastasis:lung tumor                      | 1         | 0         | 0         | 1         |
|                         | metastasis:mediastinum tumor               | 0         | 0         | 1         | 0         |
| (Digestive system)      |  |           |           |           |           |
| tongue                  | leukemic cell infiltration                 | <50><br>3 | <50><br>1 | <50><br>3 | <50><br>0 |
|                         | leukemic cell infiltration                 | <50><br>9 | <50><br>1 | <50><br>3 | <50><br>1 |
| stomach                 | leukemic cell infiltration                 | <50><br>3 | <50><br>0 | <50><br>0 | <50><br>0 |
|                         | metastasis:uterus tumor                    | 0         | 1         | 1         | 0         |
|                         | metastasis:subcutis tumor                  | 0         | 1         | 0         | 0         |
|                         | metastasis:lung tumor                      | 0         | 0         | 0         | 1         |
| small intes             | leukemic cell infiltration                 | <50><br>0 | <50><br>1 | <50><br>0 | <50><br>0 |
|                         | leukemic cell infiltration                 | <50><br>1 | <50><br>0 | <50><br>0 | <50><br>0 |
| large intes             | leukemic cell infiltration                 | <50><br>1 | <50><br>0 | <50><br>0 | <50><br>0 |
|                         | metastasis:retroperitoneum tumor           | 0         | 0         | 1         | 0         |
| < a >                   | a : Number of animals examined at the site |           |           |           |           |
| b                       | b : Number of animals with lesion          |           |           |           |           |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 9

| Organ              | Findings                                   | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|--------------------|--|---------------------------------------|---------------|---------------|----------------|----------------|
| (Digestive system) |  |                                       |               |               |                |                |
| liver              | leukemic cell infiltration                 |                                       | <50><br>13    | <50><br>8     | <50><br>10     | <50><br>4      |
|                    | metastasis:uterus tumor                    |                                       | 11            | 9             | 7              | 2              |
|                    | metastasis:subcutis tumor                  |                                       | 0             | 1             | 0              | 0              |
|                    | metastasis:spleen tumor                    |                                       | 0             | 0             | 0              | 1              |
| gall bladd         | leukemic cell infiltration                 |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>0      |
|                    |  |                                       |               |               |                |                |
| pancreas           | leukemic cell infiltration                 |                                       | <50><br>7     | <50><br>1     | <50><br>2      | <50><br>0      |
|                    | metastasis:uterus tumor                    |                                       | 1             | 1             | 1              | 0              |
|                    | metastasis:subcutis tumor                  |                                       | 0             | 1             | 0              | 0              |
| (Urinary system)   |  |                                       |               |               |                |                |
| kidney             | leukemic cell infiltration                 |                                       | <50><br>7     | <50><br>6     | <50><br>6      | <50><br>2      |
|                    | metastasis:uterus tumor                    |                                       | 0             | 2             | 2              | 0              |
|                    | metastasis:subcutis tumor                  |                                       | 0             | 1             | 0              | 0              |
|                    | metastasis:lung tumor                      |                                       | 1             | 0             | 0              | 1              |
| urin bladd         | leukemic cell infiltration                 |                                       | <50><br>7     | <50><br>2     | <50><br>7      | <50><br>3      |
|                    | metastasis:liver tumor                     |                                       | 0             | 1             | 0              | 0              |
| < a >              | a : Number of animals examined at the site |                                       |               |               |                |                |
| b                  | b : Number of animals with lesion          |                                       |               |               |                |                |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 10

| Organ   | Findings                         | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|---|----------------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Urinary system)  |                                  |                                       |               |               |                |                |
| urin bladd  | metastasis:uterus tumor          |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>0      |
|   | metastasis:subcutis tumor        |                                       | 0             | 1             | 0              | 0              |
| (Endocrine system)  |                                  |                                       |               |               |                |                |
| pituitary   | leukemic cell infiltration       |                                       | <50><br>1     | <50><br>0     | <50><br>0      | <50><br>0      |
|   | metastasis:lung tumor            |                                       | 0             | 0             | 0              | 1              |
| thyroid   | leukemic cell infiltration       |                                       | <50><br>1     | <50><br>0     | <50><br>0      | <50><br>0      |
|   | leukemic cell infiltration       |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>0      |
| adrenal   | leukemic cell infiltration       |                                       | <50><br>0     | <50><br>0     | <50><br>1      | <50><br>0      |
| (Reproductive system)   |                                  |                                       |               |               |                |                |
| ovary   | leukemic cell infiltration       |                                       | <50><br>4     | <50><br>2     | <50><br>2      | <50><br>1      |
|   | metastasis:uterus tumor          |                                       | 3             | 6             | 2              | 1              |
| uterus  | leukemic cell infiltration       |                                       | <50><br>3     | <50><br>3     | <50><br>0      | <50><br>2      |
|   | metastasis:subcutis tumor        |                                       | 0             | 1             | 0              | 0              |
|   | metastasis:retroperitoneum tumor |                                       | 0             | 0             | 1              | 0              |
| vagina  | leukemic cell infiltration       |                                       | <50><br>1     | <50><br>0     | <50><br>1      | <50><br>1      |
| < a > a : Number of animals examined at the site<br>b b : Number of animals with lesion |                                  |                                       |               |               |                |                |

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj[Cri:BDF1]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
 ALL ANIMALS (0-105W)

PAGE : 11

| Organ                            | Findings                         | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|----------------------------------|----------------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Reproductive system)            |                                  |                                       |               |               |                |                |
| vagina                           | metastasis:uterus tumor          |                                       | <50><br>1     | <50><br>2     | <50><br>0      | <50><br>0      |
| mammary gl                       | leukemic cell infiltration       |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>0      |
| (Nervous system)                 |                                  |                                       |               |               |                |                |
| brain                            | leukemic cell infiltration       |                                       | <50><br>0     | <50><br>1     | <50><br>1      | <50><br>2      |
|                                  | metastasis:Harderian gland tumor |                                       | 1             | 0             | 0              | 0              |
| spinal cord                      | leukemic cell infiltration       |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>1      |
| (Special sense organs/appendage) |                                  |                                       |               |               |                |                |
| eye                              | leukemic cell infiltration       |                                       | <50><br>2     | <50><br>0     | <50><br>0      | <50><br>0      |
|                                  | metastasis:subcutis tumor        |                                       | 0             | 1             | 0              | 0              |
|                                  | metastasis:lung tumor            |                                       | 0             | 0             | 0              | 1              |
| Harder gl                        | leukemic cell infiltration       |                                       | <50><br>5     | <50><br>1     | <50><br>3      | <50><br>3      |
|                                  | metastasis:uterus tumor          |                                       | 1             | 0             | 1              | 0              |
| (Musculoskeletal system)         |                                  |                                       |               |               |                |                |
| muscle                           | leukemic cell infiltration       |                                       | <50><br>3     | <50><br>1     | <50><br>1      | <50><br>0      |

< a > a : Number of animals examined at the site  
 b b : Number of animals with lesion

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Crlj [Crj:BDF1]  
REPORT TYPE : A1  
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)  
ALL ANIMALS (0-105W)

PAGE : 12

| Organ   | Findings                   | Group Name<br>No. of Animals on Study | Control<br>50 | 625 ppm<br>50 | 1250 ppm<br>50 | 2500 ppm<br>50 |
|---|----------------------------|---------------------------------------|---------------|---------------|----------------|----------------|
| (Musculoskeletal system)  |                            |                                       |               |               |                |                |
| muscle  | metastasis:subcutis tumor  |                                       | <50><br>0     | <50><br>1     | <50><br>0      | <50><br>0      |
|   | metastasis:lung tumor      |                                       | 0             | 0             | 0              | 1              |
| (Body cavities)   |                            |                                       |               |               |                |                |
| pleura  | metastasis:lung tumor      |                                       | <50><br>1     | <50><br>0     | <50><br>0      | <50><br>0      |
|   |                            |                                       |               |               |                |                |
| mediastinum   | leukemic cell infiltration |                                       | <50><br>8     | <50><br>8     | <50><br>5      | <50><br>3      |
|   | metastasis:subcutis tumor  |                                       | 0             | 1             | 0              | 0              |
| peritoneum  | leukemic cell infiltration |                                       | <50><br>6     | <50><br>2     | <50><br>5      | <50><br>2      |
|   | metastasis:uterus tumor    |                                       | 1             | 0             | 1              | 1              |
|   | metastasis:lung tumor      |                                       | 1             | 0             | 0              | 1              |
| < a > a : Number of animals examined at the site<br>b b : Number of animals with lesion |                            |                                       |               |               |                |                |

(JPT150)

BAIS5

TABLE R

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC  
LESIONS IN JAPAN BIOASSAY RESEARCH CENTER:  
B6D2F1/Crlj FEMALE MICE

TABLE R HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS  
IN JAPAN BIOASSAY RESEARCH CENTER : B6D2F1/Crlj FEMALE MICE

| Organs<br>Tumors                      | No. of animals<br>examined | No. of animals<br>bearing tumor | Incidence<br>(%) | Min. - Max.<br>(%) |
|---------------------------------------|----------------------------|---------------------------------|------------------|--------------------|
| Mammary gland                         | 2347                       |                                 |                  |                    |
| adenocarcinoma <sup>1)</sup>          |                            | 40                              | 1.7              | 0 - 8              |
| adenosquamous carcinoma <sup>2)</sup> |                            | 0                               | 0.0              | 0 - 0              |
| 1) + 2)                               |                            | 40                              | 1.7              | 0 - 8              |
| Uterus                                | 2345                       |                                 |                  |                    |
| Histiocytic sarcoma                   |                            | 483                             | 20.6             | 10 - 34            |

47 carcinogenicity studies examined in Japan Bioassay Research Center were used.

Study No. : 0044, 0060, 0062, 0064, 0066, 0068, 0096, 0105, 0116, 0140, 0159, 0163, 0190,  
0206, 0211, 0225, 0243, 0268, 0270, 0279, 0285, 0297, 0319, 0329, 0343, 0348,  
0366, 0372, 0402, 0406, 0418, 0422, 0438, 0449, 0458, 0462, 0498, 0515, 0561,  
0580, 0611, 0613, 0642, 0676, 0685, 0705, 0732



TABLE S 1

CAUSE OF DEATH: MALE

STUDY NO. : 0712  
ANIMAL : MOUSE B6D2F1/Cr1j [Crj:BDF1]  
SEX : MALE

COUSE OF DEATH (SUMMARY)  
(0-105W)

PAGE : 1

| Group Name                         | Control | 625 ppm | 1250 ppm | 2500 ppm |
|------------------------------------|---------|---------|----------|----------|
| Number of Dead and Moribund Animal | 26      | 18      | 19       | 11       |
| no microscop confirm               | 2       | 1       | 0        | 1        |
| renal lesion                       | 0       | 1       | 1        | 1        |
| urinary retention                  | 0       | 6       | 3        | 2        |
| hydronephrosis                     | 7       | 6       | 3        | 2        |
| tumor d:leukemia                   | 3       | 1       | 4        | 1        |
| tumor d:subcutis                   | 0       | 0       | 1        | 0        |
| tumor d:lung                       | 2       | 0       | 0        | 1        |
| tumor d:lymph node                 | 1       | 0       | 0        | 0        |
| tumor d:liver                      | 9       | 2       | 5        | 2        |
| tumor d:urin bladd                 | 1       | 0       | 0        | 0        |
| tumor d:epididymis                 | 0       | 1       | 1        | 1        |
| tumor d:muscle                     | 0       | 0       | 1        | 0        |
| tumor d:peritoneum                 | 1       | 0       | 0        | 0        |

(B10120)

BA1S5

**TABLE S 2**

**CAUSE OF DEATH: FEMALE**

STUDY NO. : 0712  
 ANIMAL : MOUSE B6D2F1/Crlj [Crl:BDF1]  
 SEX : FEMALE

COUSE OF DEATH (SUMMARY)  
 (0-105W)

PAGE : 2

| Group Name                         | Control | 625 ppm | 1250 ppm | 2500 ppm |
|------------------------------------|---------|---------|----------|----------|
| Number of Dead and Moribund Animal | 26      | 29      | 17       | 12       |
| no microscop confirm               | 0       | 1       | 3        | 1        |
| renal lesion                       | 1       | 0       | 0        | 0        |
| hydronephrosis                     | 0       | 0       | 1        | 1        |
| peritonitis                        | 1       | 0       | 0        | 0        |
| tumor d:leukemia                   | 8       | 9       | 6        | 4        |
| tumor d:subcutis                   | 0       | 2       | 0        | 1        |
| tumor d:lung                       | 1       | 1       | 0        | 1        |
| tumor d:liver                      | 2       | 2       | 0        | 1        |
| tumor d:pituitary                  | 0       | 2       | 0        | 2        |
| tumor d:uterus                     | 11      | 11      | 5        | 0        |
| tumor d:mammary gl                 | 0       | 1       | 0        | 1        |
| tumor d:Harder gl                  | 1       | 0       | 0        | 0        |
| tumor d:muscle                     | 1       | 0       | 0        | 0        |
| tumor d:mediastinum                | 0       | 0       | 1        | 0        |
| tumor d:retroperit                 | 0       | 0       | 1        | 0        |

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BAIS5