

1 - ブロモブタンのラットを用いた
吸入によるがん原性試験報告書

試験番号：0560

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TABLE A

CONCENTRATIONS OF 1 - BROMOBUTANE
IN THE INHALATION CHAMBER
OF THE 2-YEAR INHALATION STUDY

CONCENTRATIONS OF 1-BROMOBUTANE IN THE INHALATION
CHAMBER OF THE 2-YEAR INHALATION STUDY

| Group Name | Concentration(ppm) Mean \pm S.D. |
|------------|---------------------------------------|
| Control | 0.0 \pm 0.0 |
| 125 ppm | 125.2 \pm 0.4 |
| 250 ppm | 250.5 \pm 1.0 |
| 500 ppm | 500.8 \pm 1.9 |

TABLE B1

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104
SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 1

| Group Name | Animals 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 |
| 125 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 |
| 250 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 |
| 500 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(%) | | | | | | | | | | | | | | | |

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STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 2

| Group | Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | 24 | 25 | 26 | 27 |
| | 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 |
| | 125 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 |
| | 250 ppm | 50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 |
| | | | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 98.0 | 98.0 | 98.0 | 98.0 |
| | 500 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(%) | | | | | | | | | | | | | |

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BAIS4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 3

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 28 | 29 | 30 | 31 | 32 | 33 | 34 | 35 | 36 | 37 | 38 | 39 | 40 | 41 |
| 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 |
| 125 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 |
| 250 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 | 49/50 | 49/50 |
| | | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 |
| 500 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(%) | | | | | | | | | | | | | | | |

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STUDY NO. : 0560

SURVIVAL ANIMAL NUMBERS

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

PAGE : 4

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 42 | 43 | 44 | 45 | 46 | 47 | 48 | 49 | 50 | 51 | 52 | 53 | 54 | 55 |
| 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 |
| 125 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 |
| 250 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 | 49/50 | 49/50 |
| | | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 |
| 500 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(%) | | | | | | | | | | | | | | | |

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STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 5

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
| | | 56 | 57 | 58 | 59 | 60 | 61 | 62 | 63 | 64 | 65 | 66 | 67 | 68 | 69 |
| Control | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 |
| 125 ppm | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 49/50 98.0 |
| 250 ppm | 50 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 |
| 500 ppm | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 49/50 98.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | | | | | | | | |

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STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 6

| Group | Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | | 70 | 71 | 72 | 73 | 74 | 75 | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83 |
| | Control | 50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 50/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 48/50 |
| | | | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 96.0 | 96.0 | 96.0 |
| | 125 ppm | 50 | 49/50 | 49/50 | 48/50 | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 46/50 | 45/50 | 45/50 | 44/50 | 44/50 |
| | | | 98.0 | 98.0 | 96.0 | 94.0 | 94.0 | 94.0 | 94.0 | 94.0 | 92.0 | 92.0 | 90.0 | 90.0 | 88.0 | 88.0 |
| | 250 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 | 49/50 | 49/50 |
| | | | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 |
| | 500 ppm | 50 | 49/50 | 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 |
| | | | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 |
| Number of survival/ Number of effective animals | | | Survival rate(%) | | | | | | | | | | | | | |

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STUDY NO. : 0560

SURVIVAL ANIMAL NUMBERS

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

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| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 84 | 85 | 86 | 87 | 88 | 89 | 90 | 91 | 92 | 93 | 94 | 95 | 96 | 97 |
| Control | 50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 |
| | | 96.0 | 96.0 | 96.0 | 96.0 | 94.0 | 94.0 | 94.0 | 94.0 | 94.0 | 92.0 | 92.0 | 92.0 | 92.0 | 92.0 |
| 125 ppm | 50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 42/50 | 42/50 | 42/50 | 41/50 | 41/50 | 40/50 | 39/50 | 38/50 | 38/50 |
| | | 88.0 | 88.0 | 86.0 | 86.0 | 86.0 | 84.0 | 84.0 | 84.0 | 82.0 | 82.0 | 80.0 | 78.0 | 76.0 | 76.0 |
| 250 ppm | 50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 48/50 | 48/50 | 47/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 |
| | | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 96.0 | 96.0 | 94.0 | 92.0 | 92.0 | 92.0 | 92.0 | 92.0 | 92.0 |
| 500 ppm | 50 | 47/50 | 47/50 | 45/50 | 44/50 | 44/50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 43/50 | 42/50 | 42/50 | 42/50 |
| | | 94.0 | 94.0 | 90.0 | 88.0 | 88.0 | 88.0 | 88.0 | 86.0 | 86.0 | 86.0 | 86.0 | 84.0 | 84.0 | 84.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | | | | | | | | |

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STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : MALE

SURVIVAL ANIMAL NUMBERS

PAGE : 8

| Group Name | Animals At start | Administration (Weeks) | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|
| | | 98 | 99 | 100 | 101 | 102 | 103 | 104 |
| Control | 50 | 45/50 | 45/50 | 45/50 | 45/50 | 44/50 | 42/50 | 39/50 |
| | | 90.0 | 90.0 | 90.0 | 90.0 | 88.0 | 84.0 | 78.0 |
| 125 ppm | 50 | 38/50 | 38/50 | 38/50 | 38/50 | 37/50 | 36/50 | 35/50 |
| | | 76.0 | 76.0 | 76.0 | 76.0 | 74.0 | 72.0 | 70.0 |
| 250 ppm | 50 | 44/50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 41/50 |
| | | 88.0 | 88.0 | 88.0 | 86.0 | 86.0 | 86.0 | 82.0 |
| 500 ppm | 50 | 41/50 | 40/50 | 40/50 | 40/50 | 40/50 | 40/50 | 37/50 |
| | | 82.0 | 80.0 | 80.0 | 80.0 | 80.0 | 80.0 | 74.0 |
| Number of survival/ Number of effective animals | | | | | | | | |
| Survival rate(%) | | | | | | | | |

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TABLE B2

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 9

| Group Name | Animals 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 |
| 125 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 |
| 250 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 |
| 500 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(%) | | | | | | | | | | | | | | | |

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BAIS4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 10

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | 24 | 25 | 26 | 27 |
| 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 |
| 125 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 |
| 250 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 |
| 500 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. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 11

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 28 | 29 | 30 | 31 | 32 | 33 | 34 | 35 | 36 | 37 | 38 | 39 | 40 | 41 |
| 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 |
| 125 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 |
| 250 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 |
| 500 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. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 12

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 42 | 43 | 44 | 45 | 46 | 47 | 48 | 49 | 50 | 51 | 52 | 53 | 54 | 55 |
| 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 |
| 125 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 |
| 250 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 |
| 500 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. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 13

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|----------------|
| | | 56 | 57 | 58 | 59 | 60 | 61 | 62 | 63 | 64 | 65 | 66 | 67 | 68 | 69 |
| Control | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 |
| 125 ppm | 50 | 50/50 100.0 | 50/50 100.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 | 49/50 98.0 |
| 250 ppm | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 |
| 500 ppm | 50 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 50/50 100.0 | 49/50 98.0 | 49/50 98.0 | 47/50 94.0 | 47/50 94.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | | | | | | | | |

(HAN360)

BAIS4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 14

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 70 | 71 | 72 | 73 | 74 | 75 | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83 |
| Control | 50 | 50/50 | 50/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 | 49/50 |
| | | 100.0 | 100.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 | 98.0 |
| 125 ppm | 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 | 48/50 | 48/50 |
| | | 98.0 | 98.0 | 98.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 |
| 250 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 | 49/50 |
| | | 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 | 98.0 |
| 500 ppm | 50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 46/50 | 46/50 | 46/50 | 46/50 | 45/50 | 44/50 | 44/50 | 44/50 | 44/50 |
| | | 94.0 | 94.0 | 94.0 | 94.0 | 92.0 | 92.0 | 92.0 | 92.0 | 92.0 | 90.0 | 88.0 | 88.0 | 88.0 | 88.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | | | | | | | | |

(HAN360)

BAIS4

STUDY NO. : 0560

SURVIVAL ANIMAL NUMBERS

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

PAGE : 15

| Group Name | Animals At start | Administration (Weeks) | | | | | | | | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| | | 84 | 85 | 86 | 87 | 88 | 89 | 90 | 91 | 92 | 93 | 94 | 95 | 96 | 97 |
| Control | 50 | 49/50 | 47/50 | 46/50 | 46/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 | 43/50 | 43/50 | 43/50 | 42/50 | 41/50 |
| | | 98.0 | 94.0 | 92.0 | 92.0 | 90.0 | 90.0 | 90.0 | 88.0 | 88.0 | 86.0 | 86.0 | 86.0 | 84.0 | 82.0 |
| 125 ppm | 50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 48/50 | 47/50 | 47/50 | 47/50 | 46/50 | 46/50 |
| | | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 96.0 | 94.0 | 94.0 | 94.0 | 92.0 | 92.0 |
| 250 ppm | 50 | 47/50 | 47/50 | 47/50 | 47/50 | 47/50 | 46/50 | 46/50 | 46/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 | 44/50 |
| | | 94.0 | 94.0 | 94.0 | 94.0 | 94.0 | 92.0 | 92.0 | 92.0 | 90.0 | 90.0 | 90.0 | 88.0 | 88.0 | 88.0 |
| 500 ppm | 50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 44/50 | 42/50 | 42/50 |
| | | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 88.0 | 84.0 | 84.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | | | | | | | | |

(HAN360)

BAIS4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

REPORT TYPE : A1 104

SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

PAGE : 16

| Group Name | Animals At start | Administration (Weeks) | | | | | | |
|---|---------------------|------------------------|-------|-------|-------|-------|-------|-------|
| | | 98 | 99 | 100 | 101 | 102 | 103 | 104 |
| Control | 50 | 39/50 | 39/50 | 39/50 | 39/50 | 38/50 | 38/50 | 38/50 |
| | | 78.0 | 78.0 | 78.0 | 78.0 | 76.0 | 76.0 | 76.0 |
| 125 ppm | 50 | 46/50 | 45/50 | 45/50 | 45/50 | 44/50 | 44/50 | 43/50 |
| | | 92.0 | 90.0 | 90.0 | 90.0 | 88.0 | 88.0 | 86.0 |
| 250 ppm | 50 | 43/50 | 43/50 | 43/50 | 42/50 | 42/50 | 41/50 | 41/50 |
| | | 86.0 | 86.0 | 86.0 | 84.0 | 84.0 | 82.0 | 82.0 |
| 500 ppm | 50 | 42/50 | 41/50 | 40/50 | 39/50 | 38/50 | 37/50 | 37/50 |
| | | 84.0 | 82.0 | 80.0 | 78.0 | 76.0 | 74.0 | 74.0 |
| Number of survival/ Number of effective animals Survival rate(%) | | | | | | | | |

(HAN360)

BAIS4

TABLE C1

CLINICAL OBSERVATION : MALE

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 1

[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 2

[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 3

[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 4

[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 5

[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 6

[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

PAGE : 7

[illegible]

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

PAGE : 8

| Clinical sign | Group Name | Administration Week-day | | | | | |
|-------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| DEATH | Control | 2 | 2 | 2 | 2 | 3 | 3 |
| | 125 ppm | 5 | 5 | 5 | 6 | 6 | 6 |
| | 250 ppm | 4 | 4 | 4 | 4 | 4 | 4 |
| | 500 ppm | 3 | 3 | 3 | 3 | 3 | 3 |
| MORIBUND SACRIFICE | Control | 3 | 3 | 3 | 4 | 5 | 8 |
| | 125 ppm | 7 | 7 | 7 | 7 | 8 | 9 |
| | 250 ppm | 2 | 2 | 3 | 3 | 3 | 5 |
| | 500 ppm | 7 | 7 | 7 | 7 | 7 | 10 |
| LOCOMOTOR MOVEMENT DECR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| LATERAL | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| PARALYTIC GAIT | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 1 | 1 | 1 | 1 |
| WASTING | Control | 0 | 1 | 1 | 2 | 1 | 0 |
| | 125 ppm | 1 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 1 | 0 |
| SOILED | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 1 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| PILOERECTION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| LOSS OF HAIR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 10

[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

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CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 15

[illegible]

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

PAGE : 16

| Clinical sign | Group Name | Administration Week-day | | | | | |
|--------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| TRAUMA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| SOILED PERI-GENITALIA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 1 | 1 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 0 |
| EXOPHTHALMOS | Control | 0 | 0 | 0 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CATARACT | Control | 2 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| | 250 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| | 500 ppm | 5 | 5 | 5 | 5 | 5 | 4 |
| CORNEAL OPACITY | Control | 2 | 2 | 2 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GROWTH OF TEETH | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| MALOCCLUSION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| EXTERNAL MASS | Control | 8 | 8 | 8 | 8 | 8 | 7 |
| | 125 ppm | 6 | 6 | 6 | 5 | 4 | 4 |
| | 250 ppm | 9 | 9 | 9 | 9 | 9 | 8 |
| | 500 ppm | 7 | 9 | 10 | 10 | 11 | 10 |
| M. PERI-MOUTH | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

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ALL ANIMALS

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ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 23

| 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. EAR | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 2 | 2 | 2 | 1 |
| M. NECK | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. BREAST | Control | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| M. ABDOMEN | Control | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 1 | 1 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 2 | 2 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 2 | 3 | 3 | 4 | 4 | 4 | 4 |
| | 500 ppm | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 |
| M. ANTERIOR. DORSUM | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 250 ppm | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 500 ppm | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 2 | 2 | 2 |
| M. POSTERIOR DORSUM | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 2 | 1 |
| | 125 ppm | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| M. HINDLIMB | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. GENITALIA | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

PAGE : 24

| Clinical sign | Group Name | Administration Week-day | | | | | |
|---------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| M. EAR | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. PERI EAR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| M. NECK | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. BREAST | Control | 2 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 1 | 1 | 1 | 1 | 0 | 0 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| | 500 ppm | 2 | 2 | 2 | 2 | 2 | 3 |
| M. ABDOMEN | Control | 2 | 2 | 2 | 2 | 2 | 1 |
| | 125 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| | 250 ppm | 4 | 4 | 3 | 3 | 3 | 3 |
| | 500 ppm | 3 | 4 | 4 | 4 | 4 | 3 |
| M. ANTERIOR. DORSUM | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| | 250 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| | 500 ppm | 2 | 2 | 3 | 3 | 3 | 3 |
| M. POSTERIOR DORSUM | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 1 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 2 | 2 | 3 | 3 | 3 | 2 |
| | 500 ppm | 1 | 1 | 1 | 1 | 2 | 2 |
| M. HINDLIMB | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 1 | 1 | 1 | 1 | 0 |
| M. GENITALIA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

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ALL ANIMALS

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[illegible]

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

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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. SCROTUM | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 | 1 | 1 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ULCER | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 3 | 2 | 2 | 3 | 3 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| EROSION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 |
| TORTICOLLIS | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

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| Clinical sign | Group Name | Administration Week-day | | | | | |
|-------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| M. SCROTUM | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ANEMIA | Control | 0 | 0 | 0 | 2 | 2 | 0 |
| | 125 ppm | 1 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ULCER | Control | 1 | 1 | 1 | 1 | 1 | 0 |
| | 125 ppm | 4 | 4 | 4 | 3 | 3 | 3 |
| | 250 ppm | 0 | 0 | 0 | 0 | 1 | 1 |
| | 500 ppm | 1 | 1 | 3 | 3 | 3 | 2 |
| EROSION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CRUSTA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| HEMORRHAGE | Control | 1 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| TORTICOLLIS | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| PROLAPSE OF PENIS | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

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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 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

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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 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

(HAN190)

BAIS 4

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

(HAN190)

BAIS 4

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCr-j]
 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 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

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| Clinical sign | Group Name | Administration Week-day | | | | | | | | | | | | | |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | | 71-7 | 72-7 | 73-7 | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

(HAN190)

BAIS 4

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

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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 |
| IRREGULAR BREATHING | Control | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

(HAN190)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

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| Clinical sign | Group Name | Administration Week-day | | | | | |
|-------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| NOISY | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| DEEP BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| RED URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| LOOSE STOOL | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

(HAN190)

BAIS 4

TABLE C2

CLINICAL OBSERVATION : FEMALE

REPORT TYPE : A1 104

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

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ALL ANIMALS

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[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

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[illegible]

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : FEMALE

PAGE : 48

| Clinical sign | Group Name | Administration Week-day | | | | | |
|-------------------------|------------|-------------------------|-------|-------|-------|-------|-------|
| | | 99-7 | 100-7 | 101-7 | 102-7 | 103-7 | 104-7 |
| DEATH | Control | 6 | 6 | 6 | 7 | 7 | 7 |
| | 125 ppm | 3 | 3 | 3 | 4 | 4 | 4 |
| | 250 ppm | 4 | 4 | 4 | 4 | 5 | 5 |
| | 500 ppm | 2 | 3 | 3 | 3 | 4 | 4 |
| MORIBUND SACRIFICE | Control | 5 | 5 | 5 | 5 | 5 | 5 |
| | 125 ppm | 2 | 2 | 2 | 2 | 2 | 3 |
| | 250 ppm | 3 | 3 | 4 | 4 | 4 | 4 |
| | 500 ppm | 7 | 7 | 8 | 9 | 9 | 9 |
| LOCOMOTOR MOVEMENT DECR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| HUNCHBACK POSITION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ATAXIC GAIT | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GAIT | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| EXCITEMENT | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| WASTING | Control | 0 | 0 | 0 | 0 | 0 | 1 |
| | 125 ppm | 1 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| SOILED | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

REPORT TYPE : A1 104

ALL ANIMALS

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[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

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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 |
| PILOERECTION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| PROLAPSE OF ANUS | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 | 1 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LACRIMATION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| CATARACT | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 125 ppm | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GROWTH OF TEETH | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

PAGE : 52

| 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 |
| PILOERECTION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| PROLAPSE OF ANUS | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| SOILED PERI-GENITALIA | Control | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LACRIMATION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| CATARACT | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 250 ppm | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 |
| | 500 ppm | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| CORNEAL OPACITY | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GROWTH OF TEETH | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : FEMALE

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| Clinical sign | Group Name | Administration Week-day | | | | | | | | | | | | | |
|--------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | | 71-7 | 72-7 | 73-7 | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
| PILORECTION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 1 | 0 | 0 | 0 | 0 | 0 |
| FROG BELLY | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| PROLAPSE OF ANUS | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 |
| EXOPHTHALMOS | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| LACRIMATION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| CATARACT | Control | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 250 ppm | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 |
| | 500 ppm | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| CORNEAL OPACITY | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GROWTH OF TEETH | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |

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ALL ANIMALS

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[illegible]

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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 |
| PILOERECTION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| FROG BELLY | Control | 0 | 1 | 1 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| PROLAPSE OF ANUS | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 0 | 0 | 0 | 0 | 0 |
| SOILED PERI-GENITALIA | Control | 0 | 1 | 2 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 0 | 0 | 0 | 0 | 0 |
| EXOPHTHALMOS | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| LACRIMATION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CATARACT | Control | 1 | 2 | 2 | 3 | 3 | 3 |
| | 125 ppm | 3 | 3 | 3 | 3 | 3 | 3 |
| | 250 ppm | 4 | 4 | 4 | 4 | 4 | 4 |
| | 500 ppm | 4 | 4 | 4 | 4 | 4 | 4 |
| CORNEAL OPACITY | Control | 1 | 1 | 1 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ABNORMAL GROWTH OF TEETH | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

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ALL ANIMALS

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[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

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[illegible]

REPORT TYPE : A1 104

ALL ANIMALS

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[illegible]

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

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| Clinical sign | Group Name | Administration Week-day | | | | | | | | | | | | | |
|---------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | | 71-7 | 72-7 | 73-7 | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
| MALOCCLUSION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| EXTERNAL MASS | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 3 | 3 |
| | 250 ppm | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 3 | 2 |
| | 500 ppm | 1 | 1 | 2 | 2 | 4 | 4 | 5 | 5 | 4 | 3 | 3 | 3 | 3 | 3 |
| INTERNAL MASS | Control | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. MANDIBULAR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. EAR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 1 | 1 | 2 | 2 | 2 | 2 | 1 | 0 | 0 | 0 | 0 | 0 |
| M. NECK | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. BREAST | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. ABDOMEN | Control | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |

CLINICAL OBSERVATION (SUMMARY)
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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 |
| MALOCCLUSION | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| EXTERNAL MASS | Control | 1 | 1 | 2 | 2 | 2 | 4 | 4 | 5 | 6 | 6 | 6 | 6 | 6 | 6 |
| | 125 ppm | 3 | 3 | 4 | 5 | 5 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 |
| | 250 ppm | 4 | 4 | 3 | 5 | 5 | 5 | 5 | 4 | 4 | 5 | 6 | 6 | 6 | 5 |
| | 500 ppm | 3 | 3 | 4 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 4 | 4 | 4 |
| INTERNAL MASS | Control | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 0 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. MANDIBULAR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. EAR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. BREAST | Control | 0 | 0 | 1 | 1 | 1 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| M. ABDOMEN | Control | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 1 | 1 | 2 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 |
| | 250 ppm | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | 4 | 4 | 4 | 4 |
| | 500 ppm | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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 |
| MALOCCLUSION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| EXTERNAL MASS | Control | 6 | 6 | 6 | 8 | 8 | 8 |
| | 125 ppm | 8 | 8 | 8 | 8 | 8 | 8 |
| | 250 ppm | 7 | 7 | 6 | 6 | 6 | 6 |
| | 500 ppm | 3 | 3 | 3 | 2 | 2 | 3 |
| INTERNAL MASS | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. MANDIBULAR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. EAR | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. PERI EAR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. NECK | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. BREAST | Control | 3 | 3 | 3 | 5 | 5 | 5 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. ABDOMEN | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 5 | 5 | 5 | 5 | 5 | 4 |
| | 250 ppm | 5 | 5 | 5 | 5 | 5 | 5 |
| | 500 ppm | 1 | 1 | 1 | 0 | 0 | 0 |

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[illegible]

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ALL ANIMALS

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[illegible]

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ALL ANIMALS

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[illegible]

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. POSTERIOR DORSUM | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 1 | 1 | 1 | 1 | 1 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| M. GENITALIA | Control | 1 | 1 | 1 | 1 | 1 | 1 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 1 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 2 | 2 | 2 | 2 | 2 | 3 |
| ANEMIA | Control | 0 | 0 | 0 | 0 | 0 | 1 |
| | 125 ppm | 1 | 1 | 1 | 2 | 2 | 1 |
| | 250 ppm | 2 | 2 | 2 | 2 | 1 | 1 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| JAUNDICE | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| ULCER | Control | 1 | 2 | 2 | 2 | 2 | 2 |
| | 125 ppm | 3 | 3 | 3 | 4 | 4 | 3 |
| | 250 ppm | 1 | 2 | 2 | 2 | 2 | 2 |
| | 500 ppm | 2 | 2 | 2 | 2 | 2 | 2 |
| EROSION | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CRUSTA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

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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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

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ALL ANIMALS

SEX : FEMALE

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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 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

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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 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

PAGE : 76

| 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

PAGE : 77

| 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 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| BROWN URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

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| Clinical sign | Group Name | Administration Week-day | | | | | | | | | | | | | |
|-------------------------|------------|-------------------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | | 71-7 | 72-7 | 73-7 | 74-7 | 75-7 | 76-7 | 77-7 | 78-7 | 79-7 | 80-7 | 81-7 | 82-7 | 83-7 | 84-7 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 | 1 | 1 | 3 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

PAGE : 79

| 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 |
| HEMORRHAGE | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 125 ppm | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 0 | 0 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| IRREGULAR BREATHING | Control | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 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 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

PAGE : 80

| 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 |
| | 125 ppm | 1 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 1 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| A. CICATRIX | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| IRREGULAR BREATHING | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 1 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| RESPIRATORY SOUND ABNOR | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| BRADYPNEA | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| BROWN URINE | Control | 0 | 0 | 0 | 0 | 0 | 0 |
| | 125 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 250 ppm | 0 | 0 | 0 | 0 | 0 | 0 |
| | 500 ppm | 0 | 0 | 0 | 0 | 0 | 0 |

TABLE D1

BODY WEIGHT CHANGES AND SURVIVAL ANIMAL
NUMBERS : MALE

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
UNIT : g
REPORT TYPE : A1 104
SEX : MALE

MEAN BODY WEIGHTS AND SURVIVAL

PAGE : 1

| Week-Day on Study | Control | | 125 ppm | | | 250 ppm | | | 500 ppm | | |
|----------------------|----------|---------------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
| | Av. Wt. | No. of Surviv. <50> | Av. Wt. | % of cont. <50> | No. of Surviv. | Av. Wt. | % of cont. <50> | No. of Surviv. | Av. Wt. | % of cont. <50> | No. of Surviv. |
| 0-0 | 116 (50) | 50/50 | 116 (50) | 100 | 50/50 | 116 (50) | 100 | 50/50 | 116 (50) | 100 | 50/50 |
| 1-7 | 144 (50) | 50/50 | 142 (50) | 99 | 50/50 | 137 (50) | 95 | 50/50 | 132 (50) | 92 | 50/50 |
| 2-7 | 177 (50) | 50/50 | 175 (50) | 99 | 50/50 | 167 (50) | 94 | 50/50 | 159 (50) | 90 | 50/50 |
| 3-7 | 203 (50) | 50/50 | 202 (50) | 100 | 50/50 | 192 (50) | 95 | 50/50 | 179 (50) | 88 | 50/50 |
| 4-7 | 225 (50) | 50/50 | 224 (50) | 100 | 50/50 | 212 (50) | 94 | 50/50 | 198 (50) | 88 | 50/50 |
| 5-7 | 241 (50) | 50/50 | 242 (50) | 100 | 50/50 | 229 (50) | 95 | 50/50 | 213 (50) | 88 | 50/50 |
| 6-7 | 256 (50) | 50/50 | 257 (50) | 100 | 50/50 | 244 (50) | 95 | 50/50 | 225 (50) | 88 | 50/50 |
| 7-7 | 272 (50) | 50/50 | 272 (50) | 100 | 50/50 | 259 (50) | 95 | 50/50 | 238 (50) | 88 | 50/50 |
| 8-7 | 284 (50) | 50/50 | 285 (50) | 100 | 50/50 | 272 (50) | 96 | 50/50 | 250 (50) | 88 | 50/50 |
| 9-7 | 294 (50) | 50/50 | 295 (50) | 100 | 50/50 | 284 (50) | 97 | 50/50 | 260 (50) | 88 | 50/50 |
| 10-7 | 304 (50) | 50/50 | 305 (50) | 100 | 50/50 | 292 (50) | 96 | 50/50 | 267 (50) | 88 | 50/50 |
| 11-7 | 312 (50) | 50/50 | 313 (50) | 100 | 50/50 | 301 (50) | 96 | 50/50 | 272 (50) | 87 | 50/50 |
| 12-7 | 318 (50) | 50/50 | 319 (50) | 100 | 50/50 | 306 (50) | 96 | 50/50 | 277 (50) | 87 | 50/50 |
| 13-7 | 325 (50) | 50/50 | 328 (50) | 101 | 50/50 | 313 (50) | 96 | 50/50 | 283 (50) | 87 | 50/50 |
| 14-7 | 331 (50) | 50/50 | 334 (50) | 101 | 50/50 | 318 (50) | 96 | 50/50 | 288 (50) | 87 | 50/50 |
| 18-7 | 343 (50) | 50/50 | 350 (50) | 102 | 50/50 | 333 (50) | 97 | 50/50 | 303 (50) | 88 | 50/50 |
| 22-7 | 362 (50) | 50/50 | 369 (50) | 102 | 50/50 | 352 (50) | 97 | 50/50 | 317 (50) | 88 | 50/50 |
| 26-7 | 377 (50) | 50/50 | 383 (50) | 102 | 50/50 | 366 (49) | 97 | 49/50 | 332 (50) | 88 | 50/50 |
| 30-7 | 385 (50) | 50/50 | 390 (50) | 101 | 50/50 | 374 (49) | 97 | 49/50 | 338 (50) | 88 | 50/50 |
| 34-7 | 393 (50) | 50/50 | 400 (50) | 102 | 50/50 | 385 (49) | 98 | 49/50 | 347 (50) | 88 | 50/50 |
| 38-7 | 400 (50) | 50/50 | 409 (50) | 102 | 50/50 | 393 (49) | 98 | 49/50 | 355 (50) | 89 | 50/50 |
| 42-7 | 410 (50) | 50/50 | 419 (50) | 102 | 50/50 | 401 (49) | 98 | 49/50 | 361 (50) | 88 | 50/50 |
| 46-7 | 416 (50) | 50/50 | 423 (50) | 102 | 50/50 | 407 (49) | 98 | 49/50 | 366 (50) | 88 | 50/50 |
| 50-7 | 421 (50) | 50/50 | 427 (50) | 101 | 50/50 | 410 (49) | 97 | 49/50 | 370 (50) | 88 | 50/50 |
| 54-7 | 426 (50) | 50/50 | 431 (50) | 101 | 50/50 | 414 (49) | 97 | 49/50 | 371 (50) | 87 | 50/50 |
| 58-7 | 427 (50) | 50/50 | 434 (50) | 102 | 50/50 | 418 (49) | 98 | 49/50 | 375 (50) | 88 | 50/50 |
| 62-7 | 431 (50) | 50/50 | 437 (50) | 101 | 50/50 | 422 (49) | 98 | 49/50 | 381 (50) | 88 | 50/50 |
| 66-7 | 434 (50) | 50/50 | 440 (50) | 101 | 50/50 | 421 (49) | 97 | 49/50 | 378 (50) | 87 | 50/50 |
| 70-7 | 433 (50) | 50/50 | 443 (49) | 102 | 49/50 | 422 (49) | 97 | 49/50 | 378 (49) | 87 | 49/50 |
| 74-7 | 436 (50) | 50/50 | 444 (47) | 102 | 47/50 | 425 (49) | 97 | 49/50 | 378 (49) | 87 | 49/50 |
| 78-7 | 443 (49) | 49/50 | 443 (46) | 100 | 46/50 | 425 (49) | 96 | 49/50 | 378 (48) | 85 | 48/50 |
| 82-7 | 442 (48) | 48/50 | 442 (44) | 100 | 44/50 | 422 (49) | 95 | 49/50 | 374 (48) | 85 | 48/50 |
| 86-7 | 440 (48) | 48/50 | 441 (43) | 100 | 43/50 | 419 (49) | 95 | 49/50 | 373 (45) | 85 | 45/50 |
| 90-7 | 437 (47) | 47/50 | 439 (42) | 100 | 42/50 | 418 (48) | 96 | 48/50 | 372 (44) | 85 | 44/50 |
| 94-7 | 432 (46) | 46/50 | 430 (40) | 100 | 40/50 | 414 (46) | 96 | 46/50 | 364 (43) | 84 | 43/50 |
| 98-7 | 428 (45) | 45/50 | 429 (38) | 100 | 38/50 | 415 (44) | 97 | 44/50 | 363 (41) | 85 | 41/50 |
| 102-7 | 414 (44) | 44/50 | 423 (37) | 102 | 37/50 | 410 (43) | 99 | 43/50 | 355 (40) | 86 | 40/50 |
| 104-7 | 420 (39) | 39/50 | 417 (35) | 99 | 35/50 | 405 (41) | 96 | 41/50 | 350 (37) | 83 | 37/50 |

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

TABLE D2

BODY WEIGHT CHANGES AND SURVIVAL ANIMAL
NUMBERS : FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : FEMALE

MEAN BODY WEIGHTS AND SURVIVAL

PAGE : 2

| Week-Day on Study | Control | | 125 ppm | | | 250 ppm | | | 500 ppm | | |
|----------------------|----------|---------------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|----------|-----------------------|-------------------|
| | Av. Wt. | No. of Surviv. <50> | Av. Wt. | % of cont. <50> | No. of Surviv. | Av. Wt. | % of cont. <50> | No. of Surviv. | Av. Wt. | % of cont. <50> | No. of Surviv. |
| 0-0 | 96 (50) | 50/50 | 96 (50) | 100 | 50/50 | 96 (50) | 100 | 50/50 | 96 (50) | 100 | 50/50 |
| 1-7 | 110 (50) | 50/50 | 108 (50) | 98 | 50/50 | 107 (50) | 97 | 50/50 | 104 (50) | 95 | 50/50 |
| 2-7 | 123 (50) | 50/50 | 122 (50) | 99 | 50/50 | 120 (50) | 98 | 50/50 | 116 (50) | 94 | 50/50 |
| 3-7 | 134 (50) | 50/50 | 133 (50) | 99 | 50/50 | 130 (50) | 97 | 50/50 | 125 (50) | 93 | 50/50 |
| 4-7 | 143 (50) | 50/50 | 142 (50) | 99 | 50/50 | 139 (50) | 97 | 50/50 | 132 (50) | 92 | 50/50 |
| 5-7 | 152 (50) | 50/50 | 149 (50) | 98 | 50/50 | 148 (50) | 97 | 50/50 | 140 (50) | 92 | 50/50 |
| 6-7 | 156 (50) | 50/50 | 154 (50) | 99 | 50/50 | 153 (50) | 98 | 50/50 | 145 (50) | 93 | 50/50 |
| 7-7 | 162 (50) | 50/50 | 160 (50) | 99 | 50/50 | 160 (50) | 99 | 50/50 | 150 (50) | 93 | 50/50 |
| 8-7 | 165 (50) | 50/50 | 166 (50) | 101 | 50/50 | 164 (50) | 99 | 50/50 | 155 (50) | 94 | 50/50 |
| 9-7 | 169 (50) | 50/50 | 169 (50) | 100 | 50/50 | 168 (50) | 99 | 50/50 | 160 (50) | 95 | 50/50 |
| 10-7 | 173 (50) | 50/50 | 174 (50) | 101 | 50/50 | 173 (50) | 100 | 50/50 | 163 (50) | 94 | 50/50 |
| 11-7 | 177 (50) | 50/50 | 179 (50) | 101 | 50/50 | 177 (50) | 100 | 50/50 | 167 (50) | 94 | 50/50 |
| 12-7 | 178 (50) | 50/50 | 180 (50) | 101 | 50/50 | 180 (50) | 101 | 50/50 | 168 (50) | 94 | 50/50 |
| 13-7 | 180 (50) | 50/50 | 183 (50) | 102 | 50/50 | 182 (50) | 101 | 50/50 | 172 (50) | 96 | 50/50 |
| 14-7 | 183 (50) | 50/50 | 185 (50) | 101 | 50/50 | 183 (50) | 100 | 50/50 | 172 (50) | 94 | 50/50 |
| 18-7 | 188 (50) | 50/50 | 191 (50) | 102 | 50/50 | 191 (50) | 102 | 50/50 | 179 (50) | 95 | 50/50 |
| 22-7 | 195 (50) | 50/50 | 197 (50) | 101 | 50/50 | 198 (50) | 102 | 50/50 | 186 (50) | 95 | 50/50 |
| 26-7 | 201 (50) | 50/50 | 202 (50) | 100 | 50/50 | 202 (50) | 100 | 50/50 | 190 (50) | 95 | 50/50 |
| 30-7 | 204 (50) | 50/50 | 206 (50) | 101 | 50/50 | 208 (50) | 102 | 50/50 | 195 (50) | 96 | 50/50 |
| 34-7 | 209 (50) | 50/50 | 210 (50) | 100 | 50/50 | 214 (50) | 102 | 50/50 | 200 (50) | 96 | 50/50 |
| 38-7 | 215 (50) | 50/50 | 216 (50) | 100 | 50/50 | 218 (50) | 101 | 50/50 | 206 (50) | 96 | 50/50 |
| 42-7 | 221 (50) | 50/50 | 223 (50) | 101 | 50/50 | 226 (50) | 102 | 50/50 | 214 (50) | 97 | 50/50 |
| 46-7 | 225 (50) | 50/50 | 224 (50) | 100 | 50/50 | 228 (50) | 101 | 50/50 | 215 (50) | 96 | 50/50 |
| 50-7 | 228 (50) | 50/50 | 230 (50) | 101 | 50/50 | 233 (50) | 102 | 50/50 | 219 (50) | 96 | 50/50 |
| 54-7 | 235 (50) | 50/50 | 233 (50) | 99 | 50/50 | 236 (50) | 100 | 50/50 | 220 (50) | 94 | 50/50 |
| 58-7 | 238 (50) | 50/50 | 237 (49) | 100 | 49/50 | 240 (50) | 101 | 50/50 | 225 (50) | 95 | 50/50 |
| 62-7 | 242 (50) | 50/50 | 242 (49) | 100 | 49/50 | 242 (50) | 100 | 50/50 | 228 (50) | 94 | 50/50 |
| 66-7 | 249 (50) | 50/50 | 250 (49) | 100 | 49/50 | 250 (50) | 100 | 50/50 | 236 (49) | 95 | 49/50 |
| 70-7 | 255 (50) | 50/50 | 255 (49) | 100 | 49/50 | 253 (50) | 99 | 50/50 | 240 (47) | 94 | 47/50 |
| 74-7 | 261 (49) | 49/50 | 260 (48) | 100 | 48/50 | 259 (50) | 99 | 50/50 | 245 (46) | 94 | 46/50 |
| 78-7 | 266 (49) | 49/50 | 265 (48) | 100 | 48/50 | 261 (50) | 98 | 50/50 | 247 (46) | 93 | 46/50 |
| 82-7 | 269 (49) | 49/50 | 270 (48) | 100 | 48/50 | 265 (50) | 99 | 50/50 | 253 (44) | 94 | 44/50 |
| 86-7 | 273 (46) | 46/50 | 273 (48) | 100 | 48/50 | 267 (47) | 98 | 47/50 | 255 (44) | 93 | 44/50 |
| 90-7 | 279 (45) | 45/50 | 275 (48) | 99 | 48/50 | 270 (46) | 97 | 46/50 | 257 (44) | 92 | 44/50 |
| 94-7 | 277 (43) | 43/50 | 275 (47) | 99 | 47/50 | 271 (45) | 98 | 45/50 | 258 (44) | 93 | 44/50 |
| 98-7 | 283 (39) | 39/50 | 277 (46) | 98 | 46/50 | 274 (43) | 97 | 43/50 | 256 (42) | 90 | 42/50 |
| 102-7 | 284 (38) | 38/50 | 279 (44) | 98 | 44/50 | 276 (42) | 97 | 42/50 | 256 (38) | 90 | 38/50 |
| 104-7 | 283 (38) | 38/50 | 279 (43) | 99 | 43/50 | 273 (41) | 96 | 41/50 | 254 (37) | 90 | 37/50 |

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

TABLE D3

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 116± | 6 | 144± | 8 | 177± | 9 | 203± | 9 | 225± | 10 | 241± | 11 |
| 125 ppm | 116± | 6 | 142± | 8 | 175± | 10 | 202± | 10 | 224± | 10 | 242± | 12 |
| 250 ppm | 116± | 6 | 137± | 8** | 167± | 10** | 192± | 11** | 212± | 12** | 229± | 13** |
| 500 ppm | 116± | 6 | 132± | 8** | 159± | 8** | 179± | 9** | 198± | 10** | 213± | 10** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 272± | 12 | 284± | 12 | 294± | 13 | 304± | 14 | 312± | 15 | 318± | 16 |
| 125 ppm | 272± | 14 | 285± | 14 | 295± | 14 | 305± | 15 | 313± | 15 | 319± | 15 |
| 250 ppm | 259± | 14** | 272± | 15** | 284± | 16** | 292± | 16** | 301± | 17** | 306± | 17** |
| 500 ppm | 238± | 12** | 250± | 13** | 260± | 14** | 267± | 14** | 272± | 14** | 277± | 15** |

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 331± | 17 | 343± | 18 | 362± | 18 | 377± | 18 | 385± | 20 | 393± | 20 |
| 125 ppm | 334± | 16 | 350± | 17 | 369± | 19 | 383± | 21 | 390± | 21 | 400± | 22 |
| 250 ppm | 318± | 18** | 333± | 18** | 352± | 19* | 366± | 18* | 374± | 18* | 385± | 20 |
| 500 ppm | 288± | 15** | 303± | 16** | 317± | 18** | 332± | 19** | 338± | 20** | 347± | 21** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 410± | 22 | 416± | 23 | 421± | 23 | 426± | 24 | 427± | 24 | 431± | 24 |
| 125 ppm | 419± | 26 | 423± | 27 | 427± | 28 | 431± | 28 | 434± | 30 | 437± | 30 |
| 250 ppm | 401± | 20 | 407± | 19 | 410± | 20 | 414± | 22 | 418± | 21 | 422± | 23 |
| 500 ppm | 361± | 23** | 366± | 24** | 370± | 24** | 371± | 23** | 375± | 23** | 381± | 24** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 433± | 37 | 436± | 39 | 443± | 25 | 442± | 26 | 440± | 27 | 437± | 27 |
| 125 ppm | 443± | 29 | 444± | 30 | 443± | 30 | 442± | 32 | 441± | 31 | 439± | 33 |
| 250 ppm | 422± | 23** | 425± | 24* | 425± | 24** | 422± | 28** | 419± | 31** | 418± | 29** |
| 500 ppm | 378± | 23** | 378± | 24** | 378± | 22** | 374± | 26** | 373± | 29** | 372± | 24** |

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

Test of Dunnett

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
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 | 428± | 32 | 414± | 44 | 420± | 32 |
| 125 ppm | 429± | 32 | 423± | 30 | 417± | 30 |
| 250 ppm | 415± | 26 | 410± | 27 | 405± | 26 |
| 500 ppm | 363± | 25** | 355± | 29** | 350± | 29** |

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

Test of Dunnett

TABLE D4

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 96± | 5 | 110± | 6 | 123± | 6 | 134± | 6 | 143± | 7 | 152± | 8 |
| 125 ppm | 96± | 5 | 108± | 6 | 122± | 6 | 133± | 6 | 142± | 7 | 149± | 7 |
| 250 ppm | 96± | 5 | 107± | 6** | 120± | 7* | 130± | 7** | 139± | 7* | 148± | 8 |
| 500 ppm | 96± | 5 | 104± | 5** | 116± | 5** | 125± | 5** | 132± | 6** | 140± | 5** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 162± | 8 | 165± | 9 | 169± | 9 | 173± | 10 | 177± | 9 | 178± | 9 |
| 125 ppm | 160± | 8 | 166± | 9 | 169± | 9 | 174± | 9 | 179± | 9 | 180± | 10 |
| 250 ppm | 160± | 9 | 164± | 9 | 168± | 9 | 173± | 9 | 177± | 10 | 180± | 10 |
| 500 ppm | 150± | 6** | 155± | 7** | 160± | 7** | 163± | 7** | 167± | 8** | 168± | 8** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 183± | 10 | 188± | 10 | 195± | 11 | 201± | 12 | 204± | 12 | 209± | 13 |
| 125 ppm | 185± | 10 | 191± | 10 | 197± | 12 | 202± | 11 | 206± | 12 | 210± | 13 |
| 250 ppm | 183± | 11 | 191± | 11 | 198± | 12 | 202± | 13 | 208± | 12 | 214± | 13 |
| 500 ppm | 172± | 8** | 179± | 8** | 186± | 9** | 190± | 10** | 195± | 10** | 200± | 10** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 10

| Group Name | Administration | | week-day | | | | | | | | | |
|------------|----------------|-----|----------|------|------|------|------|------|------|------|------|------|
| | 42-7 | | 46-7 | | 50-7 | | 54-7 | | 58-7 | | 62-7 | |
| Control | 221± | 14 | 225± | 15 | 228± | 16 | 235± | 17 | 238± | 19 | 242± | 21 |
| 125 ppm | 223± | 14 | 224± | 14 | 230± | 16 | 233± | 16 | 237± | 17 | 242± | 17 |
| 250 ppm | 226± | 16 | 228± | 17 | 233± | 17 | 236± | 19 | 240± | 21 | 242± | 21 |
| 500 ppm | 214± | 11* | 215± | 11** | 219± | 11** | 220± | 11** | 225± | 13** | 228± | 13** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | |
| Control | 255± | 23 | 261± | 22 | 266± | 26 | 269± | 32 | 273± | 32 | 279± | 36 |
| 125 ppm | 255± | 21 | 260± | 21 | 265± | 21 | 270± | 22 | 273± | 23 | 275± | 23 |
| 250 ppm | 253± | 25 | 259± | 26 | 261± | 27 | 265± | 24 | 267± | 27 | 270± | 26 |
| 500 ppm | 240± | 14** | 245± | 14** | 247± | 17** | 253± | 16** | 255± | 18** | 257± | 19** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 283± | 23 | 284± | 22 | 283± | 23 |
| 125 ppm | 277± | 27 | 279± | 25 | 279± | 24 |
| 250 ppm | 274± | 27 | 276± | 28 | 273± | 28 |
| 500 ppm | 256± | 24** | 256± | 22** | 254± | 22** |

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

Test of Dunnett

TABLE E1

FOOD CONSUMPTION CHANGES AND SURVIVAL ANIMAL
NUMBERS : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : MALE

MEAN FOOD CONSUMPTION(FC) AND SURVIVAL

PAGE : 1

| Week-Day on Study | Control | | 125 ppm | | | 250 ppm | | | 500 ppm | | |
|----------------------|-----------|---------------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|
| | Av. FC. | No. of Surviv. <50> | Av. FC. | % of cont. <50> | No. of Surviv. | Av. FC. | % of cont. <50> | No. of Surviv. | Av. FC. | % of cont. <50> | No. of Surviv. |
| 1-7 | 13.7 (50) | 50/50 | 13.0 (50) | 95 | 50/50 | 12.3 (50) | 90 | 50/50 | 11.2 (50) | 82 | 50/50 |
| 2-7 | 15.5 (50) | 50/50 | 15.7 (50) | 101 | 50/50 | 14.7 (50) | 95 | 50/50 | 13.9 (50) | 90 | 50/50 |
| 3-7 | 16.5 (50) | 50/50 | 16.7 (50) | 101 | 50/50 | 16.0 (50) | 97 | 50/50 | 15.6 (50) | 95 | 50/50 |
| 4-7 | 16.3 (50) | 50/50 | 17.2 (50) | 106 | 50/50 | 16.7 (50) | 102 | 50/50 | 16.7 (50) | 102 | 50/50 |
| 5-7 | 16.3 (50) | 50/50 | 17.2 (50) | 106 | 50/50 | 17.2 (50) | 106 | 50/50 | 17.3 (50) | 106 | 50/50 |
| 6-7 | 16.0 (50) | 50/50 | 17.0 (50) | 106 | 50/50 | 17.2 (50) | 108 | 50/50 | 17.4 (50) | 109 | 50/50 |
| 7-7 | 16.5 (50) | 50/50 | 17.6 (50) | 107 | 50/50 | 17.7 (50) | 107 | 50/50 | 17.9 (50) | 108 | 50/50 |
| 8-7 | 16.4 (50) | 50/50 | 17.0 (50) | 104 | 50/50 | 17.1 (50) | 104 | 50/50 | 17.4 (50) | 106 | 50/50 |
| 9-7 | 16.6 (50) | 50/50 | 17.3 (50) | 104 | 50/50 | 17.5 (50) | 105 | 50/50 | 18.2 (50) | 110 | 50/50 |
| 10-7 | 16.2 (50) | 50/50 | 16.9 (50) | 104 | 50/50 | 17.0 (50) | 105 | 50/50 | 18.3 (50) | 113 | 50/50 |
| 11-7 | 16.4 (45) | 50/50 | 17.4 (50) | 106 | 50/50 | 17.4 (50) | 106 | 50/50 | 18.4 (50) | 112 | 50/50 |
| 12-7 | 16.3 (50) | 50/50 | 17.2 (50) | 106 | 50/50 | 17.1 (50) | 105 | 50/50 | 18.4 (50) | 113 | 50/50 |
| 13-7 | 16.2 (50) | 50/50 | 17.3 (50) | 107 | 50/50 | 17.1 (50) | 106 | 50/50 | 18.5 (50) | 114 | 50/50 |
| 14-7 | 16.2 (50) | 50/50 | 17.0 (50) | 105 | 50/50 | 17.0 (50) | 105 | 50/50 | 18.5 (50) | 114 | 50/50 |
| 18-7 | 16.2 (50) | 50/50 | 17.4 (50) | 107 | 50/50 | 17.2 (50) | 106 | 50/50 | 19.3 (50) | 119 | 50/50 |
| 22-7 | 16.2 (50) | 50/50 | 17.2 (50) | 106 | 50/50 | 17.0 (50) | 105 | 50/50 | 18.9 (50) | 117 | 50/50 |
| 26-7 | 16.2 (50) | 50/50 | 16.6 (50) | 102 | 50/50 | 16.6 (49) | 102 | 49/50 | 17.9 (50) | 110 | 50/50 |
| 30-7 | 16.2 (50) | 50/50 | 17.2 (50) | 106 | 50/50 | 17.2 (49) | 106 | 49/50 | 19.0 (50) | 117 | 50/50 |
| 34-7 | 16.1 (50) | 50/50 | 16.8 (50) | 104 | 50/50 | 16.9 (49) | 105 | 49/50 | 18.6 (50) | 116 | 50/50 |
| 38-7 | 16.1 (50) | 50/50 | 17.1 (50) | 106 | 50/50 | 17.2 (49) | 107 | 49/50 | 18.9 (50) | 117 | 50/50 |
| 42-7 | 16.5 (50) | 50/50 | 17.0 (50) | 103 | 50/50 | 16.9 (49) | 102 | 49/50 | 18.5 (50) | 112 | 50/50 |
| 46-7 | 16.1 (50) | 50/50 | 16.8 (50) | 104 | 50/50 | 16.7 (49) | 104 | 49/50 | 18.2 (50) | 113 | 50/50 |
| 50-7 | 16.5 (50) | 50/50 | 17.1 (50) | 104 | 50/50 | 17.2 (49) | 104 | 49/50 | 18.6 (50) | 113 | 50/50 |
| 54-7 | 16.5 (50) | 50/50 | 16.9 (50) | 102 | 50/50 | 17.1 (49) | 104 | 49/50 | 18.4 (50) | 112 | 50/50 |
| 58-7 | 17.0 (50) | 50/50 | 17.6 (50) | 104 | 50/50 | 17.7 (49) | 104 | 49/50 | 19.1 (50) | 112 | 50/50 |
| 62-7 | 16.6 (50) | 50/50 | 16.9 (50) | 102 | 50/50 | 16.9 (49) | 102 | 49/50 | 17.7 (50) | 107 | 50/50 |
| 66-7 | 16.7 (50) | 50/50 | 17.6 (50) | 105 | 50/50 | 17.2 (49) | 103 | 49/50 | 18.2 (50) | 109 | 50/50 |
| 70-7 | 16.8 (50) | 50/50 | 17.6 (49) | 105 | 49/50 | 17.1 (49) | 102 | 49/50 | 18.1 (49) | 108 | 49/50 |
| 74-7 | 16.5 (50) | 50/50 | 17.2 (47) | 104 | 47/50 | 17.1 (49) | 104 | 49/50 | 17.9 (49) | 108 | 49/50 |
| 78-7 | 16.4 (49) | 49/50 | 16.7 (46) | 102 | 46/50 | 16.9 (49) | 103 | 49/50 | 17.3 (48) | 105 | 48/50 |
| 82-7 | 16.4 (48) | 48/50 | 16.8 (44) | 102 | 44/50 | 16.7 (49) | 102 | 49/50 | 17.3 (48) | 105 | 48/50 |
| 86-7 | 16.3 (48) | 48/50 | 16.8 (43) | 103 | 43/50 | 16.6 (49) | 102 | 49/50 | 16.8 (45) | 103 | 45/50 |
| 90-7 | 17.0 (47) | 47/50 | 17.1 (42) | 101 | 42/50 | 17.0 (48) | 100 | 48/50 | 17.1 (44) | 101 | 44/50 |
| 94-7 | 16.8 (46) | 46/50 | 16.8 (40) | 100 | 40/50 | 16.6 (46) | 99 | 46/50 | 16.9 (43) | 101 | 43/50 |
| 98-7 | 17.0 (45) | 45/50 | 17.0 (38) | 100 | 38/50 | 16.7 (44) | 98 | 44/50 | 17.2 (41) | 101 | 41/50 |
| 102-7 | 15.6 (44) | 44/50 | 16.7 (37) | 107 | 37/50 | 16.9 (43) | 108 | 43/50 | 17.4 (40) | 112 | 40/50 |
| 104-7 | 16.6 (39) | 39/50 | 16.3 (35) | 98 | 35/50 | 16.5 (41) | 99 | 41/50 | 17.1 (37) | 103 | 37/50 |

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

Av. FC. : g

TABLE E2

FOOD CONSUMPTION CHANGES AND SURVIVAL ANIMAL
NUMBERS : FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : FEMALE

MEAN FOOD CONSUMPTION(FC) AND SURVIVAL

PAGE : 2

| Week-Day on Study | Control | | 125 ppm | | | 250 ppm | | | 500 ppm | | |
|----------------------|-----------|---------------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|-----------|-----------------------|-------------------|
| | Av. FC. | No. of Surviv. <50> | Av. FC. | % of cont. <50> | No. of Surviv. | Av. FC. | % of cont. <50> | No. of Surviv. | Av. FC. | % of cont. <50> | No. of Surviv. |
| 1-7 | 10.3 (50) | 50/50 | 10.1 (50) | 98 | 50/50 | 9.6 (50) | 93 | 50/50 | 9.0 (50) | 87 | 50/50 |
| 2-7 | 10.9 (50) | 50/50 | 11.1 (50) | 102 | 50/50 | 10.7 (50) | 98 | 50/50 | 10.6 (50) | 97 | 50/50 |
| 3-7 | 11.1 (50) | 50/50 | 11.4 (50) | 103 | 50/50 | 11.3 (50) | 102 | 50/50 | 11.0 (50) | 99 | 50/50 |
| 4-7 | 10.9 (50) | 50/50 | 11.3 (50) | 104 | 50/50 | 11.5 (50) | 106 | 50/50 | 11.2 (50) | 103 | 50/50 |
| 5-7 | 11.1 (50) | 50/50 | 11.4 (50) | 103 | 50/50 | 12.1 (50) | 109 | 50/50 | 11.8 (50) | 106 | 50/50 |
| 6-7 | 10.5 (50) | 50/50 | 11.0 (50) | 105 | 50/50 | 11.4 (50) | 109 | 50/50 | 11.7 (50) | 111 | 50/50 |
| 7-7 | 10.7 (50) | 50/50 | 11.2 (50) | 105 | 50/50 | 11.8 (50) | 110 | 50/50 | 11.9 (50) | 111 | 50/50 |
| 8-7 | 10.4 (50) | 50/50 | 10.7 (50) | 103 | 50/50 | 10.9 (50) | 105 | 50/50 | 11.3 (50) | 109 | 50/50 |
| 9-7 | 10.5 (50) | 50/50 | 11.2 (50) | 107 | 50/50 | 11.8 (50) | 112 | 50/50 | 12.1 (50) | 115 | 50/50 |
| 10-7 | 10.3 (50) | 50/50 | 11.0 (50) | 107 | 50/50 | 11.5 (50) | 112 | 50/50 | 12.2 (50) | 118 | 50/50 |
| 11-7 | 10.6 (50) | 50/50 | 11.5 (50) | 108 | 50/50 | 12.1 (50) | 114 | 50/50 | 12.6 (50) | 119 | 50/50 |
| 12-7 | 10.1 (50) | 50/50 | 11.2 (50) | 111 | 50/50 | 11.8 (50) | 117 | 50/50 | 12.4 (50) | 123 | 50/50 |
| 13-7 | 10.5 (50) | 50/50 | 11.3 (50) | 108 | 50/50 | 11.7 (50) | 111 | 50/50 | 12.9 (50) | 123 | 50/50 |
| 14-7 | 10.6 (50) | 50/50 | 11.2 (50) | 106 | 50/50 | 11.3 (50) | 107 | 50/50 | 12.4 (50) | 117 | 50/50 |
| 18-7 | 10.6 (50) | 50/50 | 11.3 (50) | 107 | 50/50 | 11.8 (50) | 111 | 50/50 | 13.6 (50) | 128 | 50/50 |
| 22-7 | 10.4 (50) | 50/50 | 10.9 (50) | 105 | 50/50 | 11.7 (50) | 113 | 50/50 | 13.2 (50) | 127 | 50/50 |
| 26-7 | 10.6 (50) | 50/50 | 10.5 (50) | 99 | 50/50 | 10.5 (50) | 99 | 50/50 | 11.6 (50) | 109 | 50/50 |
| 30-7 | 10.3 (50) | 50/50 | 11.1 (50) | 108 | 50/50 | 11.6 (50) | 113 | 50/50 | 13.4 (50) | 130 | 50/50 |
| 34-7 | 10.6 (50) | 50/50 | 11.0 (50) | 104 | 50/50 | 12.0 (50) | 113 | 50/50 | 12.8 (50) | 121 | 50/50 |
| 38-7 | 11.0 (50) | 50/50 | 11.3 (50) | 103 | 50/50 | 12.1 (50) | 110 | 50/50 | 13.6 (50) | 124 | 50/50 |
| 42-7 | 10.8 (50) | 50/50 | 11.2 (50) | 104 | 50/50 | 12.2 (50) | 113 | 50/50 | 13.6 (50) | 126 | 50/50 |
| 46-7 | 11.3 (50) | 50/50 | 11.1 (50) | 98 | 50/50 | 11.9 (50) | 105 | 50/50 | 12.8 (50) | 113 | 50/50 |
| 50-7 | 11.0 (50) | 50/50 | 11.6 (50) | 105 | 50/50 | 12.4 (50) | 113 | 50/50 | 13.8 (50) | 125 | 50/50 |
| 54-7 | 11.4 (50) | 50/50 | 11.5 (50) | 101 | 50/50 | 12.1 (50) | 106 | 50/50 | 13.4 (50) | 118 | 50/50 |
| 58-7 | 11.6 (50) | 50/50 | 11.9 (49) | 103 | 49/50 | 12.5 (50) | 108 | 50/50 | 14.0 (50) | 121 | 50/50 |
| 62-7 | 11.1 (50) | 50/50 | 11.6 (49) | 105 | 49/50 | 11.9 (50) | 107 | 50/50 | 12.8 (50) | 115 | 50/50 |
| 66-7 | 11.4 (50) | 50/50 | 12.1 (49) | 106 | 49/50 | 12.5 (50) | 110 | 50/50 | 13.6 (49) | 119 | 49/50 |
| 70-7 | 11.8 (50) | 50/50 | 11.9 (49) | 101 | 49/50 | 12.3 (50) | 104 | 50/50 | 13.4 (47) | 114 | 47/50 |
| 74-7 | 11.7 (49) | 49/50 | 12.0 (48) | 103 | 48/50 | 12.4 (50) | 106 | 50/50 | 13.1 (46) | 112 | 46/50 |
| 78-7 | 11.5 (49) | 49/50 | 11.8 (48) | 103 | 48/50 | 11.7 (50) | 102 | 50/50 | 12.5 (46) | 109 | 46/50 |
| 82-7 | 11.5 (49) | 49/50 | 11.9 (48) | 103 | 48/50 | 12.4 (50) | 108 | 50/50 | 13.3 (44) | 116 | 44/50 |
| 86-7 | 11.3 (46) | 46/50 | 11.9 (48) | 105 | 48/50 | 11.9 (47) | 105 | 47/50 | 12.8 (44) | 113 | 44/50 |
| 90-7 | 12.6 (44) | 45/50 | 12.2 (48) | 97 | 48/50 | 12.8 (46) | 102 | 46/50 | 13.5 (44) | 107 | 44/50 |
| 94-7 | 12.1 (43) | 43/50 | 12.2 (47) | 101 | 47/50 | 12.7 (45) | 105 | 45/50 | 13.1 (44) | 108 | 44/50 |
| 98-7 | 12.7 (39) | 39/50 | 12.2 (46) | 96 | 46/50 | 12.3 (43) | 97 | 43/50 | 12.7 (42) | 100 | 42/50 |
| 102-7 | 12.4 (38) | 38/50 | 12.5 (44) | 101 | 44/50 | 12.6 (42) | 102 | 42/50 | 13.0 (38) | 105 | 38/50 |
| 104-7 | 12.4 (38) | 38/50 | 12.1 (43) | 98 | 43/50 | 12.2 (41) | 98 | 41/50 | 13.0 (37) | 105 | 37/50 |

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

Av. FC. : g

TABLE E3

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 1

| Group Name | Administration week-day(effective) | | | | | | |
|------------|------------------------------------|-------------|-------------|-------------|-------------|-------------|-------------|
| | 1-7(6) | 2-7(7) | 3-7(7) | 4-7(7) | 5-7(7) | 6-7(7) | 7-7(7) |
| Control | 13.7± 0.8 | 15.5± 0.9 | 16.5± 1.1 | 16.3± 1.1 | 16.3± 0.9 | 16.0± 0.9 | 16.5± 0.8 |
| 125 ppm | 13.0± 0.9** | 15.7± 1.2 | 16.7± 1.0 | 17.2± 1.2** | 17.2± 1.3** | 17.0± 1.3** | 17.6± 1.3** |
| 250 ppm | 12.3± 1.0** | 14.7± 1.3** | 16.0± 1.5 | 16.7± 1.5 | 17.2± 1.6** | 17.2± 1.5** | 17.7± 1.4** |
| 500 ppm | 11.2± 0.8** | 13.9± 1.2** | 15.6± 1.3** | 16.7± 1.3 | 17.3± 1.4** | 17.4± 1.3** | 17.9± 1.3** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 2

| Group Name | Administration week-day(effective) | | | | | | |
|------------|------------------------------------|-------------|-------------|-------------|-------------|-------------|-------------|
| | 8-7(7) | 9-7(7) | 10-7(7) | 11-7(7) | 12-7(7) | 13-7(7) | 14-7(7) |
| Control | 16.4± 0.9 | 16.6± 0.9 | 16.2± 1.1 | 16.4± 1.1 | 16.3± 1.1 | 16.2± 1.0 | 16.2± 1.0 |
| 125 ppm | 17.0± 1.2** | 17.3± 1.1** | 16.9± 1.1* | 17.4± 1.2** | 17.2± 1.2** | 17.3± 1.1** | 17.0± 1.1** |
| 250 ppm | 17.1± 1.4** | 17.5± 1.5** | 17.0± 1.3** | 17.4± 1.5** | 17.1± 1.5** | 17.1± 1.4** | 17.0± 1.4** |
| 500 ppm | 17.4± 1.4** | 18.2± 1.4** | 18.3± 1.5** | 18.4± 1.5** | 18.4± 1.5** | 18.5± 1.4** | 18.5± 1.5** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 16.2± 1.1 | 16.2± 0.9 | 16.2± 0.9 | 16.2± 1.0 | 16.1± 0.9 | 16.1± 0.9 | 16.5± 0.8 |
| 125 ppm | 17.4± 1.4** | 17.2± 1.4** | 16.6± 1.3 | 17.2± 1.2** | 16.8± 1.1** | 17.1± 1.0** | 17.0± 1.1** |
| 250 ppm | 17.2± 1.5** | 17.0± 1.3** | 16.6± 0.9 | 17.2± 0.9** | 16.9± 0.9** | 17.2± 1.1** | 16.9± 0.9* |
| 500 ppm | 19.3± 1.9** | 18.9± 1.9** | 17.9± 1.7** | 19.0± 1.8** | 18.6± 1.9** | 18.9± 2.0** | 18.5± 2.0** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 16.1± 0.7 | 16.5± 0.8 | 16.5± 0.9 | 17.0± 0.9 | 16.6± 1.0 | 16.7± 0.9 | 16.8± 1.2 |
| 125 ppm | 16.8± 1.2** | 17.1± 1.2** | 16.9± 1.2 | 17.6± 1.3** | 16.9± 1.1 | 17.6± 1.1** | 17.6± 1.1** |
| 250 ppm | 16.7± 1.0* | 17.2± 0.9** | 17.1± 1.1** | 17.7± 1.0** | 16.9± 1.2 | 17.2± 1.2 | 17.1± 1.0 |
| 500 ppm | 18.2± 1.8** | 18.6± 1.8** | 18.4± 1.7** | 19.1± 2.1** | 17.7± 1.6** | 18.2± 2.1** | 18.1± 2.0** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 16.5± 1.1 | 16.4± 1.0 | 16.4± 1.2 | 16.3± 2.1 | 17.0± 1.4 | 16.8± 2.0 | 17.0± 1.6 |
| 125 ppm | 17.2± 1.1** | 16.7± 1.2 | 16.8± 1.3 | 16.8± 1.1 | 17.1± 1.6 | 16.8± 2.0 | 17.0± 1.4 |
| 250 ppm | 17.1± 1.2 | 16.9± 1.6 | 16.7± 1.3 | 16.6± 1.8 | 17.0± 1.7 | 16.6± 1.9 | 16.7± 1.3 |
| 500 ppm | 17.9± 2.0** | 17.3± 1.8 | 17.3± 2.7* | 16.8± 3.0 | 17.1± 2.1 | 16.9± 2.6 | 17.2± 2.2 |

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

Test of Dunnett

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
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 | 15.6± 3.2 | 16.6± 1.6 |
| 125 ppm | 16.7± 2.8 | 16.3± 1.5 |
| 250 ppm | 16.9± 1.5 | 16.5± 1.3 |
| 500 ppm | 17.4± 2.3* | 17.1± 2.4 |

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

Test of Dunnett

TABLE E4

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 7

| Group Name | Administration week-day(effective) | | | | | | |
|------------|------------------------------------|-----------|-----------|-------------|-------------|-------------|-------------|
| | 1-7(6) | 2-7(7) | 3-7(7) | 4-7(7) | 5-7(7) | 6-7(7) | 7-7(7) |
| Control | 10.3± 0.8 | 10.9± 0.8 | 11.1± 0.8 | 10.9± 0.7 | 11.1± 0.9 | 10.5± 0.7 | 10.7± 0.8 |
| 125 ppm | 10.1± 0.7 | 11.1± 0.9 | 11.4± 0.8 | 11.3± 0.9 | 11.4± 0.8 | 11.0± 0.9* | 11.2± 0.8* |
| 250 ppm | 9.6± 0.8** | 10.7± 0.8 | 11.3± 0.9 | 11.5± 1.0** | 12.1± 1.0** | 11.4± 0.9** | 11.8± 1.0** |
| 500 ppm | 9.0± 0.5** | 10.6± 0.6 | 11.0± 0.7 | 11.2± 0.7 | 11.8± 0.7** | 11.7± 0.9** | 11.9± 1.0** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 UNIT : g
 REPORT TYPE : A1 104
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 8

| Group Name | Administration week-day(effective) | | | | | | |
|------------|------------------------------------|-------------|-------------|-------------|-------------|-------------|-------------|
| | 8-7(7) | 9-7(7) | 10-7(7) | 11-7(7) | 12-7(7) | 13-7(7) | 14-7(7) |
| Control | 10.4± 0.9 | 10.5± 1.0 | 10.3± 0.8 | 10.6± 0.7 | 10.1± 0.7 | 10.5± 0.7 | 10.6± 0.8 |
| 125 ppm | 10.7± 1.0 | 11.2± 1.1** | 11.0± 0.8** | 11.5± 0.9** | 11.2± 1.0** | 11.3± 1.2** | 11.2± 1.0** |
| 250 ppm | 10.9± 0.8* | 11.8± 1.1** | 11.5± 0.9** | 12.1± 1.0** | 11.8± 1.0** | 11.7± 1.0** | 11.3± 0.9** |
| 500 ppm | 11.3± 0.9** | 12.1± 0.8** | 12.2± 0.9** | 12.6± 0.9** | 12.4± 1.0** | 12.9± 1.0** | 12.4± 1.0** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 10.6± 0.9 | 10.4± 0.9 | 10.6± 0.8 | 10.3± 1.0 | 10.6± 0.8 | 11.0± 1.0 | 10.8± 0.9 |
| 125 ppm | 11.3± 1.1** | 10.9± 1.0 | 10.5± 0.8 | 11.1± 1.1** | 11.0± 1.1 | 11.3± 1.0 | 11.2± 1.0 |
| 250 ppm | 11.8± 1.1** | 11.7± 1.0** | 10.5± 0.7 | 11.6± 0.7** | 12.0± 1.0** | 12.1± 1.2** | 12.2± 1.2** |
| 500 ppm | 13.6± 1.2** | 13.2± 1.2** | 11.6± 1.0** | 13.4± 1.4** | 12.8± 1.1** | 13.6± 1.3** | 13.6± 1.5** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 11.3± 0.9 | 11.0± 1.1 | 11.4± 1.1 | 11.6± 1.0 | 11.1± 1.0 | 11.4± 1.0 | 11.8± 0.8 |
| 125 ppm | 11.1± 0.9 | 11.6± 1.1* | 11.5± 1.0 | 11.9± 1.0 | 11.6± 1.0 | 12.1± 1.1** | 11.9± 1.0 |
| 250 ppm | 11.9± 1.3 | 12.4± 0.9** | 12.1± 1.2** | 12.5± 1.1** | 11.9± 1.0** | 12.5± 1.1** | 12.3± 1.1 |
| 500 ppm | 12.8± 1.3** | 13.8± 1.3** | 13.4± 1.2** | 14.0± 1.8** | 12.8± 0.9** | 13.6± 1.3** | 13.4± 1.1** |

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

Test of Dunnett

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 11.7± 1.1 | 11.5± 1.2 | 11.5± 1.2 | 11.3± 1.3 | 12.6± 1.0 | 12.1± 1.4 | 12.7± 1.3 |
| 125 ppm | 12.0± 1.0 | 11.8± 0.9 | 11.9± 1.1 | 11.9± 1.2* | 12.2± 1.3 | 12.2± 1.2 | 12.2± 1.8 |
| 250 ppm | 12.4± 1.0** | 11.7± 1.0 | 12.4± 1.0** | 11.9± 1.1* | 12.8± 1.1 | 12.7± 1.2 | 12.3± 1.1 |
| 500 ppm | 13.1± 1.1** | 12.5± 1.7** | 13.3± 1.0** | 12.8± 1.1** | 13.5± 1.0** | 13.1± 1.3** | 12.7± 1.9 |

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

Test of Dunnett

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
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 | 12.4± 1.0 | 12.4± 1.3 |
| 125 ppm | 12.5± 1.2 | 12.1± 1.3 |
| 250 ppm | 12.6± 1.2 | 12.2± 1.2 |
| 500 ppm | 13.0± 0.9 | 13.0± 1.0 |

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

Test of Dunnett

TABLE F1

HEMATOLOGY : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ³ /μl | |
|------------|-------------------|---------------------------------------|--------|--------------------|------|-----------------|------|-----------|-------|-----------|-------|--------------|-----|---------------------------------|-------|
| Control | 38 | 8.13± | 1.50 | 14.3± | 2.8 | 39.5± | 6.6 | 48.8± | 2.8 | 17.5± | 1.2 | 35.9± | 1.7 | 811± | 248 |
| 125 ppm | 34 | 9.10± | 1.22* | 15.8± | 2.0* | 43.3± | 5.1* | 47.7± | 1.7* | 17.4± | 0.6 | 36.5± | 0.8 | 813± | 119 |
| 250 ppm | 41 | 8.96± | 1.50* | 15.5± | 2.3 | 42.3± | 5.7 | 47.7± | 3.9** | 17.4± | 1.0 | 36.5± | 1.2 | 834± | 225 |
| 500 ppm | 37 | 9.26± | 1.60** | 15.5± | 2.8 | 42.8± | 6.7 | 46.4± | 2.2** | 16.8± | 0.8** | 36.1± | 1.6 | 906± | 202** |

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
MEASURE. TIME : 1
SEX : MALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | RETICULOCYTE % | |
|------------|-------------------|-------------------|-------|
| Control | 38 | 4.3± | 3.8 |
| 125 ppm | 34 | 2.9± | 2.5* |
| 250 ppm | 41 | 3.9± | 6.2** |
| 500 ppm | 37 | 3.4± | 3.6** |

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 3

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHER | |
|------------|-------------------|----------------------------|-------|------------------------|---|------------------|----|--------|---|------|---|------|---|--------|----|-------|----|
| Control | 38 | 11.86± | 22.20 | 0± | 1 | 47± | 13 | 2± | 1 | 0± | 0 | 5± | 2 | 40± | 11 | 6± | 18 |
| 125 ppm | 34 | 6.68± | 1.80 | 0± | 1 | 46± | 10 | 2± | 1 | 0± | 0 | 6± | 1 | 45± | 10 | 1± | 1 |
| 250 ppm | 41 | 7.72± | 3.85 | 0± | 1 | 46± | 9 | 2± | 1 | 0± | 0 | 6± | 2 | 43± | 10 | 3± | 13 |
| 500 ppm | 37 | 9.08± | 5.44* | 0± | 1 | 47± | 10 | 1± | 1 | 0± | 0 | 6± | 2 | 44± | 9 | 1± | 3 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS4

TABLE F2

HEMATOLOGY : FEMALE

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ³ /μl | |
|------------|-------------------|---------------------------------------|--------|--------------------|-------|-----------------|-------|-----------|-----|-----------|-----|--------------|-----|---------------------------------|------|
| Control | 38 | 7.73± | 1.28 | 14.5± | 2.3 | 39.0± | 5.7 | 50.7± | 3.1 | 18.8± | 0.7 | 37.1± | 1.4 | 683± | 226 |
| 125 ppm | 43 | 8.02± | 1.08 | 14.9± | 2.0 | 40.3± | 4.2 | 50.6± | 3.3 | 18.6± | 1.1 | 36.8± | 1.8 | 680± | 222 |
| 250 ppm | 41 | 7.92± | 1.21 | 14.9± | 2.3 | 40.1± | 5.0 | 51.3± | 5.5 | 18.9± | 1.7 | 36.8± | 2.0 | 660± | 181 |
| 500 ppm | 37 | 8.37± | 0.40** | 15.8± | 0.7** | 42.4± | 1.3** | 50.7± | 1.4 | 18.8± | 0.3 | 37.2± | 0.8 | 724± | 84** |

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
MEASURE. TIME : 1
SEX : FEMALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 5

| Group Name | NO. of Animals | RETICULOCYTE % | |
|------------|-------------------|-------------------|-----|
| Control | 38 | 3.6± | 4.9 |
| 125 ppm | 43 | 3.8± | 5.5 |
| 250 ppm | 41 | 3.8± | 5.4 |
| 500 ppm | 37 | 2.6± | 1.7 |

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

PAGE : 6

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHER | |
|------------|-------------------|----------------------------|-------|------------------------|---|------------------|------|--------|---|------|---|------|-----|--------|----|-------|----|
| Control | 38 | 5.17± | 3.73 | 1± | 1 | 44± | 14 | 2± | 1 | 0± | 0 | 5± | 2 | 47± | 14 | 2± | 7 |
| 125 ppm | 43 | 4.55± | 5.32 | 1± | 1 | 38± | 14 | 2± | 1 | 0± | 0 | 5± | 2 | 51± | 14 | 4± | 14 |
| 250 ppm | 41 | 7.90± | 18.80 | 1± | 1 | 34± | 11** | 2± | 2 | 0± | 0 | 5± | 2 | 51± | 14 | 7± | 22 |
| 500 ppm | 37 | 4.40± | 1.66 | 0± | 1 | 39± | 8 | 2± | 1 | 0± | 0 | 6± | 1** | 52± | 8 | 1± | 1 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

TABLE G1

BIOCHEMISTRY : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | TOTAL PROTEIN g/dl | | ALBUMIN g/dl | | A/G RATIO | | T-BILIRUBIN mg/dl | | GLUCOSE mg/dl | | T-CHOLESTEROL mg/dl | | TRIGLYCERIDE mg/dl | |
|------------|-------------------|-----------------------|-----|-----------------|-------|-----------|-------|----------------------|------|------------------|----|------------------------|------|-----------------------|-------|
| Control | 38 | 6.7± | 0.3 | 3.0± | 0.3 | 0.8± | 0.1 | 0.17± | 0.06 | 156± | 24 | 207± | 80 | 184± | 131 |
| 125 ppm | 34 | 6.9± | 0.3 | 3.0± | 0.2 | 0.8± | 0.1 | 0.16± | 0.04 | 158± | 19 | 249± | 70* | 213± | 99 |
| 250 ppm | 41 | 6.8± | 0.2 | 3.0± | 0.2 | 0.8± | 0.1 | 0.20± | 0.24 | 155± | 23 | 259± | 65** | 245± | 130 |
| 500 ppm | 37 | 6.7± | 0.4 | 2.8± | 0.3** | 0.7± | 0.1** | 0.18± | 0.08 | 155± | 14 | 260± | 79** | 274± | 126** |

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | PHOSPHOLIPID mg/dl | | AST IU/l | | ALT IU/l | | LDH IU/l | | ALP IU/l | | G-GTP IU/l | | CK IU/l | |
|------------|-------------------|-----------------------|------|-------------|-----|-------------|----|-------------|-----|-------------|-----|---------------|------|------------|-----|
| Control | 38 | 302± | 114 | 98± | 116 | 39± | 19 | 228± | 241 | 226± | 148 | 7± | 4 | 139± | 176 |
| 125 ppm | 34 | 351± | 96 | 69± | 18 | 31± | 6 | 173± | 56 | 166± | 58* | 9± | 5 | 99± | 26 |
| 250 ppm | 41 | 358± | 88* | 86± | 89 | 35± | 18 | 187± | 127 | 200± | 69 | 13± | 8** | 95± | 21 |
| 500 ppm | 37 | 364± | 111* | 82± | 42 | 41± | 24 | 162± | 50 | 237± | 101 | 17± | 10** | 96± | 25 |

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | UREA NITROGEN mg/dl | | CREATININE mg/dl | | SODIUM mEq/l | | POTASSIUM mEq/l | | CHLORIDE mEq/l | | CALCIUM mg/dl | | INORGANIC PHOSPHORUS mg/dl | |
|------------|-------------------|------------------------|-------|---------------------|-----|-----------------|-----|--------------------|------|-------------------|-----|------------------|-----|-------------------------------|-----|
| Control | 38 | 21.3± | 11.5 | 0.7± | 0.4 | 143± | 2 | 3.6± | 0.4 | 105± | 2 | 10.8± | 0.6 | 4.4± | 1.4 |
| 125 ppm | 34 | 20.6± | 4.6 | 0.6± | 0.1 | 142± | 1** | 3.7± | 0.3 | 107± | 1** | 10.8± | 0.4 | 4.0± | 0.7 |
| 250 ppm | 41 | 20.8± | 4.9 | 0.6± | 0.1 | 142± | 2** | 3.7± | 0.3 | 110± | 2** | 10.7± | 0.4 | 4.1± | 0.6 |
| 500 ppm | 37 | 23.0± | 6.2** | 0.6± | 0.1 | 142± | 1** | 3.8± | 0.3* | 112± | 3** | 10.9± | 0.8 | 4.4± | 0.6 |

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

Test of Dunnett

(HCL074)

BAIS 4

TABLE G2

BIOCHEMISTRY : FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of Animals | TOTAL PROTEIN g/dl | | ALBUMIN g/dl | | A/G RATIO | | T-BILIRUBIN mg/dl | | GLUCOSE mg/dl | | T-CHOLESTEROL mg/dl | | TRIGLYCERIDE mg/dl | |
|------------|-------------------|-----------------------|-----|-----------------|-------|-----------|-------|----------------------|------|------------------|----|------------------------|------|-----------------------|-----|
| Control | 38 | 7.0± | 0.4 | 3.6± | 0.3 | 1.1± | 0.2 | 0.14± | 0.07 | 143± | 18 | 140± | 26 | 87± | 59 |
| 125 ppm | 43 | 7.1± | 0.3 | 3.7± | 0.3 | 1.1± | 0.1 | 0.13± | 0.04 | 145± | 17 | 149± | 35 | 98± | 64 |
| 250 ppm | 41 | 7.1± | 0.3 | 3.7± | 0.3 | 1.1± | 0.1 | 0.15± | 0.14 | 148± | 19 | 160± | 47 | 128± | 91* |
| 500 ppm | 37 | 6.9± | 0.3 | 3.4± | 0.2** | 1.0± | 0.1** | 0.12± | 0.01 | 149± | 13 | 192± | 70** | 134± | 89* |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 5

| Group Name | NO. of Animals | PHOSPHOLIPID mg/dl | | AST IU/l | | ALT IU/l | | LDH IU/l | | ALP IU/l | | G-GTP IU/l | | CK IU/l | |
|------------|-------------------|-----------------------|-------|-------------|-----|-------------|----|-------------|-----|-------------|----|---------------|----|------------|------|
| Control | 38 | 256± | 48 | 121± | 78 | 48± | 20 | 212± | 81 | 132± | 58 | 3± | 2 | 91± | 25 |
| 125 ppm | 43 | 271± | 64 | 116± | 84 | 48± | 21 | 206± | 72 | 119± | 59 | 3± | 2 | 85± | 25 |
| 250 ppm | 41 | 286± | 73 | 141± | 146 | 56± | 46 | 211± | 112 | 131± | 75 | 4± | 5 | 91± | 70 |
| 500 ppm | 37 | 324± | 110** | 90± | 38 | 43± | 18 | 164± | 47* | 113± | 42 | 3± | 2* | 75± | 22** |

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 6

| Group Name | NO. of Animals | UREA NITROGEN mg/dl | | CREATININE mg/dl | | SODIUM mEq/l | | POTASSIUM mEq/l | | CHLORIDE mEq/l | | CALCIUM mg/dl | | INORGANIC PHOSPHORUS mg/dl | |
|------------|-------------------|------------------------|-----|---------------------|-----|-----------------|-----|--------------------|-----|-------------------|-----|------------------|-----|-------------------------------|-----|
| Control | 38 | 17.1± | 1.8 | 0.5± | 0.0 | 142± | 1 | 3.4± | 0.4 | 104± | 2 | 10.5± | 0.3 | 3.7± | 0.7 |
| 125 ppm | 43 | 17.2± | 2.1 | 0.5± | 0.1 | 142± | 1 | 3.3± | 0.4 | 107± | 2** | 10.6± | 0.4 | 3.5± | 0.7 |
| 250 ppm | 41 | 17.5± | 2.0 | 0.5± | 0.0 | 142± | 1 | 3.4± | 0.4 | 109± | 2** | 10.6± | 0.3 | 3.8± | 0.9 |
| 500 ppm | 37 | 18.0± | 2.4 | 0.5± | 0.0 | 140± | 1** | 3.4± | 0.3 | 111± | 4** | 10.5± | 0.4 | 3.9± | 0.6 |

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

Test of Dunnett

(HCL074)

BAIS 4

TABLE H1

URINALYSIS : MALE

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

URINALYSIS

PAGE : 1

| Group Name | NO. of Animals | pH | | | | | | | CHI | Protein | | | | | CHI | Glucose | | | | | CHI | Ketone body | | | | | CHI | Bilirubin | | | | 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 | 42 | 0 | 0 | 6 | 7 | 19 | 8 | 2 | | 0 | 0 | 0 | 3 | 31 | 8 | | 42 | 0 | 0 | 0 | 0 | 0 | 0 | | 41 | 1 | 0 | 0 | 0 | 0 | | 42 | 0 | 0 | 0 |
| 125 ppm | 35 | 0 | 0 | 4 | 10 | 14 | 7 | 0 | | 0 | 0 | 0 | 2 | 27 | 6 | | 35 | 0 | 0 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 |
| 250 ppm | 42 | 0 | 1 | 6 | 12 | 18 | 5 | 0 | | 0 | 0 | 0 | 1 | 25 | 16 | | 42 | 0 | 0 | 0 | 0 | 0 | 0 | | 40 | 1 | 1 | 0 | 0 | 0 | | 42 | 0 | 0 | 0 |
| 500 ppm | 40 | 0 | 1 | 8 | 13 | 14 | 3 | 1 | | 0 | 0 | 0 | 0 | 11 | 29 | ** | 40 | 0 | 0 | 0 | 0 | 0 | 0 | | 39 | 1 | 0 | 0 | 0 | 0 | | 40 | 0 | 0 | 0 |

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

Test of CHI SQUARE

(HCL101)

BAIS 4

STUDY NO. : 0560

URINALYSIS

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | Occult blood | | | | | Urobilinogen | | | | |
|------------|-------------------|--------------|---|---|----|----|--------------|---|----|----|----|
| | | - | ± | + | 2+ | 3+ | ± | + | 2+ | 3+ | 4+ |
| Control | 42 | 42 | 0 | 0 | 0 | 0 | 42 | 0 | 0 | 0 | 0 |
| 125 ppm | 35 | 35 | 0 | 0 | 0 | 0 | 35 | 0 | 0 | 0 | 0 |
| 250 ppm | 42 | 40 | 0 | 1 | 0 | 1 | 42 | 0 | 0 | 0 | 0 |
| 500 ppm | 40 | 39 | 0 | 0 | 0 | 1 | 40 | 0 | 0 | 0 | 0 |

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

Test of CHI SQUARE

(HCL101)

BAIS 4

TABLE H2

URINALYSIS : FEMALE

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

URINALYSIS

PAGE : 3

| Group Name | NO. of Animals | pH | | | | | | | CHI | Protein | | | | | | CHI | Glucose | | | | | | CHI | Ketone body | | | | | | CHI | Bilirubin | | | | 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 | 38 | 0 | 0 | 1 | 3 | 9 | 21 | 4 | | 0 | 2 | 7 | 6 | 16 | 7 | | 38 | 0 | 0 | 0 | 0 | 0 | | 27 | 11 | 0 | 0 | 0 | 0 | | 38 | 0 | 0 | 0 | |
| 125 ppm | 44 | 0 | 2 | 5 | 7 | 13 | 15 | 2 | | 0 | 4 | 6 | 10 | 18 | 6 | | 44 | 0 | 0 | 0 | 0 | 0 | | 29 | 15 | 0 | 0 | 0 | 0 | | 44 | 0 | 0 | 0 | |
| 250 ppm | 41 | 0 | 0 | 6 | 11 | 12 | 9 | 3 | ** | 0 | 2 | 5 | 6 | 21 | 7 | | 41 | 0 | 0 | 0 | 0 | 0 | | 30 | 11 | 0 | 0 | 0 | 0 | | 41 | 0 | 0 | 0 | |
| 500 ppm | 37 | 0 | 0 | 5 | 7 | 11 | 13 | 1 | | 0 | 0 | 0 | 0 | 18 | 19 | ** | 37 | 0 | 0 | 0 | 0 | 0 | | 21 | 16 | 0 | 0 | 0 | 0 | | 37 | 0 | 0 | 0 | |

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

Test of CHI SQUARE

(HCL101)

BAIS 4

STUDY NO. : 0560

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

URINALYSIS

PAGE : 4

| Group Name | NO. of Animals | Occult blood | | | | | Urobilinogen | | | | |
|------------|-------------------|--------------|---|---|----|----|--------------|---|----|----|----|
| | | - | ± | + | 2+ | 3+ | ± | + | 2+ | 3+ | 4+ |
| Control | 38 | 38 | 0 | 0 | 0 | 0 | 38 | 0 | 0 | 0 | 0 |
| 125 ppm | 44 | 41 | 2 | 0 | 1 | 0 | 44 | 0 | 0 | 0 | 0 |
| 250 ppm | 41 | 40 | 0 | 0 | 0 | 1 | 41 | 0 | 0 | 0 | 0 |
| 500 ppm | 37 | 36 | 0 | 0 | 1 | 0 | 37 | 0 | 0 | 0 | 0 |

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

Test of CHI SQUARE

(HCL101)

BAIS 4

TABLE I 1

GROSS FINDINGS : MALE

ALL ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| skin/app | nodule | | 1 | (2) | 4 | (8) | 0 | (0) | 0 | (0) |
| subcutis | edema | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | mass | | 9 | (18) | 5 | (10) | 12 | (24) | 11 | (22) |
| lung | white zone | | 2 | (4) | 3 | (6) | 2 | (4) | 1 | (2) |
| | red zone | | 1 | (2) | 1 | (2) | 0 | (0) | 0 | (0) |
| | edema | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | nodule | | 1 | (2) | 1 | (2) | 3 | (6) | 5 | (10) |
| lymph node | enlarged | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| spleen | enlarged | | 4 | (8) | 4 | (8) | 2 | (4) | 1 | (2) |
| | black zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 1 | (2) |
| heart | white zone | | 1 | (2) | 0 | (0) | 1 | (2) | 2 | (4) |
| salivary gl | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| forestomach | nodule | | 0 | (0) | 3 | (6) | 0 | (0) | 1 | (2) |
| | ulcer | | 2 | (4) | 1 | (2) | 0 | (0) | 2 | (4) |
| gl stomach | red zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | erosion | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | thick | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| small intes | thick | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| liver | enlarged | | 1 | (2) | 0 | (0) | 0 | (0) | 1 | (2) |
| | pale | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | white zone | | 1 | (2) | 2 | (4) | 0 | (0) | 1 | (2) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| liver | red zone | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 1 | (2) | 3 | (6) | 5 | (10) |
| | rough | | 1 | (2) | 0 | (0) | 2 | (4) | 0 | (0) |
| | herniation | | 6 | (12) | 9 | (18) | 6 | (12) | 3 | (6) |
| pancreas | nodule | | 1 | (2) | 1 | (2) | 1 | (2) | 0 | (0) |
| kidney | white zone | | 0 | (0) | 2 | (4) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 2 | (4) | 0 | (0) | 1 | (2) |
| | cyst | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | granular | | 10 | (20) | 12 | (24) | 15 | (30) | 9 | (18) |
| urin bladd | urine:marked retention | | 0 | (0) | 1 | (2) | 0 | (0) | 1 | (2) |
| | urine:red | | 1 | (2) | 1 | (2) | 0 | (0) | 0 | (0) |
| pituitary | enlarged | | 5 | (10) | 4 | (8) | 2 | (4) | 4 | (8) |
| | red zone | | 2 | (4) | 3 | (6) | 3 | (6) | 1 | (2) |
| | nodule | | 6 | (12) | 2 | (4) | 4 | (8) | 1 | (2) |
| thyroid | enlarged | | 7 | (14) | 4 | (8) | 5 | (10) | 2 | (4) |
| | nodule | | 0 | (0) | 2 | (4) | 3 | (6) | 1 | (2) |
| adrenal | enlarged | | 0 | (0) | 3 | (6) | 3 | (6) | 0 | (0) |
| testis | atrophic | | 1 | (2) | 0 | (0) | 1 | (2) | 0 | (0) |
| | nodule | | 36 | (72) | 40 | (80) | 41 | (82) | 40 | (80) |
| prostate | yellow zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| prep/cli gl | nodule | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| brain | red zone | | 0 | (0) | 0 | (0) | 2 | (4) | 1 | (2) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------------|------------------------------|---------|------|---------|------|---------|------|---------|-------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| periph nerv | brown zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| eye | white | | 4 | (8) | 2 | (4) | 3 | (6) | 5 | (10) |
| Harder gl | enlarged | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| Zymbal gl | nodule | | 0 | (0) | 1 | (2) | 1 | (2) | 2 | (4) |
| muscle | nodule | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| bone | nodule | | 0 | (0) | 2 | (4) | 0 | (0) | 0 | (0) |
| peritoneum | nodule | | 0 | (0) | 1 | (2) | 1 | (2) | 2 | (4) |
| retroperit | mass | | 0 | (0) | 0 | (0) | 1 | (2) | 1 | (2) |
| abdominal c | hemorrhage | | 0 | (0) | 1 | (2) | 0 | (0) | 1 | (2) |
| | ascites | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| thoracic ca | hemorrhage | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | mass | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | pleural fluid | | 1 | (2) | 2 | (4) | 1 | (2) | 0 | (0) |
| other | ear:nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 1 | (2) |
| | upper jaw:nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| whole body | anemic | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |

TABLE I 2

GROSS FINDINGS : MALE
DEAD AND MORIBUND ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 11 | (%) | 15 | (%) | 9 | (%) | 13 | (%) |
| skin/app | nodule | | 0 | (0) | 2 | (13) | 0 | (0) | 0 | (0) |
| subcutis | edema | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | mass | | 1 | (9) | 1 | (7) | 2 | (22) | 2 | (15) |
| lung | white zone | | 0 | (0) | 1 | (7) | 1 | (11) | 0 | (0) |
| | red zone | | 1 | (9) | 1 | (7) | 0 | (0) | 0 | (0) |
| | edema | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 1 | (7) | 1 | (11) | 0 | (0) |
| lymph node | enlarged | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| spleen | enlarged | | 2 | (18) | 3 | (20) | 1 | (11) | 1 | (8) |
| | black zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (8) |
| | nodule | | 1 | (9) | 0 | (0) | 0 | (0) | 1 | (8) |
| heart | white zone | | 1 | (9) | 0 | (0) | 0 | (0) | 1 | (8) |
| forestomach | nodule | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | ulcer | | 2 | (18) | 1 | (7) | 0 | (0) | 2 | (15) |
| gl stomach | erosion | | 1 | (9) | 0 | (0) | 0 | (0) | 0 | (0) |
| | thick | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (8) |
| small intes | thick | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| liver | enlarged | | 1 | (9) | 0 | (0) | 0 | (0) | 1 | (8) |
| | pale | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | white zone | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | red zone | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 2 | (15) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 11 | (%) | 15 | (%) | 9 | (%) | 13 | (%) |
| liver | herniation | | 1 | (9) | 3 | (20) | 1 | (11) | 1 | (8) |
| pancreas | nodule | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| kidney | nodule | | 0 | (0) | 2 | (13) | 0 | (0) | 0 | (0) |
| | granular | | 5 | (45) | 2 | (13) | 1 | (11) | 2 | (15) |
| urin bladd | urine:marked retention | | 0 | (0) | 1 | (7) | 0 | (0) | 1 | (8) |
| | urine:red | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| pituitary | enlarged | | 5 | (45) | 3 | (20) | 2 | (22) | 2 | (15) |
| | red zone | | 0 | (0) | 2 | (13) | 0 | (0) | 1 | (8) |
| | nodule | | 0 | (0) | 2 | (13) | 0 | (0) | 1 | (8) |
| thyroid | enlarged | | 4 | (36) | 1 | (7) | 2 | (22) | 1 | (8) |
| adrenal | enlarged | | 0 | (0) | 0 | (0) | 1 | (11) | 0 | (0) |
| testis | atrophic | | 0 | (0) | 0 | (0) | 1 | (11) | 0 | (0) |
| | nodule | | 4 | (36) | 8 | (53) | 5 | (56) | 7 | (54) |
| prostate | yellow zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (8) |
| prep/cli gl | nodule | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |
| brain | red zone | | 0 | (0) | 0 | (0) | 2 | (22) | 1 | (8) |
| periph nerv | brown zone | | 0 | (0) | 0 | (0) | 1 | (11) | 0 | (0) |
| | nodule | | 1 | (9) | 0 | (0) | 0 | (0) | 0 | (0) |
| eye | white | | 1 | (9) | 0 | (0) | 1 | (11) | 1 | (8) |
| Harder gl | enlarged | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (8) |
| Zymbal gl | nodule | | 0 | (0) | 1 | (7) | 1 | (11) | 2 | (15) |
| muscle | nodule | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|---------------|------------------------------|---------|------|---------|-------|---------|-------|---------|------|
| | | | 11 | (%) | 15 | (%) | 9 | (%) | 13 | (%) |
| bone | nodule | | 0 | (0) | 2 | (13) | 0 | (0) | 0 | (0) |
| peritoneum | nodule | | 0 | (0) | 1 | (7) | 0 | (0) | 1 | (8) |
| retroperit | mass | | 0 | (0) | 0 | (0) | 1 | (11) | 0 | (0) |
| abdominal c | hemorrhage | | 0 | (0) | 1 | (7) | 0 | (0) | 1 | (8) |
| thoracic ca | hemorrhage | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (8) |
| | pleural fluid | | 1 | (9) | 2 | (13) | 0 | (0) | 0 | (0) |
| whole body | anemic | | 0 | (0) | 1 | (7) | 0 | (0) | 0 | (0) |

(HPT080)

BAIS 4

TABLE I 3

GROSS FINDINGS : MALE
SACRIFICED ANIMALS

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 39 | (%) | 35 | (%) | 41 | (%) | 37 | (%) |
| skin/app | nodule | | 1 | (3) | 2 | (6) | 0 | (0) | 0 | (0) |
| subcutis | mass | | 8 | (21) | 4 | (11) | 10 | (24) | 9 | (24) |
| lung | white zone | | 2 | (5) | 2 | (6) | 1 | (2) | 1 | (3) |
| | nodule | | 1 | (3) | 0 | (0) | 2 | (5) | 5 | (14) |
| spleen | enlarged | | 2 | (5) | 1 | (3) | 1 | (2) | 0 | (0) |
| heart | white zone | | 0 | (0) | 0 | (0) | 1 | (2) | 1 | (3) |
| salivary gl | nodule | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| forestomach | nodule | | 0 | (0) | 2 | (6) | 0 | (0) | 1 | (3) |
| gl stomach | red zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| liver | white zone | | 1 | (3) | 1 | (3) | 0 | (0) | 1 | (3) |
| | nodule | | 0 | (0) | 1 | (3) | 3 | (7) | 3 | (8) |
| | rough | | 1 | (3) | 0 | (0) | 2 | (5) | 0 | (0) |
| | herniation | | 5 | (13) | 6 | (17) | 5 | (12) | 2 | (5) |
| pancreas | nodule | | 1 | (3) | 0 | (0) | 1 | (2) | 0 | (0) |
| kidney | white zone | | 0 | (0) | 2 | (6) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| | cyst | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| | granular | | 5 | (13) | 10 | (29) | 14 | (34) | 7 | (19) |
| urin bladd | urine:red | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| pituitary | enlarged | | 0 | (0) | 1 | (3) | 0 | (0) | 2 | (5) |
| | red zone | | 2 | (5) | 1 | (3) | 3 | (7) | 0 | (0) |
| | nodule | | 6 | (15) | 0 | (0) | 4 | (10) | 0 | (0) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

GROSS FINDINGS (SUMMARY)
 SACRIFICED ANIMALS (105W)

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 39 | (%) | 35 | (%) | 41 | (%) | 37 | (%) |
| thyroid | enlarged | | 3 | (8) | 3 | (9) | 3 | (7) | 1 | (3) |
| | nodule | | 0 | (0) | 2 | (6) | 3 | (7) | 1 | (3) |
| adrenal | enlarged | | 0 | (0) | 3 | (9) | 2 | (5) | 0 | (0) |
| testis | atrophic | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| | nodule | | 32 | (82) | 32 | (91) | 36 | (88) | 33 | (89) |
| eye | white | | 3 | (8) | 2 | (6) | 2 | (5) | 4 | (11) |
| peritoneum | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 1 | (3) |
| retroperit | mass | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| abdominal c | ascites | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| thoracic ca | mass | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| | pleural fluid | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| other | ear:nodule | | 1 | (3) | 0 | (0) | 0 | (0) | 1 | (3) |
| | upper jaw:nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |

TABLE I 4

GROSS FINDINGS : FEMALE

ALL ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|--------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| skin/app | nodule | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | erosion | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| subcutis | jaundice | | 0 | (0) | 1 | (2) | 0 | (0) | 1 | (2) |
| | mass | | 10 | (20) | 11 | (22) | 12 | (24) | 6 | (12) |
| lung | white zone | | 0 | (0) | 1 | (2) | 1 | (2) | 1 | (2) |
| | red zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 1 | (2) |
| lymph node | enlarged | | 1 | (2) | 0 | (0) | 1 | (2) | 0 | (0) |
| spleen | enlarged | | 3 | (6) | 3 | (6) | 5 | (10) | 2 | (4) |
| | atrophic | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | ulcer | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| heart | white zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| tongue | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| esophagus | invagination | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| forestomach | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | thick | | 1 | (2) | 1 | (2) | 0 | (0) | 1 | (2) |
| gl stomach | black zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | ulcer | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | erosion | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| large intes | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| liver | white zone | | 2 | (4) | 0 | (0) | 1 | (2) | 1 | (2) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 5

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|------------|------------------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| liver | red zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | brown zone | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 2 | (4) | 0 | (0) | 2 | (4) |
| | rough | | 0 | (0) | 2 | (4) | 2 | (4) | 0 | (0) |
| | nodular | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | herniation | | 15 | (30) | 8 | (16) | 9 | (18) | 8 | (16) |
| pancreas | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| kidney | granular | | 0 | (0) | 0 | (0) | 0 | (0) | 4 | (8) |
| | hydronephrosis | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| urin bladd | red zone | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | urine marked retention | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| pituitary | enlarged | | 5 | (10) | 4 | (8) | 2 | (4) | 3 | (6) |
| | red zone | | 10 | (20) | 6 | (12) | 7 | (14) | 3 | (6) |
| | black zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | nodule | | 3 | (6) | 8 | (16) | 8 | (16) | 3 | (6) |
| thyroid | enlarged | | 4 | (8) | 0 | (0) | 0 | (0) | 1 | (2) |
| | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 1 | (2) |
| adrenal | enlarged | | 1 | (2) | 0 | (0) | 0 | (0) | 1 | (2) |
| ovary | enlarged | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | cyst | | 1 | (2) | 1 | (2) | 3 | (6) | 1 | (2) |
| uterus | nodule | | 9 | (18) | 3 | (6) | 4 | (8) | 4 | (8) |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 6

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|---------------|------------------------------|---------|------|---------|------|---------|------|---------|------|
| | | | 50 | (%) | 50 | (%) | 50 | (%) | 50 | (%) |
| uterus | invagination | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | fluid:black | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| prep/cli gl | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| brain | red zone | | 2 | (4) | 0 | (0) | 0 | (0) | 0 | (0) |
| eye | turbid | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| | white | | 4 | (8) | 3 | (6) | 4 | (8) | 4 | (8) |
| Zymbal gl | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 2 | (4) |
| muscle | nodule | | 0 | (0) | 0 | (0) | 2 | (4) | 0 | (0) |
| bone | nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| retroperit | mass | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (2) |
| abdominal c | hemorrhage | | 1 | (2) | 0 | (0) | 2 | (4) | 0 | (0) |
| | mass | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |
| | ascites | | 0 | (0) | 0 | (0) | 0 | (0) | 2 | (4) |
| thoracic ca | pleural fluid | | 0 | (0) | 1 | (2) | 1 | (2) | 0 | (0) |
| other | ear:nodule | | 1 | (2) | 0 | (0) | 0 | (0) | 0 | (0) |

TABLE I 5

GROSS FINDINGS : FEMALE
DEAD AND MORIBUND ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 4

| Organ | Findings | Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|-------------|------------------------|----------------|---------|---------|---------|---------|
| | | NO. of Animals | 12 (%) | 7 (%) | 9 (%) | 13 (%) |
| skin/app | erosion | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| subcutis | jaundice | | 0 (0) | 1 (14) | 0 (0) | 1 (8) |
| | mass | | 1 (8) | 1 (14) | 3 (33) | 3 (23) |
| lung | red zone | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| lymph node | enlarged | | 1 (8) | 0 (0) | 1 (11) | 0 (0) |
| spleen | enlarged | | 3 (25) | 1 (14) | 3 (33) | 2 (15) |
| | atrophic | | 0 (0) | 1 (14) | 0 (0) | 0 (0) |
| | nodule | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | ulcer | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| esophagus | invagination | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| forestomach | thick | | 1 (8) | 0 (0) | 0 (0) | 1 (8) |
| gl stomach | black zone | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| | ulcer | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | erosion | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| large intes | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| liver | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| | rough | | 0 (0) | 2 (29) | 1 (11) | 0 (0) |
| | herniation | | 3 (25) | 0 (0) | 2 (22) | 2 (15) |
| kidney | granular | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| | hydronephrosis | | 0 (0) | 1 (14) | 0 (0) | 0 (0) |
| urin bladd | red zone | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| | urine:marked retention | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 5

| Organ | Findings | Group Name NO. of Animals | Control | 125 ppm | 250 ppm | 500 ppm |
|-------------|---------------|------------------------------|---------|---------|---------|---------|
| | | | 12 (%) | 7 (%) | 9 (%) | 13 (%) |
| pituitary | enlarged | | 4 (33) | 1 (14) | 2 (22) | 2 (15) |
| | red zone | | 1 (8) | 1 (14) | 0 (0) | 0 (0) |
| | black zone | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| | nodule | | 0 (0) | 1 (14) | 1 (11) | 0 (0) |
| thyroid | enlarged | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | nodule | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| adrenal | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| ovary | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| uterus | nodule | | 3 (25) | 1 (14) | 1 (11) | 3 (23) |
| | invagination | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| | fluid:black | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| prep/cli gl | nodule | | 0 (0) | 0 (0) | 1 (11) | 0 (0) |
| brain | red zone | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| eye | turbid | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| | white | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| Zymbal gl | nodule | | 0 (0) | 0 (0) | 1 (11) | 2 (15) |
| bone | nodule | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| retroperit | mass | | 0 (0) | 0 (0) | 0 (0) | 1 (8) |
| abdominal c | hemorrhage | | 1 (8) | 0 (0) | 2 (22) | 0 (0) |
| | mass | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | ascites | | 0 (0) | 0 (0) | 0 (0) | 2 (15) |
| thoracic ca | pleural fluid | | 0 (0) | 1 (14) | 1 (11) | 0 (0) |

TABLE I 6

GROSS FINDINGS : FEMALE
SACRIFICED ANIMALS

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|-------------|------------|------------------------------|---------|-------|---------|-------|---------|-------|---------|-------|
| | | | 38 | (%) | 43 | (%) | 41 | (%) | 37 | (%) |
| skin/app | nodule | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| subcutis | mass | | 9 | (24) | 10 | (23) | 9 | (22) | 3 | (8) |
| lung | white zone | | 0 | (0) | 1 | (2) | 1 | (2) | 1 | (3) |
| | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 1 | (3) |
| spleen | enlarged | | 0 | (0) | 2 | (5) | 2 | (5) | 0 | (0) |
| heart | white zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| tongue | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| forestomach | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| | thick | | 0 | (0) | 1 | (2) | 0 | (0) | 0 | (0) |
| liver | white zone | | 2 | (5) | 0 | (0) | 1 | (2) | 0 | (0) |
| | red zone | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | brown zone | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| | nodule | | 0 | (0) | 2 | (5) | 0 | (0) | 2 | (5) |
| | rough | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | nodular | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| | herniation | | 12 | (32) | 8 | (19) | 7 | (17) | 6 | (16) |
| pancreas | nodule | | 0 | (0) | 0 | (0) | 1 | (2) | 0 | (0) |
| kidney | granular | | 0 | (0) | 0 | (0) | 0 | (0) | 3 | (8) |
| pituitary | enlarged | | 1 | (3) | 3 | (7) | 0 | (0) | 1 | (3) |
| | red zone | | 9 | (24) | 5 | (12) | 7 | (17) | 3 | (8) |
| | nodule | | 3 | (8) | 7 | (16) | 7 | (17) | 3 | (8) |
| thyroid | enlarged | | 3 | (8) | 0 | (0) | 0 | (0) | 1 | (3) |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control | | 125 ppm | | 250 ppm | | 500 ppm | |
|---------|------------|------------------------------|---------|-------|---------|------|---------|-------|---------|-------|
| | | | 38 | (%) | 43 | (%) | 41 | (%) | 37 | (%) |
| thyroid | nodule | | 0 | (0) | 0 | (0) | 0 | (0) | 1 | (3) |
| adrenal | enlarged | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| ovary | enlarged | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| | cyst | | 1 | (3) | 1 | (2) | 3 | (7) | 1 | (3) |
| uterus | nodule | | 6 | (16) | 2 | (5) | 3 | (7) | 1 | (3) |
| brain | red zone | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |
| eye | white | | 3 | (8) | 3 | (7) | 4 | (10) | 4 | (11) |
| muscle | nodule | | 0 | (0) | 0 | (0) | 2 | (5) | 0 | (0) |
| other | ear:nodule | | 1 | (3) | 0 | (0) | 0 | (0) | 0 | (0) |

(HPT080)

BAIS 4

TABLE J1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE
UNIT: g

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

PAGE : 1

| Group Name | NO. of Animals | Body Weight | ADRENALS | TESTES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|-------------|--------------|---------------|---------------|--------------|--------------|
| Control | 39 | 394± 34 | 0.071± 0.013 | 3.177± 1.347 | 1.217± 0.125 | 1.390± 0.227 | 2.761± 0.447 |
| 125 ppm | 35 | 390± 29 | 0.106± 0.174 | 3.681± 1.275 | 1.207± 0.092 | 1.405± 0.359 | 2.880± 0.337 |
| 250 ppm | 41 | 380± 26 | 0.098± 0.151 | 3.297± 1.276 | 1.191± 0.088 | 1.381± 0.141 | 2.843± 0.236 |
| 500 ppm | 37 | 326± 28** | 0.067± 0.018 | 4.008± 1.605* | 1.165± 0.141* | 1.321± 0.095 | 2.752± 0.264 |

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

Test of Dunnett

(HCL040)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE
UNIT: g

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

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRATN | |
|------------|-------------------|--------|---------|---------|---------|--------|---------|
| Control | 39 | 1.337± | 1.360 | 11.551± | 1.295 | 2.032± | 0.044 |
| 125 ppm | 35 | 0.993± | 0.400* | 12.328± | 1.684 | 2.043± | 0.044 |
| 250 ppm | 41 | 1.240± | 1.647* | 12.787± | 1.806** | 2.027± | 0.043 |
| 500 ppm | 37 | 0.916± | 0.178** | 12.564± | 2.184* | 1.978± | 0.047** |

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

Test of Dunnett

(HCL040)

BAIS 4

TABLE J2

ORGAN WEIGHT, ABSOLUTE : FEMALE

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
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 | 38 | 264± 22 | 0.090± 0.135 | 0.240± 0.725 | 0.859± 0.061 | 0.928± 0.065 | 1.720± 0.131 |
| 125 ppm | 43 | 260± 24 | 0.068± 0.008 | 0.115± 0.027 | 0.857± 0.073 | 0.938± 0.157 | 1.713± 0.127 |
| 250 ppm | 41 | 255± 27 | 0.068± 0.013 | 0.139± 0.126 | 0.882± 0.086 | 0.990± 0.248 | 1.709± 0.128 |
| 500 ppm | 37 | 237± 22** | 0.065± 0.005 | 0.118± 0.020 | 0.875± 0.060 | 0.947± 0.049 | 1.758± 0.104 |

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

Test of Dunnett

(HCL040)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

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

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRAIN | |
|------------|-------------------|--------|-------|--------|--------|--------|---------|
| Control | 38 | 0.624± | 0.289 | 6.690± | 1.131 | 1.862± | 0.040 |
| 125 ppm | 43 | 0.617± | 0.401 | 6.759± | 1.061 | 1.836± | 0.036** |
| 250 ppm | 41 | 0.974± | 1.775 | 6.904± | 1.462 | 1.829± | 0.044** |
| 500 ppm | 37 | 0.526± | 0.088 | 7.039± | 0.689* | 1.787± | 0.029** |

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

Test of Dunnett

(HCL040)

BAIS 4

TABLE K1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE
 UNIT: %

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

PAGE : 1

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | TESTES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|--------------------|----------------|----------------|----------------|----------------|----------------|
| Control | 39 | 394± 34 | 0.018± 0.004 | 0.805± 0.334 | 0.311± 0.042 | 0.357± 0.081 | 0.709± 0.159 |
| 125 ppm | 35 | 390± 29 | 0.027± 0.047 | 0.941± 0.309 | 0.310± 0.027 | 0.361± 0.091 | 0.740± 0.088* |
| 250 ppm | 41 | 380± 26 | 0.026± 0.043 | 0.868± 0.327 | 0.314± 0.027 | 0.364± 0.042** | 0.749± 0.061** |
| 500 ppm | 37 | 326± 28** | 0.021± 0.005** | 1.237± 0.496** | 0.360± 0.051** | 0.408± 0.038** | 0.848± 0.093** |

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE
UNIT: %

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

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|---------------|----------------|----------------|
| Control | 39 | 0.349± 0.410 | 2.954± 0.467 | 0.519± 0.044 |
| 125 ppm | 35 | 0.254± 0.095* | 3.161± 0.371 | 0.527± 0.041 |
| 250 ppm | 41 | 0.330± 0.456 | 3.368± 0.459** | 0.535± 0.035 |
| 500 ppm | 37 | 0.283± 0.064 | 3.850± 0.517** | 0.611± 0.048** |

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

Test of Dunnett

(HCL042)

BAIS 4

TABLE K2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

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

PAGE : 3

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | OVARIES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|--------------------|--------------|--------------|----------------|----------------|----------------|
| Control | 38 | 264± 22 | 0.034± 0.049 | 0.094± 0.293 | 0.326± 0.028 | 0.353± 0.035 | 0.653± 0.053 |
| 125 ppm | 43 | 260± 24 | 0.026± 0.003 | 0.045± 0.012 | 0.330± 0.030 | 0.363± 0.069 | 0.661± 0.046 |
| 250 ppm | 41 | 255± 27 | 0.027± 0.006 | 0.056± 0.054 | 0.349± 0.052* | 0.392± 0.113** | 0.675± 0.064 |
| 500 ppm | 37 | 237± 22** | 0.028± 0.004 | 0.050± 0.009 | 0.373± 0.044** | 0.403± 0.039** | 0.749± 0.091** |

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

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

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|--------------|----------------|----------------|
| Control | 38 | 0.238± 0.116 | 2.532± 0.366 | 0.709± 0.056 |
| 125 ppm | 43 | 0.240± 0.163 | 2.599± 0.375 | 0.711± 0.064 |
| 250 ppm | 41 | 0.392± 0.724 | 2.719± 0.586* | 0.724± 0.073 |
| 500 ppm | 37 | 0.224± 0.041 | 2.995± 0.378** | 0.762± 0.078** |

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

Test of Dunnett

(HCL042)

BAIS 4

TABLE L1

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE
ALL ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|----------------------------------|--|--|---------------|--------|-------|-------|---------------|--------|-------|-------|---------------|--------|-------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Integumentary system/appandage} | | | | | | | | | | | | | | | | | | |
| skin/app | scab | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 1 | 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) |
| subcutis | inflammation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 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) | (2) | (0) | (0) | (0) |
| | epidermal cyst | | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | hyperplasia:gland | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 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) | (2) | (0) | (0) | (0) |
| | eosinophilic change:olfactory epithelium | | 34 | 14 | 0 | 0 | 33 | 10 | 0 | 0 | 16 | 25 | 4 | 0 ** | 10 | 34 | 6 | 0 ** |
| | | | (68) | (28) | (0) | (0) | (66) | (20) | (0) | (0) | (32) | (50) | (8) | (0) | (20) | (68) | (12) | (0) |
| | eosinophilic change:respiratory epithelium | | 26 | 0 | 0 | 0 | 28 | 0 | 0 | 0 | 31 | 0 | 0 | 0 | 44 | 0 | 0 | 0 ** |
| | | | (52) | (0) | (0) | (0) | (56) | (0) | (0) | (0) | (62) | (0) | (0) | (0) | (88) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|----------------------|---|--|---------------|-------|------|------|---------------|-------|------|------|---------------|-------|------|------|---------------|-------|------|------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | inflammation:foreign body | | 8 | 7 | 0 | 0 | 13 | 6 | 0 | 0 | 9 | 8 | 0 | 0 | 16 | 10 | 0 | 0 |
| | | | (16) | (14) | (0) | (0) | (26) | (12) | (0) | (0) | (18) | (16) | (0) | (0) | (32) | (20) | (0) | (0) |
| | inflammation:respiratory epithelium | | 0 | 1 | 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) |
| | inflammation:olfactory epithelium | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| lung | respiratory metaplasia:olfactory epithelium | | 4 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 8 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| | respiratory metaplasia:gland | | 10 | 0 | 0 | 0 | 12 | 0 | 0 | 0 | 8 | 1 | 0 | 0 | 15 | 0 | 0 | 0 |
| | | | (20) | (0) | (0) | (0) | (24) | (0) | (0) | (0) | (16) | (2) | (0) | (0) | (30) | (0) | (0) | (0) |
| | squamous cell metaplasia:respiratory epithelium | | 3 | 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) | (0) | (0) | (0) | (0) |
| | atrophy:olfactory epithelium | | 0 | 0 | 0 | 0 | 0 | 0 | 3 | 0 | 1 | 1 | 0 | 0 | 2 | 2 | 1 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (6) | (0) | (2) | (2) | (0) | (0) | (4) | (4) | (2) | (0) |
| lung | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | congestion | | 0 | 1 | 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) |

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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 3

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|------------------------|---------------------------------------|--|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| lung | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | edema | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | inflammation | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) |
| | inflammatory infiltration | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| | accumulation of foamy cells | | 0 | 0 | 0 | 0 | 1 | 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) |
| | bronchiolar-alveolar cell hyperplasia | | 0 | 0 | 0 | 0 | 4 | 2 | 0 | 0 * | 4 | 0 | 0 | 0 | 2 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (8) | (4) | (0) | (0) | (8) | (0) | (0) | (0) | (4) | (2) | (0) | (0) |
| | inflammation:foreign body | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| bone marrow | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | hemorrhage | | 0 | 0 | 0 | 0 | 1 | 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) |

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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 4

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 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

increased hematopoiesis

50

50

50

50

5

0

0

0

(10)

(0)

(0)

(0)

3

2

0

0

(6)

(4)

(0)

(0)

7

0

0

0

(14)

(0)

(0)

(0)

5

0

0

0

(10)

(0)

(0)

(0)

granulopoiesis:increased

50

50

50

50

1

0

0

0

(2)

(0)

(0)

(0)

0

0

0

0

(0)

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(0)

(0)

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(0)

0

0

0

0

(0)

(0)

(0)

(0)

xanthogranuloma

50

50

50

50

0

0

0

0

(0)

(0)

(0)

(0)

0

0

1

0

(0)

(0)

(2)

(0)

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(0)

(0)

(0)

(0)

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0

(0)

(0)

(0)

(0)

lymph node

lymphadenitis

50

50

50

50

0

1

0

0

(0)

(2)

(0)

(0)

0

0

0

0

(0)

(0)

(0)

(0)

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0

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0

(0)

(0)

(0)

(0)

0

0

0

0

(0)

(0)

(0)

(0)

spleen

congestion

50

50

50

50

2

3

0

0

(4)

(6)

(0)

(0)

0

2

0

0

(0)

(4)

(0)

(0)

0

1

0

0

(0)

(2)

(0)

(0)

1

0

0

0

(2)

(0)

(0)

(0)

hemorrhage

50

50

50

50

0

1

0

0

(0)

(2)

(0)

(0)

0

0

0

0

(0)

(0)

(0)

(0)

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0

0

0

(0)

(0)

(0)

(0)

0

0

0

0

(0)

(0)

(0)

(0)

deposit of hemosiderin

50

50

50

50

4

0

0

0

(8)

(0)

(0)

(0)

3

3

0

0

(6)

(6)

(0)

(0)

3

0

1

0

(6)

(0)

(2)

(0)

4

3

0

0

(8)

(6)

(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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 5

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------------|--|--|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| spleen | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | fibrosis:focal | | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | increased extramedullary hematopoiesis | | 7 | 3 | 0 | 0 | 4 | 2 | 2 | 0 | 1 | 4 | 0 | 0 | 2 | 3 | 1 | 0 |
| | | | (14) | (6) | (0) | (0) | (8) | (4) | (4) | (0) | (2) | (8) | (0) | (0) | (4) | (6) | (2) | (0) |
| {Circulatory system} | | | | | | | | | | | | | | | | | | |
| heart | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | myocardial fibrosis | | 29 | 1 | 0 | 0 | 23 | 1 | 0 | 0 | 20 | 1 | 0 | 0 | 20 | 2 | 0 | 0 |
| | | | (58) | (2) | (0) | (0) | (46) | (2) | (0) | (0) | (40) | (2) | (0) | (0) | (40) | (4) | (0) | (0) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| oral cavity | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | squamous cell hyperplasia | | 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) | (2) | (0) | (0) | (0) |
| esophagus | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | inflammation | | 0 | 1 | 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) |

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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 6

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|--------------------|----------------------------------|--|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| stomach | | | | | | | | | | | | | | | | | | |
| | ulcer:forestomach | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 2 | 0 | 2 | 1 | 2 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 4 | 1 |
| | | | (4) | (0) | (4) | (2) | (4) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (2) | (8) | (2) |
| | hyperplasia:forestomach | | 2 | 1 | 0 | 0 | 3 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 3 | 2 | 0 | 0 |
| | | | (4) | (2) | (0) | (0) | (6) | (2) | (0) | (0) | (0) | (0) | (2) | (0) | (6) | (4) | (0) | (0) |
| | erosion:glandular stomach | | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 2 | 1 | 0 | 0 | 7 | 0 | 0 | 0 |
| | | | (6) | (0) | (0) | (0) | (8) | (0) | (0) | (0) | (4) | (2) | (0) | (0) | (14) | (0) | (0) | (0) |
| | ulcer:glandular stomach | | 0 | 3 | 0 | 0 | 1 | 0 | 1 | 0 | 2 | 0 | 0 | 0 | 3 | 1 | 0 | 0 |
| | | | (0) | (6) | (0) | (0) | (2) | (0) | (2) | (0) | (4) | (0) | (0) | (0) | (6) | (2) | (0) | (0) |
| | hyperplasia:glandular stomach | | 3 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 4 | 0 | 0 | 0 |
| | | | (6) | (0) | (0) | (0) | (10) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| | mineralization:glandular stomach | | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | dilated glands | | 1 | 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) |
| small intes | | | | | | | | | | | | | | | | | | |
| | erosion | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (8) | (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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 7

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 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} | | | | | | | | | | | | | | | | | | |
| liver | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | herniation | 6 (12) | 0 (0) | 0 (0) | 0 (0) | 9 (18) | 0 (0) | 0 (0) | 0 (0) | 6 (12) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | hemorrhage | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | necrosis:central | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) |
| | necrosis:focal | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change | 0 (0) | 2 (4) | 1 (2) | 0 (0) | 1 (2) | 1 (2) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change:central | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change:peripheral | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | degeneration:central | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 8

| Organ | Findings | Group Name No. of Animals on Study | | | | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|------------------------|---------------------------------------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|
| | | Grade | | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | | | | |
| liver | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | granulation | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (6) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | clear cell focus | 7 | 1 | 0 | 0 | 11 | 3 | 0 | 0 | 8 | 0 | 0 | 0 | 6 | 0 | 0 | 0 | 6 | 0 | 0 | 0 |
| | | (14) | (2) | (0) | (0) | (22) | (6) | (0) | (0) | (16) | (0) | (0) | (0) | (12) | (0) | (0) | (0) | (12) | (0) | (0) | (0) |
| | acidophilic cell focus | 1 | 1 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (2) | (2) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | basophilic cell focus | 5 | 0 | 0 | 0 | 3 | 1 | 0 | 0 | 5 | 3 | 0 | 0 | 5 | 2 | 0 | 0 | 5 | 2 | 0 | 0 |
| | | (10) | (0) | (0) | (0) | (6) | (2) | (0) | (0) | (10) | (6) | (0) | (0) | (10) | (4) | (0) | (0) | (10) | (4) | (0) | (0) |
| | spongiosis hepatitis | 3 | 1 | 0 | 0 | 4 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (6) | (2) | (0) | (0) | (8) | (0) | (0) | (0) | (8) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | bile duct hyperplasia | 7 | 42 | 0 | 0 | 9 | 40 | 0 | 0 | 2 | 47 | 0 | 0 | 5 | 45 | 0 | 0 | 5 | 45 | 0 | 0 |
| | | (14) | (84) | (0) | (0) | (18) | (80) | (0) | (0) | (4) | (94) | (0) | (0) | (10) | (90) | (0) | (0) | (10) | (90) | (0) | (0) |
| pancreas | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | atrophy | 9 | 8 | 1 | 0 | 5 | 4 | 0 | 0 | 9 | 2 | 2 | 0 | 5 | 5 | 0 | 0 | 5 | 5 | 0 | 0 |
| | | (18) | (16) | (2) | (0) | (10) | (8) | (0) | (0) | (18) | (4) | (4) | (0) | (10) | (10) | (0) | (0) | (10) | (10) | (0) | (0) |
| | islet cell hyperplasia | 1 | 2 | 0 | 0 | 2 | 1 | 2 | 0 | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (2) | (4) | (0) | (0) | (4) | (2) | (4) | (0) | (2) | (2) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 9

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------|------------------------|--|---------|--------|--------|--------|---------|--------|--------|-------|---------|--------|--------|-------|---------|--------|--------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | ectopic tissue | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | cyst | | 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) | (2) | (0) | (0) | (0) |
| | scar | | 0 | 0 | 0 | 0 | 1 | 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) |
| | chronic nephropathy | | 11 | 22 | 11 | 5 | 16 | 14 | 17 | 0 * | 8 | 18 | 22 | 0 * | 8 | 25 | 16 | 1 |
| | | | (22) | (44) | (22) | (10) | (32) | (28) | (34) | (0) | (16) | (36) | (44) | (0) | (16) | (50) | (32) | (2) |
| | hydronephrosis | | 0 | 0 | 0 | 0 | 1 | 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) |
| | tubular necrosis | | 0 | 1 | 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) |
| | papillary necrosis | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | mineralization:papilla | | 1 | 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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 10

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 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 |
| Organ | Findings | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| <hr/> | | | | | | | | | | | | | | | | | | |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | mineralization:pelvis | | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) |
| | | | | | | | | | | | | | | | | | | |
| | urothelial hyperplasia:pelvis | | 1 | 2 | 0 | 0 | 2 | 0 | 0 | 0 | 2 | 1 | 1 | 0 | 2 | 2 | 0 | 0 |
| | | | (2) | (4) | (0) | (0) | (4) | (0) | (0) | (0) | (4) | (2) | (2) | (0) | (4) | (4) | (0) | (0) |
| | atypical tubule hyperplasia | | 0 | 1 | 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) |
| urin bladd | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | dilatation | | 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) | (2) | (0) | (0) | (0) |
| | | | | | | | | | | | | | | | | | | |
| | inflammation | | 0 | 0 | 0 | 0 | 1 | 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) |
| <hr/> | | | | | | | | | | | | | | | | | | |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| pituitary | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | cyst | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (2) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 11

| Organ | Findings | Group Name No. of Animals on Study | | | | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|--------------------|---------------------------|---------------------------------------|-------------|------------|------------|---------------|--------------|------------|------------|---------------|-------------|------------|------------|---------------|--------------|------------|------------|---------------|------------|------------|------------|
| | | Grade | | | | | | | | | | | | | | | | | | | |
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| (Endocrine system) | | | | | | | | | | | | | | | | | | | | | |
| pituitary | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | hyperplasia | 2 (4) | 9 (18) | 0 (0) | 0 (0) | 3 (6) | 10 (20) | 0 (0) | 0 (0) | 2 (4) | 6 (12) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) |
| | Rathke pouch | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 1 (2) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| thyroid | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | follicular hyperplasia | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | C-cell hyperplasia | 14 (28) | 4 (8) | 1 (2) | 0 (0) | 13 (26) | 3 (6) | 2 (4) | 0 (0) | 13 (26) | 6 (12) | 0 (0) | 0 (0) | 0 (0) | 14 (28) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| adrenal | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | osseous metaplasia | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | hyperplasia:medulla | 1 (2) | 5 (10) | 0 (0) | 0 (0) | 1 (2) | 3 (6) | 0 (0) | 0 (0) | 1 (2) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | focal fatty change:cortex | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 12

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|-----------------------|-------------------------------|--|---------------|-------|-------|-------|---------------|--------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| testis | | | | | | | | | | | | | | | | | | |
| | mineralization | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | arteritis | | 0 | 1 | 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) |
| | interstitial cell hyperplasia | | 15 | 1 | 0 | 0 | 9 | 1 | 0 | 0 | 7 | 0 | 0 | 0 | 8 | 0 | 0 | 0 |
| | | | (30) | (2) | (0) | (0) | (18) | (2) | (0) | (0) | (14) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| prostate | | | | | | | | | | | | | | | | | | |
| | inflammation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 2 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) |
| | hyperplasia | | 12 | 1 | 0 | 0 | 7 | 5 | 0 | 0 | 6 | 2 | 0 | 0 | 6 | 1 | 0 | 0 |
| | | | (24) | (2) | (0) | (0) | (14) | (10) | (0) | (0) | (12) | (4) | (0) | (0) | (12) | (2) | (0) | (0) |
| {Nervous system} | | | | | | | | | | | | | | | | | | |
| brain | | | | | | | | | | | | | | | | | | |
| | necrosis:focal | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 1 | 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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 13

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|----------------------------------|-------------------------------|--|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Nervous system} | | | | | | | | | | | | | | | | | | |
| brain | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | mineralization | | 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) | (2) | (0) | (0) |
| | dilatation:cerebral ventricle | | 0 | 1 | 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) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | cataract | | 3 | 2 | 0 | 0 | 1 | 1 | 0 | 0 | 3 | 1 | 0 | 0 | 4 | 1 | 0 | 0 |
| | | | (6) | (4) | (0) | (0) | (2) | (2) | (0) | (0) | (6) | (2) | (0) | (0) | (8) | (2) | (0) | (0) |
| | retinal atrophy | | 2 | 3 | 2 | 0 | 0 | 0 | 1 | 1 | 0 | 1 | 1 | 0 | 0 | 1 | 2 | 2 |
| | | | (4) | (6) | (4) | (0) | (0) | (0) | (2) | (2) | (0) | (2) | (2) | (0) | (0) | (2) | (4) | (4) |
| | keratitis | | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | iritis | | 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) | (2) | (0) | (0) |
| | degeneration:cornea | | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 1 | 0 |
| | | | (6) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (4) | (0) | (2) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 14

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|---------------------------------|--|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | squamous cell metaplasia:cornea | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 1 | 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) |
| Harder gl | lymphocytic infiltration | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 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) | (2) | (0) | (0) | (0) |
| nasolacr d | inflammation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (2) | (0) | (0) |
| {Body cavities} | | | | | | | | | | | | | | | | | | |
| peritoneum | abscess | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 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) | (2) | (0) | (0) | (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 L2

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE
DEAD AND MORIBUND ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|----------------------|---|--|---------------|------|------|------|---------------|-------|-------|------|--------------|-------|------|------|---------------|-------|------|------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | eosinophilic change:olfactory epithelium | | 8 | 1 | 0 | 0 | 10 | 1 | 0 | 0 | 3 | 1 | 0 | 0 | 8 | 5 | 0 | 0 |
| | | | (73) | (9) | (0) | (0) | (67) | (7) | (0) | (0) | (33) | (11) | (0) | (0) | (62) | (38) | (0) | (0) |
| | eosinophilic change:respiratory epithelium | | 3 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 11 | 0 | 0 | 0 * |
| | | | (27) | (0) | (0) | (0) | (33) | (0) | (0) | (0) | (33) | (0) | (0) | (0) | (85) | (0) | (0) | (0) |
| | inflammation:foreign body | | 0 | 1 | 0 | 0 | 2 | 2 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 2 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (13) | (13) | (0) | (0) | (22) | (0) | (0) | (0) | (8) | (15) | (0) | (0) |
| | inflammation:respiratory epithelium | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | respiratory metaplasia:olfactory epithelium | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (15) | (0) | (0) | (0) |
| | respiratory metaplasia:gland | | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 5 | 0 | 0 | 0 |
| | | | (27) | (0) | (0) | (0) | (27) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (38) | (0) | (0) | (0) |
| | squamous cell metaplasia:respiratory epithelium | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (18) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | atrophy:olfactory epithelium | | 0 | 0 | 0 | 0 | 0 | 0 | 3 | 0 | 1 | 0 | 0 | 0 | 1 | 2 | 1 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (20) | (0) | (11) | (0) | (0) | (0) | (8) | (15) | (8) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|------------------------|---------------------------------------|--|---------------|-------|-------|-------|---------------|--------|-------|-------|--------------|-------|--------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| lung | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | congestion | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | edema | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | inflammation | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) |
| | inflammatory infiltration | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | bronchiolar-alveolar cell hyperplasia | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (8) | (8) | (0) | (0) |
| | inflammation:foreign body | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| bone marrow | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | increased hematopoiesis | | 2 | 0 | 0 | 0 | 2 | 2 | 0 | 0 | 4 | 0 | 0 | 0 | 4 | 0 | 0 | 0 |
| | | | (18) | (0) | (0) | (0) | (13) | (13) | (0) | (0) | (44) | (0) | (0) | (0) | (31) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 3

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|-------|----------|--|---------------|-----|-----|-----|---------------|-----|-----|-----|--------------|-----|-----|-----|---------------|-----|-----|-----|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |

{Hematopoietic system}

| | | | | | | | | | | | | | | | | | | |
|--------|--|--|--------|-------|-------|-------|--------|--------|--------|-------|--------|--------|--------|-------|--------|--------|-------|-------|
| spleen | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | hemorrhage | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | deposit of hemosiderin | | 4 | 0 | 0 | 0 | 2 | 3 | 0 | 0 | 2 | 0 | 1 | 0 | 4 | 3 | 0 | 0 |
| | | | (36) | (0) | (0) | (0) | (13) | (20) | (0) | (0) | (22) | (0) | (11) | (0) | (31) | (23) | (0) | (0) |
| | fibrosis:focal | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | increased extramedullary hematopoiesis | | 4 | 1 | 0 | 0 | 1 | 2 | 2 | 0 | 0 | 3 | 0 | 0 | 2 | 1 | 1 | 0 |
| | | | (36) | (9) | (0) | (0) | (7) | (13) | (13) | (0) | (0) | (33) | (0) | (0) | (15) | (8) | (8) | (0) |

{Circulatory system}

| | | | | | | | | | | | | | | | | | | |
|-------|---------------------|--|--------|-------|-------|-------|--------|-------|-------|-------|--------|-------|-------|-------|--------|-------|-------|-------|
| heart | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | myocardial fibrosis | | 7 | 1 | 0 | 0 | 7 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 7 | 1 | 0 | 0 |
| | | | (64) | (9) | (0) | (0) | (47) | (0) | (0) | (0) | (56) | (0) | (0) | (0) | (54) | (8) | (0) | (0) |

{Digestive system}

| | | | | | | | | | | | | | | | | | | |
|-----------|--------------|--|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| esophagus | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | inflammation | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 4

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|--------------------|----------------------------------|--|---------------|--------|--------|-------|---------------|-------|-------|-------|--------------|--------|--------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| stomach | | | | | | | | | | | | | | | | | | |
| | ulcer:forestomach | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 0 | 2 | 1 | 2 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 4 | 1 |
| | | | (18) | (0) | (18) | (9) | (13) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (8) | (31) | (8) |
| | hyperplasia:forestomach | | 2 | 1 | 0 | 0 | 2 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 2 | 2 | 0 | 0 |
| | | | (18) | (9) | (0) | (0) | (13) | (7) | (0) | (0) | (0) | (0) | (11) | (0) | (15) | (15) | (0) | (0) |
| | erosion:glandular stomach | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (18) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (15) | (0) | (0) | (0) |
| | ulcer:glandular stomach | | 0 | 2 | 0 | 0 | 1 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 2 | 1 | 0 | 0 |
| | | | (0) | (18) | (0) | (0) | (7) | (0) | (7) | (0) | (11) | (0) | (0) | (0) | (15) | (8) | (0) | (0) |
| | hyperplasia:glandular stomach | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | mineralization:glandular stomach | | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | dilated glands | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| small intes | | | | | | | | | | | | | | | | | | |
| | erosion | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (20) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 5

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------|-------------------------|-----------|-----------|-----------|-----------|------------|------------|-----------|-----------|------------|-----------|-----------|-----------|-----------|------------|-----------|-----------|
| | | No. of Animals on Study | 11 | | | | 15 | | | | 9 | | | | 13 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | herniation | | 1 (9) | 0 (0) | 0 (0) | 0 (0) | 3 (20) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | hemorrhage | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (7) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | necrosis:central | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (7) | 2 (13) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (15) | 0 (0) | 0 (0) |
| | necrosis:focal | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change | | 0 (0) | 1 (9) | 1 (9) | 0 (0) | 1 (7) | 1 (7) | 1 (7) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change:peripheral | | 0 (0) | 1 (9) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | degeneration:central | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (7) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | clear cell focus | | 1 (9) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 6

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|--------------------|------------------------|--|---------------|--------|--------|--------|---------------|--------|--------|-------|--------------|--------|--------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | acidophilic cell focus | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | basophilic cell focus | | 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) | (8) | (0) | (0) | (0) |
| | spongiosis hepatitis | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | bile duct hyperplasia | | 5 | 5 | 0 | 0 | 7 | 7 | 0 | 0 | 1 | 7 | 0 | 0 | 3 | 10 | 0 | 0 |
| | | | (45) | (45) | (0) | (0) | (47) | (47) | (0) | (0) | (11) | (78) | (0) | (0) | (23) | (77) | (0) | (0) |
| pancreas | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | atrophy | | 2 | 1 | 0 | 0 | 2 | 1 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 2 | 0 | 0 |
| | | | (18) | (9) | (0) | (0) | (13) | (7) | (0) | (0) | (22) | (0) | (0) | (0) | (0) | (15) | (0) | (0) |
| | islet cell hyperplasia | | 0 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (7) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | chronic nephropathy | | 2 | 1 | 4 | 3 | 7 | 2 | 3 | 0 | 4 | 2 | 1 | 0 | 5 | 4 | 4 | 0 |
| | | | (18) | (9) | (36) | (27) | (47) | (13) | (20) | (0) | (44) | (22) | (11) | (0) | (38) | (31) | (31) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 7

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------|-------------------------------|-------------------------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | No. of Animals on Study | 11 | | | | 15 | | | | 9 | | | | 13 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| Organ_____ | Findings_____ | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | hydronephrosis | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | tubular necrosis | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | papillary necrosis | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | mineralization:papilla | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | |
| | mineralization:pelvis | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| | urothelial hyperplasia:pelvis | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (9) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | atypical tubule hyperplasia | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | urin bladd | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | dilatation | | 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) | (8) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 8

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 11 | | | | 125 ppm 15 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|-----------------------|------------------------|--|---------------|--------|-------|-------|---------------|--------|-------|-------|--------------|--------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| urin bladd | inflammation | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| pituitary | hyperplasia | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 3 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (0) | (27) | (0) | (0) | (0) | (13) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (8) | (0) | (0) |
| thyroid | follicular hyperplasia | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 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) | (15) | (0) | (0) | (0) |
| | C-cell hyperplasia | | 3 | 0 | 0 | 0 | 2 | 1 | 1 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (27) | (0) | (0) | (0) | (13) | (7) | (7) | (0) | (22) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| adrenal | hyperplasia:medulla | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (9) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| testis | mineralization | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | | 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) | (8) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 9

| Organ | Findings | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|-----------------------|-------------------------------|---------------------------------------|------|------|------|---------|------|------|------|---------|------|------|------|---------|------|------|------|
| | | Group Name No. of Animals on Study | | | | 11 | | | | 15 | | | | 9 | | | |
| | | Grade | | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Reproductive system} | | | | | | | | | | | | | | | | | |
| testis | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | arteritis | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | interstitial cell hyperplasia | 6 | 1 | 0 | 0 | 6 | 1 | 0 | 0 | 4 | 0 | 0 | 0 | 5 | 0 | 0 | 0 |
| | | (55) | (9) | (0) | (0) | (40) | (7) | (0) | (0) | (44) | (0) | (0) | (0) | (38) | (0) | (0) | (0) |
| prostate | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | inflammation | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | (9) | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) |
| | hyperplasia | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| {Nervous system} | | | | | | | | | | | | | | | | | |
| brain | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | necrosis:focal | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | mineralization | 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) | (8) | (0) | (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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 10

| Organ | Findings | Group Name No. of Animals on Study | | | | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|---------------------------------|---------------------------------------|-------|-------|-------|---------|-------|-------|-------|---------|--------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | Grade | | | | 11 | | | | 15 | | | | 9 | | | | 13 | | | |
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Nervous system} | | | | | | | | | | | | | | | | | | | | | |
| brain | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | | | | | |
| | dilatation:cerebral ventricle | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | | | | |
| eye | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | | | | | |
| | cataract | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | retinal atrophy | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | (0) | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) |
| | keratitis | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (9) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| | degeneration:cornea | 0 | 0 | 0 | 0 | 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) | (0) | (0) | (0) | (8) | (0) |
| | squamous cell metaplasia:cornea | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| nasolacr d | | <11> | | | | <15> | | | | < 9> | | | | <13> | | | | | | | |
| | inflammation | 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) | (0) | (0) | (0) | (11) | (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 : Number of animals with lesion
(c) c : b / a * 100
Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 11

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------|---------------|-------------------------|---------|-----|-----|-----|---------|-----|-----|-----|---------|-----|-----|-----|---------|-----|-----|-----|
| | | No. of Animals on Study | 11 | | | | 15 | | | | 9 | | | | 13 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |

{Body cavities}

| | | | | | | | | | | | | | | | | | |
|------------|---------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| peritoneum | abscess | <11> | | | | <15> | | | | < 9> | | | | <13> | | | |
| | | 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) | (8) | (0) | (0) | (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

(HPT150)

BAIS4

TABLE L3

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE
SACRIFICED ANIMALS

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 1

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|--|-------------------------|---------|-------|-------|--------|---------|-------|-------|--------|---------|--------|-------|--------|---------|--------|-------|-------|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Integumentary system/appandage} | | | | | | | | | | | | | | | | | | |
| skin/app | scab | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| subcutis | inflammation | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | | 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) | (3) | (0) | (0) | (0) | (0) |
| | epidermal cyst | | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | hyperplasia:gland | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | | 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) | (3) | (0) | (0) | (0) | (0) |
| | eosinophilic change:olfactory epithelium | | 26 | 13 | 0 | 0 | 23 | 9 | 0 | 0 | 13 | 24 | 4 | 0 ** | 2 | 29 | 6 | 0 ** |
| | | (67) | (33) | (0) | (0) | (66) | (26) | (0) | (0) | (32) | (59) | (10) | (0) | (5) | (78) | (16) | (0) | (0) |
| | eosinophilic change:respiratory epithelium | | 23 | 0 | 0 | 0 | 23 | 0 | 0 | 0 | 28 | 0 | 0 | 0 | 33 | 0 | 0 | 0 ** |
| | | (59) | (0) | (0) | (0) | (66) | (0) | (0) | (0) | (68) | (0) | (0) | (0) | (89) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------|---|-------------------------|---------|-------|------|------|---------|-------|------|------|---------|-------|------|------|---------|-------|------|------|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | inflammation:foreign body | | 8 | 6 | 0 | 0 | 11 | 4 | 0 | 0 | 7 | 8 | 0 | 0 | 15 | 8 | 0 | 0 |
| | | | (21) | (15) | (0) | (0) | (31) | (11) | (0) | (0) | (17) | (20) | (0) | (0) | (41) | (22) | (0) | (0) |
| | inflammation:olfactory epithelium | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| | respiratory metaplasia:olfactory epithelium | | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 6 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| | respiratory metaplasia:gland | | 7 | 0 | 0 | 0 | 8 | 0 | 0 | 0 | 7 | 1 | 0 | 0 | 10 | 0 | 0 | 0 |
| | | | (18) | (0) | (0) | (0) | (23) | (0) | (0) | (0) | (17) | (2) | (0) | (0) | (27) | (0) | (0) | (0) |
| | squamous cell metaplasia:respiratory epithelium | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | atrophy:olfactory epithelium | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (3) | (0) | (0) | (0) |
| lung | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | accumulation of foamy cells | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | bronchiolar-alveolar cell hyperplasia | | 0 | 0 | 0 | 0 | 3 | 2 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (9) | (6) | (0) | (0) | (7) | (0) | (0) | (0) | (3) | (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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 3

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------------|---------------------------|-------------------------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| lung | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | inflammation:foreign body | | 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) | (3) | (0) | (0) | (0) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| bone marrow | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | hemorrhage | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | increased hematopoiesis | | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| | granulopoiesis:increased | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | xanthogranuloma | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| lymph node | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | lymphadenitis | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (3) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 4

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 39 | | | | 125 ppm 35 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|------------------------|--|--|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| spleen | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | congestion | | 2 | 3 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (5) | (8) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (3) | (0) | (0) | (0) |
| | deposit of hemosiderin | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | fibrosis:focal | | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | increased extramedullary hematopoiesis | | 3 | 2 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 2 | 0 | 0 |
| | | | (8) | (5) | (0) | (0) | (9) | (0) | (0) | (0) | (2) | (2) | (0) | (0) | (0) | (5) | (0) | (0) |
| {Circulatory system} | | | | | | | | | | | | | | | | | | |
| heart | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | myocardial fibrosis | | 22 | 0 | 0 | 0 | 16 | 1 | 0 | 0 | 15 | 1 | 0 | 0 | 13 | 1 | 0 | 0 |
| | | | (56) | (0) | (0) | (0) | (46) | (3) | (0) | (0) | (37) | (2) | (0) | (0) | (35) | (3) | (0) | (0) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| oral cavity | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | squamous cell hyperplasia | | 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) | (3) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 5

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------------|-------------------------|---------|-------|-------|--------|---------|-------|-------|--------|---------|-------|-------|--------|---------|-------|-------|-----|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| stomach | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | hyperplasia:forestomach | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | |
| | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | |
| | erosion:glandular stomach | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | |
| | | (3) | (0) | (0) | (0) | (9) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | |
| small intes | ulcer:glandular stomach | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | |
| | | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | |
| | hyperplasia:glandular stomach | 2 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | |
| | | (5) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | |
| | erosion | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | | |
| liver | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | herniation | 5 | 0 | 0 | 0 | 6 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | |
| | | (13) | (0) | (0) | (0) | (17) | (0) | (0) | (0) | (12) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | |
| | fatty change | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | (0) | (3) | (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 : 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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 6

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|------------------------|-------------------------|---------|-------|------|------|---------|-------|------|------|---------|-------|------|------|---------|-------|------|------|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | | | | | | | | | | | | | | | | |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | fatty change:central | | 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) | (3) | (0) | (0) | (0) |
| | granulation | | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | clear cell focus | | 6 | 1 | 0 | 0 | 11 | 3 | 0 | 0 | 7 | 0 | 0 | 0 | 6 | 0 | 0 | 0 |
| | | | (15) | (3) | (0) | (0) | (31) | (9) | (0) | (0) | (17) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| | acidophilic cell focus | | 1 | 1 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (3) | (3) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | |
| | basophilic cell focus | | 5 | 0 | 0 | 0 | 3 | 1 | 0 | 0 | 5 | 3 | 0 | 0 | 4 | 2 | 0 | 0 |
| | | | (13) | (0) | (0) | (0) | (9) | (3) | (0) | (0) | (12) | (7) | (0) | (0) | (11) | (5) | (0) | (0) |
| | spongiosis hepatis | | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 4 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (10) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| | bile duct hyperplasia | | 2 | 37 | 0 | 0 | 2 | 33 | 0 | 0 | 1 | 40 | 0 | 0 | 2 | 35 | 0 | 0 |
| | | | (5) | (95) | (0) | (0) | (6) | (94) | (0) | (0) | (2) | (98) | (0) | (0) | (5) | (95) | (0) | (0) |
| pancreas | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | atrophy | | 7 | 7 | 1 | 0 | 3 | 3 | 0 | 0 | 7 | 2 | 2 | 0 | 5 | 3 | 0 | 0 |
| | | | (18) | (18) | (3) | (0) | (9) | (9) | (0) | (0) | (17) | (5) | (5) | (0) | (14) | (8) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 7

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 39 | | | | 125 ppm 35 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|--------------------|-------------------------------|--|---------------|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| pancreas | islet cell hyperplasia | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | | | 1 | 2 | 0 | 0 | 1 | 1 | 1 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (5) | (0) | (0) | (3) | (3) | (3) | (0) | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | ectopic tissue | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | cyst | | 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) | (3) | (0) | (0) | (0) |
| | scar | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | chronic nephropathy | | 9 | 21 | 7 | 2 | 9 | 12 | 14 | 0 | 4 | 16 | 21 | 0 ** | 3 | 21 | 12 | 1 |
| | | | (23) | (54) | (18) | (5) | (26) | (34) | (40) | (0) | (10) | (39) | (51) | (0) | (8) | (57) | (32) | (3) |
| | mineralization:pelvis | | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| | urothelial hyperplasia:pelvis | | 0 | 2 | 0 | 0 | 1 | 0 | 0 | 0 | 2 | 1 | 1 | 0 | 2 | 2 | 0 | 0 |
| | | | (0) | (5) | (0) | (0) | (3) | (0) | (0) | (0) | (5) | (2) | (2) | (0) | (5) | (5) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 8

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|------------------------|-------------------------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|-------|-------|-------|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| pituitary | | <39> | | | | | | | | | | | | | | | | |
| | cyst | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) |
| | hyperplasia | | 2 | 6 | 0 | 0 | 3 | 8 | 0 | 0 | 2 | 5 | 0 | 0 | 0 | 0 | 0 | 0 * |
| | | | (5) | (15) | (0) | (0) | (9) | (23) | (0) | (0) | (5) | (12) | (0) | (0) | (0) | (0) | (0) | (0) |
| | Rathke pouch | | 0 | 0 | 0 | 0 | 2 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (6) | (3) | (0) | (0) | (2) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| thyroid | | <39> | | | | | | | | | | | | | | | | |
| | follicular hyperplasia | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | C-cell hyperplasia | | 11 | 4 | 1 | 0 | 11 | 2 | 1 | 0 | 11 | 6 | 0 | 0 | 13 | 2 | 0 | 0 |
| | | | (28) | (10) | (3) | (0) | (31) | (6) | (3) | (0) | (27) | (15) | (0) | (0) | (35) | (5) | (0) | (0) |
| adrenal | | <39> | | | | | | | | | | | | | | | | |
| | osseous metaplasia | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | hyperplasia:medulla | | 1 | 4 | 0 | 0 | 1 | 2 | 0 | 0 | 1 | 2 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (3) | (10) | (0) | (0) | (3) | (6) | (0) | (0) | (2) | (5) | (0) | (0) | (0) | (3) | (0) | (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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 9

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 39 | | | | 125 ppm 35 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|----------------------------------|-------------------------------|--|---------------|-------|-------|-------|---------------|--------|-------|-------|---------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| adrenal | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | focal fatty change:cortex | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| testis | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | mineralization | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | interstitial cell hyperplasia | | 9 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (23) | (0) | (0) | (0) | (9) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| prostate | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | inflammation | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| | hyperplasia | | 12 | 1 | 0 | 0 | 6 | 5 | 0 | 0 | 6 | 2 | 0 | 0 | 5 | 1 | 0 | 0 |
| | | | (31) | (3) | (0) | (0) | (17) | (14) | (0) | (0) | (15) | (5) | (0) | (0) | (14) | (3) | (0) | (0) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | |
| | cataract | | 2 | 2 | 0 | 0 | 1 | 1 | 0 | 0 | 2 | 1 | 0 | 0 | 3 | 1 | 0 | 0 |
| | | | (5) | (5) | (0) | (0) | (3) | (3) | (0) | (0) | (5) | (2) | (0) | (0) | (8) | (3) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 10

| Organ_____ | Findings_____ | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | | |
|----------------------------------|--------------------------|-------------------------|---------|-----|-----|-----|---------|-----|-----|-----|---------|-----|-----|-----|---------|-----|-----|-----|----|
| | | No. of Animals on Study | 39 | | | | 35 | | | | 41 | | | | 37 | | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | |
| <hr/> | | | | | | | | | | | | | | | | | | | |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | | |
| eye | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | | |
| | retinal atrophy | | 2 | 2 | 2 | 0 | 0 | 0 | 1 | 1 | 0 | 1 | 1 | 0 | 0 | 0 | 2 | 2 | |
| | | (| 5) | (| 5) | (| 0) | (| 0) | (| 3) | (| 3) | (| 0) | (| 2) | (| 5) |
| | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | |
| | keratitis | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | (| 0) | (| 3) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) |
| | iritis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | |
| | | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 3) | (| 0) |
| | degeneration:cornea | | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | |
| | | (| 8) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 5) | (| 0) | (| 0) |
| Harder gl | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | | |
| | lymphocytic infiltration | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | |
| | | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 0) | (| 3) | (| 0) | (| 0) |
| nasolacr d | | | <39> | | | | <35> | | | | <41> | | | | <37> | | | | |
| | inflammation | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (| 0) | (| 0) | (| 0) | (| 0) | (| 3) | (| 0) | (| 0) | (| 3) | (| 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 L4

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : FEMALE
ALL ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 15

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|----------------------------------|---|--|---------------|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Integumentary system/appandage} | | | | | | | | | | | | | | | | | | |
| subcutis | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | abscess | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | goblet cell hyperplasia | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (6) | (0) | (0) |
| | eosinophilic change:olfactory epithelium | | 1 | 28 | 20 | 0 | 1 | 31 | 17 | 0 | 0 | 28 | 22 | 0 | 1 | 21 | 28 | 0 |
| | | | (2) | (56) | (40) | (0) | (2) | (62) | (34) | (0) | (0) | (56) | (44) | (0) | (2) | (42) | (56) | (0) |
| | eosinophilic change:respiratory epithelium | | 37 | 7 | 0 | 0 | 41 | 7 | 0 | 0 | 38 | 11 | 0 | 0 | 37 | 11 | 0 | 0 |
| | | | (74) | (14) | (0) | (0) | (82) | (14) | (0) | (0) | (76) | (22) | (0) | (0) | (74) | (22) | (0) | (0) |
| | inflammation:foreign body | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) |
| | inflammation:respiratory epithelium | | 1 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (2) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (2) | (0) | (0) | (0) |
| | respiratory metaplasia:olfactory epithelium | | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (10) | (0) | (0) | (0) | (6) | (0) | (0) | (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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 16

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------|---------------------------------------|--|---------|------|------|------|---------|------|------|------|---------|------|------|------|---------|------|------|------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | respiratory metaplasia:gland | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 14 | 0 | 0 | 0 | 15 | 2 | 0 | 0 | 16 | 0 | 0 | 0 | 16 | 1 | 0 | 0 |
| | | | (28) | (0) | (0) | (0) | (30) | (4) | (0) | (0) | (32) | (0) | (0) | (0) | (32) | (2) | (0) | (0) |
| | atrophy:olfactory epithelium | | 0 | 0 | 0 | 0 | 2 | 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) |
| larynx | inflammation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 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) | (2) | (0) | (0) |
| lung | edema | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | inflammatory infiltration | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | accumulation of foamy cells | | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (2) | (0) | (0) |
| | bronchiolar-alveolar cell hyperplasia | | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (4) | (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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 17

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------------|---------------------------|-------------------------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|
| | | No. of Animals on Study | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| (Respiratory system) | | | | | | | | | | | | | | | | | | |
| lung | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | inflammation:foreign body | | 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) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| (Hematopoietic system) | | | | | | | | | | | | | | | | | | |
| bone marrow | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | granulation | | 0 | 0 | 0 | 0 | 3 | 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) |
| | increased hematopoiesis | | 6 | 1 | 0 | 0 | 4 | 1 | 0 | 0 | 4 | 2 | 0 | 0 | 6 | 1 | 0 | 0 |
| | | | (12) | (2) | (0) | (0) | (8) | (2) | (0) | (0) | (8) | (4) | (0) | (0) | (12) | (2) | (0) | (0) |
| spleen | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | atrophy | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | congestion | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (2) | (2) | (0) |
| | deposit of hemosiderin | | 28 | 6 | 0 | 0 | 30 | 7 | 0 | 0 | 25 | 12 | 0 | 0 | 33 | 9 | 1 | 0 |
| | | | (56) | (12) | (0) | (0) | (60) | (14) | (0) | (0) | (50) | (24) | (0) | (0) | (66) | (18) | (2) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 18

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 50 | | | | 125 ppm 50 | | | | 250 ppm 50 | | | | 500 ppm 50 | | | |
|------------------------|--|--|---------------|------|------|------|---------------|------|------|------|---------------|------|------|------|---------------|------|------|------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| spleen | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | fibrosis:focal | | 0 | 0 | 0 | 0 | 2 | 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) |
| | increased extramedullary hematopoiesis | | 8 | 3 | 1 | 0 | 2 | 4 | 0 | 0 | 3 | 4 | 1 | 0 | 4 | 4 | 2 | 0 |
| | | | (16) | (6) | (2) | (0) | (4) | (8) | (0) | (0) | (6) | (8) | (2) | (0) | (8) | (8) | (4) | (0) |
| {Circulatory system} | | | | | | | | | | | | | | | | | | |
| heart | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | necrosis:focal | | 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) | (2) | (0) | (0) | (0) |
| | myocardial fibrosis | | 6 | 0 | 0 | 0 | 11 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 8 | 0 | 0 | 0 |
| | | | (12) | (0) | (0) | (0) | (22) | (0) | (0) | (0) | (10) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| esophagus | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | inflammation | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 19

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------------|--|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| stomach | inflammation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | ulcer:forestomach | | 1 | 0 | 0 | 1 | 0 | 2 | 0 | 0 | 0 | 1 | 1 | 0 | 3 | 1 | 0 | 0 |
| | | | (2) | (0) | (0) | (2) | (0) | (4) | (0) | (0) | (0) | (2) | (2) | (0) | (6) | (2) | (0) | (0) |
| | hyperplasia:forestomach | | 2 | 0 | 0 | 0 | 1 | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (2) | (2) | (2) | (0) | (0) | (2) | (0) | (0) | (6) | (0) | (0) | (0) |
| small intes | erosion | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | ulcer:glandular stomach | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (4) | (0) | (0) | (4) | (0) | (0) | (0) |
| | hyperplasia:glandular stomach | | 1 | 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) |
| liver | herniation | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 15 | 0 | 0 | 0 | 8 | 0 | 0 | 0 | 9 | 0 | 0 | 0 | 8 | 0 | 0 | 0 |
| | | | (30) | (0) | (0) | (0) | (16) | (0) | (0) | (0) | (18) | (0) | (0) | (0) | (16) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 20

| Organ | Findings | Group Name No. of Animals on Study | | | | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|--------------------------|---------------------------------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|--------------|------------|------------|------------|
| | | Grade | | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | | | | |
| liver | | <50> | | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | peliosis-like lesion | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | necrosis:central | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 4 (8) | 0 (0) | 0 (0) | 0 (0) |
| | necrosis:focal | 3 (6) | 2 (4) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 1 (2) | 0 (0) | 0 (0) |
| | fatty change | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | fatty change:central | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) |
| | lymphocytic infiltration | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | granulation | 7 (14) | 3 (6) | 0 (0) | 0 (0) | 5 (10) | 1 (2) | 0 (0) | 0 (0) | 5 (10) | 0 (0) | 0 (0) | 0 (0) | 5 (10) | 0 (0) | 0 (0) | 0 (0) | 10 (20) | 0 (0) | 0 (0) | 0 (0) |
| | inflammatory cell nest | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (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 : Number of animals with lesion

(c) c : b / a * 100

Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 21

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|------------------------|--|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | fibrosis | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | clear cell focus | | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (2) | (2) | (0) | (0) | (2) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (0) | (2) | (0) | (0) |
| pancreas | basophilic cell focus | | 17 | 5 | 0 | 0 | 21 | 4 | 0 | 0 | 22 | 0 | 0 | 0 | 9 | 1 | 0 | 0 * |
| | | | (34) | (10) | (0) | (0) | (42) | (8) | (0) | (0) | (44) | (0) | (0) | (0) | (18) | (2) | (0) | (0) |
| | bile duct hyperplasia | | 7 | 2 | 0 | 0 | 6 | 1 | 0 | 0 | 3 | 1 | 0 | 0 | 1 | 1 | 0 | 0 |
| | | | (14) | (4) | (0) | (0) | (12) | (2) | (0) | (0) | (6) | (2) | (0) | (0) | (2) | (2) | (0) | (0) |
| | atrophy | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| kidney | | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 4 | 1 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (8) | (2) | (0) | (0) | (6) | (0) | (0) | (0) |
| | islet cell hyperplasia | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | chronic nephropathy | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | | 39 | 7 | 0 | 0 | 39 | 7 | 0 | 0 | 38 | 6 | 1 | 0 | 36 | 10 | 3 | 0 |
| | | | (78) | (14) | (0) | (0) | (78) | (14) | (0) | (0) | (76) | (12) | (2) | (0) | (72) | (20) | (6) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 22

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------|-------------------------------|-------------------------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | No. of Animals on Study | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | hydronephrosis | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | tubular necrosis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | mineralization:papilla | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) |
| | mineralization:pelvis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| | desquamation:pelvis | | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 2 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (4) | (2) | (0) | (0) | (2) | (0) | (0) | (0) |
| | urothelial hyperplasia:pelvis | | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (2) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| | atypical tubule hyperplasia | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | |
| urin bladd | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| dilatation | | 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) | (2) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 23

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------------|--|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| urin bladd | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | transitional cell hyperplasia | | 0 | 1 | 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) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| pituitary | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | angiectasis | | 1 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) |
| | cyst | | 4 | 4 | 0 | 0 | 4 | 0 | 0 | 0 | 6 | 1 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (8) | (8) | (0) | (0) | (8) | (0) | (0) | (0) | (12) | (2) | (0) | (0) | (6) | (0) | (0) | (0) |
| | hyperplasia | | 7 | 7 | 0 | 0 | 11 | 6 | 0 | 0 | 5 | 6 | 0 | 0 | 7 | 6 | 0 | 0 |
| | | | (14) | (14) | (0) | (0) | (22) | (12) | (0) | (0) | (10) | (12) | (0) | (0) | (14) | (12) | (0) | (0) |
| | Rathke pouch | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (4) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (4) | (0) | (0) | (0) | (2) | (0) | (0) | (0) |
| thyroid | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | follicular hyperplasia | | 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) | (2) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 24

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|-----------------------|---------------------------|--|---------|------|------|------|---------|------|------|------|---------|-------|------|------|---------|------|------|------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| thyroid | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | C-cell hyperplasia | | 13 | 3 | 0 | 0 | 13 | 3 | 1 | 0 | 8 | 5 | 0 | 0 | 11 | 4 | 0 | 0 |
| | | | (26) | (6) | (0) | (0) | (26) | (6) | (2) | (0) | (16) | (10) | (0) | (0) | (22) | (8) | (0) | (0) |
| adrenal | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | peliosis-like lesion | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| | hyperplasia:cortical cell | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) |
| | hyperplasia:medulla | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (2) | (0) | (0) | (2) | (0) | (0) | (0) |
| | focal fatty change:cortex | | 8 | 4 | 0 | 0 | 5 | 4 | 0 | 0 | 5 | 2 | 0 | 0 | 10 | 3 | 0 | 0 |
| | | | (16) | (8) | (0) | (0) | (10) | (8) | (0) | (0) | (10) | (4) | (0) | (0) | (20) | (6) | (0) | (0) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| ovary | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | cyst | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (0) | (2) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 25

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|--------------------------------|--|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| uterus | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | cystic endometrial hyperplasia | | 5 | 2 | 0 | 0 | 4 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 6 | 1 | 0 | 0 |
| | | | (10) | (4) | (0) | (0) | (8) | (0) | (0) | (0) | (2) | (4) | (0) | (0) | (12) | (2) | (0) | (0) |
| {Nervous system} | | | | | | | | | | | | | | | | | | |
| brain | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | hemorrhage | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | cataract | | 2 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 1 | 3 | 0 | 0 |
| | | | (4) | (2) | (0) | (0) | (0) | (6) | (0) | (0) | (0) | (8) | (0) | (0) | (2) | (6) | (0) | (0) |
| | retinal atrophy | | 1 | 0 | 2 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 3 | 1 | 0 | 0 | 3 | 1 |
| | | | (2) | (0) | (4) | (0) | (0) | (0) | (2) | (4) | (0) | (0) | (6) | (2) | (0) | (0) | (6) | (2) |
| | keratitis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 26

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|--------------------------|-------------------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|
| | | No. of Animals on Study | 50 | | | | 50 | | | | 50 | | | | 50 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | degeneration:cornea | | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| Harder gl | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | degeneration | | 4 (8) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 3 (6) | 0 (0) | 0 (0) | 0 (0) |
| | lymphocytic infiltration | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| nasolacr d | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | inflammation | | 1 (2) | 5 (10) | 0 (0) | 0 (0) | 0 (0) | 5 (10) | 0 (0) | 0 (0) | 0 (0) | 5 (10) | 1 (2) | 0 (0) | 0 (0) | 6 (12) | 0 (0) | 0 (0) |
| {Musculoskeletal system} | | | | | | | | | | | | | | | | | | |
| muscle | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | fibrosis | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| bone | | | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | osteosclerosis | | 3 (6) | 0 (0) | 0 (0) | 0 (0) | 2 (4) | 2 (4) | 0 (0) | 0 (0) | 3 (6) | 2 (4) | 2 (4) | 0 (0) | 1 (2) | 2 (4) | 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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 27

| Organ | Findings | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 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 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |

{Body cavities}

| | | | | | | | | | | | | | | | | | |
|------------|--------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| peritoneum | inflammation | <50> | | | | <50> | | | | <50> | | | | <50> | | | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 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) |

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)

BAIS4

TABLE L5

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : FEMALE
DEAD AND MORIBUND ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 12

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 12 | | | | 125 ppm 7 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|------------------------|--|--|---------------|-------------|------------|------------|--------------|-------------|------------|-------------|--------------|-------------|-------------|------------|---------------|-------------|-------------|------------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | eosinophilic change:olfactory epithelium | | 1 (8) | 9 (75) | 1 (8) | 0 (0) | 1 (14) | 5 (71) | 0 (0) | 0 (0) | 0 (0) | 8 (89) | 1 (11) | 0 (0) | 1 (8) | 9 (69) | 3 (23) | 0 (0) |
| | eosinophilic change:respiratory epithelium | | 7 (58) | 0 (0) | 0 (0) | 0 (0) | 5 (71) | 0 (0) | 0 (0) | 0 (0) | 9 (100) | 0 (0) | 0 (0) | 0 (0) | 9 (69) | 2 (15) | 0 (0) | 0 (0) |
| | respiratory metaplasia:gland | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (14) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 4 (31) | 0 (0) | 0 (0) | 0 (0) |
| lung | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | edema | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | inflammatory infiltration | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | accumulation of foamy cells | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (8) | 0 (0) | 0 (0) |
| | bronchiolar-alveolar cell hyperplasia | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (11) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| bone marrow | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | increased hematopoiesis | | 2 (17) | 1 (8) | 0 (0) | 0 (0) | 1 (14) | 0 (0) | 0 (0) | 0 (0) | 3 (33) | 1 (11) | 0 (0) | 0 (0) | 5 (38) | 1 (8) | 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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 13

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 12 | | | | 125 ppm 7 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|------------------------|--|--|---------------|--------|-------|-------|--------------|--------|--------|-------|--------------|--------|--------|-------|---------------|--------|--------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| spleen | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | atrophy | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | congestion | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (8) | (0) |
| | deposit of hemosiderin | | 2 | 5 | 0 | 0 | 3 | 0 | 0 | 0 | 2 | 2 | 0 | 0 | 4 | 3 | 1 | 0 |
| | | | (17) | (42) | (0) | (0) | (43) | (0) | (0) | (0) | (22) | (22) | (0) | (0) | (31) | (23) | (8) | (0) |
| | increased extramedullary hematopoiesis | | 1 | 1 | 1 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 1 | 0 | 1 | 3 | 2 | 0 |
| | | | (8) | (8) | (8) | (0) | (0) | (29) | (0) | (0) | (0) | (11) | (11) | (0) | (8) | (23) | (15) | (0) |
| {Circulatory system} | | | | | | | | | | | | | | | | | | |
| heart | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | necrosis:focal | | 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) | (8) | (0) | (0) | (0) |
| | myocardial fibrosis | | 4 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (33) | (0) | (0) | (0) | (43) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (23) | (0) | (0) | (0) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| esophagus | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | inflammation | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 14

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|---------------------------|--|---------|------|------|------|---------|-------|------|------|---------|-------|-------|------|---------|------|------|------|
| | | | 12 | | | | 7 | | | | 9 | | | | 13 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| stomach | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | ulcer:forestomach | | 1 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 1 | 1 | 0 | 3 | 1 | 0 | 0 |
| | | | (8) | (0) | (0) | (8) | (0) | (14) | (0) | (0) | (0) | (11) | (11) | (0) | (23) | (8) | (0) | (0) |
| | hyperplasia:forestomach | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (23) | (0) | (0) | (0) |
| | erosion:glandular stomach | | 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) | (8) | (0) | (0) | (0) | |
| | ulcer:glandular stomach | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (22) | (0) | (0) | (15) | (0) | (0) | (0) |
| small intes | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | erosion | | 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) | (8) | (0) | (0) |
| liver | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | herniation | | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (25) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (22) | (0) | (0) | (0) | (15) | (0) | (0) | (0) |
| | necrosis:central | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 4 | 0 | 0 | 0 | |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (22) | (0) | (0) | (31) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 15

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 12 | | | | 125 ppm 7 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|--------------------|------------------------|--|---------------|-------|------|------|--------------|-------|------|------|--------------|------|------|------|---------------|------|------|------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | | | | | | | | | | | | | | | | |
| | necrosis:focal | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (0) |
| | fatty change | | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | fatty change:central | | 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) | (8) | (0) | (0) |
| | granulation | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | inflammatory cell nest | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | fibrosis | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | clear cell focus | | 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) | (8) | (0) | (0) |
| | basophilic cell focus | | 2 | 1 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (17) | (8) | (0) | (0) | (29) | (0) | (0) | (0) | (11) | (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 : 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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 16

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 12 | | | | 125 ppm 7 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|--------------------|-----------------------|--|---------------|--------|-------|-------|--------------|--------|-------|-------|--------------|-------|-------|-------|---------------|-------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | bile duct hyperplasia | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| pancreas | atrophy | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | chronic nephropathy | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 5 | 3 | 0 | 0 | 2 | 1 | 0 | 0 | 5 | 0 | 0 | 0 | 12 | 0 | 0 | 0 * |
| | | | (42) | (25) | (0) | (0) | (29) | (14) | (0) | (0) | (56) | (0) | (0) | (0) | (92) | (0) | (0) | (0) |
| | hydronephrosis | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | tubular necrosis | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (22) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| urin bladd | dilatation | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 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) | (8) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 17

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------------|--|---------|-------|-------|-------|---------|--------|-------|-------|---------|--------|-------|-------|---------|-------|-------|-------|
| | | | 12 | | | | 7 | | | | 9 | | | | 13 | | | |
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| urin bladd | transitional cell hyperplasia | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| pituitary | cyst | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (8) | (0) | (0) | (0) |
| | hyperplasia | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 2 | 1 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (29) | (0) | (0) | (0) | (11) | (11) | (0) | (0) | (15) | (8) | (0) | (0) |
| thyroid | C-cell hyperplasia | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 1 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (17) | (8) | (0) | (0) | (0) | (14) | (0) | (0) | (11) | (11) | (0) | (0) | (15) | (0) | (0) | (0) |
| adrenal | peliosis-like lesion | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 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) | (11) | (0) | (0) | (0) | (0) | (0) | (0) |
| | focal fatty change:cortex | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 18

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|--------------------------------|-------------------------|---------|------|------|------|---------|------|------|------|---------|-------|------|------|---------|------|------|------|
| | | No. of Animals on Study | 12 | | | | 7 | | | | 9 | | | | 13 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| uterus | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | cystic endometrial hyperplasia | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (23) | (0) | (0) | (0) |
| {Nervous system} | | | | | | | | | | | | | | | | | | |
| brain | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | hemorrhage | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | keratitis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (0) | (8) | (0) | (0) |
| | degeneration:cornea | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| Harder gl | | | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | degeneration | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (17) | (0) | (0) | (0) | (14) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (8) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 19

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 12 | | | | 125 ppm 7 | | | | 250 ppm 9 | | | | 500 ppm 13 | | | |
|-------|----------|--|---------------|-----|-----|-----|--------------|-----|-----|-----|--------------|-----|-----|-----|---------------|-----|-----|-----|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |

{Special sense organs/appendage}

| | | | | | | | | | | | | | | | | | |
|------------|--------------|-------|-------|-------|-------|-------|--------|-------|-------|-------|-------|--------|-------|-------|--------|-------|-------|
| nasolacr d | inflammation | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | 1 | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 2 | 0 | 0 |
| | | (8) | (8) | (0) | (0) | (0) | (29) | (0) | (0) | (0) | (0) | (11) | (0) | (0) | (15) | (0) | (0) |

{Musculoskeletal system}

| | | | | | | | | | | | | | | | | | |
|------|----------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| bone | osteosclerosis | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | 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) | (8) | (0) | (0) | (0) |

{Body cavities}

| | | | | | | | | | | | | | | | | | |
|------------|--------------|-------|-------|-------|-------|-------|-------|--------|-------|-------|-------|-------|-------|-------|-------|-------|-------|
| peritoneum | inflammation | <12> | | | | < 7> | | | | < 9> | | | | <13> | | | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (0) | (0) | (14) | (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 : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Chi Square

(HPT150)

BAIS4

TABLE L6

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : FEMALE
SACRIFICED ANIMALS

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 11

| Organ | Findings | Control No. of Animals on Study Grade | | | | 125 ppm 43 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|----------------------------------|---|---|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|---------------|--------|--------|-------|
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Integumentary system/appandage} | | | | | | | | | | | | | | | | | |
| subcutis | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | abscess | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| {Respiratory system} | | | | | | | | | | | | | | | | | |
| nasal cavit | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | goblet cell hyperplasia | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 |
| | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (8) | (0) | (0) |
| | eosinophilic change:olfactory epithelium | 0 | 19 | 19 | 0 | 0 | 26 | 17 | 0 | 0 | 20 | 21 | 0 | 0 | 12 | 25 | 0 |
| | | (0) | (50) | (50) | (0) | (0) | (60) | (40) | (0) | (0) | (49) | (51) | (0) | (0) | (32) | (68) | (0) |
| | eosinophilic change:respiratory epithelium | 30 | 7 | 0 | 0 | 36 | 7 | 0 | 0 | 29 | 11 | 0 | 0 | 28 | 9 | 0 | 0 |
| | | (79) | (18) | (0) | (0) | (84) | (16) | (0) | (0) | (71) | (27) | (0) | (0) | (76) | (24) | (0) | (0) |
| | inflammation:foreign body | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | (5) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) |
| | inflammation:respiratory epithelium | 1 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | (3) | (0) | (0) | (0) | (2) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (3) | (0) | (0) | (0) |
| | respiratory metaplasia:olfactory epithelium | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | (3) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (12) | (0) | (0) | (0) | (8) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 12

| Organ | Findings | Group Name No. of Animals on Study Grade | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------------|---------------------------------------|--|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-------------|-----------|-----------|-----------|-----------|
| | | | 38 | | | | 43 | | | | 41 | | | | 37 | | | |
| | | | 1 (%) | 2 (%) | 3 (%) | 4 (%) | 1 (%) | 2 (%) | 3 (%) | 4 (%) | 1 (%) | 2 (%) | 3 (%) | 4 (%) | 1 (%) | 2 (%) | 3 (%) | 4 (%) |
| {Respiratory system} | | | | | | | | | | | | | | | | | | |
| nasal cavit | respiratory metaplasia:gland | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 14 (37) | 0 (0) | 0 (0) | 0 (0) | 14 (33) | 2 (5) | 0 (0) | 0 (0) | 15 (37) | 0 (0) | 0 (0) | 0 (0) | 12 (32) | 1 (3) | 0 (0) | 0 (0) | |
| | atrophy:olfactory epithelium | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | |
| larynx | inflammation | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (3) | 0 (0) | 0 (0) | |
| lung | accumulation of foamy cells | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 1 (3) | 0 (0) | 0 (0) | 0 (0) | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (3) | 0 (0) | 0 (0) | 0 (0) | |
| | bronchiolar-alveolar cell hyperplasia | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 1 (3) | 0 (0) | 0 (0) | 0 (0) | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | inflammation:foreign body | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| {Hematopoietic system} | | | | | | | | | | | | | | | | | | |
| bone marrow | granulation | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 3 (7) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 13

| Organ | Findings | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|------------------------|--|-------------------------|-------------|-----------|-----------|-----------|-------------|------------|-----------|-----------|-------------|-------------|-----------|-------------|-------------|------------|-----------|--------------|
| | | No. of Animals on Study | 38 | | | | 43 | | | | 41 | | | | 37 | | | |
| | | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| (Hematopoietic system) | | | | | | | | | | | | | | | | | | |
| bone marrow | increased hematopoiesis | | 4 (11) | 0 (0) | 0 (0) | 0 (0) | 3 (7) | 1 (2) | 0 (0) | 0 (0) | 1 (2) | 1 (2) | 0 (0) | 0 (0) | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| | | | | | | | | | | | | | | | | | | |
| spleen | congestion | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 1 (2) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | deposit of hemosiderin | | 26 (68) | 1 (3) | 0 (0) | 0 (0) | 27 (63) | 7 (16) | 0 (0) | 0 (0) | 23 (56) | 10 (24) | 0 (0) | 0 * (0) | 29 (78) | 6 (16) | 0 (0) | 0 ** (0) |
| | fibrosis:focal | | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) | 0 (0) |
| | increased extramedullary hematopoiesis | | 7 (18) | 2 (5) | 0 (0) | 0 (0) | 2 (5) | 2 (5) | 0 (0) | 0 (0) | 3 (7) | 3 (7) | 0 (0) | 0 (0) | 3 (8) | 1 (3) | 0 (0) | 0 (0) |
| | | | | | | | | | | | | | | | | | | |
| (Circulatory system) | | | | | | | | | | | | | | | | | | |
| heart | myocardial fibrosis | | 2 (5) | 0 (0) | 0 (0) | 0 (0) | 8 (19) | 0 (0) | 0 (0) | 0 (0) | 4 (10) | 0 (0) | 0 (0) | 0 (0) | 5 (14) | 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 : Number of animals with lesion
(c) c : b / a * 100
Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 14

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|--------------------|-------------------------------|-------------------------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|---------|-------|-------|-------|
| | | No. of Animals on Study | 38 | | | | 43 | | | | 41 | | | | 37 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| (Digestive system) | | | | | | | | | | | | | | | | | | |
| stomach | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | inflammation | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | ulcer:forestomach | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| | hyperplasia:forestomach | | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (2) | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | erosion:glandular stomach | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | hyperplasia:glandular stomach | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| liver | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | herniation | | 12 | 0 | 0 | 0 | 8 | 0 | 0 | 0 | 7 | 0 | 0 | 0 | 6 | 0 | 0 | 0 |
| | | | (32) | (0) | (0) | (0) | (19) | (0) | (0) | (0) | (17) | (0) | (0) | (0) | (16) | (0) | (0) | (0) |
| | peliosis-like lesion | | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | necrosis:focal | | 1 | 2 | 0 | 0 | 2 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (3) | (5) | (0) | (0) | (5) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (5) | (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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 15

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 38 | | | | 125 ppm 43 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|--------------------|--------------------------|--|---------------|--------|-------|-------|---------------|--------|-------|-------|---------------|--------|-------|-------|---------------|--------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Digestive system} | | | | | | | | | | | | | | | | | | |
| liver | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | lymphocytic infiltration | | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (5) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | granulation | | 7 | 3 | 0 | 0 | 4 | 1 | 0 | 0 | 4 | 0 | 0 | 0 | 10 | 0 | 0 | 0 |
| | | | (18) | (8) | (0) | (0) | (9) | (2) | (0) | (0) | (10) | (0) | (0) | (0) | (27) | (0) | (0) | (0) |
| | clear cell focus | | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (3) | (3) | (0) | (0) | (2) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| | basophilic cell focus | | 15 | 4 | 0 | 0 | 19 | 4 | 0 | 0 | 21 | 0 | 0 | 0 | 9 | 1 | 0 | 0 |
| | | | (39) | (11) | (0) | (0) | (44) | (9) | (0) | (0) | (51) | (0) | (0) | (0) | (24) | (3) | (0) | (0) |
| | bile duct hyperplasia | | 5 | 2 | 0 | 0 | 5 | 1 | 0 | 0 | 2 | 1 | 0 | 0 | 1 | 1 | 0 | 0 |
| | | | (13) | (5) | (0) | (0) | (12) | (2) | (0) | (0) | (5) | (2) | (0) | (0) | (3) | (3) | (0) | (0) |
| pancreas | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | atrophy | | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 4 | 1 | 0 | 0 | 3 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (10) | (2) | (0) | (0) | (8) | (0) | (0) | (0) |
| | islet cell hyperplasia | | 0 | 0 | 0 | 0 | 0 | 1 | 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) |
| {Urinary system} | | | | | | | | | | | | | | | | | | |
| kidney | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | chronic nephropathy | | 34 | 4 | 0 | 0 | 37 | 6 | 0 | 0 | 33 | 6 | 1 | 0 | 24 | 10 | 3 | 0 * |
| | | | (89) | (11) | (0) | (0) | (86) | (14) | (0) | (0) | (80) | (15) | (2) | (0) | (65) | (27) | (8) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 16

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 38 | | | | 125 ppm 43 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|-------|----------|--|---------------|-----|-----|-----|---------------|-----|-----|-----|---------------|-----|-----|-----|---------------|-----|-----|-----|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |

{Urinary system}

| | | | | | | | | | | | | | | | | | | |
|--------|-------------------------------|--|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|--------|-------|-------|-------|
| kidney | mineralization:papilla | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) |
| | mineralization:pelvis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (11) | (0) | (0) | (0) |
| | desquamation:pelvis | | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 2 | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (5) | (2) | (0) | (0) | (3) | (0) | (0) | (0) |
| | urothelial hyperplasia:pelvis | | 0 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| | | | (0) | (3) | (0) | (0) | (2) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (3) | (0) | (0) | (0) |
| | atypical tubule hyperplasia | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |

{Endocrine system}

| | | | | | | | | | | | | | | | | | | |
|-----------|-------------|--|--------|--------|-------|-------|-------|-------|-------|-------|--------|-------|-------|-------|-------|-------|-------|-------|
| pituitary | angiectasis | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | | | 1 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (3) | (0) | (0) |
| | cyst | | 4 | 4 | 0 | 0 | 3 | 0 | 0 | 0 | 5 | 1 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (11) | (11) | (0) | (0) | (7) | (0) | (0) | (0) | (12) | (2) | (0) | (0) | (5) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 17

| Organ | Findings | Group Name No. of Animals on Study Grade | | | | Control 38 | | | | 125 ppm 43 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|--------------------|---------------------------|--|-------|------|------|---------------|-------|------|------|---------------|-------|------|------|---------------|-------|------|------|---------------|-----|-----|-----|
| | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Endocrine system} | | | | | | | | | | | | | | | | | | | | | |
| pituitary | hyperplasia | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 5 | 7 | 0 | 0 | 9 | 6 | 0 | 0 | 4 | 5 | 0 | 0 | 5 | 5 | 0 | 0 | | | | |
| | | (13) | (18) | (0) | (0) | (21) | (14) | (0) | (0) | (10) | (12) | (0) | (0) | (14) | (14) | (0) | (0) | | | | |
| | Rathke pouch | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | | | | |
| | | (5) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (3) | (0) | (0) | (0) | | | | |
| thyroid | follicular hyperplasia | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 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) | (3) | (0) | (0) | (0) | | | | |
| | C-cell hyperplasia | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 11 | 2 | 0 | 0 | 13 | 2 | 1 | 0 | 7 | 4 | 0 | 0 | 9 | 4 | 0 | 0 | | | | |
| | | (29) | (5) | (0) | (0) | (30) | (5) | (2) | (0) | (17) | (10) | (0) | (0) | (24) | (11) | (0) | (0) | | | | |
| adrenal | peliosis-like lesion | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | | | |
| | | (0) | (3) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | | | | |
| | hyperplasia:cortical cell | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 | 0 | | | | |
| | | (0) | (0) | (0) | (0) | (0) | (2) | (0) | (0) | (2) | (2) | (0) | (0) | (0) | (0) | (0) | (0) | | | | |
| | hyperplasia:medulla | <38> | | | | <43> | | | | <41> | | | | <37> | | | | | | | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 1 | 0 | 0 | 0 | | | | |
| | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (2) | (2) | (0) | (0) | (3) | (0) | (0) | (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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 18

| | | Group Name | Control | | | | 125 ppm | | | | 250 ppm | | | | 500 ppm | | | |
|----------------------------------|--------------------------------|-------------------------|---------|-------|------|------|---------|------|------|------|---------|-------|------|------|---------|------|------|------|
| | | No. of Animals on Study | 38 | | | | 43 | | | | 41 | | | | 37 | | | |
| Organ | Findings | Grade | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | | | | | | | | | | | | | | | | |
| {Endocrine system} | | | | | | | | | | | | | | | | | | |
| adrenal | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | focal fatty change:cortex | | 6 | 4 | 0 | 0 | 5 | 4 | 0 | 0 | 5 | 2 | 0 | 0 | 10 | 2 | 0 | 0 |
| | | | (16) | (11) | (0) | (0) | (12) | (9) | (0) | (0) | (12) | (5) | (0) | (0) | (27) | (5) | (0) | (0) |
| | | | | | | | | | | | | | | | | | | |
| {Reproductive system} | | | | | | | | | | | | | | | | | | |
| ovary | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | cyst | | 1 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| | | | (3) | (0) | (0) | (0) | (2) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (0) | (3) | (0) | (0) |
| uterus | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | cystic endometrial hyperplasia | | 3 | 2 | 0 | 0 | 4 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 3 | 1 | 0 | 0 |
| | | | (8) | (5) | (0) | (0) | (9) | (0) | (0) | (0) | (2) | (5) | (0) | (0) | (8) | (3) | (0) | (0) |
| | | | | | | | | | | | | | | | | | | |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | cataract | | 2 | 1 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 4 | 0 | 0 | 1 | 3 | 0 | 0 |
| | | | (5) | (3) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (10) | (0) | (0) | (3) | (8) | (0) | (0) |
| | retinal atrophy | | 1 | 0 | 2 | 0 | 0 | 0 | 1 | 2 | 0 | 0 | 3 | 1 | 0 | 0 | 3 | 1 |
| | | | (3) | (0) | (5) | (0) | (0) | (0) | (2) | (5) | (0) | (0) | (7) | (2) | (0) | (0) | (8) | (3) |

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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 19

| Organ | Findings | Group Name No. of Animals on Study Grade | Control 38 | | | | 125 ppm 43 | | | | 250 ppm 41 | | | | 500 ppm 37 | | | |
|----------------------------------|--------------------------|--|---------------|--------|-------|-------|---------------|-------|-------|-------|---------------|--------|-------|-------|---------------|--------|-------|-------|
| | | | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 | 1 | 2 | 3 | 4 |
| | | | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) | (%) |
| {Special sense organs/appendage} | | | | | | | | | | | | | | | | | | |
| eye | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | degeneration:cornea | | 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) | (3) | (0) | (0) | (0) |
| Harder gl | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | degeneration | | 2 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 |
| | | | (5) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) |
| | lymphocytic infiltration | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| nasolacr d | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | inflammation | | 0 | 4 | 0 | 0 | 0 | 3 | 0 | 0 | 0 | 5 | 0 | 0 | 0 | 4 | 0 | 0 |
| | | | (0) | (11) | (0) | (0) | (0) | (7) | (0) | (0) | (0) | (12) | (0) | (0) | (0) | (11) | (0) | (0) |
| {Musculoskeletal system} | | | | | | | | | | | | | | | | | | |
| muscle | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | fibrosis | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (0) | (5) | (0) | (0) | (0) | (0) | (0) | (0) | (0) |
| bone | | | <38> | | | | <43> | | | | <41> | | | | <37> | | | |
| | osteosclerosis | | 3 | 0 | 0 | 0 | 2 | 2 | 0 | 0 | 3 | 2 | 2 | 0 | 0 | 2 | 0 | 0 |
| | | | (8) | (0) | (0) | (0) | (5) | (5) | (0) | (0) | (7) | (5) | (5) | (0) | (0) | (5) | (0) | (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 M1

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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 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 | 125 ppm | 250 ppm | 500 ppm |
|--------------------|-------------------------------------|------------|---------|---------|---------|---------|
| 0 - 52 | NO. OF EXAMINED ANIMALS | | 0 | 0 | 1 | 0 |
| | NO. OF ANIMALS WITH TUMORS | | 0 | 0 | 1 | 0 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 0 | 0 | 1 | 0 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 0 | 0 | 0 | 0 |
| | NO. OF BENIGN TUMORS | | 0 | 0 | 0 | 0 |
| | NO. OF MALIGNANT TUMORS | | 0 | 0 | 1 | 0 |
| | NO. OF TOTAL TUMORS | | 0 | 0 | 1 | 0 |
| 53 - 78 | NO. OF EXAMINED ANIMALS | | 1 | 4 | 0 | 2 |
| | NO. OF ANIMALS WITH TUMORS | | 1 | 4 | 0 | 1 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 0 | 3 | 0 | 1 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 1 | 1 | 0 | 0 |
| | NO. OF BENIGN TUMORS | | 1 | 3 | 0 | 1 |
| | NO. OF MALIGNANT TUMORS | | 1 | 2 | 0 | 0 |
| | NO. OF TOTAL TUMORS | | 2 | 5 | 0 | 1 |
| 79 - 104 | NO. OF EXAMINED ANIMALS | | 10 | 11 | 8 | 11 |
| | NO. OF ANIMALS WITH TUMORS | | 10 | 11 | 8 | 10 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 2 | 2 | 2 | 1 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 8 | 9 | 6 | 9 |
| | NO. OF BENIGN TUMORS | | 14 | 22 | 11 | 16 |
| | NO. OF MALIGNANT TUMORS | | 7 | 7 | 6 | 7 |
| | NO. OF TOTAL TUMORS | | 21 | 29 | 17 | 23 |
| 105 - 105 | NO. OF EXAMINED ANIMALS | | 39 | 35 | 41 | 37 |
| | NO. OF ANIMALS WITH TUMORS | | 39 | 35 | 41 | 37 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 13 | 11 | 12 | 15 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 26 | 24 | 29 | 22 |
| | NO. OF BENIGN TUMORS | | 65 | 65 | 78 | 67 |
| | NO. OF MALIGNANT TUMORS | | 11 | 10 | 11 | 4 |
| | NO. OF TOTAL TUMORS | | 76 | 75 | 89 | 71 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 2

| Time-related Weeks | Items | Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|-----------------------|-------------------------------------|------------|---------|---------|---------|---------|
| 0 - 105 | NO. OF EXAMINED ANIMALS | | 50 | 50 | 50 | 50 |
| | NO. OF ANIMALS WITH TUMORS | | 50 | 50 | 50 | 48 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 15 | 16 | 15 | 17 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 35 | 34 | 35 | 31 |
| | NO. OF BENIGN TUMORS | | 80 | 90 | 89 | 84 |
| | NO. OF MALIGNANT TUMORS | | 19 | 19 | 18 | 11 |
| | NO. OF TOTAL TUMORS | | 99 | 109 | 107 | 95 |

(HPT070)

BAIS4

TABLE M2

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

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
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 | 125 ppm | 250 ppm | 500 ppm |
|-----------------------|-------------------------------------|------------|---------|---------|---------|---------|
| 0 - 52 | NO. OF EXAMINED ANIMALS | | 0 | 0 | 0 | 0 |
| | NO. OF ANIMALS WITH TUMORS | | 0 | 0 | 0 | 0 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 0 | 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 | | 0 | 0 | 0 | 0 |
| | NO. OF TOTAL TUMORS | | 0 | 0 | 0 | 0 |
| 53 - 78 | NO. OF EXAMINED ANIMALS | | 1 | 2 | 0 | 4 |
| | NO. OF ANIMALS WITH TUMORS | | 1 | 1 | 0 | 4 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 1 | 1 | 0 | 3 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 0 | 0 | 0 | 1 |
| | NO. OF BENIGN TUMORS | | 0 | 1 | 0 | 0 |
| | NO. OF MALIGNANT TUMORS | | 1 | 0 | 0 | 5 |
| | NO. OF TOTAL TUMORS | | 1 | 1 | 0 | 5 |
| 79 - 104 | NO. OF EXAMINED ANIMALS | | 11 | 5 | 9 | 9 |
| | NO. OF ANIMALS WITH TUMORS | | 10 | 4 | 9 | 8 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 7 | 2 | 5 | 6 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 3 | 2 | 4 | 2 |
| | NO. OF BENIGN TUMORS | | 8 | 3 | 7 | 4 |
| | NO. OF MALIGNANT TUMORS | | 5 | 3 | 6 | 6 |
| | NO. OF TOTAL TUMORS | | 13 | 6 | 13 | 10 |
| 105 - 105 | NO. OF EXAMINED ANIMALS | | 38 | 43 | 41 | 37 |
| | NO. OF ANIMALS WITH TUMORS | | 24 | 27 | 25 | 16 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 13 | 21 | 17 | 11 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 11 | 6 | 8 | 5 |
| | NO. OF BENIGN TUMORS | | 30 | 33 | 31 | 18 |
| | NO. OF MALIGNANT TUMORS | | 7 | 3 | 5 | 4 |
| | NO. OF TOTAL TUMORS | | 37 | 36 | 36 | 22 |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 4

| Time-related Weeks | Items | Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|-----------------------|-------------------------------------|------------|---------|---------|---------|---------|
| 0 - 105 | NO. OF EXAMINED ANIMALS | | 50 | 50 | 50 | 50 |
| | NO. OF ANIMALS WITH TUMORS | | 35 | 32 | 34 | 28 |
| | NO. OF ANIMALS WITH SINGLE TUMORS | | 21 | 24 | 22 | 20 |
| | NO. OF ANIMALS WITH MULTIPLE TUMORS | | 14 | 8 | 12 | 8 |
| | NO. OF BENIGN TUMORS | | 38 | 37 | 38 | 22 |
| | NO. OF MALIGNANT TUMORS | | 13 | 6 | 11 | 15 |
| | NO. OF TOTAL TUMORS | | 51 | 43 | 49 | 37 |

(HPT070)

BAIS4

TABLE N1

HISTOPATHOLOGICAL FINDINGS :
NEOPLASTIC LESIONS : MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|----------------------------------|------------------------------|---------------------------------------|---------------|---------------|---------------|---------------|
| {Integumentary system/appandage} | | | | | | |
| skin/app | | | <50> | <50> | <50> | <50> |
| | squamous cell papilloma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| | trichoepithelioma | | 1 (2%) | 0 (0%) | 1 (2%) | 1 (2%) |
| | keratoacanthoma | | 0 (0%) | 2 (4%) | 0 (0%) | 1 (2%) |
| | sebaceous adenoma | | 0 (0%) | 2 (4%) | 0 (0%) | 0 (0%) |
| | squamous cell carcinoma | | 0 (0%) | 0 (0%) | 1 (2%) | 1 (2%) |
| subcutis | | | <50> | <50> | <50> | <50> |
| | fibroma | | 3 (6%) | 1 (2%) | 5 (10%) | 5 (10%) |
| | hemangioma | | 1 (2%) | 0 (0%) | 1 (2%) | 1 (2%) |
| | histiocytic sarcoma | | 0 (0%) | 1 (2%) | 0 (0%) | 0 (0%) |
| {Respiratory system} | | | | | | |
| nasal cavit | | | <50> | <50> | <50> | <50> |
| | adenoma | | 0 (0%) | 1 (2%) | 1 (2%) | 0 (0%) |
| | chondroma | | 1 (2%) | 0 (0%) | 0 (0%) | 0 (0%) |
| | osteosarcoma | | 1 (2%) | 0 (0%) | 0 (0%) | 0 (0%) |
| lung | | | <50> | <50> | <50> | <50> |
| | bronchiolar-alveolar adenoma | | 2 (4%) | 1 (2%) | 3 (6%) | 6 (12%) |

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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|------------------------|--------------------------------|---------------------------------------|------------------|------------------|-----------------|-----------------|
| {Respiratory system} | | | | | | |
| lung | bronchiolar-alveolar carcinoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 2 (4%) | <50> 1 (2%) |
| {Hematopoietic system} | | | | | | |
| spleen | mononuclear cell leukemia | | <50> 7 (14%) | <50> 4 (8%) | <50> 2 (4%) | <50> 1 (2%) |
| | hemangiosarcoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| {Digestive system} | | | | | | |
| stomach | squamous cell papilloma | | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) | <50> 1 (2%) |
| | squamous cell carcinoma | | 0 (0%) | 2 (4%) | 0 (0%) | 1 (2%) |
| large intes | adenoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |
| liver | hepatocellular adenoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 3 (6%) | <50> 3 (6%) |
| | hemangiosarcoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| | hepatocellular carcinoma | | 0 (0%) | 0 (0%) | 1 (2%) | 0 (0%) |
| pancreas | islet cell adenoma | | <50> 1 (2%) | <50> 7 (14%) | <50> 2 (4%) | <50> 1 (2%) |
| | acinar cell adenoma | | 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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 3

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|--------------------|-----------------------------|---------------------------------------|-------------------|-------------------|------------------|------------------|
| {Digestive system} | | | | | | |
| pancreas | islet cell adenocarcinoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |
| {Urinary system} | | | | | | |
| kidney | transitional cell papilloma | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) |
| | nephroblastoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) |
| urin bladd | transitional cell papilloma | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 2 (4%) |
| {Endocrine system} | | | | | | |
| pituitary | adenoma | | <50> 12 (24%) | <50> 9 (18%) | <50> 9 (18%) | <50> 5 (10%) |
| thyroid | C-cell adenoma | | <50> 7 (14%) | <50> 11 (22%) | <50> 9 (18%) | <50> 6 (12%) |
| | follicular adenoma | | <50> 0 (0%) | <50> 2 (4%) | <50> 2 (4%) | <50> 2 (4%) |
| | C-cell carcinoma | | <50> 6 (12%) | <50> 1 (2%) | <50> 3 (6%) | <50> 1 (2%) |
| | follicular adenocarcinoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) |
| adrenal | pheochromocytoma | | <50> 4 (8%) | <50> 4 (8%) | <50> 1 (2%) | <50> 2 (4%) |
| | pheochromocytoma:malignant | | <50> 0 (0%) | <50> 1 (2%) | <50> 3 (6%) | <50> 0 (0%) |

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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 4

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|----------------------------------|------------------------------|---------------------------------------|-------------------|-------------------|-------------------|-------------------|
| {Reproductive system} | | | | | | |
| testis | interstitial cell tumor | | <50> 42 (84%) | <50> 44 (88%) | <50> 46 (92%) | <50> 43 (86%) |
| prostate | adenoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |
| mammary gl | fibroadenoma | | <50> 1 (2%) | <50> 2 (4%) | <50> 1 (2%) | <50> 0 (0%) |
| | adenocarcinoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| prep/cli gl | adenoma | | <50> 1 (2%) | <50> 2 (4%) | <50> 2 (4%) | <50> 3 (6%) |
| {Nervous system} | | | | | | |
| brain | malignant reticulosis | | <50> 0 (0%) | <50> 1 (2%) | <50> 1 (2%) | <50> 0 (0%) |
| | glioma | | <50> 0 (0%) | <50> 3 (6%) | <50> 1 (2%) | <50> 1 (2%) |
| {Special sense organs/appendage} | | | | | | |
| Harder gl | adenocarcinoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |
| Zymbal gl | Zmbal gland tumor:benign | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| | Zymbal gland tumor:malignant | | <50> 0 (0%) | <50> 1 (2%) | <50> 1 (2%) | <50> 2 (4%) |
| {Musculoskeletal system} | | | | | | |
| bone | osteosarcoma | | <50> 0 (0%) | <50> 2 (4%) | <50> 0 (0%) | <50> 0 (0%) |

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

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 5

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|-----------------|--|---------------------------------------|-----------------|-----------------|-----------------|-----------------|
| {Body cavities} | | | | | | |
| peritoneum | mesothelioma | | <50> 2 (4%) | <50> 1 (2%) | <50> 1 (2%) | <50> 0 (0%) |
| retroperit | histiocytic sarcoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) |
| adipose | lipoma | | <50> 2 (4%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| | | | | | | |
| < a > | a : Number of animals examined at the site | | | | | |
| b (c) | b : Number of animals with neoplasm c : b / a * 100 | | | | | |

(HPT085)

BAIS4

TABLE N2

**HISTOPATHOLOGICAL FINDINGS :
NEOPLASTIC LESIONS : FEMALE**

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 6

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|----------------------------------|--------------------------------|---------------------------------------|------------------|------------------|------------------|-----------------|
| {Integumentary system/appandage} | | | | | | |
| skin/app | keratoacanthoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 1 (2%) | <50> 0 (0%) |
| subcutis | fibroma | | <50> 1 (2%) | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) |
| {Respiratory system} | | | | | | |
| lung | bronchiolar-alveolar adenoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 1 (2%) | <50> 2 (4%) |
| | bronchiolar-alveolar carcinoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| {Hematopoietic system} | | | | | | |
| lymph node | histiocytic sarcoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| spleen | mononuclear cell leukemia | | <50> 5 (10%) | <50> 5 (10%) | <50> 6 (12%) | <50> 1 (2%) |
| {Digestive system} | | | | | | |
| tongue | squamous cell papilloma | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) |
| stomach | squamous cell papilloma | | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) |
| | squamous cell carcinoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| large intes | adenocarcinoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 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. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 7

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|--------------------|-----------------------------|---------------------------------------|-------------------|-------------------|-------------------|------------------|
| {Digestive system} | | | | | | |
| liver | hepatocellular adenoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 1 (2%) | <50> 1 (2%) |
| | hepatocellular carcinoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| pancreas | islet cell adenoma | | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) |
| | islet cell adenocarcinoma | | 0 (0%) | 0 (0%) | 2 (4%) | 0 (0%) |
| {Urinary system} | | | | | | |
| urin bladd | transitional cell papilloma | | <50> 0 (0%) | <50> 0 (0%) | <50> 2 (4%) | <50> 0 (0%) |
| | transitional cell carcinoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| {Endocrine system} | | | | | | |
| pituitary | adenoma | | <50> 13 (26%) | <50> 14 (28%) | <50> 13 (26%) | <50> 7 (14%) |
| | adenocarcinoma | | 1 (2%) | 0 (0%) | 0 (0%) | 1 (2%) |
| thyroid | C-cell adenoma | | <50> 6 (12%) | <50> 6 (12%) | <50> 2 (4%) | <50> 3 (6%) |
| | C-cell carcinoma | | 1 (2%) | 0 (0%) | 0 (0%) | 1 (2%) |
| adrenal | pheochromocytoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |

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

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 8

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|----------------------------------|-----------------------------|---------------------------------------|------------------|-----------------|------------------|-----------------|
| {Endocrine system} | | | | | | |
| adrenal | pheochromocytoma:malignant | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) |
| {Reproductive system} | | | | | | |
| ovary | granulosa-theca cell tumor | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| uterus | endometrial stromal polyp | | <50> 5 (10%) | <50> 3 (6%) | <50> 5 (10%) | <50> 3 (6%) |
| | adenocarcinoma | | 0 (0%) | 0 (0%) | 1 (2%) | 0 (0%) |
| | endometrial stromal sarcoma | | 3 (6%) | 1 (2%) | 1 (2%) | 3 (6%) |
| mammary gl | adenoma | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 0 (0%) |
| | fibroadenoma | | 8 (16%) | 5 (10%) | 5 (10%) | 1 (2%) |
| | adenocarcinoma | | 0 (0%) | 0 (0%) | 0 (0%) | 1 (2%) |
| prep/cli gl | adenoma | | <50> 2 (4%) | <50> 4 (8%) | <50> 6 (12%) | <50> 3 (6%) |
| {Nervous system} | | | | | | |
| periph nerv | histiocytic sarcoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| {Special sense organs/appendage} | | | | | | |
| Zymbal gl | Zmbal gland tumor:benign | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 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. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 9

| Organ | Findings | Group Name No. of animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|----------------------------------|------------------------------|---------------------------------------|-----------------|-----------------|-----------------|-----------------|
| {Special sense organs/appendage} | | | | | | |
| Zymbal gl | Zymbal gland tumor:malignant | | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) | <50> 1 (2%) |
| {Musculoskeletal system} | | | | | | |
| bone | osteoma | | <50> 1 (2%) | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) |
| vertebra | chordoma:malignant | | <50> 0 (0%) | <50> 0 (0%) | <50> 0 (0%) | <50> 1 (2%) |

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

(HPT085)

BAIS4

TABLE O1

NEOPLASTIC LESIONS-INCIDENCE
AND STATISTICAL ANALYSIS : MALE

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 1

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|---|-------------|------------|-------------|-------------|
| SITE : subcutis TUMOR : fibroma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 3/50(6.0) | 1/50(2.0) | 5/50(10.0) | 5/50(10.0) |
| Adjusted rates(b) | 7.69 | 2.86 | 9.76 | 13.51 |
| Terminal rates(c) | 3/39(7.7) | 1/35(2.9) | 4/41(9.8) | 5/37(13.5) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.3942 | | | |
| Prevalence method(d) | P = 0.1098 | | | |
| Combined analysis(d) | P = 0.1088 | | | |
| Cochran-Armitage test(e) | P = 0.2232 | | | |
| Fisher Exact test(e) | | P = 0.3087 | P = 0.3575 | P = 0.3575 |
| SITE : lung TUMOR : bronchiolar-alveolar adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 2/50(4.0) | 1/50(2.0) | 3/50(6.0) | 6/50(12.0) |
| Adjusted rates(b) | 5.13 | 2.86 | 6.98 | 15.00 |
| Terminal rates(c) | 2/39(5.1) | 1/35(2.9) | 2/41(4.9) | 5/37(13.5) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.0252* | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.0441* | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.5000 | P = 0.1343 |
| SITE : lung TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 3/50(6.0) | 1/50(2.0) | 5/50(10.0) | 7/50(14.0) |
| Adjusted rates(b) | 7.69 | 2.86 | 11.63 | 17.50 |
| Terminal rates(c) | 3/39(7.7) | 1/35(2.9) | 4/41(9.8) | 6/37(16.2) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.0292* | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.0526 | | | |
| Fisher Exact test(e) | | P = 0.3087 | P = 0.3575 | P = 0.1589 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 2

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|-------------|------------|------------|-------------|
| SITE : spleen TUMOR : mononuclear cell leukemia | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 7/50(14.0) | 4/50(8.0) | 2/50(4.0) | 1/50(2.0) |
| Adjusted rates(b) | 12.82 | 5.71 | 2.44 | 0.0 |
| Terminal rates(c) | 5/39(12.8) | 2/35(5.7) | 1/41(2.4) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.7385 | | | |
| Prevalence method(d) | P = 0.9963 | | | |
| Combined analysis(d) | P = 0.9916 | | | |
| Cochran-Armitage test(e) | P = 0.0192* | | | |
| Fisher Exact test(e) | | P = 0.2623 | P = 0.0798 | P = 0.0297* |
| SITE : stomach TUMOR : squamous cell papilloma, squamous cell carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 3/50(6.0) | 0/50(0.0) | 2/50(4.0) |
| Adjusted rates(b) | 0.0 | 6.00 | 0.0 | 4.88 |
| Terminal rates(c) | 0/39(0.0) | 2/35(5.7) | 0/41(0.0) | 1/37(2.7) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.2412 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.4908 | | | |
| Fisher Exact test(e) | | P = 0.1212 | P = N. C. | P = 0.2475 |
| SITE : liver TUMOR : hepatocellular adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 1/50(2.0) | 3/50(6.0) | 3/50(6.0) |
| Adjusted rates(b) | 0.0 | 2.86 | 7.32 | 7.50 |
| Terminal rates(c) | 0/39(0.0) | 1/35(2.9) | 3/41(7.3) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.0455* | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.0791 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.1212 | P = 0.1212 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 3

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|-------------|-------------|------------|------------|
| SITE : liver TUMOR : hepatocellular adenoma, hepatocellular carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 1/50(2.0) | 4/50(8.0) | 3/50(6.0) |
| Adjusted rates(b) | 0.0 | 2.86 | 9.76 | 7.50 |
| Terminal rates(c) | 0/39(0.0) | 1/35(2.9) | 4/41(9.8) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.0497* | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.0877 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.0587 | P = 0.1212 |
| SITE : pancreas TUMOR : islet cell adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 1/50(2.0) | 7/50(14.0) | 2/50(4.0) | 1/50(2.0) |
| Adjusted rates(b) | 2.27 | 17.50 | 4.88 | 2.50 |
| Terminal rates(c) | 0/39(0.0) | 6/35(17.1) | 2/41(4.9) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.7934 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.3728 | | | |
| Fisher Exact test(e) | | P = 0.0297* | P = 0.5000 | P = 0.7525 |
| SITE : pancreas TUMOR : islet cell adenoma, islet cell adenocarcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 2/50(4.0) | 7/50(14.0) | 3/50(6.0) | 1/50(2.0) |
| Adjusted rates(b) | 4.55 | 17.50 | 7.32 | 2.50 |
| Terminal rates(c) | 1/39(2.6) | 6/35(17.1) | 3/41(7.3) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.8528 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.2648 | | | |
| Fisher Exact test(e) | | P = 0.0798 | P = 0.5000 | P = 0.5000 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 4

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|--------------|--------------|-------------|-------------|
| SITE : pituitary gland TUMOR : adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 12/50(24.0) | 9/50(18.0) | 9/50(18.0) | 5/50(10.0) |
| Adjusted rates(b) | 17.95 | 15.38 | 19.57 | 7.32 |
| Terminal rates(c) | 7/39(17.9) | 4/35(11.4) | 7/41(17.1) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.9168 | | | |
| Prevalence method(d) | P = 0.8786 | | | |
| Combined analysis(d) | P = 0.9607 | | | |
| Cochran-Armitage test(e) | P = 0.0730 | | | |
| Fisher Exact test(e) | | P = 0.3121 | P = 0.3121 | P = 0.0542 |
| SITE : thyroid TUMOR : C-cell adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 7/50(14.0) | 11/50(22.0) | 9/50(18.0) | 6/50(12.0) |
| Adjusted rates(b) | 17.95 | 27.78 | 20.45 | 13.64 |
| Terminal rates(c) | 7/39(17.9) | 9/35(25.7) | 8/41(19.5) | 5/37(13.5) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ——— | | | |
| Prevalence method(d) | P = 0.7099 | | | |
| Combined analysis(d) | P = ——— | | | |
| Cochran-Armitage test(e) | P = 0.5407 | | | |
| Fisher Exact test(e) | | P = 0.2178 | P = 0.3929 | P = 0.5000 |
| SITE : thyroid TUMOR : C-cell carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 6/50(12.0) | 1/50(2.0) | 3/50(6.0) | 1/50(2.0) |
| Adjusted rates(b) | 5.13 | 2.86 | 4.88 | 0.0 |
| Terminal rates(c) | 2/39(5.1) | 1/35(2.9) | 2/41(4.9) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.8887 | | | |
| Prevalence method(d) | P = 0.8717 | | | |
| Combined analysis(d) | P = 0.9570 | | | |
| Cochran-Armitage test(e) | P = 0.0836 | | | |
| Fisher Exact test(e) | | P = 0.0559 | P = 0.2435 | P = 0.0559 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 5

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|--------------|--------------|--------------|-------------|
| SITE : thyroid TUMOR : C-cell adenoma,C-cell carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 13/50(26.0) | 12/50(24.0) | 12/50(24.0) | 7/50(14.0) |
| Adjusted rates(b) | 23.08 | 30.56 | 25.00 | 13.95 |
| Terminal rates(c) | 9/39(23.1) | 10/35(28.6) | 10/41(24.4) | 5/37(13.5) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.8887 | | | |
| Prevalence method(d) | P = 0.8525 | | | |
| Combined analysis(d) | P = 0.9287 | | | |
| Cochran-Armitage test(e) | P = 0.1335 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.5000 | P = 0.1054 |
| SITE : thyroid TUMOR : follicular adenoma,follicular adenocarcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 3/50(6.0) | 2/50(4.0) | 2/50(4.0) |
| Adjusted rates(b) | 0.0 | 7.14 | 4.88 | 5.41 |
| Terminal rates(c) | 0/39(0.0) | 2/35(5.7) | 2/41(4.9) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.2219 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.4744 | | | |
| Fisher Exact test(e) | | P = 0.1212 | P = 0.2475 | P = 0.2475 |
| SITE : adrenal gland TUMOR : pheochromocytoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 4/50(8.0) | 4/50(8.0) | 1/50(2.0) | 2/50(4.0) |
| Adjusted rates(b) | 8.70 | 10.81 | 2.44 | 5.41 |
| Terminal rates(c) | 2/39(5.1) | 3/35(8.6) | 1/41(2.4) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.8442 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.2709 | | | |
| Fisher Exact test(e) | | P = 0.6425 | P = 0.1811 | P = 0.3389 |

STUDY No. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
SEX : MALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 6

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|--------------|--------------|--------------|--------------|
| SITE : adrenal gland TUMOR : pheochromocytoma:malignant | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 1/50(2.0) | 3/50(6.0) | 0/50(0.0) |
| Adjusted rates(b) | 0.0 | 2.86 | 4.88 | 0.0 |
| Terminal rates(c) | 0/39(0.0) | 1/35(2.9) | 2/41(4.9) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.4074 | | | |
| Prevalence method(d) | P = 0.5147 | | | |
| Combined analysis(d) | P = 0.4826 | | | |
| Cochran-Armitage test(e) | P = 1.0000 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.1212 | P = N. C. |
| SITE : adrenal gland TUMOR : pheochromocytoma, pheochromocytoma:malignant | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 4/50(8.0) | 5/50(10.0) | 4/50(8.0) | 2/50(4.0) |
| Adjusted rates(b) | 8.70 | 13.51 | 7.32 | 5.41 |
| Terminal rates(c) | 2/39(5.1) | 4/35(11.4) | 3/41(7.3) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.4074 | | | |
| Prevalence method(d) | P = 0.8223 | | | |
| Combined analysis(d) | P = 0.8008 | | | |
| Cochran-Armitage test(e) | P = 0.3406 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.6425 | P = 0.3389 |
| SITE : testis TUMOR : interstitial cell tumor | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 42/50(84.0) | 44/50(88.0) | 46/50(92.0) | 43/50(86.0) |
| Adjusted rates(b) | 94.87 | 97.37 | 97.73 | 97.50 |
| Terminal rates(c) | 37/39(94.9) | 34/35(97.1) | 40/41(97.6) | 36/37(97.3) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ——— | | | |
| Prevalence method(d) | P = 0.2809 | | | |
| Combined analysis(d) | P = ——— | | | |
| Cochran-Armitage test(e) | P = 0.8003 | | | |
| Fisher Exact test(e) | | P = 0.3871 | P = 0.1783 | P = 0.5000 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 7

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|------------|------------|------------|------------|
| SITE : preputial/clitoral gland TUMOR : adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 1/50(2.0) | 2/50(4.0) | 2/50(4.0) | 3/50(6.0) |
| Adjusted rates(b) | 2.38 | 2.86 | 4.88 | 5.41 |
| Terminal rates(c) | 0/39(0.0) | 1/35(2.9) | 2/41(4.9) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.2456 | | | |
| Prevalence method(d) | P = 0.2389 | | | |
| Combined analysis(d) | P = 0.1663 | | | |
| Cochran-Armitage test(e) | P = 0.3291 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.5000 | P = 0.3087 |
| SITE : brain TUMOR : glioma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 0/50(0.0) | 3/50(6.0) | 1/50(2.0) | 1/50(2.0) |
| Adjusted rates(b) | 0.0 | 8.57 | 0.0 | 0.0 |
| Terminal rates(c) | 0/39(0.0) | 3/35(8.6) | 0/41(0.0) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.1385 | | | |
| Prevalence method(d) | P = 0.7956 | | | |
| Combined analysis(d) | P = 0.4569 | | | |
| Cochran-Armitage test(e) | P = 0.9390 | | | |
| Fisher Exact test(e) | | P = 0.1212 | P = 0.5000 | P = 0.5000 |

(HPT360A)

BAIS4

- (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.

TABLE O2

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

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 8

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|--------------|--------------|--------------|-------------|
| SITE : lung | | | | |
| TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 1/50(2.0) | 0/50(0.0) | 1/50(2.0) | 3/50(6.0) |
| Adjusted rates(b) | 2.33 | 0.0 | 2.44 | 8.11 |
| Terminal rates(c) | 0/38(0.0) | 0/43(0.0) | 1/41(2.4) | 3/37(8.1) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.0595 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.1079 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.7525 | P = 0.3087 |
| SITE : spleen | | | | |
| TUMOR : mononuclear cell leukemia | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 5/50(10.0) | 5/50(10.0) | 6/50(12.0) | 1/50(2.0) |
| Adjusted rates(b) | 7.89 | 6.98 | 7.32 | 0.0 |
| Terminal rates(c) | 3/38(7.9) | 3/43(7.0) | 3/41(7.3) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.6530 | | | |
| Prevalence method(d) | P = 0.9415 | | | |
| Combined analysis(d) | P = 0.9206 | | | |
| Cochran-Armitage test(e) | P = 0.1336 | | | |
| Fisher Exact test(e) | | P = 0.6297 | P = 0.5000 | P = 0.1022 |
| SITE : pituitary gland | | | | |
| TUMOR : adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 13/50(26.0) | 14/50(28.0) | 13/50(26.0) | 7/50(14.0) |
| Adjusted rates(b) | 23.68 | 27.91 | 26.83 | 16.22 |
| Terminal rates(c) | 9/38(23.7) | 12/43(27.9) | 11/41(26.8) | 6/37(16.2) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.9002 | | | |
| Prevalence method(d) | P = 0.8232 | | | |
| Combined analysis(d) | P = 0.9256 | | | |
| Cochran-Armitage test(e) | P = 0.1081 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.5900 | P = 0.1054 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 9

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|--------------|--------------|--------------|-------------|
| SITE : pituitary gland TUMOR : adenoma, adenocarcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 14/50(28.0) | 14/50(28.0) | 13/50(26.0) | 8/50(16.0) |
| Adjusted rates(b) | 26.32 | 27.91 | 26.83 | 16.22 |
| Terminal rates(c) | 10/38(26.3) | 12/43(27.9) | 11/41(26.8) | 6/37(16.2) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.7569 | | | |
| Prevalence method(d) | P = 0.8733 | | | |
| Combined analysis(d) | P = 0.9123 | | | |
| Cochran-Armitage test(e) | P = 0.1264 | | | |
| Fisher Exact test(e) | | P = 0.5880 | P = 0.5000 | P = 0.1135 |
| SITE : thyroid TUMOR : C-cell adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 6/50(12.0) | 6/50(12.0) | 2/50(4.0) | 3/50(6.0) |
| Adjusted rates(b) | 13.16 | 13.33 | 4.88 | 8.11 |
| Terminal rates(c) | 5/38(13.2) | 5/43(11.6) | 2/41(4.9) | 3/37(8.1) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.8950 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.1840 | | | |
| Fisher Exact test(e) | | P = 0.6202 | P = 0.1343 | P = 0.2435 |
| SITE : thyroid TUMOR : C-cell adenoma, C-cell carcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 7/50(14.0) | 6/50(12.0) | 2/50(4.0) | 4/50(8.0) |
| Adjusted rates(b) | 15.79 | 13.33 | 4.88 | 10.81 |
| Terminal rates(c) | 6/38(15.8) | 5/43(11.6) | 2/41(4.9) | 4/37(10.8) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.8633 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.2372 | | | |
| Fisher Exact test(e) | | P = 0.5000 | P = 0.0798 | P = 0.2623 |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 10

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|--|-------------|-------------|-------------|-------------|
| SITE : uterus TUMOR : endometrial stromal polyp | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 5/50(10.0) | 3/50(6.0) | 5/50(10.0) | 3/50(6.0) |
| Adjusted rates(b) | 13.16 | 6.98 | 11.90 | 8.11 |
| Terminal rates(c) | 5/38(13.2) | 3/43(7.0) | 4/41(9.8) | 3/37(8.1) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = ----- | | | |
| Prevalence method(d) | P = 0.6791 | | | |
| Combined analysis(d) | P = ----- | | | |
| Cochran-Armitage test(e) | P = 0.5970 | | | |
| Fisher Exact test(e) | | P = 0.3575 | P = 0.6297 | P = 0.3575 |
| SITE : uterus TUMOR : endometrial stromal sarcoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 3/50(6.0) | 1/50(2.0) | 1/50(2.0) | 3/50(6.0) |
| Adjusted rates(b) | 0.0 | 0.0 | 2.27 | 2.13 |
| Terminal rates(c) | 0/38(0.0) | 0/43(0.0) | 0/41(0.0) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.6383 | | | |
| Prevalence method(d) | P = 0.1356 | | | |
| Combined analysis(d) | P = 0.3917 | | | |
| Cochran-Armitage test(e) | P = 0.8073 | | | |
| Fisher Exact test(e) | | P = 0.3087 | P = 0.3087 | P = 0.6611 |
| SITE : mammary gland TUMOR : fibroadenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 8/50(16.0) | 5/50(10.0) | 5/50(10.0) | 1/50(2.0) |
| Adjusted rates(b) | 18.42 | 11.63 | 10.00 | 0.0 |
| Terminal rates(c) | 7/38(18.4) | 5/43(11.6) | 3/41(7.3) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.3865 | | | |
| Prevalence method(d) | P = 0.9958 | | | |
| Combined analysis(d) | P = 0.9907 | | | |
| Cochran-Armitage test(e) | P = 0.0202* | | | |
| Fisher Exact test(e) | | P = 0.2768 | P = 0.2768 | P = 0.0154* |

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 11

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|---|-------------|-------------|-------------|-------------|
| SITE : mammary gland TUMOR : adenoma, fibroadenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 8/50(16.0) | 5/50(10.0) | 6/50(12.0) | 1/50(2.0) |
| Adjusted rates(b) | 18.42 | 11.63 | 12.00 | 0.0 |
| Terminal rates(c) | 7/38(18.4) | 5/43(11.6) | 4/41(9.8) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.3865 | | | |
| Prevalence method(d) | P = 0.9938 | | | |
| Combined analysis(d) | P = 0.9877 | | | |
| Cochran-Armitage test(e) | P = 0.0257* | | | |
| Fisher Exact test(e) | | P = 0.2768 | P = 0.3871 | P = 0.0154* |
| SITE : mammary gland TUMOR : adenoma, fibroadenoma, adenocarcinoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 8/50(16.0) | 5/50(10.0) | 6/50(12.0) | 2/50(4.0) |
| Adjusted rates(b) | 18.42 | 11.63 | 12.00 | 0.0 |
| Terminal rates(c) | 7/38(18.4) | 5/43(11.6) | 4/41(9.8) | 0/37(0.0) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.1467 | | | |
| Prevalence method(d) | P = 0.9933 | | | |
| Combined analysis(d) | P = 0.9607 | | | |
| Cochran-Armitage test(e) | P = 0.0669 | | | |
| Fisher Exact test(e) | | P = 0.2768 | P = 0.3871 | P = 0.0458* |

(HPT360A)

BAIS4

STUDY No. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS—INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 12

| Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|---------------------------------|------------|------------|-------------|------------|
| SITE : preputial/clitoral gland | | | | |
| TUMOR : adenoma | | | | |
| Tumor rate | | | | |
| Overall rates(a) | 2/50(4.0) | 4/50(8.0) | 6/50(12.0) | 3/50(6.0) |
| Adjusted rates(b) | 5.26 | 9.09 | 9.76 | 5.41 |
| Terminal rates(c) | 2/38(5.3) | 3/43(7.0) | 4/41(9.8) | 2/37(5.4) |
| Statistical analysis | | | | |
| Peto test | | | | |
| Standard method(d) | P = 0.1474 | | | |
| Prevalence method(d) | P = 0.5361 | | | |
| Combined analysis(d) | P = 0.3365 | | | |
| Cochran-Armitage test(e) | P = 0.7508 | | | |
| Fisher Exact test(e) | | P = 0.3389 | P = 0.1343 | P = 0.5000 |

(HPT360A)

BAIS4

- (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 P1

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR :

MALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

| Group Name No. of Animals on Study | | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|---------------------------------------|----------------------------------|---------------|---------------|---------------|---------------|
| Organ | Findings | | | | |
| {Respiratory system} | | | | | |
| larynx | | <50> | <50> | <50> | <50> |
| | metastasis:thyroid tumor | 1 | 0 | 1 | 0 |
| trachea | | <50> | <50> | <50> | <50> |
| | metastasis:thyroid tumor | 1 | 0 | 1 | 0 |
| lung | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 4 | 3 | 2 | 1 |
| | metastasis:subcutis tumor | 0 | 1 | 0 | 0 |
| | metastasis:mammary gland tumor | 1 | 0 | 0 | 0 |
| | metastasis:zympal gland tumor | 0 | 1 | 0 | 0 |
| | metastasis:retroperitoneum tumor | 0 | 0 | 0 | 1 |
| {Hematopoietic system} | | | | | |
| bone marrow | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 4 | 3 | 1 | 1 |
| | metastasis:retroperitoneum tumor | 0 | 0 | 0 | 1 |
| lymph node | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 1 | 2 | 1 | 0 |
| | metastasis:subcutis tumor | 0 | 1 | 0 | 0 |
| {Circulatory system} | | | | | |
| heart | | <50> | <50> | <50> | <50> |
| | metastasis:subcutis tumor | 0 | 1 | 0 | 0 |

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

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

| Organ | Findings | Group Name No. of Animals on Study | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|--------------------|--|---------------------------------------|---------------|---------------|---------------|---------------|
| {Digestive system} | | | | | | |
| esophagus | | | <50> | <50> | <50> | <50> |
| | metastasis:thyroid tumor | | 0 | 0 | 1 | 0 |
| liver | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 5 | 3 | 2 | 1 |
| | metastasis:subcutis tumor | | 0 | 1 | 0 | 0 |
| | metastasis:retroperitoneum tumor | | 0 | 0 | 0 | 1 |
| pancreas | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 0 | 1 | 0 | 0 |
| | metastasis:retroperitoneum tumor | | 0 | 0 | 0 | 1 |
| {Urinary system} | | | | | | |
| kidney | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 1 | 1 | 0 | 0 |
| | metastasis:subcutis tumor | | 0 | 1 | 0 | 0 |
| | metastasis:retroperitoneum tumor | | 0 | 0 | 0 | 1 |
| {Endocrine system} | | | | | | |
| parathyroid | | | <50> | <50> | <50> | <50> |
| | metastasis:thyroid tumor | | 0 | 0 | 1 | 0 |
| adrenal | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 1 | 0 | 0 | 0 |
| {Nervous system} | | | | | | |
| brain | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 1 | 1 | 1 | 1 |
| < a > | a : Number of animals examined at the site | | | | | |
| b | b : Number of animals with lesion | | | | | |

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 3

| Group Name No. of Animals on Study | | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|---------------------------------------|----------------------------------|--|---------------|---------------|---------------|
| Organ | Findings | | | | |
| {Nervous system} | | | | | |
| brain | metastasis:bone tumor | <50> 0 | <50> 1 | <50> 0 | <50> 0 |
| | metastasis:zymbal gland tumor | 0 | 1 | 0 | 0 |
| spinal cord | leukemic cell infiltration | <50> 1 | <50> 0 | <50> 0 | <50> 0 |
| {Musculoskeletal system} | | | | | |
| muscle | leukemic cell infiltration | <50> 1 | <50> 0 | <50> 0 | <50> 0 |
| | metastasis:subcutis tumor | <50> 0 | <50> 1 | <50> 0 | <50> 0 |
| {Body cavities} | | | | | |
| pleura | metastasis:retroperitoneum tumor | <50> 0 | <50> 0 | <50> 0 | <50> 1 |
| | metastasis:retroperitoneum tumor | <50> 0 | <50> 0 | <50> 0 | <50> 1 |
| < a > | | a : Number of animals examined at the site | | | |
| b | | b : Number of animals with lesion | | | |

(JPT150)

BAIS4

TABLE P2

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR :

FEMALE

STUDY NO. : 0560
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 4

| Group Name No. of Animals on Study | | Control 50 | 125 ppm 50 | 250 ppm 50 | 500 ppm 50 |
|---|----------------------------|---------------|---------------|---------------|---------------|
| Organ | Findings | | | | |
| {Respiratory system} | | | | | |
| lung | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 3 | 3 | 6 | 1 |
| | metastasis:uterus tumor | 0 | 1 | 0 | 0 |
| | metastasis:vertebra tumor | 0 | 0 | 0 | 1 |
| {Hematopoietic system} | | | | | |
| bone marrow | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 2 | 2 | 1 | 1 |
| lymph node | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 1 | 1 | 2 | 0 |
| {Digestive system} | | | | | |
| liver | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 2 | 3 | 6 | 1 |
| pancreas | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 0 | 1 | 0 | 0 |
| {Urinary system} | | | | | |
| kidney | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | 1 | 0 | 2 | 0 |
| urin bladd | | <50> | <50> | <50> | <50> |
| | metastasis:uterus tumor | 0 | 1 | 0 | 0 |
| {Reproductive system} | | | | | |
| ovary | | <50> | <50> | <50> | <50> |
| | metastasis:uterus tumor | 0 | 0 | 0 | 1 |
| < a > a : Number of animals examined at the site b b : Number of animals with lesion | | | | | |

STUDY NO. : 0560
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 5

| | | Group Name | Control | 125 ppm | 250 ppm | 500 ppm |
|------------------|-------------------------------|-------------------------|---------|---------|---------|---------|
| | | No. of Animals on Study | 50 | 50 | 50 | 50 |
| Organ | Findings | | | | | |
| {Nervous system} | | | | | | |
| brain | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 1 | 0 | 0 | 0 |
| | metastasis:pituitary tumor | | 1 | 0 | 0 | 1 |
| | metastasis:zymbal gland tumor | | 0 | 0 | 0 | 1 |
| spinal cord | | | <50> | <50> | <50> | <50> |
| | leukemic cell infiltration | | 1 | 0 | 0 | 0 |
| {Body cavities} | | | | | | |
| peritoneum | | | <50> | <50> | <50> | <50> |
| | metastasis:uterus tumor | | 1 | 0 | 0 | 0 |

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

(JPT150)

BAIS4

TABLE Q

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC
LESIONS IN JAPAN BIOASSAY RESEARCH CENTER :
F344/DuCr1Cr1j MALE RATS

TABLE Q HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS
IN JAPAN BIOASSAY RESEARCH CENTER : F344/DuCr1Cr1j MALE RATS

| Organs Tumors | No. of animals examined | No. of animals bearing tumor | Incidence (%) | Min. - Max. (%) |
|--------------------------------|----------------------------|---------------------------------|------------------|--------------------|
| Lung | 2199 | | | |
| Bronchio-alveolar adenoma 1) | | 84 | 3.8 | 0 - 12 |
| Bronchio-alveolar carcinoma 2) | | 19 | 0.9 | 0 - 8 |
| 1)+2) | | 103 | 4.7 | 0 - 14 |
| Liver | 2199 | | | |
| Hepatocellular adenoma 1) | | 41 | 1.9 | 0 - 8 |
| Hepatocellular carcinoma 2) | | 7 | 0.3 | 0 - 2 |
| 1)+2) | | 47 | 2.1 | 0 - 8 |
| Pncreas | 2199 | | | |
| Islet cell adenoma | | 46 | 2.1 | 0 - 14 |

Forty four carcinogenicity studies examined in Japan Bioassay Research Center were used.

Study No. : 0043, 0059, 0061, 0063, 0065, 0067, 0095, 0104, 0115, 0130, 0141, 0158, 0162, 0189,
0205, 0210, 0224, 0242, 0246, 0267, 0269, 0278, 0284, 0288, 0294, 0296, 0318, 0328,
0342, 0347, 0365, 0371, 0396, 0399, 0401, 0407, 0417, 0421, 0437, 0448, 0457, 0461,
0497, 0535

TABLE R

CAUSE OF DEATH OF RATS IN THE 2-YEAR
INHALATION STUDY OF 1 - BROMOBUTANE

TABLE R CAUSE OF DEATH OF RATS IN THE 2-YEAR INHALATION STUDY
OF 1-BROMOBUTANE

| Group name | Male | | | | Female | | | |
|------------------------------------|---------|---------|---------|---------|---------|---------|---------|---------|
| | Control | 125 ppm | 250 ppm | 500 ppm | Control | 125 ppm | 250 ppm | 500 ppm |
| Number of dead or moribund animals | 11 | 15 | 9 | 13 | 12 | 7 | 9 | 13 |
| Urinary system lesion | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 |
| Chronic nephropathy | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 |
| Peritonitis | 0 | 0 | 0 | 0 | 0 | 1 | 0 | 0 |
| Tumor death : leukemia | 2 | 2 | 1 | 1 | 2 | 2 | 3 | 1 |
| skin / appendage | 0 | 1 | 1 | 0 | 0 | 0 | 0 | 0 |
| subcutis | 0 | 1 | 1 | 1 | 0 | 0 | 0 | 0 |
| lymph node | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 |
| spleen | 0 | 0 | 0 | 1 | 0 | 0 | 0 | 0 |
| kidney | 0 | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| urinary bladder | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| pituitary gland | 5 | 3 | 0 | 2 | 4 | 2 | 2 | 2 |
| thyroid | 4 | 0 | 1 | 1 | 0 | 0 | 0 | 0 |
| adrenal gland | 0 | 0 | 1 | 0 | 0 | 0 | 0 | 0 |
| uterus | — | — | — | — | 3 | 1 | 0 | 2 |
| mammary gland | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 2 |
| preputial/clitoral gland | 0 | 1 | 0 | 1 | 0 | 0 | 2 | 1 |
| brain | 0 | 0 | 2 | 1 | 0 | 0 | 0 | 0 |
| Zymbal gland | 0 | 1 | 1 | 2 | 0 | 0 | 1 | 2 |
| vertebra | — | — | — | — | 0 | 0 | 0 | 1 |
| bone | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 |
| No microscopical confirmation | 0 | 1 | 1 | 2 | 1 | 1 | 1 | 1 |