

プロピオノニトリルのラットを用いた
吸入によるがん原性試験報告書

試験番号：0535

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APPENDIX A 1

IDENTITY OF PROPIONONITRILE IN THE 2-YEAR INHALATION STUDY

IDENTITY OF PROPIONONITRILE IN THE 2-YEAR INHALATION STUDY

Test Substance : Propiononitrile (Wako Pure Chemical Industries, Ltd.)

A. Lot No. : PKK4727

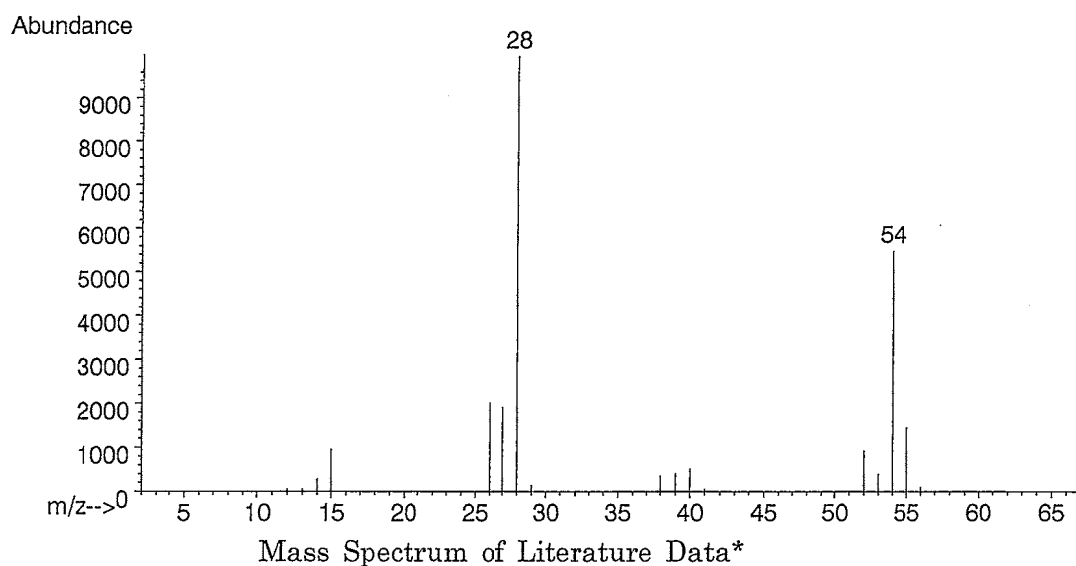
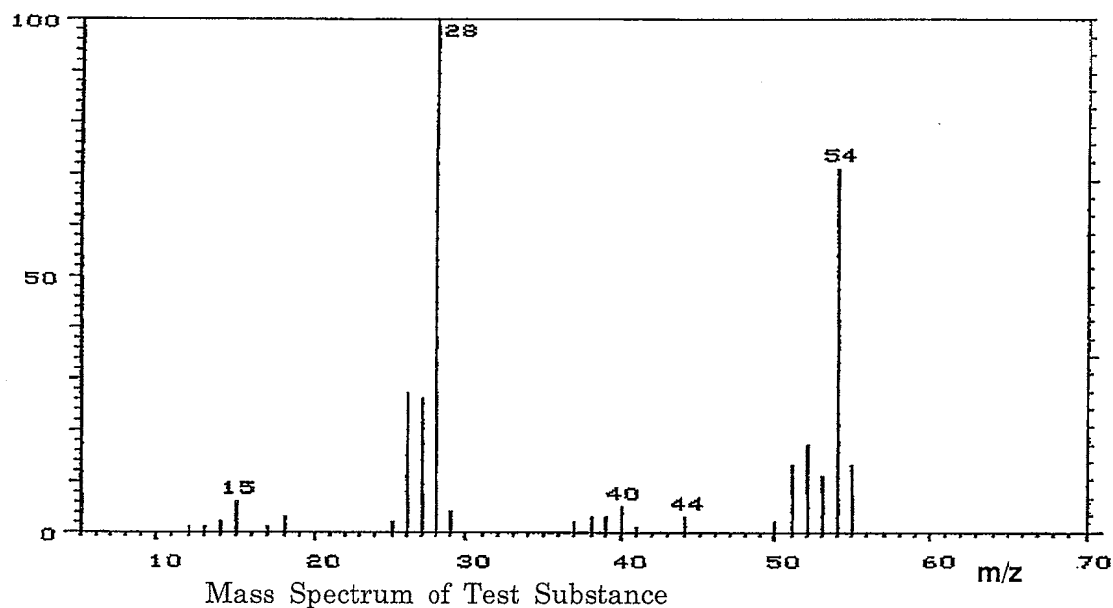
1. Spectral Data

Mass Spectrometry

Instrument : Hitachi M-80B Mass Spectrometer

Ionization : EI (Electron Ionization)

Ionization Voltage : 70eV



Result: The mass spectrum was consistent with literature spectrum.

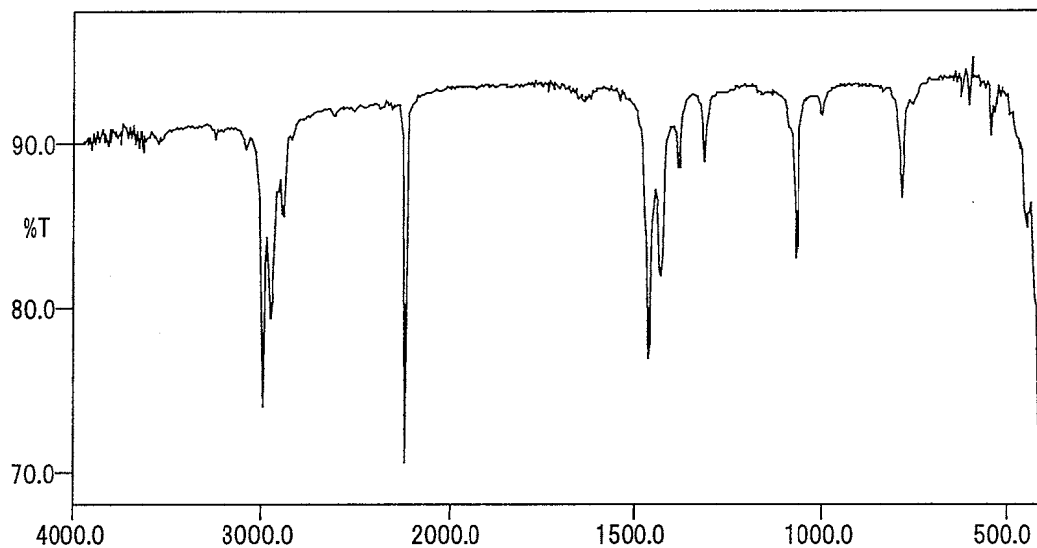
(*McLafferty FW, ed. 1994. Wiley Registry of Mass Spectral Data. 6th ed.
New York, NY: John Wiley and Sons.)

Infrared Spectrometry

Instrument : Shimadzu FTIR-8200PC Infrared Spectrometer

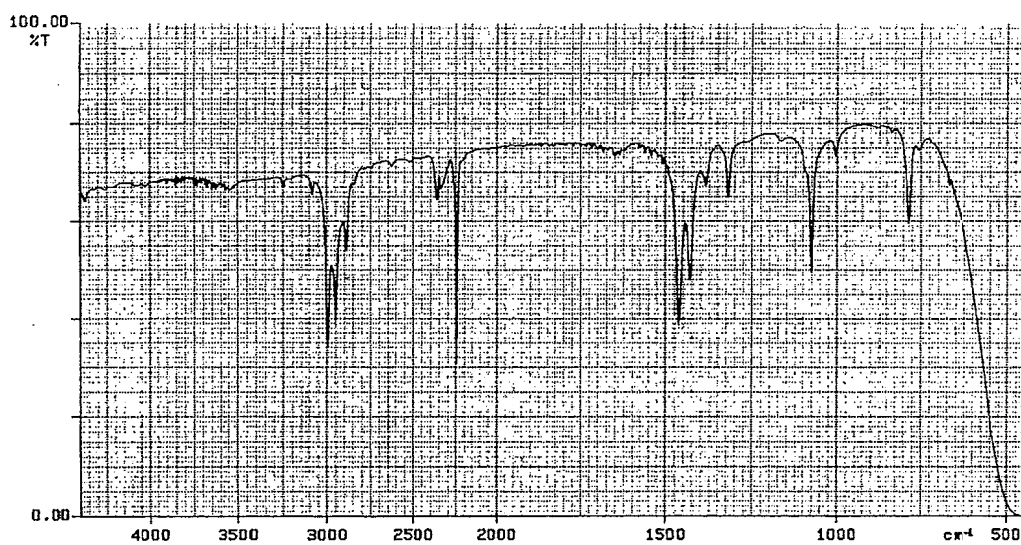
Cell : KBr Liquid Cell

Resolution : 4 cm^{-1}



Infrared Spectrum of Test Substance

Wavenumbers cm^{-1}



Infrared Spectrum of Literature Data*

Wavenumbers

Result: The infrared spectrum was consistent with literature spectrum.

(*Performed by Wako Pure Chemical Industries, Ltd.)

2. Conclusion: The test substance was identified as propiononitrile by mass spectrum and infrared spectrum.

B. Lot No. : CEL7045

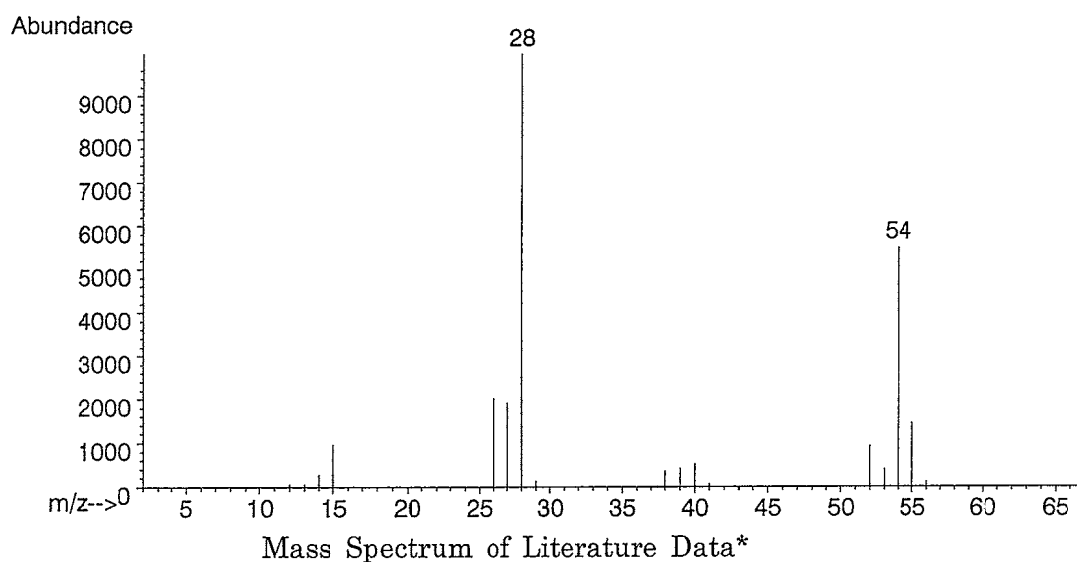
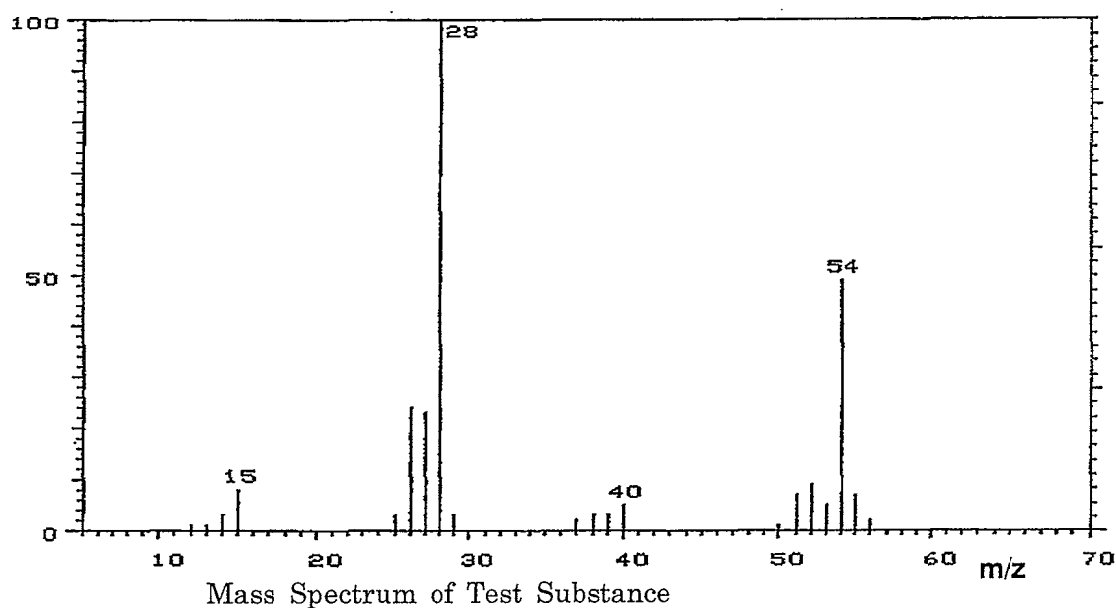
1. Spectral Data

Mass Spectrometry

Instrument : Hitachi M-80B Mass Spectrometer

Ionization : EI (Electron Ionization)

Ionization Voltage : 70eV



Result: The mass spectrum was consistent with literature spectrum.

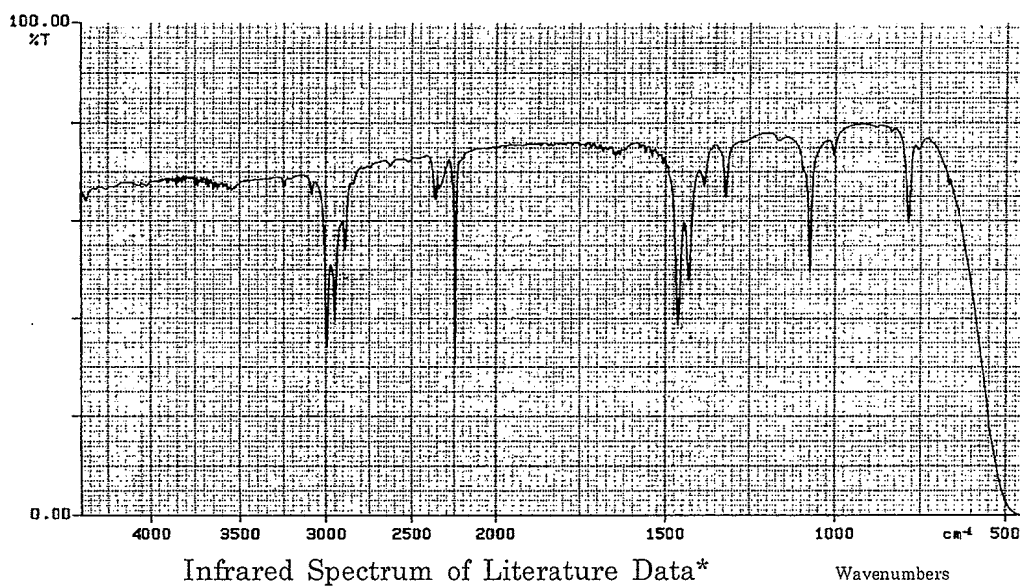
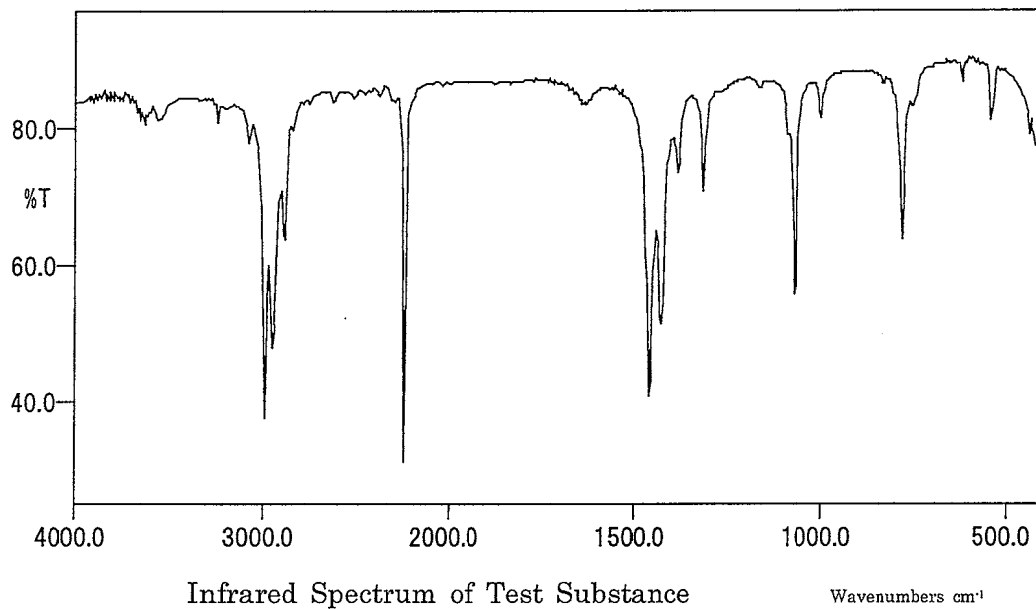
(*McLafferty FW, ed. 1994. Wiley Registry of Mass Spectral Data. 6th ed.
New York, NY: John Wiley and Sons.)

Infrared Spectrometry

Instrument : Shimadzu FTIR-8200PC Infrared Spectrometer

Cell : KBr Liquid Cell

Resolution : 4 cm^{-1}



Result: The infrared spectrum was consistent with literature spectrum.

(*Performed by Wako Pure Chemical Industries, Ltd.)

2. Conclusion: The test substance was identified as propiononitrile by mass spectrum and infrared spectrum.

C. Lot No. : SDM0881

1. Spectral Data

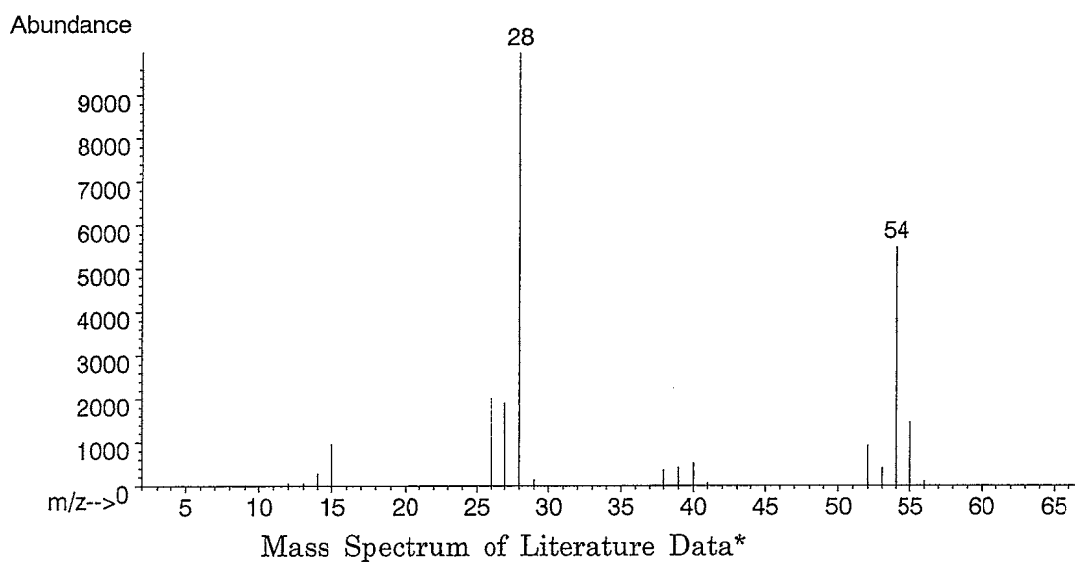
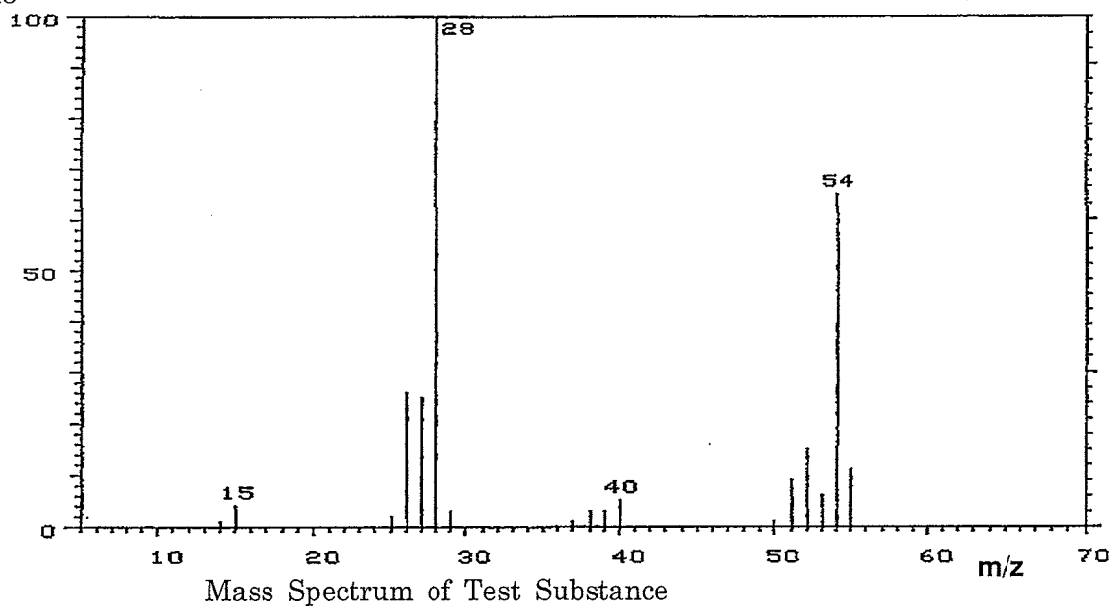
Mass Spectrometry

Instrument : Hitachi M-80B Mass Spectrometer

Ionization : EI (Electron Ionization)

Ionization Voltage : 70eV

75 148



Result: The mass spectrum was consistent with literature spectrum.

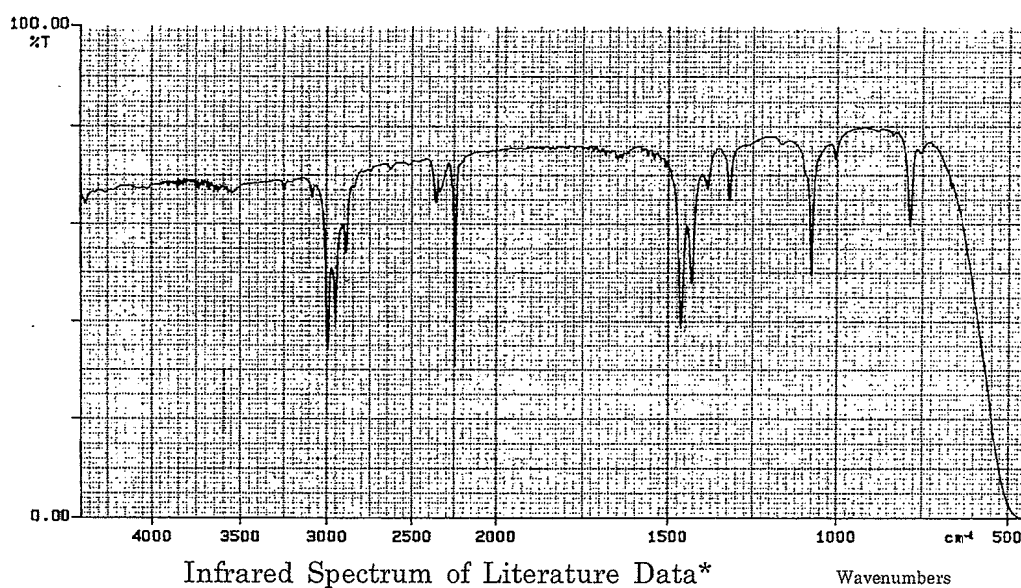
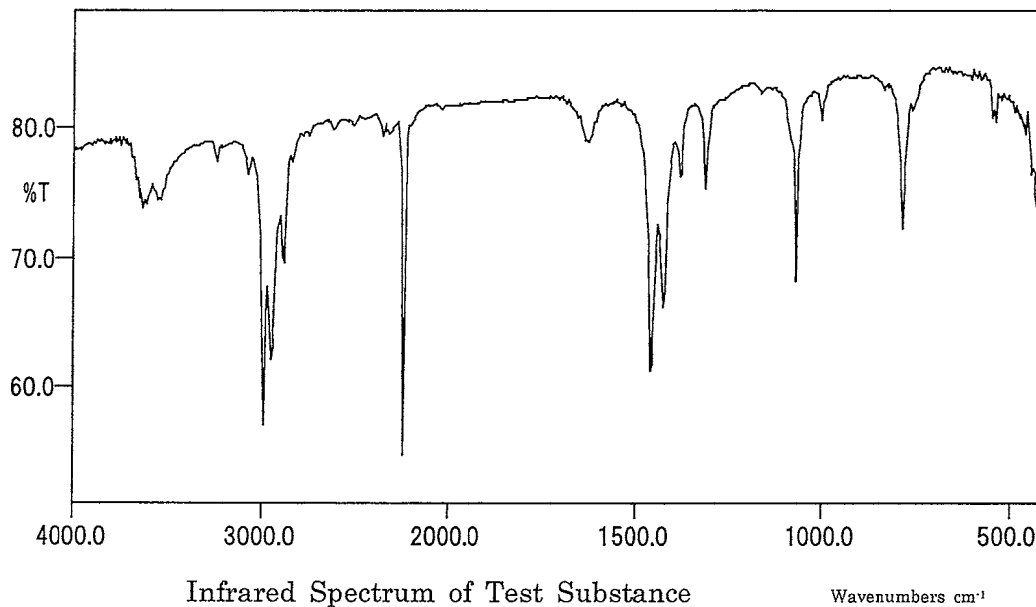
(*McLafferty FW, ed. 1994. Wiley Registry of Mass Spectral Data. 6th ed.
New York, NY: John Wiley and Sons.)

Infrared Spectrometry

Instrument : Shimadzu FTIR-8200PC Infrared Spectrometer

Cell : KBr Liquid Cell

Resolution : 4 cm^{-1}



Result: The infrared spectrum was consistent with literature spectrum.

(*Performed by Wako Pure Chemical Industries, Ltd.)

2. Conclusion: The test substance was identified as propiononitrile by mass spectrum and infrared spectrum.

APPENDIX A 2

STABILITY OF PROPIONONITRILE IN THE 2-YEAR INHALATION STUDY

STABILITY OF PROPIONONITRILE IN THE 2-YEAR INHALATION STUDY

Test Substance : Propiononitrile (Wako Pure Chemical Industries, Ltd.)

A. Lot No. : PKK4727

1. Gas Chromatography

Instrument : Hewlett Packard 5890A Gas Chromatograph

Column : Methyl Silicone (0.53 mm ϕ \times 60 m)

Column Temperature : 80° C

Flow Rate : 10 mL/min

Detector : FID (Flame Ionization Detector)

Injection Volume : 1 μ L

Date Analyzed	Peak No. (min)	Retention Time (%)	Area
2003.11.28	1	3.770	100
2004.11.10	1	3.768	100

Result: Gas chromatography indicated one major peak (peak No.1) analyzed on 2003.11.28 and one major peak (peak No.1) analyzed on 2004.11.10. No new trace impurity peak in the test substance analyzed on 2004.11.10 was detected.

2. Conclusion: The test substance was stable for the period that the test substance had been used for the study.

B. Lot No. : CEL7045

1. Gas Chromatography

Instrument : Hewlett Packard 5890A Gas Chromatograph

Column : Methyl Silicone (0.53 mm ϕ \times 60 m)

Column Temperature : 80° C

Flow Rate : 10 mL/min

Detector : FID (Flame Ionization Detector)

Injection Volume : 1 μ L

Date Analyzed	Peak No. (min)	Retention Time (%)	Area
2004.11.05	1	3.768	100
2005.06.24	1	3.783	100

Result: Gas chromatography indicated one major peak (peak No.1) analyzed on 2004.11.5 and one major peak (peak No.1) analyzed on 2005.6.24. No new trace impurity peak in the test substance analyzed on 2005.6.24 was detected.

2. Conclusion: The test substance was stable for the period that the test substance had been used for the study.

C. Lot No. : SDM0881

1. Gas Chromatography

Instrument : Hewlett Packard 5890A Gas Chromatograph

Column : Methyl Silicone (0.53 mm ϕ \times 60 m)

Column Temperature : 80° C

Flow Rate : 10 mL/min

Detector : FID (Flame Ionization Detector)

Injection Volume : 1 μ L

Date Analyzed	Peak No. (min)	Retention Time (%)	Area
2005.06.17	1	3.771	100
2005.12.05	1	3.769	100

Result: Gas chromatography indicated one major peak (peak No.1) analyzed on 2005.6.17 and one major peak (peak No.1) analyzed on 2005.12.5. No new trace impurity peak in the test substance analyzed on 2005.12.5 was detected.

2. Conclusion: The test substance was stable for the period that the test substance had been used for the study.

APPENDIX B

ENVIRONMENTAL CONDITIONS OF INHALATION CHAMBER IN THE 2-YEAR INHALATION STUDY OF PROPIONONITRILE

ENVIRONMENTAL CONDITIONS OF INHALATION CHAMBER IN THE 2-YEAR
INHALATION STUDY OF PROPIONONITRILE

Group Name	Temperature (°C) Mean ± S.D.	Humidity (%) Mean ± S.D.	Ventilation Rate (L/min) Mean ± S.D.	Air Change (time/h) Mean
Control	23.1 ± 0.2	57.0 ± 1.0	1520.9 ± 12.2	12.0
25 ppm	23.1 ± 0.1	55.4 ± 1.3	1521.3 ± 12.6	12.0
50 ppm	23.2 ± 0.2	55.3 ± 1.3	1516.2 ± 11.7	12.0
100 ppm	23.1 ± 0.2	57.3 ± 0.9	1515.4 ± 11.7	12.0

APPENDIX C 1

CLINICAL OBSERVATION : MALE

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 1

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REPORT TYPE : A1 104

ALL ANIMALS

PAGE : 2

[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 3

[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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Clinical sign	Group Name	Administration Week-day				46-7	47-7	48-7	49-7	50-7	51-7	52-7	53-7	54-7	55-7	56-7
		43-7	44-7	45-7												
DEATH	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	1	1	1	1	1	1	1	1	1	1	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	1	1	1	1	1	1	1	1
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ATAXIC GAIT	Control	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
PARALYTIC GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
TUMBLE	Control	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ABNORMAL GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
WASTING	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
PILOERECTION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

PAGE : 6

[illegible]

STUDY NO. : 0535
 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
DEATH	Control	2	2	2	2	2	2	2	2	2	2	2	2	3	3
	25 ppm	1	1	1	1	1	1	1	2	2	2	3	4	4	4
	50 ppm	3	3	3	3	3	3	3	3	4	4	4	4	4	4
	100 ppm	1	1	3	3	4	4	4	4	4	4	4	5	6	8
MORIBUND SACRIFICE	Control	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	25 ppm	1	1	1	1	1	1	1	1	1	1	2	2	2	2
	50 ppm	3	3	3	3	3	3	3	3	3	3	3	3	3	3
	100 ppm	1	1	1	1	1	1	1	2	3	3	3	3	3	3
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
ATAXIC GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
PARALYTIC GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	1	1	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
TUMBLE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ABNORMAL GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	1	1	1	1	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
WASTING	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	1	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	1	0	0	0	0	0	0
PILORECTION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	1	0	0	0	0	0	0	0

STUDY NO. : 0535
 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
DEATH	Control	3	4	4	4	4	4
	25 ppm	4	4	4	4	4	4
	50 ppm	4	4	4	4	5	5
	100 ppm	9	9	9	9	9	10
MORIBUND SACRIFICE	Control	1	1	1	1	2	2
	25 ppm	2	2	2	2	2	2
	50 ppm	3	3	3	3	3	3
	100 ppm	4	4	4	4	4	4
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ATAXIC GAIT	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
PARALYTIC GAIT	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
TUMBLE	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ABNORMAL GAIT	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	1	1	1	0	0
	100 ppm	0	0	0	0	0	0
WASTING	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	1	1	2
	100 ppm	0	0	0	0	0	0
PILORECTION	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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ALL ANIMALS

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CLINICAL OBSERVATION (SUMMARY)
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]

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1 104

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

PAGE : 15

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
TRAUMA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	1	1	1	1	1	1	1	1	1	1	0
FROG BELLY	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	1	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
SOILED PERI-GENITALIA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
EXOPHTHALMOS	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	100 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	1	1	1	1	0	0
	25 ppm	2	3	3	3	3	3	3	3	3	3	3	3	3	3
	50 ppm	2	2	2	2	3	3	3	3	3	3	3	3	3	3
	100 ppm	1	1	1	1	2	2	2	2	2	2	2	2	2	2
CORNEAL OPACITY	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
ANTERIOR CHAMBER OPACITY	Control	0	0	0	0	0	0	1	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	1	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	1	1	1	1	1	1	1	1	1	1	0
EXTERNAL MASS	Control	3	3	3	4	6	6	7	6	7	7	7	7	6	6
	25 ppm	3	4	5	6	6	6	7	7	8	8	9	8	8	9
	50 ppm	0	2	2	4	4	4	4	5	5	6	5	5	5	5
	100 ppm	8	8	7	9	9	9	8	9	8	10	10	11	11	10
INTERNAL MASS	Control	0	0	0	0	0	0	0	0	0	1	1	1	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	1	1	1	2	1	1

STUDY NO. : 0535
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
TRAUMA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
FROG BELLY	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	1	1	0	0
	100 ppm	0	0	0	0	0	0
SOILED PERI-GENITALIA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
EXOPHTHALMOS	Control	0	0	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	1	1	1	1	1	1
	100 ppm	0	0	0	0	0	0
CATARACT	Control	0	0	1	1	1	1
	25 ppm	3	3	3	3	3	4
	50 ppm	3	3	3	3	3	3
	100 ppm	1	1	1	1	1	1
CORNEAL OPACITY	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ANTERIOR CHAMBER OPACITY	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
EXTERNAL MASS	Control	6	7	8	9	9	11
	25 ppm	9	8	9	9	9	10
	50 ppm	5	6	7	7	7	7
	100 ppm	9	10	11	11	13	12
INTERNAL MASS	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

CLINICAL OBSERVATION (SUMMARY)
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[illegible]

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STUDY NO. : 0535
 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													
		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
M. NOSE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M. PERI-MOUTH	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	1	1	1	1	1	1	1	1	1	1	1	1	1	1
M. EAR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M. FORELIMB	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M. ABDOMEN	Control	0	0	0	0	0	0	0	0	0	0	0	1	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	1	1	1	1	1	1	1

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[illegible]

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[illegible]

STUDY NO. : 0535
 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. NOSE	Control	0	0	0	0	0	0
	25 ppm	1	1	1	1	1	1
	50 ppm	0	0	0	0	0	0
	100 ppm	1	1	1	1	1	1
M. PERI-MOUTH	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. MANDIBULAR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	1	1	1	1	1	1
M. EAR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. PERI EAR	Control	0	0	0	0	0	0
	25 ppm	1	1	1	1	1	1
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. NECK	Control	0	0	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. FORELIMB	Control	1	1	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	1	1	1	1	1	1
M. BREAST	Control	1	2	2	3	3	3
	25 ppm	2	2	3	3	3	3
	50 ppm	1	1	2	2	2	2
	100 ppm	1	1	2	2	3	2
M. ABDOMEN	Control	2	2	2	2	2	2
	25 ppm	4	4	4	4	4	3
	50 ppm	1	1	1	1	1	1
	100 ppm	3	3	3	3	4	4

CLINICAL OBSERVATION (SUMMARY)
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[illegible]

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[illegible]

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[illegible]

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[illegible]

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[illegible]

STUDY NO. : 0535
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				74-7	75-7	76-7	77-7	78-7	79-7	80-7	81-7	82-7	83-7	84-7
		71-7	72-7	73-7												
M. ANTERIOR. DORSUM	Control	1	1	2		2	2	2	2	2	2	2	2	1	1	1
	25 ppm	0	0	1		1	1	1	1	1	1	1	1	1	1	1
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0		0	1	1	1	1	1	1	1	1	1	1
M. POSTERIOR DORSUM	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	1	1	1		1	1	1	1	1	2	2	2	2	2	2
M. HINDLIMB	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
M. GENITALIA	Control	0	0	1		1	1	1	1	1	1	1	1	1	1	1
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
M. SCROTUM	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
ANEMIA	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	1	1
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	1	0	0
	100 ppm	0	0	0		0	0	0	0	0	0	1	0	0	0	0
JAUNDICE	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	1	1
	100 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
ULCER	Control	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0		0	0	0	0	0	0	0	0	0	0	0
	100 ppm	1	1	1		1	1	1	1	1	0	0	0	0	0	0

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[illegible]

STUDY NO. : 0535
 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. ANTERIOR. DORSUM	Control	1	1	1	1	1	2
	25 ppm	1	1	1	1	1	2
	50 ppm	3	3	3	3	3	3
	100 ppm	1	1	1	1	1	1
M. POSTERIOR DORSUM	Control	0	0	1	1	1	0
	25 ppm	1	0	0	0	0	0
	50 ppm	0	1	1	1	1	1
	100 ppm	1	2	2	2	2	2
M. HINDLIMB	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	1
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. GENITALIA	Control	1	1	1	1	1	2
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. SCROTUM	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ANEMIA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
JAUNDICE	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ULCER	Control	0	1	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	1	1	1	1	1	1
	100 ppm	1	1	2	2	2	2
CRUSTA	Control	1	1	1	1	1	0
	25 ppm	0	0	0	0	0	1
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

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[illegible]

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[illegible]

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STUDY NO. : 0535
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
HEMORRHAGE	Control	0	0	0	0	0	0
	25 ppm	0	1	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ABNORMAL TESTIS	Control	2	2	2	2	2	2
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
PROLAPSE OF PENIS	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
IRREGULAR BREATHING	Control	0	0	0	1	0	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
RESPIRATORY SOUND ABNOR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
TACHYPNEA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
DEEP BREATHING	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
LOOSE STOOL	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
SUBNORMAL TEMP	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

APPENDIX C 2

CLINICAL OBSERVATION : FEMALE

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[illegible]

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[illegible]

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STUDY NO. : 0535
 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
DEATH	Control	5	5	6	6	6	7
	25 ppm	1	2	2	3	3	3
	50 ppm	7	9	9	9	9	10
	100 ppm	6	6	6	6	6	6
MORIBUND SACRIFICE	Control	4	4	4	4	6	6
	25 ppm	2	2	2	2	2	2
	50 ppm	4	4	4	4	4	4
	100 ppm	3	3	4	5	5	5
LOCOMOTOR MOVEMENT DECR	Control	0	0	1	1	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
HUNCHBACK POSITION	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	1	1	1	1
ATAXIC GAIT	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
PARALYTIC GAIT	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
WASTING	Control	0	0	1	1	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	0	0	0	1
	100 ppm	1	1	2	1	1	2
SOILED	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
PILOBRECTION	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	0	0	0	0
	100 ppm	1	1	1	1	1	1

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
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[illegible]

CLINICAL OBSERVATION (SUMMARY)

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[illegible]

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

STUDY NO. : 0535
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
TRAUMA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	100 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	1	0	0	0	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	1	1	1
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EXOPHTHALMOS	Control	0	1	1	1	1	1	1	2	2	2	1	1	1	1
	25 ppm	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CATARACT	Control	3	3	3	3	3	3	3	3	3	3	3	3	3	3
	25 ppm	4	4	4	4	4	4	4	4	4	4	4	4	5	5
	50 ppm	3	3	4	4	5	5	5	5	5	5	5	5	5	5
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CONJUNCTIVAL EDEMA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANTERIOR CHAMBER OPACITY	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CORNEAL EDEMA	Control	0	0	0	0	0	0	0	0	0	1	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EXTERNAL MASS	Control	0	0	0	0	0	0	0	0	0	0	0	1	1	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	1	1	2	1	1	2	2	1	1
	100 ppm	0	0	0	0	0	1	1	2	2	2	2	2	2	2

STUDY NO. : 0535
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													
		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
TRAUMA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 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
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	1	1	1	1	1	2	2	1	1	1	1	0	1	1
	100 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	1	0	0	0	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	1	0	0	0	1	1	0	0	0	0	0	0	1
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	1	0
EXOPHTHALMOS	Control	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	25 ppm	1	1	1	1	1	1	1	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CATARACT	Control	3	3	3	3	3	3	3	3	3	3	3	3	3	3
	25 ppm	5	5	5	5	5	5	5	4	4	4	4	4	4	4
	50 ppm	5	5	5	6	6	6	6	6	6	6	6	6	6	6
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CONJUNCTIVAL EDEMA	Control	1	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANTERIOR CHAMBER OPACITY	Control	0	0	0	0	0	0	0	0	0	0	0	0	1	1
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CORNEAL EDEMA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EXTERNAL MASS	Control	2	3	3	3	3	3	3	3	3	4	3	3	3	3
	25 ppm	0	0	0	1	2	3	3	3	3	4	5	5	5	6
	50 ppm	1	1	1	1	1	1	1	0	0	1	2	2	2	2
	100 ppm	4	4	4	4	4	4	5	5	6	7	8	9	8	7

STUDY NO. : 0535
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
TRAUMA	Control	1	1	1	1	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
FROG BELLY	Control	1	1	2	2	1	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	0	1	1	1
	100 ppm	0	0	0	0	1	1
SOILED PERI-GENITALIA	Control	0	1	1	1	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	1	0	0	0
	100 ppm	0	1	1	0	0	0
EXOPHTHALMOS	Control	1	1	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
CATARACT	Control	3	3	4	4	4	4
	25 ppm	4	4	4	4	4	4
	50 ppm	6	5	5	5	5	5
	100 ppm	0	0	0	0	0	0
CONJUNCTIVAL EDEMA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
ANTERIOR CHAMBER OPACITY	Control	1	1	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
CORNEAL EDEMA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
EXTERNAL MASS	Control	3	3	4	5	5	5
	25 ppm	6	6	7	6	6	7
	50 ppm	2	2	2	2	3	4
	100 ppm	6	6	6	6	6	7

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

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[illegible]

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[illegible]

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[illegible]

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CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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STUDY NO. : 0535
 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
INTERNAL MASS	Control	2	2	2	2	1	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	1	2	2	2	2
	100 ppm	2	2	2	1	1	1
M. NOSE	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	1	1	1	1	1	1
M. EYE	Control	1	1	1	1	1	1
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. PERI-MOUTH	Control	0	0	0	1	1	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. PERI EAR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. FORELIMB	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
M. BREAST	Control	0	0	1	1	1	1
	25 ppm	3	2	3	3	3	3
	50 ppm	1	1	1	1	1	1
	100 ppm	3	3	4	4	4	4
M. ABDOMEN	Control	2	2	2	2	2	3
	25 ppm	2	3	3	2	2	3
	50 ppm	1	1	1	1	1	2
	100 ppm	1	1	1	1	1	3
M. ANTERIOR. DORSUM	Control	0	0	0	0	0	0
	25 ppm	1	1	1	1	1	1
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

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[illegible]

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[illegible]

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

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[illegible]

STUDY NO. : 0535
 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. GENITALIA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	1
	100 ppm	1	1	0	0	0	0
ANEMIA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	1
	100 ppm	1	2	1	0	0	1
JAUNDICE	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	1	1	1
	100 ppm	0	0	0	0	0	0
HEMORRHAGE	Control	0	0	1	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	1	0	0	0
	100 ppm	0	0	0	0	1	1
IRREGULAR BREATHING	Control	0	0	0	1	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	1	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
RESPIRATORY SOUND ABNOR	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
BRADYPNEA	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0
DEEP BREATHING	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	1	1	1
BROWN URINE	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

STUDY NO. : 0535
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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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													
		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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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													
		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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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													
		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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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
SUBNORMAL TEMP	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0	0	0	0	0	1	0	0	0
	100 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

(HAN190)

BAIS 4

STUDY NO. : 0535
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
SUBNORMAL TEMP	Control	0	0	0	0	0	0
	25 ppm	0	0	0	0	0	0
	50 ppm	0	0	0	0	0	0
	100 ppm	0	0	0	0	0	0

(HAN190)

BAIS 4

APPENDIX D 1

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0535
 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	122±	5	151±	6	182±	8	206±	10	225±	10	243±	12
25 ppm	122±	5	150±	7	181±	8	205±	9	226±	10	242±	11
50 ppm	122±	5	150±	7	181±	9	205±	10	225±	10	243±	11
100 ppm	121±	5	146±	9**	170±	9**	190±	11**	209±	12**	227±	13**

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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		13-7	
Control	271±	13	284±	13	294±	14	302±	15	310±	16	316±	15	321±	16		
25 ppm	268±	12	282±	14	292±	14	300±	14	307±	14	313±	15	319±	15		
50 ppm	270±	13	282±	14	292±	14	301±	14	309±	14	314±	15	321±	15		
100 ppm	250±	16**	261±	16**	270±	16**	277±	17**	286±	17**	291±	17**	297±	17**		

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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									
	17-7		21-7		25-7		29-7		33-7		37-7	
Control	341±	16	356±	18	368±	20	381±	21	390±	23	391±	24
25 ppm	334±	19	352±	18	364±	19	376±	21	386±	25	388±	23
50 ppm	340±	16	353±	16	366±	18	376±	19	386±	21	388±	20
100 ppm	315±	17**	329±	17**	342±	16**	351±	18**	359±	22**	362±	21**

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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									
	45-7		49-7		53-7		54-7		58-7		62-7	
Control	395±	25	409±	26	416±	27	416±	27	418±	29	423±	28
25 ppm	392±	25	405±	27	410±	25	411±	26	415±	26	418±	26
50 ppm	392±	21	405±	21	411±	22	412±	22	416±	22	417±	23
100 ppm	367±	21**	379±	23**	385±	24**	386±	24**	389±	24**	393±	23**
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett												

(HAN260)

BAIS 4

STUDY NO. : 0535
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	428±	29	430±	28	430±	26	428±	26	427±	26	425±	26
25 ppm	423±	27	426±	27	425±	27	422±	27	422±	33	420±	47
50 ppm	420±	21	422±	19	420±	22	415±	29	418±	19	417±	20
100 ppm	398±	26**	401±	25**	402±	24**	399±	27**	399±	27**	395±	31**

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
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	416±	27	412±	33	408±	32
25 ppm	414±	24	410±	24	406±	25
50 ppm	407±	22	400±	27	395±	28
100 ppm	383±	31**	382±	27**	377±	28**
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett						

(HAN260)

BAIS 4

APPENDIX D 2

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0535
 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	95±	3	109±	4	123±	5	132±	5	140±	6	148±	7
25 ppm	95±	3	109±	5	123±	6	133±	7	142±	7	148±	8
50 ppm	95±	3	109±	5	123±	6	132±	7	141±	8	148±	9
100 ppm	95±	3	108±	4	120±	5*	129±	6	138±	7	146±	7
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett												

(HAN260)

BAIS 4

STUDY NO. : 0535
 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	157±	9	162±	9	166±	9	170±	9	174±	9	176±	10
25 ppm	159±	9	164±	10	170±	11	173±	11	177±	11	178±	10
50 ppm	158±	10	162±	11	166±	11	169±	11	173±	11	176±	12
100 ppm	155±	8	160±	9	164±	9	167±	9	171±	10	173±	10

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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									
	17-7		21-7		25-7		29-7		33-7		37-7	
Control	186±	10	190±	11	196±	12	199±	11	200±	11	209±	12
25 ppm	190±	12	194±	12	200±	12	203±	13	202±	13	212±	14
50 ppm	188±	13	190±	13	195±	14	197±	13	197±	12	207±	13
100 ppm	183±	9	188±	10	194±	10	197±	11	195±	10	206±	10

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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									
	45-7		49-7		53-7		54-7		58-7		62-7	
Control	215±	13	218±	13	221±	14	221±	14	224±	15	228±	15
25 ppm	219±	14	222±	15	225±	16	226±	16	230±	16	236±	18*
50 ppm	212±	15	215±	15	219±	15	219±	14	221±	14	224±	18
100 ppm	212±	12	215±	13	217±	13	217±	13	221±	13	225±	14
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett												

(HAN260)

BAIS 4

STUDY NO. : 0535
 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	238±	18	243±	20	246±	21	249±	22	253±	22	257±	23
25 ppm	246±	21	250±	22	254±	22	257±	23	260±	24	264±	25
50 ppm	233±	19	238±	20	243±	20	245±	21	249±	23	254±	28
100 ppm	234±	15	239±	16	244±	17	246±	17	250±	17	253±	18

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
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	257±	27	258±	28	259±	21
25 ppm	269±	27	272±	27	271±	28
50 ppm	260±	23	261±	26	259±	27
100 ppm	254±	24	257±	27	256±	25
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett						

(HAN260)

BAIS 4

APPENDIX E 1

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0535

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCr1j]

UNIT : g

REPORT TYPE : A1 104

SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)

ALL ANIMALS

PAGE : 1

Group Name	Administration week-day(effective)						
	1-7(4)	2-7(7)	3-7(7)	4-7(7)	5-7(7)	6-7(7)	7-7(7)
Control	14.4± 0.9	15.7± 1.1	16.8± 1.2	16.9± 1.2	16.9± 1.2	16.6± 1.1	16.7± 1.0
25 ppm	14.3± 0.9	15.7± 1.0	16.9± 1.1	16.7± 1.0	16.9± 0.9	16.4± 0.9	16.5± 1.1
50 ppm	14.1± 1.1	15.4± 1.3	16.6± 1.1	16.7± 1.1	17.0± 1.0	16.6± 1.0	16.3± 1.2
100 ppm	13.2± 1.3**	13.9± 1.0**	14.8± 1.0**	15.5± 0.9**	16.4± 0.9*	15.4± 1.0**	15.0± 1.1**
Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Dunnett							

(HAN260)

BAIS 4

STUDY NO. : 0535
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 8-7(7)	week-day(effective) 9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	17-7(7)
Control	17.2± 1.0	16.9± 1.2	16.9± 1.1	16.4± 1.1	16.1± 0.9	15.9± 0.9	16.0± 1.1
25 ppm	17.3± 1.0	16.8± 1.1	16.8± 0.9	16.2± 1.1	15.9± 1.0	15.7± 1.0	15.5± 1.4
50 ppm	16.9± 1.0	16.7± 1.1	16.8± 1.0	16.4± 1.0	16.2± 1.1	16.0± 0.9	15.9± 1.0
100 ppm	15.9± 1.1**	15.5± 1.2**	15.9± 1.0**	15.5± 1.0**	15.3± 1.1**	15.0± 1.0**	15.2± 1.0**

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

Test of Dunnett

STUDY NO. : 0535
 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 21-7(7)	week-day(effective) 25-7(7)	29-7(7)	33-7(7)	37-7(7)	41-7(7)	45-7(7)
Control	16.3± 1.1	16.0± 1.1	16.6± 1.2	16.4± 1.2	15.4± 1.0	15.7± 1.1	15.7± 1.5
25 ppm	16.0± 0.9	15.8± 1.1	16.5± 1.2	16.3± 1.2	15.0± 0.9	15.6± 1.1	15.9± 1.2
50 ppm	16.0± 0.8	15.7± 1.0	16.4± 1.1	16.2± 1.0	15.5± 1.0	15.6± 0.9	15.6± 1.0
100 ppm	15.3± 1.0**	15.3± 0.9**	15.8± 1.1**	15.2± 1.2**	15.0± 0.9*	15.3± 1.1	15.0± 1.0**

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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)						
	50-7(7)	53-7(7)	54-7(7)	58-7(7)	62-7(7)	66-7(7)	70-7(7)
Control	16.5± 1.0	16.5± 0.9	16.5± 1.0	16.2± 1.1	17.1± 0.9	16.9± 0.9	17.2± 1.0
25 ppm	16.4± 1.2	16.5± 1.1	16.4± 1.1	16.3± 0.9	16.9± 1.2	16.8± 1.1	17.0± 1.1
50 ppm	16.3± 0.9	16.1± 0.9	16.2± 1.1	16.3± 0.8	16.6± 1.1	16.6± 1.1	17.0± 1.0
100 ppm	15.5± 1.0**	15.8± 0.9**	15.9± 0.9**	15.7± 1.1*	16.3± 1.0**	16.1± 0.9**	16.6± 1.1

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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	16.7± 1.1	16.7± 1.4	16.3± 1.0	16.6± 1.1	16.2± 1.2	16.3± 1.2	16.5± 1.3
25 ppm	16.5± 1.1	16.4± 1.1	16.1± 1.0	16.3± 1.9	16.3± 2.1	16.4± 2.8	16.7± 1.3
50 ppm	16.5± 0.9	16.6± 1.1	16.1± 1.9	16.7± 1.0	16.8± 1.2*	16.7± 1.1	16.9± 1.5
100 ppm	16.1± 1.3	16.4± 1.3	15.9± 1.3	16.2± 1.0	15.9± 1.5	15.4± 1.9*	15.5± 2.4

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
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	16.5±	2.1	16.5±	2.2
25 ppm	16.7±	1.2	16.7±	1.3
50 ppm	16.4±	2.0	16.9±	2.1
100 ppm	16.2±	1.7	15.9±	1.2*
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett				

(HAN260)

BAIS 4

APPENDIX E 2

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0535
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(4)	2-7(7)	3-7(7)	4-7(7)	5-7(7)	6-7(7)	7-7(7)
Control	10.5± 0.6	11.0± 0.8	11.0± 0.7	11.2± 0.8	11.5± 0.9	11.1± 1.0	10.8± 0.9
25 ppm	10.2± 0.7*	11.1± 0.8	11.3± 0.8	11.4± 0.7	11.6± 0.8	11.2± 0.9	11.3± 1.0*
50 ppm	10.1± 0.7**	11.0± 0.9	11.0± 0.8	11.3± 0.9	11.7± 1.0	10.9± 0.7	10.9± 1.0
100 ppm	10.0± 0.7**	10.5± 0.8**	10.9± 0.7	11.1± 0.8	11.8± 0.9	10.6± 0.9*	10.7± 0.9
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett							

(HAN260)

BAIS 4

STUDY NO. : 0535
 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 8-7(7)	week-day(effective) 9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	17-7(7)
Control	10.9± 1.5	10.9± 0.9	10.7± 0.7	11.0± 0.9	10.4± 0.8	10.7± 0.7	10.9± 0.7
25 ppm	10.9± 1.0	11.5± 1.4*	11.1± 1.2	11.2± 0.8	10.5± 0.8	11.3± 1.1*	11.2± 1.3
50 ppm	10.7± 0.9	10.9± 1.1	10.8± 0.9	10.8± 0.9	10.6± 0.9	11.0± 1.0	11.0± 1.0
100 ppm	10.6± 0.9	10.8± 0.9	10.4± 0.9	11.0± 1.2	10.4± 0.9	10.7± 1.0	10.7± 0.9
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett							

(HAN260)

BAIS 4

STUDY NO. : 0535
 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 21-7(7)	week-day(effective) 25-7(7)	29-7(7)	33-7(7)	37-7(7)	41-7(7)	45-7(7)
Control	10.4± 0.8	11.1± 1.1	10.8± 0.7	10.5± 0.9	12.4± 1.7	11.1± 0.9	11.3± 0.9
25 ppm	10.6± 0.9	11.4± 1.2	11.1± 0.9	10.5± 0.9	12.5± 1.5	11.3± 0.9	11.5± 0.9
50 ppm	10.3± 0.8	10.7± 1.3	10.6± 0.9	10.4± 0.8	12.1± 1.5	11.0± 1.1	10.8± 1.1*
100 ppm	10.4± 0.8	10.7± 1.2	11.1± 1.0	10.0± 0.8*	12.4± 1.4	11.5± 1.2	11.1± 0.8

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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)						
	50-7(7)	53-7(7)	54-7(7)	58-7(7)	62-7(7)	66-7(7)	70-7(7)
Control	10.5± 0.8	11.2± 0.7	10.8± 0.8	11.0± 0.8	11.5± 0.8	11.6± 0.9	11.7± 0.8
25 ppm	10.8± 0.8	11.4± 0.9	11.2± 0.9*	11.3± 0.8	11.8± 1.0	11.8± 1.1	12.0± 1.2
50 ppm	10.1± 0.6*	11.1± 0.9	10.8± 0.7	10.6± 0.7*	11.5± 0.9	11.2± 0.9	11.6± 1.0
100 ppm	10.5± 0.8	10.8± 0.7*	10.7± 0.8	10.8± 0.7	11.4± 0.9	11.5± 0.9	11.8± 0.8

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
 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	11.5± 1.1	11.5± 1.1	11.4± 1.1	11.8± 1.1	12.0± 1.0	11.7± 1.5	11.9± 1.6
25 ppm	11.8± 1.0	12.0± 0.9	11.8± 1.0	12.1± 1.2	12.2± 1.1	12.2± 0.9	12.5± 1.0
50 ppm	11.5± 1.1	11.6± 1.2	11.6± 1.3	12.0± 1.7	12.0± 1.9	11.9± 1.3	12.6± 1.4
100 ppm	11.4± 1.1	11.7± 1.0	11.6± 1.1	11.9± 0.9	11.8± 1.0	11.6± 1.9	12.0± 2.0

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

Test of Dunnett

(HAN260)

BAIS 4

STUDY NO. : 0535
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	11.6± 1.9	12.2± 1.0
25 ppm	12.4± 1.3	12.4± 1.3
50 ppm	12.3± 1.7	12.0± 1.9
100 ppm	12.3± 1.3	11.9± 1.7
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett		

(HAN260)

BAIS 4

APPENDIX F 1

HEMATOLOGY : MALE

STUDY NO. : 0535
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	43	8.02±	1.48	13.5±	2.8	39.6±	6.9	49.5±	3.1	16.8±	1.5	33.9±	1.9	955±	328
25 ppm	44	8.19±	1.59	13.6±	3.0	39.8±	7.3	48.8±	3.1	16.5±	1.5	33.8±	1.8	997±	365
50 ppm	42	8.49±	1.65	14.3±	2.8	41.3±	7.1	49.4±	6.6	17.0±	2.3	34.3±	1.4	909±	377
100 ppm	36	8.16±	1.48	13.7±	2.9	40.2±	7.3	49.3±	2.9	16.8±	1.5	33.9±	1.8	934±	315

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0535

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)

ALL ANIMALS (105W)

PAGE : 2

Group Name	NO. of Animals	RETICULOCYTE %	
Control	43	5.0±	4.0
25 ppm	44	4.9±	3.9
50 ppm	42	4.9±	7.1
100 ppm	36	4.5±	3.1

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0535
 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	43	5.97±	2.03	0±	1	46±	9	2±	1	0±	0	5±	2	45±	10	2±	5
25 ppm	44	6.24±	2.70	1±	1	49±	10	2±	1	0±	0	5±	2	43±	11	1±	3
50 ppm	42	7.90±	10.13	0±	1	48±	10	2±	1	0±	0	5±	2	41±	11	3±	14
100 ppm	36	5.27±	1.78	1±	1	46±	7	2±	1	0±	0	5±	2	45±	8	2±	2

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

APPENDIX F 2

HEMATOLOGY : FEMALE

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
MEASURE. TIME : 1
SEX : FEMALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

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	37	8.23±	0.88	15.1±	1.7	42.1±	4.0	51.3±	2.1	18.3±	0.5	35.7±	1.1	651±	175
25 ppm	45	8.19±	0.94	14.9±	1.7	41.9±	3.7	51.4±	3.1	18.2±	1.0	35.4±	1.6	675±	185
50 ppm	36	7.90±	1.43	14.5±	2.4	40.7±	5.9	52.7±	7.9	18.5±	1.7	35.4±	1.7	636±	207
100 ppm	39	7.94±	1.16	14.8±	2.1	41.9±	5.0	53.4±	5.0**	18.6±	0.9*	35.0±	2.0**	644±	109

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0535
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	37	3.1±	4.2
25 ppm	45	3.3±	3.8
50 ppm	36	4.8±	8.0*
100 ppm	39	3.9±	3.5**

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
MEASURE. TIME : 1
SEX : FEMALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 6

Group Name	NO. of Animals	WBC 10 ³ /μl		Differential N-BAND		WBC (%) N-SEG		EOSINO		BASO		MONO		LYMPHO		OTHER	
Control	37	2.99±	1.41	1±	1	38±	10	2±	1	0±	0	5±	2	54±	10	1±	1
25 ppm	45	8.08±	34.45	0±	1	41±	11	2±	1	0±	0	5±	1	49±	11	3±	14
50 ppm	36	5.26±	11.34	1±	1	40±	11	2±	1	0±	0	5±	2	46±	13	6±	19
100 ppm	39	6.08±	12.05	0±	1	39±	12	2±	1	0±	0	4±	2	49±	15	6±	21

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

APPENDIX G 1

BIOCHEMISTRY : MALE

STUDY NO. : 0535

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (105W)

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	43	6.6±	0.3	3.0±	0.3	0.8±	0.1	0.17±	0.05	162±	21	162±	40	86±	43
25 ppm	44	6.6±	0.3	3.0±	0.3	0.8±	0.1	0.16±	0.03	166±	19	160±	50	81±	57
50 ppm	42	6.7±	0.3	3.0±	0.2	0.8±	0.1	0.30±	0.66	163±	23	169±	53	103±	104
100 ppm	36	6.8±	0.5	3.1±	0.3	0.8±	0.1	0.17±	0.05	163±	27	168±	48	81±	42

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

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	43	232±	49	105±	56	44±	20	181±	46	247±	70	9±	5	121±	37
25 ppm	44	230±	59	109±	42	43±	14	205±	57	254±	82	9±	3	136±	42
50 ppm	42	247±	74	117±	108	46±	21	196±	73	251±	110	8±	4	135±	47
100 ppm	36	245±	64	115±	40	52±	29	191±	55	267±	220	11±	7	132±	44

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

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	43	19.2±	3.1	0.6±	0.1	142±	1	3.8±	0.3	105±	2	10.5±	0.2	4.2±	0.5
25 ppm	44	19.8±	3.4	0.6±	0.1	142±	2	3.8±	0.3	105±	1	10.4±	0.3	4.0±	0.5
50 ppm	42	19.1±	3.3	0.6±	0.1	141±	1	3.7±	0.3	105±	2	10.4±	0.4	4.0±	0.7
100 ppm	36	19.8±	3.2	0.6±	0.1	142±	2	3.7±	0.3	105±	2	10.5±	0.4	4.1±	0.5

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

APPENDIX G 2

BIOCHEMISTRY : FEMALE

STUDY NO. : 0535
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	37	6.9±	0.4	3.6±	0.3	1.1±	0.1	0.13±	0.02	154±	15	125±	23	39±	21
25 ppm	45	6.8±	0.4	3.6±	0.3	1.1±	0.1	0.15±	0.11	155±	15	129±	26	46±	22
50 ppm	36	6.8±	0.5	3.6±	0.4	1.1±	0.1	0.23±	0.52	150±	17	129±	26	57±	45
100 ppm	39	6.9±	0.5	3.6±	0.4	1.1±	0.1	0.16±	0.16	154±	17	141±	35	65±	87

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

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	37	221±	39	130±	75	60±	32	206±	77	132±	35	3±	2	98±	24
25 ppm	45	227±	38	154±	220	63±	67	287±	514	150±	62	3±	2	116±	73
50 ppm	36	231±	46	151±	104	62±	34	233±	112	171±	129	4±	5	118±	64
100 ppm	39	250±	57*	137±	91	60±	33	240±	168	158±	104	3±	1	105±	28

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

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	37	17.5±	1.7	0.5±	0.0	141±	2	3.5±	0.3	103±	2	10.5±	0.4	3.9±	0.6
25 ppm	45	17.8±	1.7	0.5±	0.1	141±	1	3.4±	0.3	103±	2	10.5±	0.4	3.8±	0.8
50 ppm	36	18.0±	2.0	0.5±	0.1	141±	2	3.4±	0.4	102±	2	10.5±	0.3	4.1±	0.7
100 ppm	39	18.5±	3.2	0.5±	0.1	141±	1	3.4±	0.4	103±	2	10.6±	0.4	3.9±	0.9

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

APPENDIX H 1

URINALYSIS : MALE

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 MEASURE. TIME : 1
 SEX : MALE

URINALYSIS

REPORT TYPE : A1

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	44	0	0	3	3	16	20	2		0	0	0	7	33	4		44	0	0	0	0	0		41	3	0	0	0	0		43	1	0	0	
25 ppm	44	0	0	0	3	23	17	1		0	0	1	4	36	3		44	0	0	0	0	0		42	2	0	0	0	0		43	1	0	0	
50 ppm	42	0	0	2	3	16	20	1		0	0	1	6	28	7		42	0	0	0	0	0		42	0	0	0	0	0		41	1	0	0	
100 ppm	37	0	1	2	2	18	13	1		0	0	2	9	22	4		37	0	0	0	0	0		37	0	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. : 0535

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

URINALYSIS

PAGE : 2

Group Name	NO. of Animals	Occult blood					CHI	Urobilinogen					CHI
		—	±	+	2+	3+		±	+	2+	3+	4+	
Control	44	41	1	0	0	2		44	0	0	0	0	
25 ppm	44	43	0	0	1	0		44	0	0	0	0	
50 ppm	42	41	1	0	0	0		42	0	0	0	0	
100 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)

BAIS4

APPENDIX H 2

URINALYSIS : FEMALE

STUDY NO. : 0535

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	40	0	2	2	6	9	13	8		2	10	11	8	7	2		40	0	0	0	0	0		25	15	0	0	0	0		40	0	0	0	
25 ppm	45	0	3	1	3	9	17	12		2	5	14	18	4	2		45	0	0	0	0	0		32	13	0	0	0	0		45	0	0	0	
50 ppm	37	0	2	3	4	4	16	8		2	4	9	14	8	0		37	0	0	0	0	0		26	11	0	0	0	0		36	0	0	1	
100 ppm	39	0	2	1	2	5	18	11		1	3	9	12	11	3		39	0	0	0	0	0		17	22	0	0	0	0		38	1	0	0	

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

Test of CHI SQUARE

(HCL101)

BAIS 4

STUDY NO. : 0535

URINALYSIS

ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

PAGE : 4

Group Name	NO. of Animals	Occult blood					CHI	Urobilinogen					CHI
		—	±	+	2+	3+		±	+	2+	3+	4+	
Control	40	39	1	0	0	0		40	0	0	0	0	
25 ppm	45	43	1	1	0	0		45	0	0	0	0	
50 ppm	37	33	0	0	1	3		36	0	0	1	0	
100 ppm	39	37	0	1	0	1		39	0	0	0	0	

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

Test of CHI SQUARE

(HCL101)

BAIS 4

APPENDIX I 1

GROSS FINDINGS : MALE

ALL ANIMALS

STUDY NO. : 0535
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		25 ppm		50 ppm		100 ppm	
			50	(%)	50	(%)	50	(%)	50	(%)
skin/app	nodule		4	(8)	3	(6)	3	(6)	4	(8)
	scab		1	(2)	1	(2)	0	(0)	0	(0)
subcutis	edema		0	(0)	1	(2)	1	(2)	0	(0)
	jaundice		0	(0)	0	(0)	2	(4)	1	(2)
	mass		8	(16)	9	(18)	5	(10)	10	(20)
lung	white zone		5	(10)	2	(4)	3	(6)	3	(6)
	red zone		0	(0)	2	(4)	3	(6)	0	(0)
	nodule		2	(4)	0	(0)	0	(0)	2	(4)
	voluminus		0	(0)	1	(2)	0	(0)	0	(0)
lymph node	enlarged		0	(0)	0	(0)	2	(4)	1	(2)
thymus	nodule		0	(0)	0	(0)	1	(2)	0	(0)
spleen	enlarged		7	(14)	4	(8)	5	(10)	4	(8)
	white zone		1	(2)	1	(2)	0	(0)	0	(0)
heart	white zone		0	(0)	0	(0)	0	(0)	1	(2)
	nodule		0	(0)	1	(2)	0	(0)	0	(0)
tongue	nodule		0	(0)	1	(2)	0	(0)	0	(0)
salivary gl	nodule		0	(0)	1	(2)	0	(0)	0	(0)
forestomach	nodule		1	(2)	0	(0)	1	(2)	0	(0)
	ulcer		1	(2)	0	(0)	1	(2)	3	(6)
gl stomach	red zone		0	(0)	0	(0)	0	(0)	1	(2)
	nodule		0	(0)	2	(4)	0	(0)	0	(0)
	ulcer		1	(2)	0	(0)	0	(0)	0	(0)

STUDY NO. : 0535
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				25 ppm				50 ppm				100 ppm			
			50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)
liver	enlarged		0	(0)	4	(8)	0	(0)	0	(0)	0	(0)	0	(0)				
	nodule		2	(4)	1	(2)	1	(2)	1	(2)	0	(0)	0	(0)				
	rough		1	(2)	1	(2)	2	(4)	0	(0)	1	(2)	0	(0)				
	granular		0	(0)	0	(0)	0	(0)	0	(0)	1	(2)	0	(0)				
	nodular		0	(0)	1	(2)	0	(0)	0	(0)	0	(0)	0	(0)				
	herniation		7	(14)	3	(6)	2	(4)	5	(10)								
kidney	white zone		0	(0)	0	(0)	0	(0)	1	(2)	0	(0)	0	(0)				
	nodule		1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)				
	cyst		1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)				
	granular		1	(2)	3	(6)	5	(10)	3	(6)								
urethra	nodule		0	(0)	0	(0)	1	(2)	0	(0)								
pituitary	enlarged		0	(0)	0	(0)	1	(2)	3	(6)								
	red zone		2	(4)	1	(2)	2	(4)	1	(2)								
	nodule		0	(0)	4	(8)	2	(4)	4	(8)								
thyroid	enlarged		1	(2)	2	(4)	6	(12)	2	(4)								
	nodule		1	(2)	1	(2)	0	(0)	1	(2)								
adrenal	enlarged		0	(0)	1	(2)	1	(2)	3	(6)								
	nodule		0	(0)	0	(0)	1	(2)	0	(0)								
testis	nodule		43	(86)	49	(98)	45	(90)	43	(86)								
brain	red zone		0	(0)	0	(0)	1	(2)	0	(0)								
	nodule		2	(4)	0	(0)	0	(0)	0	(0)								
spinal cord	red zone		0	(0)	0	(0)	1	(2)	0	(0)								

STUDY NO. : 0535
 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				25 ppm				50 ppm				100 ppm			
			50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)	50	(%)
periph nerv	nodule		2	(4)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)
eye	turbid		0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	1	(2)	1	(2)
	white		2	(4)	4	(8)	3	(6)	3	(6)	3	(6)	3	(6)	2	(4)	2	(4)
	red		1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)
	exophthalmos		0	(0)	0	(0)	1	(2)	1	(2)	1	(2)	1	(2)	0	(0)	0	(0)
Zymbal gl	nodule		0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	2	(4)	2	(4)
bone	nodule		1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)
peritoneum	nodule		1	(2)	2	(4)	1	(2)	1	(2)	1	(2)	1	(2)	2	(4)	2	(4)
retroperit	mass		0	(0)	0	(0)	1	(2)	1	(2)	1	(2)	1	(2)	0	(0)	0	(0)
abdominal c	hemorrhage		1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)
	ascites		2	(4)	3	(6)	2	(4)	2	(4)	2	(4)	2	(4)	2	(4)	2	(4)
adipose	nodule		0	(0)	0	(0)	1	(2)	1	(2)	1	(2)	1	(2)	0	(0)	0	(0)
thoracic ca	pleural fluid		1	(2)	2	(4)	2	(4)	2	(4)	2	(4)	2	(4)	2	(4)	2	(4)
other	forelimb:nodule		0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	1	(2)	1	(2)
	upper jaw:nodule		0	(0)	0	(0)	1	(2)	1	(2)	1	(2)	1	(2)	0	(0)	0	(0)
	nose:nodule		0	(0)	1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	1	(2)	1	(2)
whole body	anemic		0	(0)	1	(2)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)	0	(0)

APPENDIX I 2

GROSS FINDINGS : MALE DEAD AND MORIBUND ANIMALS

STUDY NO. : 0535
 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	25 ppm	50 ppm	100 ppm
			6 (%)	6 (%)	8 (%)	14 (%)
skin/app	nodule		0 (0)	1 (17)	0 (0)	0 (0)
subcutis	edema		0 (0)	1 (17)	1 (13)	0 (0)
	jaundice		0 (0)	0 (0)	2 (25)	1 (7)
	mass		1 (17)	1 (17)	1 (13)	3 (21)
lung	white zone		0 (0)	0 (0)	0 (0)	1 (7)
	red zone		0 (0)	2 (33)	3 (38)	0 (0)
	nodule		0 (0)	0 (0)	0 (0)	1 (7)
	voluminus		0 (0)	1 (17)	0 (0)	0 (0)
lymph node	enlarged		0 (0)	0 (0)	1 (13)	1 (7)
spleen	enlarged		2 (33)	3 (50)	3 (38)	3 (21)
heart	white zone		0 (0)	0 (0)	0 (0)	1 (7)
tongue	nodule		0 (0)	1 (17)	0 (0)	0 (0)
forestomach	ulcer		1 (17)	0 (0)	1 (13)	3 (21)
gl stomach	red zone		0 (0)	0 (0)	0 (0)	1 (7)
	nodule		0 (0)	1 (17)	0 (0)	0 (0)
	ulcer		1 (17)	0 (0)	0 (0)	0 (0)
liver	enlarged		0 (0)	4 (67)	0 (0)	0 (0)
	rough		0 (0)	1 (17)	0 (0)	0 (0)
	granular		0 (0)	0 (0)	0 (0)	1 (7)
	nodular		0 (0)	1 (17)	0 (0)	0 (0)
	herniation		1 (17)	1 (17)	0 (0)	2 (14)
kidney	granular		0 (0)	0 (0)	1 (13)	0 (0)

STUDY NO. : 0535
 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	25 ppm	50 ppm	100 ppm
			6 (%)	6 (%)	8 (%)	14 (%)
pituitary	enlarged		0 (0)	0 (0)	1 (13)	2 (14)
	red zone		0 (0)	0 (0)	1 (13)	0 (0)
thyroid	enlarged		0 (0)	0 (0)	1 (13)	0 (0)
adrenal	enlarged		0 (0)	0 (0)	1 (13)	2 (14)
testis	nodule		2 (33)	5 (83)	3 (38)	7 (50)
brain	red zone		0 (0)	0 (0)	1 (13)	0 (0)
	nodule		2 (33)	0 (0)	0 (0)	0 (0)
spinal cord	red zone		0 (0)	0 (0)	1 (13)	0 (0)
periph nerv	nodule		2 (33)	0 (0)	0 (0)	0 (0)
eye	turbid		0 (0)	0 (0)	0 (0)	1 (7)
	white		1 (17)	0 (0)	0 (0)	1 (7)
	red		1 (17)	0 (0)	0 (0)	0 (0)
	exophthalmos		0 (0)	0 (0)	1 (13)	0 (0)
Zymbal gl	nodule		0 (0)	0 (0)	0 (0)	2 (14)
bone	nodule		1 (17)	0 (0)	0 (0)	0 (0)
peritoneum	nodule		0 (0)	0 (0)	0 (0)	1 (7)
retroperit	mass		0 (0)	0 (0)	1 (13)	0 (0)
abdominal c	hemorrhage		1 (17)	0 (0)	0 (0)	0 (0)
	ascites		1 (17)	1 (17)	1 (13)	1 (7)
adipose	nodule		0 (0)	0 (0)	1 (13)	0 (0)
thoracic ca	pleural fluid		0 (0)	1 (17)	2 (25)	1 (7)
other	forelimb:nodule		0 (0)	0 (0)	0 (0)	1 (7)

STUDY NO. : 0535
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	Control	25 ppm	50 ppm	100 ppm
		NO. of Animals	6 (%)	6 (%)	8 (%)	14 (%)
whole body	anemic		0 (0)	1 (17)	0 (0)	0 (0)

(HPT080)

BAIS 4

APPENDIX I 3

GROSS FINDINGS : MALE SACRIFICED ANIMALS

STUDY NO. : 0535
 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		25 ppm		50 ppm		100 ppm	
			44	(%)	44	(%)	42	(%)	36	(%)
skin/app	nodule		4	(9)	2	(5)	3	(7)	4	(11)
	scab		1	(2)	1	(2)	0	(0)	0	(0)
subcutis	mass		7	(16)	8	(18)	4	(10)	7	(19)
lung	white zone		5	(11)	2	(5)	3	(7)	2	(6)
	nodule		2	(5)	0	(0)	0	(0)	1	(3)
lymph node	enlarged		0	(0)	0	(0)	1	(2)	0	(0)
thymus	nodule		0	(0)	0	(0)	1	(2)	0	(0)
spleen	enlarged		5	(11)	1	(2)	2	(5)	1	(3)
	white zone		1	(2)	1	(2)	0	(0)	0	(0)
heart	nodule		0	(0)	1	(2)	0	(0)	0	(0)
salivary gl	nodule		0	(0)	1	(2)	0	(0)	0	(0)
forestomach	nodule		1	(2)	0	(0)	1	(2)	0	(0)
gl stomach	nodule		0	(0)	1	(2)	0	(0)	0	(0)
liver	nodule		2	(5)	1	(2)	1	(2)	0	(0)
	rough		1	(2)	0	(0)	2	(5)	0	(0)
	herniation		6	(14)	2	(5)	2	(5)	3	(8)
kidney	white zone		0	(0)	0	(0)	0	(0)	1	(3)
	nodule		1	(2)	0	(0)	0	(0)	0	(0)
	cyst		1	(2)	0	(0)	0	(0)	0	(0)
	granular		1	(2)	3	(7)	4	(10)	3	(8)
urethra	nodule		0	(0)	0	(0)	1	(2)	0	(0)
pituitary	enlarged		0	(0)	0	(0)	0	(0)	1	(3)

STUDY NO. : 0535
 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	25 ppm	50 ppm	100 ppm
			44 (%)	44 (%)	42 (%)	36 (%)
pituitary	red zone		2 (5)	1 (2)	1 (2)	1 (3)
	nodule		0 (0)	4 (9)	2 (5)	4 (11)
thyroid	enlarged		1 (2)	2 (5)	5 (12)	2 (6)
	nodule		1 (2)	1 (2)	0 (0)	1 (3)
adrenal	enlarged		0 (0)	1 (2)	0 (0)	1 (3)
	nodule		0 (0)	0 (0)	1 (2)	0 (0)
testis	nodule		41 (93)	44 (100)	42 (100)	36 (100)
eye	white		1 (2)	4 (9)	3 (7)	1 (3)
peritoneum	nodule		1 (2)	2 (5)	1 (2)	1 (3)
abdominal c	ascites		1 (2)	2 (5)	1 (2)	1 (3)
thoracic ca	pleural fluid		1 (2)	1 (2)	0 (0)	1 (3)
other	upper jaw:nodule		0 (0)	0 (0)	1 (2)	0 (0)
	nose:nodule		0 (0)	1 (2)	0 (0)	1 (3)

APPENDIX I 4

GROSS FINDINGS : FEMALE

ALL ANIMALS

STUDY NO. : 0535
 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	25 ppm	50 ppm	100 ppm
			50 (%)	50 (%)	50 (%)	50 (%)
subcutis	jaundice		2 (4)	0 (0)	1 (2)	3 (6)
	mass		6 (12)	11 (22)	9 (18)	9 (18)
lung	white zone		1 (2)	1 (2)	1 (2)	1 (2)
	red zone		1 (2)	0 (0)	0 (0)	0 (0)
	nodule		1 (2)	1 (2)	1 (2)	1 (2)
lymph node	enlarged		3 (6)	2 (4)	1 (2)	2 (4)
spleen	enlarged		8 (16)	3 (6)	7 (14)	9 (18)
	nodule		0 (0)	0 (0)	1 (2)	0 (0)
heart	white zone		0 (0)	0 (0)	0 (0)	1 (2)
forestomach	nodule		1 (2)	1 (2)	0 (0)	0 (0)
	ulcer		2 (4)	1 (2)	2 (4)	1 (2)
	erosion		0 (0)	1 (2)	0 (0)	0 (0)
gl stomach	nodule		0 (0)	0 (0)	1 (2)	0 (0)
	ulcer		1 (2)	1 (2)	0 (0)	0 (0)
	erosion		0 (0)	0 (0)	0 (0)	1 (2)
cecum	dilated		0 (0)	0 (0)	1 (2)	0 (0)
liver	enlarged		1 (2)	0 (0)	0 (0)	2 (4)
	white zone		3 (6)	1 (2)	1 (2)	2 (4)
	red zone		1 (2)	1 (2)	0 (0)	0 (0)
	nodule		0 (0)	0 (0)	1 (2)	1 (2)
	rough		2 (4)	2 (4)	4 (8)	6 (12)
	herniation		12 (24)	9 (18)	8 (16)	9 (18)

STUDY NO. : 0535
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		25 ppm		50 ppm		100 ppm	
			50	(%)	50	(%)	50	(%)	50	(%)
kidney	white zone		1	(2)	0	(0)	0	(0)	1	(2)
	nodule		0	(0)	0	(0)	1	(2)	1	(2)
	cyst		0	(0)	1	(2)	0	(0)	0	(0)
	granular		0	(0)	0	(0)	1	(2)	0	(0)
	hydronephrosis		0	(0)	0	(0)	2	(4)	0	(0)
urin bladd	red zone		0	(0)	0	(0)	1	(2)	0	(0)
	urine:marked retention		0	(0)	0	(0)	1	(2)	0	(0)
pituitary	enlarged		3	(6)	3	(6)	5	(10)	2	(4)
	red zone		3	(6)	7	(14)	2	(4)	5	(10)
	black zone		1	(2)	0	(0)	0	(0)	0	(0)
	nodule		6	(12)	3	(6)	3	(6)	5	(10)
	cyst		0	(0)	0	(0)	0	(0)	1	(2)
thyroid	enlarged		0	(0)	0	(0)	1	(2)	2	(4)
	nodule		0	(0)	1	(2)	1	(2)	1	(2)
adrenal	enlarged		1	(2)	0	(0)	1	(2)	1	(2)
	nodule		0	(0)	0	(0)	0	(0)	1	(2)
ovary	enlarged		0	(0)	1	(2)	1	(2)	0	(0)
	nodule		0	(0)	0	(0)	0	(0)	1	(2)
	cyst		3	(6)	6	(12)	2	(4)	1	(2)
uterus	black zone		0	(0)	0	(0)	1	(2)	0	(0)
	nodule		8	(16)	2	(4)	9	(18)	9	(18)
	cyst		0	(0)	1	(2)	0	(0)	0	(0)

STUDY NO. : 0535
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		25 ppm		50 ppm		100 ppm	
			50	(%)	50	(%)	50	(%)	50	(%)
prep/cli gl	nodule		0	(0)	0	(0)	0	(0)	1	(2)
brain	red zone		0	(0)	0	(0)	1	(2)	0	(0)
	hemorrhage		0	(0)	0	(0)	1	(2)	0	(0)
	deformed		0	(0)	1	(2)	0	(0)	0	(0)
spinal cord	red zone		0	(0)	0	(0)	0	(0)	1	(2)
eye	turbid		1	(2)	0	(0)	0	(0)	0	(0)
	white		4	(8)	5	(10)	7	(14)	0	(0)
Zymbal gl	nodule		0	(0)	0	(0)	0	(0)	2	(4)
bone	nodule		0	(0)	0	(0)	1	(2)	0	(0)
mediastinum	mass		0	(0)	0	(0)	0	(0)	2	(4)
peritoneum	nodule		1	(2)	1	(2)	0	(0)	2	(4)
	adhesion		0	(0)	0	(0)	1	(2)	0	(0)
abdominal c	ascites		2	(4)	0	(0)	0	(0)	1	(2)
thoracic ca	pleural fluid		1	(2)	0	(0)	0	(0)	3	(6)
other	eye lid:nodule		1	(2)	0	(0)	0	(0)	0	(0)
	upper jaw:nodule		1	(2)	0	(0)	0	(0)	0	(0)
whole body	anemic		1	(2)	0	(0)	0	(0)	1	(2)

APPENDIX I 5

GROSS FINDINGS : FEMALE DEAD AND MORIBUND ANIMALS

STUDY NO. : 0535
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 NO. of Animals	Control	25 ppm	50 ppm	100 ppm
			13 (%)	5 (%)	14 (%)	11 (%)
subcutis	jaundice		2 (15)	0 (0)	0 (0)	3 (27)
	mass		2 (15)	2 (40)	3 (21)	2 (18)
lung	white zone		0 (0)	0 (0)	1 (7)	0 (0)
	red zone		1 (8)	0 (0)	0 (0)	0 (0)
	nodule		1 (8)	0 (0)	1 (7)	0 (0)
lymph node	enlarged		3 (23)	2 (40)	1 (7)	2 (18)
spleen	enlarged		8 (62)	2 (40)	4 (29)	7 (64)
heart	white zone		0 (0)	0 (0)	0 (0)	1 (9)
forestomach	nodule		1 (8)	0 (0)	0 (0)	0 (0)
	ulcer		2 (15)	1 (20)	1 (7)	1 (9)
	erosion		0 (0)	1 (20)	0 (0)	0 (0)
gl stomach	ulcer		1 (8)	1 (20)	0 (0)	0 (0)
	erosion		0 (0)	0 (0)	0 (0)	1 (9)
liver	enlarged		1 (8)	0 (0)	0 (0)	2 (18)
	white zone		1 (8)	0 (0)	1 (7)	1 (9)
	nodule		0 (0)	0 (0)	1 (7)	0 (0)
	rough		0 (0)	0 (0)	2 (14)	4 (36)
	herniation		6 (46)	1 (20)	4 (29)	1 (9)
kidney	white zone		1 (8)	0 (0)	0 (0)	1 (9)
	nodule		0 (0)	0 (0)	1 (7)	1 (9)
	granular		0 (0)	0 (0)	1 (7)	0 (0)
	hydronephrosis		0 (0)	0 (0)	2 (14)	0 (0)

STUDY NO. : 0535
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				25 ppm				50 ppm				100 ppm			
			13	(%)	5	(%)	14	(%)	11	(%)	14	(%)	11	(%)	11	(%)	11	(%)
urin bladd	red zone		0	(0)	0	(0)	1	(7)	0	(0)								
	urine:marked retention		0	(0)	0	(0)	1	(7)	0	(0)								
pituitary	enlarged		2	(15)	2	(40)	1	(7)	0	(0)								
	red zone		0	(0)	1	(20)	1	(7)	2	(18)								
	black zone		1	(8)	0	(0)	0	(0)	0	(0)								
	nodule		1	(8)	0	(0)	0	(0)	2	(18)								
	cyst		0	(0)	0	(0)	0	(0)	1	(9)								
adrenal	enlarged		0	(0)	0	(0)	0	(0)	1	(9)								
	nodule		0	(0)	0	(0)	0	(0)	1	(9)								
ovary	enlarged		0	(0)	0	(0)	1	(7)	0	(0)								
	cyst		1	(8)	1	(20)	0	(0)	0	(0)								
uterus	nodule		4	(31)	0	(0)	2	(14)	2	(18)								
prep/cli gl	nodule		0	(0)	0	(0)	0	(0)	1	(9)								
brain	red zone		0	(0)	0	(0)	1	(7)	0	(0)								
	hemorrhage		0	(0)	0	(0)	1	(7)	0	(0)								
	deformed		0	(0)	1	(20)	0	(0)	0	(0)								
spinal cord	red zone		0	(0)	0	(0)	0	(0)	1	(9)								
eye	white		0	(0)	1	(20)	2	(14)	0	(0)								
Zymbal gl	nodule		0	(0)	0	(0)	0	(0)	2	(18)								
bone	nodule		0	(0)	0	(0)	1	(7)	0	(0)								
mediastinum	mass		0	(0)	0	(0)	0	(0)	2	(18)								
peritoneum	nodule		1	(8)	0	(0)	0	(0)	1	(9)								

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 6

Organ	Findings	Group Name NO. of Animals	Control	25 ppm	50 ppm	100 ppm
			13 (%)	5 (%)	14 (%)	11 (%)
abdominal c	ascites		2 (15)	0 (0)	0 (0)	1 (9)
thoracic ca	pleural fluid		1 (8)	0 (0)	0 (0)	2 (18)
whole body	anemic		1 (8)	0 (0)	0 (0)	1 (9)

(HPT080)

BAIS 4

APPENDIX I 6

GROSS FINDINGS : FEMALE SACRIFICED ANIMALS

APPENDIX I 6

GROSS FINDINGS : FEMALE

SACRIFICED ANIMALS

STUDY NO. : 0535
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		25 ppm		50 ppm		100 ppm	
			37	(%)	45	(%)	36	(%)	39	(%)
subcutis	jaundice		0	(0)	0	(0)	1	(3)	0	(0)
	mass		4	(11)	9	(20)	6	(17)	7	(18)
lung	white zone		1	(3)	1	(2)	0	(0)	1	(3)
	nodule		0	(0)	1	(2)	0	(0)	1	(3)
spleen	enlarged		0	(0)	1	(2)	3	(8)	2	(5)
	nodule		0	(0)	0	(0)	1	(3)	0	(0)
forestomach	nodule		0	(0)	1	(2)	0	(0)	0	(0)
	ulcer		0	(0)	0	(0)	1	(3)	0	(0)
gl stomach	nodule		0	(0)	0	(0)	1	(3)	0	(0)
cecum	dilated		0	(0)	0	(0)	1	(3)	0	(0)
liver	white zone		2	(5)	1	(2)	0	(0)	1	(3)
	red zone		1	(3)	1	(2)	0	(0)	0	(0)
	nodule		0	(0)	0	(0)	0	(0)	1	(3)
	rough		2	(5)	2	(4)	2	(6)	2	(5)
	herniation		6	(16)	8	(18)	4	(11)	8	(21)
kidney	cyst		0	(0)	1	(2)	0	(0)	0	(0)
pituitary	enlarged		1	(3)	1	(2)	4	(11)	2	(5)
	red zone		3	(8)	6	(13)	1	(3)	3	(8)
	nodule		5	(14)	3	(7)	3	(8)	3	(8)
thyroid	enlarged		0	(0)	0	(0)	1	(3)	2	(5)
	nodule		0	(0)	1	(2)	1	(3)	1	(3)
adrenal	enlarged		1	(3)	0	(0)	1	(3)	0	(0)

STUDY NO. : 0535
 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		25 ppm		50 ppm		100 ppm	
			37	(%)	45	(%)	36	(%)	39	(%)
ovary	enlarged		0	(0)	1	(2)	0	(0)	0	(0)
	nodule		0	(0)	0	(0)	0	(0)	1	(3)
	cyst		2	(5)	5	(11)	2	(6)	1	(3)
uterus	black zone		0	(0)	0	(0)	1	(3)	0	(0)
	nodule		4	(11)	2	(4)	7	(19)	7	(18)
	cyst		0	(0)	1	(2)	0	(0)	0	(0)
eye	turbid		1	(3)	0	(0)	0	(0)	0	(0)
	white		4	(11)	4	(9)	5	(14)	0	(0)
peritoneum	nodule		0	(0)	1	(2)	0	(0)	1	(3)
	adhesion		0	(0)	0	(0)	1	(3)	0	(0)
thoracic ca	pleural fluid		0	(0)	0	(0)	0	(0)	1	(3)
other	eye lid:nodule		1	(3)	0	(0)	0	(0)	0	(0)
	upper jaw:nodule		1	(3)	0	(0)	0	(0)	0	(0)

APPENDIX J 1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0535
 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	44	382± 31	0.077±	0.011	3.467±	1.347	1.227±	0.125	1.434±	0.346	2.710±	0.320
25 ppm	44	380± 24	0.108±	0.203	3.718±	1.344	1.242±	0.117	1.339±	0.100	2.649±	0.214
50 ppm	42	367± 28*	0.081±	0.036	3.647±	1.509	1.211±	0.114	1.341±	0.152	2.704±	0.355
100 ppm	36	352± 29**	0.079±	0.029	4.003±	1.392	1.170±	0.113	1.332±	0.158	2.592±	0.199
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett												

(HCL040)

BAIS 4

STUDY NO. : 0535
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		BRAIN	
Control	44	1.253±	0.743	10.727±	1.801	2.074±	0.046
25 ppm	44	1.091±	0.478	10.903±	1.375	2.069±	0.043
50 ppm	42	1.256±	1.349	10.879±	1.498	2.057±	0.044
100 ppm	36	1.053±	0.358	10.537±	1.089	2.057±	0.046
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett							
(HCL040)							

BAIS 4

APPENDIX J 2

ORGAN WEIGHT, ABSOLUTE : FEMALE

STUDY NO. : 0535
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	37	240±	21	0.092±	0.086	0.152±	0.123	0.822±	0.063	0.922±	0.085	1.659±	0.098
25 ppm	45	252±	27	0.081±	0.008	0.240±	0.696	0.881±	0.073**	0.964±	0.130	1.741±	0.175
50 ppm	36	241±	27	0.082±	0.019	0.149±	0.125	0.878±	0.079**	0.989±	0.168	1.715±	0.139
100 ppm	39	238±	24	0.081±	0.010	0.129±	0.059	0.854±	0.076	0.994±	0.232	1.722±	0.148

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

Test of Dunnett

(HCL040)

BAIS 4

STUDY NO. : 0535
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	37	0.553±	0.250	6.101±	0.822	1.862±	0.034
25 ppm	45	0.681±	0.851	6.477±	0.992	1.873±	0.040
50 ppm	36	1.017±	1.818*	6.543±	0.876*	1.859±	0.042
100 ppm	39	0.995±	1.866*	6.711±	1.338*	1.866±	0.049

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

(HCL040)

BAIS 4

APPENDIX K 1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0535
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	44	382± 31	0.020± 0.003	0.910± 0.343	0.323± 0.043	0.379± 0.112	0.714± 0.112
25 ppm	44	380± 24	0.030± 0.064	0.983± 0.366	0.328± 0.034	0.354± 0.032	0.700± 0.062
50 ppm	42	367± 28*	0.023± 0.014	0.997± 0.419	0.332± 0.039	0.368± 0.054	0.740± 0.103
100 ppm	36	352± 29**	0.023± 0.008	1.134± 0.376	0.335± 0.044	0.381± 0.059	0.741± 0.079*

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0535
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	44	0.331± 0.207	2.801± 0.319	0.546± 0.042
25 ppm	44	0.288± 0.123	2.871± 0.296	0.547± 0.036
50 ppm	42	0.345± 0.384	2.972± 0.413	0.564± 0.047
100 ppm	36	0.301± 0.108	3.002± 0.298*	0.588± 0.051**
Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett				
(HCL042)				BAIS 4

APPENDIX K 2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0535
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	37	240± 21	0.038± 0.032	0.063± 0.048	0.343± 0.030	0.387± 0.050	0.693± 0.054
25 ppm	45	252± 27	0.032± 0.004	0.098± 0.290	0.351± 0.034	0.386± 0.076	0.693± 0.059
50 ppm	36	241± 27	0.034± 0.007	0.062± 0.049	0.368± 0.054*	0.418± 0.118	0.719± 0.091
100 ppm	39	238± 24	0.035± 0.007	0.054± 0.024	0.361± 0.045	0.422± 0.116	0.729± 0.096

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0535
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	37	0.232± 0.108	2.542± 0.295	0.780± 0.067
25 ppm	45	0.275± 0.363	2.569± 0.305	0.750± 0.075
50 ppm	36	0.473± 1.037	2.737± 0.452**	0.781± 0.088
100 ppm	39	0.432± 0.874	2.824± 0.505**	0.791± 0.082

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

Test of Dunnett

(HCL042)

BAIS 4

APPENDIX L 1

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE
ALL ANIMALS

STUDY NO. : 0535
 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				25 ppm				50 ppm				100 ppm			
			50				50				50				50			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Integumentary system/appandage}																		
skin/app			<50>				<50>				<50>				<50>			
	inflammation		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)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	squamous cell hyperplasia		0	0	0	0	1	1	0	0	0	1	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	epidermal cyst		1	0	0	0	0	0	0	0	0	2	0	0	0	2	0	0
			(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)
{Respiratory system}																		
nasal cavit			<50>				<50>				<50>				<50>			
	thrombus		1	0	0	0	1	0	0	0	0	0	0	0	3	0	0	0
			(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	goblet cell 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)
	eosinophilic change:olfactory epithelium		18	17	0	0	13	21	3	0	15	20	3	0	20	20	0	0
			(36)	(34)	(0)	(0)	(26)	(42)	(6)	(0)	(30)	(40)	(6)	(0)	(40)	(40)	(0)	(0)
	eosinophilic change:respiratory epithelium		4	0	0	0	6	0	0	0	4	0	0	0	3	0	0	0
			(8)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(8)	(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 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. : 0535
 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				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																					
nasal cavit		<50>				<50>				<50>				<50>				<50>			
	inflammation:foreign body	7 (14)	4 (8)	0 (0)	0 (0)	7 (14)	0 (0)	0 (0)	0 (0)	10 (20)	2 (4)	0 (0)	0 (0)	4 (8)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	inflammation:respiratory epithelium	0 (0)	1 (2)	0 (0)	0 (0)	1 (2)	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)
	respiratory metaplasia:olfactory epithelium	0 (0)	0 (0)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	3 (6)	0 (0)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	respiratory metaplasia:gland	7 (14)	0 (0)	0 (0)	0 (0)	5 (10)	0 (0)	0 (0)	0 (0)	5 (10)	0 (0)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	squamous cell metaplasia:respiratory epithelium	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)
nasopharynx		<50>				<50>				<50>				<50>				<50>			
	inflammation	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)	0 (0)	0 (0)	0 (0)	0 (0)
lung		<50>				<50>				<50>				<50>				<50>			
	congestion	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)	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. : 0535
 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				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																					
lung		<50>				<50>				<50>				<50>				<50>			
	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)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	inflammatory infiltration	0	0	0	0	1	0	0	0	1	0	0	0	1	0	0	0	2	0	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	granulomatous inflammation	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	accumulation of foamy cells	0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	bronchiolar-alveolar cell hyperplasia	3	2	0	0	2	1	0	0	0	0	0	0	0	0	0	0	4	0	0	0
		(6)	(4)	(0)	(0)	(4)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
{Hematopoietic system}																					
bone marrow		<50>				<50>				<50>				<50>				<50>			
	granulation	1	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	increased hematopoiesis	3	0	0	0	3	0	0	0	2	0	0	0	2	0	0	0	2	0	0	0
		(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(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

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCrI CrIj[F344/DuCrI]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 4

Organ	Findings	Group Name No. of Animals on Study				Control 50				25 ppm 50				50 ppm 50				100 ppm 50			
		Grade																			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																					
bone marrow		<50>				<50>				<50>				<50>				<50>			
	granulopoiesis:increased	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)	(2)	(0)	(0)	(0)
spleen		<50>				<50>				<50>				<50>				<50>			
	congestion	8	0	0	0	9	0	0	0	7	0	0	0	5	0	0	0	10	0	0	0
		(16)	(0)	(0)	(0)	(18)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(10)	(0)	(0)	(0)
	deposit of hemosiderin	19	2	0	0	21	1	0	0	20	3	0	0	20	5	0	0	40	10	0	0
		(38)	(4)	(0)	(0)	(42)	(2)	(0)	(0)	(40)	(6)	(0)	(0)	(40)	(10)	(0)	(0)	(40)	(10)	(0)	(0)
	granulation	0	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	fibrosis:focal	1	0	0	0	0	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	increased extramedullary hematopoiesis	5	2	1	0	5	4	0	0	2	3	0	0	4	5	1	0	8	10	2	0
		(10)	(4)	(2)	(0)	(10)	(8)	(0)	(0)	(4)	(6)	(0)	(0)	(8)	(10)	(2)	(0)	(8)	(10)	(2)	(0)
{Circulatory system}																					
heart		<50>				<50>				<50>				<50>				<50>			
	thrombus	0	1	0	0	0	0	0	0	1	0	0	0	0	1	0	0	0	2	0	0
		(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(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 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. : 0535
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				Control 50				25 ppm 50				50 ppm 50				100 ppm 50				
		Grade				1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Circulatory system}																						
heart		<50>				<50>				<50>				<50>				<50>				
	myocardial fibrosis	22 (44)	1 (2)	0 (0)	0 (0)	21 (42)	0 (0)	0 (0)	0 (0)	23 (46)	1 (2)	0 (0)	0 (0)	26 (52)	1 (2)	0 (0)	0 (0)	0 (0)				
	subendocardial fibrosis	1 (2)	0 (0)	0 (0)	0 (0)	1 (2)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)				
{Digestive system}																						
tooth		<50>				<50>				<50>				<50>				<50>				
	epidermal cyst	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)	0 (0)	0 (0)	0 (0)	0 (0)				
tongue		<50>				<50>				<50>				<50>				<50>				
	inflammation	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)				
	squamous cell hyperplasia	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)				
	arteritis	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)	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

STUDY NO. : 0535
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				Control				25 ppm				50 ppm				100 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}																					
stomach		<50>				<50>				<50>				<50>				<50>			
	ulcer:forestomach	0	1	0	0	2	0	0	0	1	0	0	0	3	0	0	0	3	0	0	0
		(0)	(2)	(0)	(0)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	hyperplasia:forestomach	0	0	0	0	1	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	erosion:glandular stomach	0	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0	2	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	ulcer:glandular stomach	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia:glandular stomach	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
liver		<50>				<50>				<50>				<50>				<50>			
	herniation	7	0	0	0	3	0	0	0	3	0	0	0	5	0	0	0	5	0	0	0
		(14)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(10)	(0)	(0)	(0)
	necrosis:central	0	0	0	0	1	1	0	0	0	0	0	0	1	1	0	0	2	2	0	0
		(0)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(2)	(2)	(0)	(0)
	necrosis:focal	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

STUDY NO. : 0535
 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 No. of Animals on Study				Control				25 ppm				50 ppm				100 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>			
	degeneration:central	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)	(2)	(0)	(0)
	granulation	1	1	0	0	3	0	0	0	4	0	0	0	0	0	0	0	0	0	0	0
		(2)	(2)	(0)	(0)	(6)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	inflammatory cell nest	0	0	0	0	1	1	0	0	0	0	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)	(0)
	clear cell focus	4	3	0	0	5	1	0	0	3	1	0	0	2	3	0	0	2	3	0	0
		(8)	(6)	(0)	(0)	(10)	(2)	(0)	(0)	(6)	(2)	(0)	(0)	(4)	(6)	(0)	(0)	(4)	(6)	(0)	(0)
	acidophilic cell focus	1	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)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	basophilic cell focus	1	1	0	0	1	1	0	0	4	2	0	0	0	1	0	0	0	1	0	0
		(2)	(2)	(0)	(0)	(2)	(2)	(0)	(0)	(8)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	mixed cell focus	0	0	0	0	0	0	0	0	1	1	0	0	0	0	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)
	spongiosis hepatitis	3	0	0	0	3	0	0	0	3	0	0	0	0	1	0	0	0	1	0	0
		(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(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 ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 8

		Group Name	Control				25 ppm				50 ppm				100 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		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
liver			<50>				<50>				<50>				<50>			
	bile duct hyperplasia		2	48	0	0	1	48	0	0	3	46	0	0	5	43	0	0
			(4)	(96)	(0)	(0)	(2)	(96)	(0)	(0)	(6)	(92)	(0)	(0)	(10)	(86)	(0)	(0)
pancreas			<50>				<50>				<50>				<50>			
	atrophy		17	6	0	0	18	6	0	0	17	5	0	0	11	5	0	0
			(34)	(12)	(0)	(0)	(36)	(12)	(0)	(0)	(34)	(10)	(0)	(0)	(22)	(10)	(0)	(0)
	arteritis		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)
	islet cell hyperplasia		0	3	0	0	0	1	0	0	1	0	0	0	0	1	0	0
			(0)	(6)	(0)	(0)	(0)	(2)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(2)	(0)	(0)
{Urinary system}																		
kidney			<50>				<50>				<50>				<50>			
	fatty change		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)
	cyst		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. : 0535
 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				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Urinary system}																					
kidney		<50>				<50>				<50>				<50>				<50>			
	scar	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)	1 (2)	0 (0)	0 (0)
	chronic nephropathy	15 (30)	29 (58)	0 (0)	0 (0)	21 (42)	25 (50)	1 (2)	0 (0)	10 (20)	30 (60)	4 (8)	0 (0)	21 (42)	23 (46)	2 (4)	0 (0)	21 (42)	23 (46)	2 (4)	0 (0)
	tubular necrosis	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)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	papillary necrosis	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)
	mineralization:papilla	1 (2)	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)	0 (0)	0 (0)	0 (0)	0 (0)
	mineralization:pelvis	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)	1 (2)	2 (4)	0 (0)	0 (0)
	transitional cell 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)	3 (6)	0 (0)	0 (0)	0 (0)	3 (6)	0 (0)	0 (0)	0 (0)
	atypical tubule hyperplasia	1 (2)	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)	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

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 10

Organ	Findings	Group Name No. of Animals on Study Grade	Control				25 ppm				50 ppm				100 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>				<49>				<49>			
	papillary hyperplasia:transitional epithelium		0	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)
{Endocrine system}																		
pituitary			<50>				<50>				<50>				<50>			
	angiectasis		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)
	cyst		1	0	0	0	0	0	0	0	2	0	0	0	1	0	0	0
			(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	hyperplasia		4	3	0	0	4	2	0	0	6	3	0	0	3	4	0	0
			(8)	(6)	(0)	(0)	(8)	(4)	(0)	(0)	(12)	(6)	(0)	(0)	(6)	(8)	(0)	(0)
	Rathke pouch		2	0	0	0	0	0	0	0	1	0	0	0	1	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
thyroid			<50>				<50>				<50>				<50>			
	C-cell hyperplasia		9	4	0	0	12	4	0	0	12	6	0	0	9	5	0	0
			(18)	(8)	(0)	(0)	(24)	(8)	(0)	(0)	(24)	(12)	(0)	(0)	(18)	(10)	(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. : 0535
 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				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Endocrine system}																					
adrenal																					
	hyperplasia:cortical cell	<50>				<50>				<50>				<50>				<50>			
		0	0	0	0	0	0	0	0	1	0	0	0	0	1	0	0	0	1	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	hyperplasia:medulla	3	4	0	0	7	0	0	0	4	0	0	0	2	0	0	0	2	0	0	0
		(6)	(8)	(0)	(0)	(14)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	focal fatty change:cortex	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
{Reproductive system}																					
testis																					
	mineralization	<50>				<50>				<50>				<50>				<50>			
		2	0	0	0	0	0	0	0	2	0	0	0	1	0	0	0	2	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	arteritis	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	interstitial cell hyperplasia	6	0	0	0	7	0	0	0	9	0	0	0	3	0	0	0	6	0	0	0
		(12)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(18)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
prostate																					
	inflammation	<50>				<50>				<49>				<49>				<49>			
		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)	(2)	(0)	(0)	(0)	(2)	(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. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 12

		Group Name	Control				25 ppm				50 ppm				100 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
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
prostate	hyperplasia		<50>				<50>				<49>				<49>			
			8	1	0	0	7	2	0	0	9	1	0	0	7	0	0	0
			(16)	(2)	(0)	(0)	(14)	(4)	(0)	(0)	(18)	(2)	(0)	(0)	(14)	(0)	(0)	(0)
mammary gl	galactoceles		<50>				<50>				<50>				<50>			
			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)
{Special sense organs/appendage}																		
eye	hemorrhage		<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)
	cataract		2	0	0	0	4	0	0	0	3	0	0	0	2	0	0	0
			(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	retinal atrophy		18	1	0	0	16	4	0	0	13	5	0	0	12	1	0	0
			(36)	(2)	(0)	(0)	(32)	(8)	(0)	(0)	(26)	(10)	(0)	(0)	(24)	(2)	(0)	(0)
	keratitis		1	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
			(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(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 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. : 0535
 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				Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study				50				50				50				50			
		Grade				1 2 3 4				1 2 3 4				1 2 3 4				1 2 3 4			
		(%) (%) (%) (%)				(%) (%) (%) (%)				(%) (%) (%) (%)				(%) (%) (%) (%)				(%) (%) (%) (%)			

{Special sense organs/appendage}

eye	squamous cell metaplasia:cornea	<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)

{Musculoskeletal system}

muscle	mineralization	<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)

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

(HPT150)

BAIS4

APPENDIX L 2

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

STUDY NO. : 0535
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	Control				25 ppm				50 ppm				100 ppm			
		Group Name				6				8				14			
		No. of Animals on Study				Grade				Grade				Grade			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																	
nasal cavit	thrombus	< 6>				< 6>				< 8>				<14>			
		0	0	0	0	1	0	0	0	0	0	0	0	3	0	0	0
		(0)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(21)	(0)	(0)	(0)
	eosinophilic change:olfactory epithelium	1	2	0	0	0	2	0	0	0	1	0	0	3	4	0	0
		(17)	(33)	(0)	(0)	(0)	(33)	(0)	(0)	(0)	(13)	(0)	(0)	(21)	(29)	(0)	(0)
	eosinophilic change:respiratory epithelium	0	0	0	0	1	0	0	0	1	0	0	0	1	0	0	0
		(0)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(13)	(0)	(0)	(0)	(7)	(0)	(0)	(0)
	inflammation:foreign body	0	1	0	0	1	0	0	0	1	0	0	0	1	1	0	0
		(0)	(17)	(0)	(0)	(17)	(0)	(0)	(0)	(13)	(0)	(0)	(0)	(7)	(7)	(0)	(0)
	respiratory metaplasia:olfactory epithelium	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(13)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
nasopharynx	inflammation	< 6>				< 6>				< 8>				<14>			
		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)	(7)	(0)	(0)
lung	congestion	< 6>				< 6>				< 8>				<14>			
		0	0	0	0	1	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)

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. : 0535
 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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	6				6				8				14			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
lung			< 6>				< 6>				< 8>				<14>			
	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)	(13)	(0)	(0)	(0)	(0)	(0)	(0)
	inflammatory infiltration		0	0	0	0	1	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(0)	(0)	(0)
	bronchiolar-alveolar 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)	(7)	(0)	(0)	(0)
{Hematopoietic system}																		
bone marrow			< 6>				< 6>				< 8>				<14>			
	increased hematopoiesis		1	0	0	0	2	0	0	0	1	0	0	0	1	0	0	0
			(17)	(0)	(0)	(0)	(33)	(0)	(0)	(0)	(13)	(0)	(0)	(0)	(7)	(0)	(0)	(0)
	granulopoiesis:increased		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)	(7)	(0)	(0)	(0)
spleen			< 6>				< 6>				< 8>				<14>			
	deposit of hemosiderin		1	2	0	0	0	1	0	0	0	3	0	0	2	4	0	0
			(17)	(33)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(38)	(0)	(0)	(14)	(29)	(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. : 0535
 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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	6				6				8				14			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																		
spleen			< 6>				< 6>				< 8>				<14>			
	increased extramedullary hematopoiesis		1	0	0	0	1	0	0	0	0	1	0	0	2	4	0	0
			(17)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(0)	(13)	(0)	(0)	(14)	(29)	(0)	(0)
{Circulatory system}																		
heart			< 6>				< 6>				< 8>				<14>			
	thrombus		0	0	0	0	0	0	0	0	1	0	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(13)	(0)	(0)	(0)	(0)	(7)	(0)	(0)
	myocardial fibrosis		3	0	0	0	2	0	0	0	4	0	0	0	8	0	0	0
			(50)	(0)	(0)	(0)	(33)	(0)	(0)	(0)	(50)	(0)	(0)	(0)	(57)	(0)	(0)	(0)
{Digestive system}																		
tongue			< 6>				< 6>				< 8>				<14>			
	inflammation		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)	(7)	(0)	(0)	(0)
	squamous cell hyperplasia		0	0	0	0	1	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)

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. : 0535
 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	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	6				6				8				14			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)

{Digestive system}

stomach

ulcer:forestomach

0100

(0) (17) (0) (0)

2000

(33) (0) (0) (0)

1000

(13) (0) (0) (0)

3000

(21) (0) (0) (0)

hyperplasia:forestomach

0000

(0) (0) (0) (0)

1000

(17) (0) (0) (0)

1000

(13) (0) (0) (0)

0000

(0) (0) (0) (0)

erosion:glandular stomach

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

2000

(14) (0) (0) (0)

ulcer:glandular stomach

1000

(17) (0) (0) (0)

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

liver

herniation

1000

(17) (0) (0) (0)

1000

(17) (0) (0) (0)

0000

(0) (0) (0) (0)

2000

(14) (0) (0) (0)

necrosis:central

0000

(0) (0) (0) (0)

1100

(17) (17) (0) (0)

0000

(0) (0) (0) (0)

1100

(7) (7) (0) (0)

degeneration:central

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

0000

(0) (0) (0) (0)

0100

(0) (7) (0) (0)

bile duct hyperplasia

2400

(33) (67) (0) (0)

1400

(17) (67) (0) (0)

2500

(25) (63) (0) (0)

3900

(21) (64) (0) (0)

Grade 1 : Slight 2 : Moderate 3 : Marked 4 : Severe
 < a > a : Number of animals examined at the site
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 (c) c : b / a * 100
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 SEX : MALE

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

PAGE : 5

Organ	Findings	Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	6				6				8				14			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
pancreas			< 6>				< 6>				< 8>				<14>			
	atrophy		0	1	0	0	1	1	0	0	1	1	0	0	1	1	0	0
			(0)	(17)	(0)	(0)	(17)	(17)	(0)	(0)	(13)	(13)	(0)	(0)	(7)	(7)	(0)	(0)
	islet cell 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)	(13)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Urinary system}																		
kidney			< 6>				< 6>				< 8>				<14>			
	scar		1	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)	(0)
	chronic nephropathy		2	0	0	0	5	0	0	0	2	0	1	0	8	2	0	0
			(33)	(0)	(0)	(0)	(83)	(0)	(0)	(0)	(25)	(0)	(13)	(0)	(57)	(14)	(0)	(0)
	tubular necrosis		0	0	0	0	0	1	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)
	mineralization:pelvis		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)	(7)	(7)	(0)	(0)
	transitional cell hyperplasia		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)	(14)	(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. : 0535
 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 6				25 ppm 6				50 ppm 8				100 ppm 14			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
<hr/>																		
{Urinary system}																		
urin bladd			< 6>				< 6>				< 7>				<14>			
	papillary hyperplasia:transitional epithelium		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)	(7)	(0)	(0)
{Endocrine system}																		
pituitary			< 6>				< 6>				< 8>				<14>			
	hyperplasia		0	0	0	0	0	1	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(17)	(0)	(0)	(13)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
thyroid			< 6>				< 6>				< 8>				<14>			
	C-cell hyperplasia		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)	(7)	(7)	(0)	(0)
adrenal			< 6>				< 6>				< 8>				<14>			
	hyperplasia:medulla		0	0	0	0	2	0	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(33)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Reproductive system}																		
testis			< 6>				< 6>				< 8>				<14>			
	mineralization		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)	(7)	(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. : 0535
 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

Organ	Findings	Group Name No. of Animals on Study				Control 6				25 ppm 6				50 ppm 8				100 ppm 14				
		Grade				1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																						
testis		< 6>				< 6>				< 8>				<14>								
	interstitial cell hyperplasia	0 (0)	0 (0)	0 (0)	0 (0)	2 (33)	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 (7)	0 (0)	0 (0)	0 (0)
{Special sense organs/appendage}																						
eye		< 6>				< 6>				< 8>				<14>								
	hemorrhage	1 (17)	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)	0 (0)
	cataract	1 (17)	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 (7)	0 (0)	0 (0)	0 (0)
	retinal atrophy	0 (0)	1 (17)	0 (0)	0 (0)	1 (17)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (13)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (7)	0 (0)	0 (0)	0 (0)
	keratitis	1 (17)	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 (7)	0 (0)	0 (0)
	squamous cell metaplasia:cornea	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 (7)	0 (0)	0 (0)	0 (0)
{Musculoskeletal system}																						
muscle		< 6>				< 6>				< 8>				<14>								
	mineralization	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 (13)	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

APPENDIX L 3

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE
SACRIFICED ANIMALS

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : MALE

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

PAGE : 1

Organ_____	Findings_____	Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	44				44				42				36			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Integumentary system/appandage}																		
skin/app			<44>				<44>				<42>				<36>			
	inflammation		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)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	squamous cell hyperplasia		0	0	0	0	1	1	0	0	0	1	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(3)	(0)	(0)
	epidermal cyst		1	0	0	0	0	0	0	0	0	2	0	0	0	2	0	0
			(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(5)	(0)	(0)	(0)	(6)	(0)	(0)
{Respiratory system}																		
nasal cavit			<44>				<44>				<42>				<36>			
	thrombus		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)
	goblet cell 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)
	eosinophilic change:olfactory epithelium		17	15	0	0	13	19	3	0	15	19	3	0	17	16	0	0
			(39)	(34)	(0)	(0)	(30)	(43)	(7)	(0)	(36)	(45)	(7)	(0)	(47)	(44)	(0)	(0)
	eosinophilic change:respiratory epithelium		4	0	0	0	5	0	0	0	3	0	0	0	2	0	0	0
			(9)	(0)	(0)	(0)	(11)	(0)	(0)	(0)	(7)	(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 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. : 0535
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 No. of Animals on Study				Control 44				25 ppm 44				50 ppm 42				100 ppm 36			
		Grade																			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																					
nasal cavit		<44>				<44>				<42>				<36>							
	inflammation:foreign body	7 (16)	3 (7)	0 (0)	0 (0)	6 (14)	0 (0)	0 (0)	0 (0)	9 (21)	2 (5)	0 (0)	0 (0)	3 (8)	0 (0)	0 (0)	0 (0)				
	inflammation:respiratory epithelium	0 (0)	1 (2)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (3)	0 (0)	0 (0)	0 (0)				
	respiratory metaplasia:olfactory epithelium	0 (0)	0 (0)	0 (0)	0 (0)	1 (2)	0 (0)	0 (0)	0 (0)	2 (5)	0 (0)	0 (0)	0 (0)	1 (3)	0 (0)	0 (0)	0 (0)				
	respiratory metaplasia:gland	7 (16)	0 (0)	0 (0)	0 (0)	5 (11)	0 (0)	0 (0)	0 (0)	5 (12)	0 (0)	0 (0)	0 (0)	1 (3)	0 (0)	0 (0)	0 (0)				
	squamous cell metaplasia:respiratory epithelium	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)				
lung		<44>				<44>				<42>				<36>							
	inflammatory infiltration	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 (3)	0 (0)	0 (0)	0 (0)				
	granulomatous inflammation	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)				
	accumulation of foamy cells	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)				

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. : 0535
 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				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	44				44				42				36			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
lung			<44>				<44>				<42>				<36>			
	bronchiolar-alveolar cell hyperplasia		3	2	0	0	2	1	0	0	0	0	0	0	3	0	0	0
			(7)	(5)	(0)	(0)	(5)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
{Hematopoietic system}																		
bone marrow			<44>				<44>				<42>				<36>			
	granulation		1	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
			(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	increased hematopoiesis		2	0	0	0	1	0	0	0	1	0	0	0	1	0	0	0
			(5)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
spleen			<44>				<44>				<42>				<36>			
	congestion		8	0	0	0	9	0	0	0	7	0	0	0	5	0	0	0
			(18)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(17)	(0)	(0)	(0)	(14)	(0)	(0)	(0)
	deposit of hemosiderin		18	0	0	0	21	0	0	0	20	0	0	0	18	1	0	0
			(41)	(0)	(0)	(0)	(48)	(0)	(0)	(0)	(48)	(0)	(0)	(0)	(50)	(3)	(0)	(0)
	granulation		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)	(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 ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Chi Square

STUDY NO. : 0535
 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 44				25 ppm 44				50 ppm 42				100 ppm 36			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4				
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)				
{Hematopoietic system}																					
spleen		<44>				<44>				<42>				<36>							
	fibrosis:focal	1 (2)	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)				
	increased extramedullary hematopoiesis	4 (9)	2 (5)	1 (2)	0 (0)	4 (9)	4 (9)	0 (0)	0 (0)	2 (5)	2 (5)	0 (0)	0 (0)	2 (6)	1 (3)	1 (3)	0 (0)				
{Circulatory system}																					
heart		<44>				<44>				<42>				<36>							
	thrombus	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)				
	myocardial fibrosis	19 (43)	1 (2)	0 (0)	0 (0)	19 (43)	0 (0)	0 (0)	0 (0)	19 (45)	1 (2)	0 (0)	0 (0)	18 (50)	1 (3)	0 (0)	0 (0)				
	subendocardial fibrosis	1 (2)	0 (0)	0 (0)	0 (0)	1 (2)	1 (2)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (3)	0 (0)	0 (0)	0 (0)				
{Digestive system}																					
tooth		<44>				<44>				<42>				<36>							
	epidermal cyst	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)	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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 5

Organ	Findings	Control				25 ppm				50 ppm				100 ppm			
		Group Name				44				42				36			
		No. of Animals on Study				Grade				Grade				Grade			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																	
tongue		<44>				<44>				<42>				<36>			
	arteritis	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)	(0)	(0)	(0)	(0)
stomach		<44>				<44>				<42>				<36>			
	hyperplasia:glandular stomach	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)
liver		<44>				<44>				<42>				<36>			
	herniation	6	0	0	0	2	0	0	0	3	0	0	0	3	0	0	0
		(14)	(0)	(0)	(0)	(5)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
	necrosis:focal	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)
	granulation	1	1	0	0	3	0	0	0	4	0	0	0	0	0	0	0
		(2)	(2)	(0)	(0)	(7)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	inflammatory cell nest	0	0	0	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)
	clear cell focus	4	3	0	0	5	1	0	0	3	1	0	0	2	3	0	0
		(9)	(7)	(0)	(0)	(11)	(2)	(0)	(0)	(7)	(2)	(0)	(0)	(6)	(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. : 0535
 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				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	44				44				42				36			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
liver			<44>				<44>				<42>				<36>			
	acidophilic cell focus		1	0	0	0	1	0	0	0	1	0	0	0	0	0	0	0
			(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	basophilic cell focus		1	1	0	0	1	1	0	0	4	2	0	0	0	1	0	0
			(2)	(2)	(0)	(0)	(2)	(2)	(0)	(0)	(10)	(5)	(0)	(0)	(0)	(3)	(0)	(0)
	mixed cell focus		0	0	0	0	0	0	0	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)
	spongiosis hepatitis		3	0	0	0	3	0	0	0	3	0	0	0	0	1	0	0
			(7)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(0)	(3)	(0)	(0)
	bile duct hyperplasia		0	44	0	0	0	44	0	0	1	41	0	0	2	34	0	0
			(0)	(100)	(0)	(0)	(0)	(100)	(0)	(0)	(2)	(98)	(0)	(0)	(6)	(94)	(0)	(0)
pancreas			<44>				<44>				<42>				<36>			
	atrophy		17	5	0	0	17	5	0	0	16	4	0	0	10	4	0	0
			(39)	(11)	(0)	(0)	(39)	(11)	(0)	(0)	(38)	(10)	(0)	(0)	(28)	(11)	(0)	(0)
	arteritis		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)
	islet cell hyperplasia		0	3	0	0	0	1	0	0	0	0	0	0	0	1	0	0
			(0)	(7)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 7

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	44				44				42				36			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Urinary system}																		
kidney			<44>				<44>				<42>				<36>			
	fatty change		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)
	cyst		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)
	scar		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)
	chronic nephropathy		13 (30)	29 (66)	0 (0)	0 (0)	16 (36)	25 (57)	1 (2)	0 (0)	8 (19)	30 (71)	3 (7)	0 (0)	13 (36)	21 (58)	2 (6)	0 (0)
	papillary necrosis		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)
	mineralization:papilla		1 (2)	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)
	mineralization:pelvis		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)	1 (3)	0 (0)	0 (0)
transitional cell 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)	1 (3)	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. : 0535
 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				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	44				44				42				36			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Urinary system}																		
kidney			<44>				<44>				<42>				<36>			
	atypical tubule hyperplasia	1	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
urin bladd			<44>				<44>				<42>				<35>			
	papillary hyperplasia:transitional epithelium	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Endocrine system}																		
pituitary			<44>				<44>				<42>				<36>			
	angiectasis	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)	(3)	(0)	(0)	(0)
	cyst	1	0	0	0	0	0	0	0	0	2	0	0	0	1	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(5)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
	hyperplasia	4	3	0	0	4	1	0	0	5	3	0	0	3	4	0	0	
		(9)	(7)	(0)	(0)	(9)	(2)	(0)	(0)	(12)	(7)	(0)	(0)	(8)	(11)	(0)	(0)	
	Rathke pouch	2	0	0	0	0	0	0	0	1	0	0	0	1	0	0	0	
		(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 9

Organ	Findings	Control				25 ppm				50 ppm				100 ppm			
		44				44				42				36			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Endocrine system}																	
thyroid		<44>				<44>				<42>				<36>			
	C-cell hyperplasia	9	4	0	0	12	4	0	0	12	6	0	0	8	4	0	0
		(20)	(9)	(0)	(0)	(27)	(9)	(0)	(0)	(29)	(14)	(0)	(0)	(22)	(11)	(0)	(0)
adrenal		<44>				<44>				<42>				<36>			
	hyperplasia:cortical cell	0	0	0	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)	(3)	(0)	(0)
	hyperplasia:medulla	3	4	0	0	5	0	0	0	4	0	0	0	2	0	0	0
		(7)	(9)	(0)	(0)	(11)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	focal fatty change:cortex	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)
{Reproductive system}																	
testis		<44>				<44>				<42>				<36>			
	mineralization	2	0	0	0	0	0	0	0	2	0	0	0	0	0	0	0
		(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	arteritis	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 10

Organ	Findings	Control				25 ppm				50 ppm				100 ppm			
		44				44				42				36			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																	
testis		<44>				<44>				<42>				<36>			
	interstitial cell hyperplasia	6	0	0	0	5	0	0	0	9	0	0	0	2	0	0	0
		(14)	(0)	(0)	(0)	(11)	(0)	(0)	(0)	(21)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
prostate		<44>				<44>				<42>				<35>			
	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)	(2)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia	8	1	0	0	7	2	0	0	9	1	0	0	7	0	0	0
		(18)	(2)	(0)	(0)	(16)	(5)	(0)	(0)	(21)	(2)	(0)	(0)	(20)	(0)	(0)	(0)
mammary gl		<44>				<44>				<42>				<36>			
	galactoceles	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)
{Special sense organs/appendage}																	
eye		<44>				<44>				<42>				<36>			
	cataract	1	0	0	0	4	0	0	0	3	0	0	0	1	0	0	0
		(2)	(0)	(0)	(0)	(9)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
	retinal atrophy	18	0	0	0	15	4	0	0	13	4	0	0	11	1	0	0
		(41)	(0)	(0)	(0)	(34)	(9)	(0)	(0)	(31)	(10)	(0)	(0)	(31)	(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

APPENDIX L 4

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : FEMALE
ALL ANIMALS

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 14

Organ	Findings	Group Name No. of Animals on Study				Control 50				25 ppm 50				50 ppm 50				100 ppm 50			
		Grade																			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Integumentary system/appandage}																					
subcutis		<50>				<50>				<50>				<50>				<50>			
	adhesion	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	inflammation	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)	(2)	(0)	(0)
{Respiratory system}																					
nasal cavit		<50>				<50>				<50>				<50>				<50>			
	thrombus	1	0	0	0	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	goblet cell hyperplasia	1	0	0	0	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	eosinophilic change:olfactory epithelium	6	32	11	0	2	38	8	0	6	29	9	0	4	35	7	0	8	70	14	0
		(12)	(64)	(22)	(0)	(4)	(76)	(16)	(0)	(12)	(58)	(18)	(0)	(8)	(70)	(14)	(0)	(8)	(70)	(14)	(0)
	eosinophilic change:respiratory epithelium	11	0	0	0	14	0	0	0	13	0	0	0	8	0	0	0	16	0	0	0
		(22)	(0)	(0)	(0)	(28)	(0)	(0)	(0)	(26)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
	inflammation:foreign body	2	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(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 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. : 0535
 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				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																					
nasal cavit		<50>				<50>				<50>				<50>				<50>			
	inflammation:respiratory epithelium	0	1	0	0	0	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)
	respiratory metaplasia:gland	3	0	0	0	3	0	0	0	5	0	0	0	3	0	0	0	3	0	0	0
		(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
{Hematopoietic system}																					
larynx		<50>				<50>				<50>				<50>				<50>			
	inflammation	0	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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
lung		<50>				<50>				<50>				<50>				<50>			
	inflammatory infiltration	1	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0	2	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(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)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Hematopoietic system}																					
bone marrow		<50>				<50>				<50>				<50>				<50>			
	granulation	4	1	0	0	4	0	0	0	6	1	0	0	2	1	0	0	4	2	0	0
		(8)	(2)	(0)	(0)	(8)	(0)	(0)	(0)	(12)	(2)	(0)	(0)	(4)	(2)	(0)	(0)	(4)	(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. : 0535
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				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																					
bone marrow		<50>				<50>				<50>				<50>				<50>			
	increased hematopoiesis	3	0	0	0	2	0	0	0	6	0	0	0	3	0	0	0	3	0	0	0
		(6)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	granulopoiesis:increased	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
spleen		<50>				<50>				<50>				<50>				<50>			
	congestion	2	0	0	0	4	0	0	0	3	0	0	0	3	0	0	0	3	0	0	0
		(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	deposit of hemosiderin	11	26	0	0	17	24	0	0	15	17	0	0	8	24	0	0	8	24	0	0
		(22)	(52)	(0)	(0)	(34)	(48)	(0)	(0)	(30)	(34)	(0)	(0)	(16)	(48)	(0)	(0)	(16)	(48)	(0)	(0)
	increased extramedullary hematopoiesis	7	2	3	0	5	1	0	0	5	5	1	0	8	3	1	0	8	3	1	0
		(14)	(4)	(6)	(0)	(10)	(2)	(0)	(0)	(10)	(10)	(2)	(0)	(16)	(6)	(2)	(0)	(16)	(6)	(2)	(0)
{Circulatory system}																					
heart		<50>				<50>				<50>				<50>				<50>			
	thrombus	1	1	0	0	0	0	0	0	0	0	0	0	1	2	0	0	1	2	0	0
		(2)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(4)	(0)	(0)	(2)	(4)	(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. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 17

Organ	Findings	Group Name No. of Animals on Study				Control				25 ppm				50 ppm				100 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
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Circulatory system}																					
heart		<50>				<50>				<50>				<50>				<50>			
	inflammatory cell nest	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
		(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	myocardial fibrosis	12	0	0	0	8	0	0	0	7	0	0	0	8	0	0	0	8	0	0	0
		(24)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
	subendocardial fibrosis	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Digestive system}																					
stomach		<50>				<50>				<50>				<50>				<50>			
	erosion:forestomach	0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0	2	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	ulcer:forestomach	2	2	0	0	0	1	0	0	1	2	0	0	1	3	0	0	2	6	0	0
		(4)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(2)	(4)	(0)	(0)	(2)	(6)	(0)	(0)	(2)	(6)	(0)	(0)
	hyperplasia:forestomach	2	1	0	0	1	2	0	0	1	0	0	0	0	1	0	0	0	2	0	0
		(4)	(2)	(0)	(0)	(2)	(4)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	erosion:glandular stomach	1	0	0	0	0	0	0	0	0	0	0	0	3	0	0	0	6	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(6)	(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 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. : 0535
 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				Control				25 ppm				50 ppm				100 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}																					
stomach	ulcer:glandular stomach	<50>				0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia:glandular stomach	<50>				0	0	0	0	0	0	0	0	2	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
large intes	inflammation	<50>				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)	(2)	(0)	(0)	(0)	(0)	(0)	(0)
liver	herniation	<50>				11	0	0	0	9	0	0	0	7	0	0	0	9	0	0	0
		(22)	(0)	(0)	(0)	(22)	(0)	(0)	(0)	(18)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(18)	(0)	(0)	(0)
	necrosis:central	<50>				1	0	0	0	0	0	0	0	1	1	0	0	1	0	0	0
		(2)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(2)	(0)	(0)	(0)
	necrosis:focal	<50>				0	0	0	0	0	1	0	0	0	0	0	0	0	1	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)
	fatty change	<50>				2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 19

		Group Name	Control				25 ppm				50 ppm				100 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	
{Digestive system}																			
liver			<50>				<50>				<50>				<50>				
	degeneration: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)	(2)	(0)	(0)	(0)	
	granulation		8	3	0	0	8	5	0	0	6	3	0	0	8	4	0	0	
			(16)	(6)	(0)	(0)	(16)	(10)	(0)	(0)	(12)	(6)	(0)	(0)	(16)	(8)	(0)	(0)	
	fibrosis		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)	
	clear cell focus		2	0	0	0	0	1	0	0	0	0	0	0	0	1	0	0	
		(4)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)		
	acidophilic 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)	(2)	(0)	(0)	
	basophilic cell focus		23	3	0	0	20	1	0	0	18	1	0	0	21	1	0	0	
			(46)	(6)	(0)	(0)	(40)	(2)	(0)	(0)	(36)	(2)	(0)	(0)	(42)	(2)	(0)	(0)	
	bile duct hyperplasia		8	1	0	0	4	7	0	0	7	4	0	0	3	3	0	0	
			(16)	(2)	(0)	(0)	(8)	(14)	(0)	(0)	(14)	(8)	(0)	(0)	(6)	(6)	(0)	(0)	
	pancreas			<50>				<50>				<50>				<50>			
		atrophy		1	4	0	0	4	0	0	0	3	1	0	0	5	0	0	0 *
			(2)	(8)	(0)	(0)	(8)	(0)	(0)	(0)	(6)	(2)	(0)	(0)	(10)	(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. : 0535
 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 Grade	Control				25 ppm				50 ppm				100 ppm			
			50				50				50				50			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
pancreas	islet cell hyperplasia		<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)
{Urinary system}																		
kidney	cyst		<50>				<50>				<50>				<50>			
			0	0	0	0	0	1	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)
	deposit of hemosiderin		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 cell nest		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)
	chronic nephropathy		12	2	0	0	17	2	0	0	18	1	0	0	18	3	0	0
			(24)	(4)	(0)	(0)	(34)	(4)	(0)	(0)	(36)	(2)	(0)	(0)	(36)	(6)	(0)	(0)
	tubular necrosis		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)
	mineralization:papilla		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)	(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. : 0535
 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	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study				50				50				50			
		Grade				1				1				1			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Urinary system}																	
kidney	mineralization:pelvis	<50>				<50>				<50>				<50>			
		2	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	desquamation:pelvis	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)
	transitional 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)
	atypical tubule hyperplasia	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)	(4)	(0)	(0)	(0)
	dilated pelvis	2	0	0	0	0	0	0	0	0	2	0	0	0	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)
urin bladd	nodular hyperplasia:transitional epithelium	<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)
{Endocrine system}																	
pituitary	angiectasis	<50>				<50>				<50>				<50>			
		3	0	0	0	0	1	0	0	0	0	0	0	2	0	0	0
		(6)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 22

Organ	Findings	Group Name No. of Animals on Study				Control 50				25 ppm 50				50 ppm 50				100 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	cyst	<50>				<50>				<50>				<50>				<50>			
		4	0	0	0	3	0	0	0	1	0	0	0	3	0	0	0	3	0	0	0
		(8)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(6)	(0)	(0)	(0)
	hyperplasia	7	4	0	0	6	2	0	0	5	4	0	0	9	5	0	0	9	5	0	0
		(14)	(8)	(0)	(0)	(12)	(4)	(0)	(0)	(10)	(8)	(0)	(0)	(18)	(10)	(0)	(0)	(18)	(10)	(0)	(0)
	Rathke pouch	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
thyroid	C-cell hyperplasia	<50>				<50>				<50>				<50>				<50>			
		7	5	0	0	5	5	0	0	6	3	0	0	5	3	0	0	5	3	0	0
		(14)	(10)	(0)	(0)	(10)	(10)	(0)	(0)	(12)	(6)	(0)	(0)	(10)	(6)	(0)	(0)	(10)	(6)	(0)	(0)
adrenal	hyperplasia:cortical cell	<50>				<50>				<50>				<50>				<50>			
		0	0	0	0	1	0	0	0	2	0	0	0	0	1	0	0	0	1	0	0
		(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(0)	(0)
	hyperplasia:medulla	0	0	0	0	0	1	0	0	0	0	0	0	1	2	0	0	1	2	0	0
		(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(2)	(4)	(0)	(0)	(2)	(4)	(0)	(0)
	focal fatty change:cortex	4	0	0	0	1	1	0	0	4	1	0	0	4	0	0	0	4	0	0	0
		(8)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(8)	(2)	(0)	(0)	(8)	(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. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 23

		Group Name	Control				25 ppm				50 ppm				100 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
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
ovary	cyst		<50>				<50>				<50>				<50>			
		2	0	0	0	6	0	0	0	2	0	0	0	1	0	0	0	
		(4)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	
uterus	fibrosis		<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)	
	hyperplasia:epithelium		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)	(4)	(0)	(0)
		hyperplasia:gland	0	0	0	0	1	0	0	0	1	1	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(2)	(0)	(0)	(0)
		cystic endometrial hyperplasia	1	0	0	0	4	0	0	0	0	0	0	0	4	0	0	0
			(2)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
mammary gl	galactocoele		<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)
{Special sense organs/appendage}																		
eye	cataract		<50>				<50>				<50>				<50>			
		5	0	0	0	4	0	0	0	7	0	0	0	0	0	0	0	
		(10)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(14)	(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. : 0535
 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	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study				50				50				50			
		Grade				1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Special sense organs/appendage}																	
eye	retinal atrophy	14 (28)	5 (10)	0 (0)	0 (0)	15 (30)	3 (6)	0 (0)	0 (0)	8 (16)	5 (10)	0 (0)	0 (0)	11 (22)	0 (0)	0 (0)	0 * (0)
nasolacr d	inflammation	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)
{Musculoskeletal system}																	
muscle	mineralization	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	2 (4)	0 (0)	0 (0)	0 (0)
bone	osteosclerosis	4 (8)	0 (0)	0 (0)	0 (0)	4 (8)	3 (6)	0 (0)	0 (0)	4 (8)	1 (2)	0 (0)	0 (0)	5 (10)	1 (2)	0 (0)	0 (0)
{Body cavities}																	
mediastinum	inflammatory infiltration	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)	1 (2)	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. : 0535
 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				25 ppm				50 ppm				100 ppm			
			50				50				50				50			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)

{Body cavities}

peritoneum	adhesion	<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)

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

APPENDIX L 5

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

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 8

Organ_____	Findings_____	Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
<hr/>																		
{Integumentary system/appandage}																		
subcutis																		
			<13>				< 5>				<14>				<11>			
adhesion			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)
inflammation			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)	(9)	(0)	(0)
{Respiratory system}																		
nasal cavit																		
			<13>				< 5>				<14>				<11>			
thrombus			1	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(0)	(0)	(0)	(0)
eosinophilic change:olfactory epithelium			4	8	1	0	0	4	0	0	2	5	1	0	1	5	1	0
			(31)	(62)	(8)	(0)	(0)	(80)	(0)	(0)	(14)	(36)	(7)	(0)	(9)	(45)	(9)	(0)
eosinophilic change:respiratory epithelium			1	0	0	0	1	0	0	0	2	0	0	0	0	0	0	0
			(8)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
inflammation:respiratory epithelium			0	1	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
respiratory metaplasia:gland			0	0	0	0	1	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(7)	(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. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 9

Organ	Findings	Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
larynx			<13>				< 5>				<14>				<11>			
	inflammation		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)	(9)	(0)	(0)	(0)	
lung			<13>				< 5>				<14>				<11>			
	inflammatory infiltration		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)	(9)	(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)	(7)	(0)	(0)	(0)	(0)	(0)	(0)	
{Hematopoietic system}																		
bone marrow			<13>				< 5>				<14>				<11>			
	increased hematopoiesis		3	0	0	0	0	0	0	0	5	0	0	0	2	0	0	0
			(23)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(36)	(0)	(0)	(0)	(18)	(0)	(0)	
	granulopoiesis:increased		1	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)		
spleen			<13>				< 5>				<14>				<11>			
	deposit of hemosiderin		1	3	0	0	0	3	0	0	4	2	0	0	0	3	0	0
			(8)	(23)	(0)	(0)	(0)	(60)	(0)	(0)	(29)	(14)	(0)	(0)	(0)	(27)	(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. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 10

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																		
spleen			<13>				< 5>				<14>				<11>			
	increased extramedullary hematopoiesis		4	1	3	0	0	0	0	0	0	4	1	0 *	0	2	1	0
			(31)	(8)	(23)	(0)	(0)	(0)	(0)	(0)	(0)	(29)	(7)	(0)	(0)	(18)	(9)	(0)
{Circulatory system}																		
heart			<13>				< 5>				<14>				<11>			
	thrombus		1	1	0	0	0	0	0	0	0	0	0	0	1	2	0	0
			(8)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(18)	(0)	(0)
	inflammatory cell nest		0	1	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
	myocardial fibrosis		3	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(23)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
{Digestive system}																		
stomach			<13>				< 5>				<14>				<11>			
	erosion:forestomach		0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(18)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 11

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
stomach			<13>				< 5>				<14>				<11>			
	ulcer:forestomach		2	2	0	0	0	1	0	0	1	1	0	0	1	3	0	0
			(15)	(15)	(0)	(0)	(0)	(20)	(0)	(0)	(7)	(7)	(0)	(0)	(9)	(27)	(0)	(0)
	hyperplasia:forestomach		2	1	0	0	1	0	0	0	0	0	0	0	0	1	0	0
			(15)	(8)	(0)	(0)	(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(9)	(0)	(0)
	erosion:glandular stomach		1	0	0	0	0	0	0	0	0	0	0	0	3	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(27)	(0)	(0)	(0)
	ulcer:glandular stomach		0	0	0	0	1	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)
	hyperplasia: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)	(7)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
liver			<13>				< 5>				<14>				<11>			
	herniation		5	0	0	0	1	0	0	0	4	0	0	0	1	0	0	0
			(38)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(29)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
	necrosis:central		1	0	0	0	0	0	0	0	1	1	0	0	1	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(7)	(0)	(0)	(9)	(0)	(0)	(0)
	necrosis:focal		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)	(9)	(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. : 0535
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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
liver			<13>				< 5>				<14>				<11>			
	fatty change		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)
	degeneration: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)	(9)	(0)	(0)	(0)
	basophilic cell focus		2	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
			(15)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	bile duct hyperplasia		2	0	0	0	1	1	0	0	1	1	0	0	1	2	0	0
			(15)	(0)	(0)	(0)	(20)	(20)	(0)	(0)	(7)	(7)	(0)	(0)	(9)	(18)	(0)	(0)
pancreas			<13>				< 5>				<14>				<11>			
	atrophy		0	1	0	0	1	0	0	0	0	0	0	0	2	0	0	0
			(0)	(8)	(0)	(0)	(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(18)	(0)	(0)	(0)
{Urinary system}																		
kidney			<13>				< 5>				<14>				<11>			
	deposit of hemosiderin		0	0	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)

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. : 0535
 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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
Organ_____	Findings_____		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
<hr/>																		
{Urinary system}																		
kidney																		
			<13>				< 5>				<14>				<11>			
	inflammatory cell nest		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)
	chronic nephropathy		0	1	0	0	1	0	0	0	1	1	0	0	0	0	0	0
			(0)	(8)	(0)	(0)	(20)	(0)	(0)	(0)	(7)	(7)	(0)	(0)	(0)	(0)	(0)	(0)
	tubular necrosis		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)	(9)	(0)	(0)	(0)
	mineralization:pelvis		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)	(9)	(0)	(0)	(0)
	transitional cell hyperplasia		0	0	0	0	0	1	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)
	dilated pelvis		2	0	0	0	0	0	0	0	0	2	0	0	0	0	0	0
			(15)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(0)	(0)	(0)
urin bladd																		
			<13>				< 5>				<14>				<11>			
	nodular hyperplasia:transitional epithelium		0	0	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)
{Endocrine system}																		
pituitary																		
			<13>				< 5>				<14>				<11>			
	angiectasis		2	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0
			(15)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(18)	(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. : 0535
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	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Endocrine system}																		
pituitary	cyst		<13>				< 5>				<14>				<11>			
			1	0	0	0	1	0	0	0	1	0	0	0	1	0	0	0
			(8)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
	hyperplasia		0	1	0	0	0	0	0	0	1	0	0	0	1	1	0	0
			(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(9)	(9)	(0)	(0)
	Rathke pouch		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)
thyroid	C-cell hyperplasia		<13>				< 5>				<14>				<11>			
			0	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(0)	(0)	(0)
adrenal	hyperplasia:cortical cell		<13>				< 5>				<14>				<11>			
			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)	(9)	(0)	(0)
	focal fatty change:cortex		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)
{Reproductive system}																		
ovary	cyst		<13>				< 5>				<14>				<11>			
			0	0	0	0	1	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 : Number of animals with lesion
(c) c : b / a * 100
Significant difference ; * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0535
 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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
uterus			<13>				< 5>				<14>				<11>			
	cystic endometrial hyperplasia		0	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(18)	(0)	(0)	(0)
{Special sense organs/appendage}																		
eye			<13>				< 5>				<14>				<11>			
	cataract		0	0	0	0	1	0	0	0	2	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(14)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	retinal atrophy		0	0	0	0	0	1	0	0	0	1	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(0)	(0)	(0)
nasolacr d			<13>				< 5>				<14>				<11>			
	inflammation		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)
{Musculoskeletal system}																		
muscle			<13>				< 5>				<14>				<11>			
	mineralization		0	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(18)	(0)	(0)	(0)

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. : 0535
 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

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	13				5				14				11			
Organ	Findings	Grade	<u>1</u>	<u>2</u>	<u>3</u>	<u>4</u>	<u>1</u>	<u>2</u>	<u>3</u>	<u>4</u>	<u>1</u>	<u>2</u>	<u>3</u>	<u>4</u>	<u>1</u>	<u>2</u>	<u>3</u>	<u>4</u>
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Musculoskeletal system}																		
bone			<13>				< 5>				<14>				<11>			
	osteosclerosis		1	0	0	0	0	0	0	0	1	0	0	0	1	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(9)	(0)	(0)	(0)
{Body cavities}																		
mediastinum			<13>				< 5>				<14>				<11>			
	inflammatory infiltration		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)	(9)	(9)	(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

APPENDIX L 6

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : FEMALE
SACRIFICED ANIMALS

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 11

Organ	Findings	Group Name No. of Animals on Study Grade	Control 37				25 ppm 45				50 ppm 36				100 ppm 39			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
nasal cavit			<37>				<45>				<36>				<39>			
	goblet cell hyperplasia		1	0	0	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)	(3)	(0)	(0)	(0)
	eosinophilic change:olfactory epithelium		2	24	10	0	2	34	8	0	4	24	8	0	3	30	6	0
			(5)	(65)	(27)	(0)	(4)	(76)	(18)	(0)	(11)	(67)	(22)	(0)	(8)	(77)	(15)	(0)
lung	eosinophilic change:respiratory epithelium		10	0	0	0	13	0	0	0	11	0	0	0	8	0	0	0
			(27)	(0)	(0)	(0)	(29)	(0)	(0)	(0)	(31)	(0)	(0)	(0)	(21)	(0)	(0)	(0)
	inflammation:foreign body		2	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
	respiratory metaplasia:gland		3	0	0	0	2	0	0	0	4	0	0	0	3	0	0	0
			(8)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(11)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
bone marrow			<37>				<45>				<36>				<39>			
	inflammatory infiltration		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)
	granulation		4	1	0	0	4	0	0	0	6	1	0	0	2	1	0	0
			(11)	(3)	(0)	(0)	(9)	(0)	(0)	(0)	(17)	(3)	(0)	(0)	(5)	(3)	(0)	(0)
{Hematopoietic system}																		
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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 12

Organ	Findings	Control				25 ppm				50 ppm				100 ppm			
		Group Name				37				45				36			
		No. of Animals on Study				Grade				Grade				Grade			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																	
bone marrow		<37>				<45>				<36>				<39>			
	increased hematopoiesis	0	0	0	0	2	0	0	0	1	0	0	0	1	0	0	0
		(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(3)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
spleen		<37>				<45>				<36>				<39>			
	congestion	2	0	0	0	4	0	0	0	3	0	0	0	3	0	0	0
		(5)	(0)	(0)	(0)	(9)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
	deposit of hemosiderin	10	23	0	0	17	21	0	0	11	15	0	0	8	21	0	0
		(27)	(62)	(0)	(0)	(38)	(47)	(0)	(0)	(31)	(42)	(0)	(0)	(21)	(54)	(0)	(0)
	increased extramedullary hematopoiesis	3	1	0	0	5	1	0	0	5	1	0	0	8	1	0	0
		(8)	(3)	(0)	(0)	(11)	(2)	(0)	(0)	(14)	(3)	(0)	(0)	(21)	(3)	(0)	(0)
{Circulatory system}																	
heart		<37>				<45>				<36>				<39>			
	myocardial fibrosis	9	0	0	0	8	0	0	0	7	0	0	0	7	0	0	0
		(24)	(0)	(0)	(0)	(18)	(0)	(0)	(0)	(19)	(0)	(0)	(0)	(18)	(0)	(0)	(0)
	subendocardial fibrosis	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. : 0535
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				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	37				45				36				39			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
stomach			<37>				<45>				<36>				<39>			
	ulcer:forestomach		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)	(0)	(0)
	hyperplasia:forestomach		0	0	0	0	0	2	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(3)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia:glandular stomach		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
large intes			<37>				<45>				<36>				<39>			
	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)	(3)	(0)	(0)	(0)	(0)	(0)	(0)
liver			<37>				<45>				<36>				<39>			
	herniation		6	0	0	0	8	0	0	0	3	0	0	0	8	0	0	0
			(16)	(0)	(0)	(0)	(18)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(21)	(0)	(0)	(0)
	necrosis:focal		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)
	fatty change		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)

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. : 0535
 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				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	37				45				36				39			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
liver			<37>				<45>				<36>				<39>			
	granulation		8	3	0	0	8	5	0	0	6	3	0	0	8	4	0	0
			(22)	(8)	(0)	(0)	(18)	(11)	(0)	(0)	(17)	(8)	(0)	(0)	(21)	(10)	(0)	(0)
	fibrosis		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)	(0)	(0)	(0)
	clear cell focus		2	0	0	0	0	1	0	0	0	0	0	0	0	1	0	0
		(5)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	
	acidophilic 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)	(3)	(0)	(0)
	basophilic cell focus		21	3	0	0	19	1	0	0	18	1	0	0	21	1	0	0
			(57)	(8)	(0)	(0)	(42)	(2)	(0)	(0)	(50)	(3)	(0)	(0)	(54)	(3)	(0)	(0)
	bile duct hyperplasia		6	1	0	0	3	6	0	0	6	3	0	0	2	1	0	0
			(16)	(3)	(0)	(0)	(7)	(13)	(0)	(0)	(17)	(8)	(0)	(0)	(5)	(3)	(0)	(0)
pancreas			<37>				<45>				<36>				<39>			
	atrophy		1	3	0	0	3	0	0	0	3	1	0	0	3	0	0	0
			(3)	(8)	(0)	(0)	(7)	(0)	(0)	(0)	(8)	(3)	(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)

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. : 0535
 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	Control				25 ppm				50 ppm				100 ppm				
		No. of Animals on Study	37				45				36				39				
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	
<hr/>																			
{Urinary system}																			
kidney			<37>				<45>				<36>				<39>				
	cyst		0	0	0	0	0	1	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	(0)
	chronic nephropathy		12	1	0	0	16	2	0	0	17	0	0	0	18	3	0	0	
			(32)	(3)	(0)	(0)	(36)	(4)	(0)	(0)	(47)	(0)	(0)	(0)	(46)	(8)	(0)	(0)	
	mineralization:papilla		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)	(0)	(0)	(0)	(0)	
mineralization:pelvis		2	0	0	0	0	0	0	0	1	0	0	0	1	0	0	0		
		(5)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	(0)	(3)	(0)	(0)	(0)		
desquamation:pelvis		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)		
atypical tubule hyperplasia		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)	(5)	(0)	(0)	(0)		

{Endocrine system}

pituitary			<37>				<45>				<36>				<39>							
	angiectasis		1	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0				
			(3)	(0)	(0)	(0)	(0)	(2)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 16

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	37				45				36				39			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
{Endocrine system}																		
pituitary			<37>				<45>				<36>				<39>			
	cyst		3	0	0	0	2	0	0	0	0	0	0	0	2	0	0	0
			(8)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(5)	(0)	(0)	(0)
	hyperplasia		7	3	0	0	6	2	0	0	4	4	0	0	8	4	0	0
			(19)	(8)	(0)	(0)	(13)	(4)	(0)	(0)	(11)	(11)	(0)	(0)	(21)	(10)	(0)	(0)
thyroid			<37>				<45>				<36>				<39>			
	C-cell hyperplasia		7	5	0	0	5	5	0	0	6	2	0	0	5	3	0	0
			(19)	(14)	(0)	(0)	(11)	(11)	(0)	(0)	(17)	(6)	(0)	(0)	(13)	(8)	(0)	(0)
adrenal			<37>				<45>				<36>				<39>			
	hyperplasia:cortical cell		0	0	0	0	1	0	0	0	2	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(6)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia:medulla		0	0	0	0	0	1	0	0	0	0	0	0	1	2	0	0
			(0)	(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(5)	(0)	(0)
	focal fatty change:cortex		3	0	0	0	1	1	0	0	4	1	0	0	4	0	0	0
			(8)	(0)	(0)	(0)	(2)	(2)	(0)	(0)	(11)	(3)	(0)	(0)	(10)	(0)	(0)	(0)
{Reproductive system}																		
ovary			<37>				<45>				<36>				<39>			
	cyst		2	0	0	0	5	0	0	0	2	0	0	0	1	0	0	0
			(5)	(0)	(0)	(0)	(11)	(0)	(0)	(0)	(6)	(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 ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Chi Square

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 17

		Group Name	Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study	37				45				36				39			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
uterus			<37>				<45>				<36>				<39>			
	fibrosis		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)	(0)	(0)	(0)
	hyperplasia:epithelium		0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(3)	(0)	(0)	(0)	(5)	(0)	(0)	(0)
	hyperplasia:gland		0	0	0	0	1	0	0	0	1	1	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(2)	(0)	(0)	(0)	(3)	(3)	(0)	(0)	(3)	(0)	(0)	(0)
	cystic endometrial hyperplasia		1	0	0	0	4	0	0	0	0	0	0	0	2	0	0	0
			(3)	(0)	(0)	(0)	(9)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(5)	(0)	(0)	(0)
mammary gl			<37>				<45>				<36>				<39>			
	galactoceles		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)
{Special sense organs/appendage}																		
eye			<37>				<45>				<36>				<39>			
	cataract		5	0	0	0	3	0	0	0	5	0	0	0	0	0	0	0
		(14)	(0)	(0)	(0)	(7)	(0)	(0)	(0)	(14)	(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. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 18

Organ_____	Findings_____	Group Name				Control				25 ppm				50 ppm				100 ppm			
		No. of Animals on Study				37				45				36				39			
		Grade																			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4				
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)			

{Special sense organs/appendage}

eye	retinal atrophy	<37>				<45>				<36>				<39>			
		14	5	0	0	15	2	0	0	8	4	0	0	11	0	0	0 *
		(38)	(14)	(0)	(0)	(33)	(4)	(0)	(0)	(22)	(11)	(0)	(0)	(28)	(0)	(0)	(0)

{Musculoskeletal system}

bone	osteosclerosis	<37>				<45>				<36>				<39>			
		3	0	0	0	4	3	0	0	3	1	0	0	4	1	0	0
		(8)	(0)	(0)	(0)	(9)	(7)	(0)	(0)	(8)	(3)	(0)	(0)	(10)	(3)	(0)	(0)

{Body cavities}

peritoneum	adhesion	<37>				<45>				<36>				<39>			
		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)	(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

(HPT150)

BAIS4

APPENDIX M 1

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

STUDY NO. : 0535
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	25 ppm	50 ppm	100 ppm
0 - 52	NO. OF EXAMINED ANIMALS		1	0	1	0
	NO. OF ANIMALS WITH TUMORS		1	0	1	0
	NO. OF ANIMALS WITH SINGLE TUMORS		1	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		1	0	1	0
	NO. OF TOTAL TUMORS		1	0	1	0
53 - 78	NO. OF EXAMINED ANIMALS		1	1	2	0
	NO. OF ANIMALS WITH TUMORS		1	1	2	0
	NO. OF ANIMALS WITH SINGLE TUMORS		1	1	1	0
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	1	0
	NO. OF BENIGN TUMORS		0	0	1	0
	NO. OF MALIGNANT TUMORS		1	1	2	0
	NO. OF TOTAL TUMORS		1	1	3	0
79 - 104	NO. OF EXAMINED ANIMALS		4	5	5	14
	NO. OF ANIMALS WITH TUMORS		4	5	5	14
	NO. OF ANIMALS WITH SINGLE TUMORS		1	0	1	2
	NO. OF ANIMALS WITH MULTIPLE TUMORS		3	5	4	12
	NO. OF BENIGN TUMORS		4	6	7	18
	NO. OF MALIGNANT TUMORS		3	4	3	9
	NO. OF TOTAL TUMORS		7	10	10	27
105 - 105	NO. OF EXAMINED ANIMALS		44	44	42	36
	NO. OF ANIMALS WITH TUMORS		44	44	41	36
	NO. OF ANIMALS WITH SINGLE TUMORS		16	21	17	15
	NO. OF ANIMALS WITH MULTIPLE TUMORS		28	23	24	21
	NO. OF BENIGN TUMORS		77	73	67	63
	NO. OF MALIGNANT TUMORS		9	7	8	4
	NO. OF TOTAL TUMORS		86	80	75	67

STUDY NO. : 0535
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	25 ppm	50 ppm	100 ppm
0 - 105	NO. OF EXAMINED ANIMALS		50	50	50	50
	NO. OF ANIMALS WITH TUMORS		50	50	49	50
	NO. OF ANIMALS WITH SINGLE TUMORS		19	22	20	17
	NO. OF ANIMALS WITH MULTIPLE TUMORS		31	28	29	33
	NO. OF BENIGN TUMORS		81	79	75	81
	NO. OF MALIGNANT TUMORS		14	12	14	13
	NO. OF TOTAL TUMORS		95	91	89	94
(HPT070)						BAIS4

APPENDIX M 2

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

STUDY NO. : 0535
 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	25 ppm	50 ppm	100 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	0	4	2
	NO. OF ANIMALS WITH TUMORS		1	0	3	2
	NO. OF ANIMALS WITH SINGLE TUMORS		1	0	3	2
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	0	0
	NO. OF BENIGN TUMORS		0	0	0	1
	NO. OF MALIGNANT TUMORS		1	0	3	1
	NO. OF TOTAL TUMORS		1	0	3	2
79 - 104	NO. OF EXAMINED ANIMALS		12	5	10	9
	NO. OF ANIMALS WITH TUMORS		12	5	9	9
	NO. OF ANIMALS WITH SINGLE TUMORS		6	3	8	5
	NO. OF ANIMALS WITH MULTIPLE TUMORS		6	2	1	4
	NO. OF BENIGN TUMORS		9	4	2	8
	NO. OF MALIGNANT TUMORS		10	3	8	6
	NO. OF TOTAL TUMORS		19	7	10	14
105 - 105	NO. OF EXAMINED ANIMALS		37	45	36	39
	NO. OF ANIMALS WITH TUMORS		22	25	26	26
	NO. OF ANIMALS WITH SINGLE TUMORS		17	12	15	12
	NO. OF ANIMALS WITH MULTIPLE TUMORS		5	13	11	14
	NO. OF BENIGN TUMORS		27	34	32	37
	NO. OF MALIGNANT TUMORS		5	5	9	7
	NO. OF TOTAL TUMORS		32	39	41	44

STUDY NO. : 0535
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	25 ppm	50 ppm	100 ppm
0 - 105	NO. OF EXAMINED ANIMALS		50	50	50	50
	NO. OF ANIMALS WITH TUMORS		35	30	38	37
	NO. OF ANIMALS WITH SINGLE TUMORS		24	15	26	19
	NO. OF ANIMALS WITH MULTIPLE TUMORS		11	15	12	18
	NO. OF BENIGN TUMORS		36	38	34	46
	NO. OF MALIGNANT TUMORS		16	8	20	14
	NO. OF TOTAL TUMORS		52	46	54	60
(HPT070)			BAIS4			

APPENDIX N 1

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : MALE

STUDY NO. : 0535
 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	25 ppm 50	50 ppm 50	100 ppm 50
{Integumentary system/appandage}						
skin/app			<50>	<50>	<50>	<50>
	trichoepithelioma		1 (2%)	1 (2%)	0 (0%)	0 (0%)
	keratoacanthoma		1 (2%)	0 (0%)	0 (0%)	2 (4%)
	trichoepithelioma:malignant		1 (2%)	1 (2%)	1 (2%)	2 (4%)
subcutis			<50>	<50>	<50>	<50>
	fibroma		1 (2%)	8 (16%)	2 (4%)	3 (6%)
	leiomyoma		0 (0%)	0 (0%)	0 (0%)	1 (2%)
	fibrosarcoma		1 (2%)	0 (0%)	1 (2%)	2 (4%)
{Respiratory system}						
lung			<50>	<50>	<50>	<50>
	bronchiolar-alveolar adenoma		6 (12%)	3 (6%)	2 (4%)	4 (8%)
	bronchiolar-alveolar carcinoma		1 (2%)	0 (0%)	0 (0%)	0 (0%)
{Hematopoietic system}						
bone marrow			<50>	<50>	<50>	<50>
	xanthoma		0 (0%)	0 (0%)	0 (0%)	1 (2%)
thymus			<50>	<50>	<50>	<50>
	thymoma:benign		0 (0%)	0 (0%)	1 (2%)	0 (0%)
spleen			<50>	<50>	<50>	<50>
	mononuclear cell leukemia		5 (10%)	5 (10%)	6 (12%)	4 (8%)

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

STUDY NO. : 0535
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	25 ppm 50	50 ppm 50	100 ppm 50
{Digestive system}						
oral cavity			<50>	<50>	<50>	<50>
	squamous cell papilloma		0 (0%)	0 (0%)	2 (4%)	0 (0%)
salivary gl			<50>	<50>	<50>	<50>
	adenoma		0 (0%)	1 (2%)	0 (0%)	0 (0%)
stomach			<50>	<50>	<50>	<50>
	squamous cell papilloma		1 (2%)	0 (0%)	0 (0%)	0 (0%)
	squamous cell carcinoma		0 (0%)	0 (0%)	1 (2%)	0 (0%)
	osteosarcoma		0 (0%)	1 (2%)	0 (0%)	0 (0%)
liver			<50>	<50>	<50>	<50>
	hepatocellular adenoma		0 (0%)	0 (0%)	1 (2%)	1 (2%)
	hepatocellular carcinoma		1 (2%)	1 (2%)	0 (0%)	0 (0%)
pancreas			<50>	<50>	<50>	<50>
	islet cell adenoma		3 (6%)	3 (6%)	2 (4%)	0 (0%)
{Urinary system}						
kidney			<50>	<50>	<50>	<50>
	renal cell adenoma		1 (2%)	0 (0%)	0 (0%)	0 (0%)
	hemangiosarcoma		0 (0%)	0 (0%)	1 (2%)	0 (0%)
{Endocrine system}						
pituitary			<50>	<50>	<50>	<50>
	adenoma		6 (12%)	4 (8%)	7 (14%)	8 (16%)

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

STUDY NO. : 0535
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	25 ppm 50	50 ppm 50	100 ppm 50
{Endocrine system}						
thyroid			<50>	<50>	<50>	<50>
	C-cell adenoma		3 (6%)	6 (12%)	8 (16%)	6 (12%)
	follicular adenoma		0 (0%)	0 (0%)	1 (2%)	0 (0%)
	C-cell carcinoma		1 (2%)	1 (2%)	1 (2%)	0 (0%)
adrenal			<50>	<50>	<50>	<50>
	pheochromocytoma		2 (4%)	2 (4%)	3 (6%)	3 (6%)
	cortical adenoma		1 (2%)	0 (0%)	0 (0%)	0 (0%)
	pheochromocytoma:malignant		0 (0%)	1 (2%)	1 (2%)	1 (2%)
{Reproductive system}						
testis			<50>	<50>	<50>	<50>
	interstitial cell tumor		47 (94%)	49 (98%)	44 (88%)	48 (96%)
mammary gl			<50>	<50>	<50>	<50>
	adenoma		0 (0%)	1 (2%)	0 (0%)	0 (0%)
	fibroadenoma		3 (6%)	0 (0%)	0 (0%)	2 (4%)
prep/cli gl			<50>	<50>	<50>	<50>
	adenoma		3 (6%)	1 (2%)	1 (2%)	2 (4%)
{Nervous system}						
brain			<50>	<50>	<50>	<50>
	malignant reticulosis		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. : 0535
 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	25 ppm 50	50 ppm 50	100 ppm 50
{Nervous system}						
brain	glioma		<50> 1 (2%)	<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)
periph nerv	schwannoma:malignant		<50> 1 (2%)	<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)
{Special sense organs/appendage}						
Zymbal gl	squamous cell carcinoma		<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)	<50> 2 (4%)
{Musculoskeletal system}						
bone	osteoma		<50> 2 (4%)	<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)
	osteosarcoma		1 (2%)	0 (0%)	0 (0%)	0 (0%)
vertebra	chordoma		<50> 0 (0%)	<50> 0 (0%)	<50> 1 (2%)	<50> 0 (0%)
{Body cavities}						
peritoneum	mesothelioma		<50> 1 (2%)	<50> 2 (4%)	<50> 1 (2%)	<50> 2 (4%)

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

(HPT085)

BAIS4

APPENDIX N 2

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : FEMALE

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 5

Organ	Findings	Group Name No. of animals on Study	Control 50	25 ppm 50	50 ppm 50	100 ppm 50
{Integumentary system/appandage}						
subcutis	fibroma		<50> 1 (2%)	<50> 1 (2%)	<50> 0 (0%)	<50> 2 (4%)
{Respiratory system}						
lung	bronchiolar-alveolar adenoma		<50> 0 (0%)	<50> 2 (4%)	<50> 0 (0%)	<50> 1 (2%)
	bronchiolar-alveolar carcinoma		0 (0%)	1 (2%)	0 (0%)	1 (2%)
{Hematopoietic system}						
spleen	mononuclear cell leukemia		<50> 8 (16%)	<50> 4 (8%)	<50> 9 (18%)	<50> 9 (18%)
{Digestive system}						
oral cavity	squamous cell papilloma		<50> 1 (2%)	<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)
liver	hepatocellular adenoma		<50> 2 (4%)	<50> 1 (2%)	<50> 0 (0%)	<50> 1 (2%)
pancreas	islet cell adenoma		<50> 2 (4%)	<50> 1 (2%)	<50> 0 (0%)	<50> 1 (2%)
{Urinary system}						
kidney	mesenchymoma:malignant		<50> 0 (0%)	<50> 0 (0%)	<50> 0 (0%)	<50> 1 (2%)
urin bladd	transitional cell papilloma		<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. : 0535
 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	25 ppm 50	50 ppm 50	100 ppm 50
{Endocrine system}						
pituitary	adenoma		<50> 11 (22%)	<50> 12 (24%)	<50> 10 (20%)	<50> 8 (16%)
	adenocarcinoma		2 (4%)	0 (0%)	1 (2%)	1 (2%)
thyroid	C-cell adenoma		<50> 2 (4%)	<50> 2 (4%)	<50> 1 (2%)	<50> 4 (8%)
	follicular adenoma		0 (0%)	0 (0%)	0 (0%)	1 (2%)
	C-cell carcinoma		0 (0%)	0 (0%)	2 (4%)	0 (0%)
adrenal	pheochromocytoma		<50> 0 (0%)	<50> 1 (2%)	<50> 2 (4%)	<50> 2 (4%)
	cortical adenoma		1 (2%)	0 (0%)	0 (0%)	1 (2%)
	pheochromocytoma:malignant		1 (2%)	0 (0%)	0 (0%)	0 (0%)
{Reproductive system}						
ovary	hemangioma		<50> 0 (0%)	<50> 0 (0%)	<50> 1 (2%)	<50> 0 (0%)
	sertoli cell tumor		0 (0%)	0 (0%)	0 (0%)	1 (2%)
	adenocarcinoma		0 (0%)	1 (2%)	0 (0%)	0 (0%)
uterus	endometrial stromal polyp		<50> 11 (22%)	<50> 10 (20%)	<50> 12 (24%)	<50> 14 (28%)

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

(HPT085)

BAIS4

STUDY NO. : 0535
 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	25 ppm 50	50 ppm 50	100 ppm 50
{Reproductive system}						
uterus	adenocarcinoma		<50> 1 (2%)	<50> 0 (0%)	<50> 2 (4%)	<50> 0 (0%)
	endometrial stromal sarcoma		4 (8%)	0 (0%)	2 (4%)	2 (4%)
mammary gl	fibroadenoma		<50> 4 (8%)	<50> 8 (16%)	<50> 6 (12%)	<50> 6 (12%)
	adenocarcinoma		0 (0%)	1 (2%)	2 (4%)	0 (0%)
prep/cli gl	adenoma		<50> 1 (2%)	<50> 0 (0%)	<50> 1 (2%)	<50> 2 (4%)
{Nervous system}						
brain	glioma		<50> 0 (0%)	<50> 1 (2%)	<50> 1 (2%)	<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%)
{Musculoskeletal system}						
bone	osteosarcoma		<50> 0 (0%)	<50> 0 (0%)	<50> 1 (2%)	<50> 0 (0%)
{Body cavities}						
peritoneum	hemangioma		<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

APPENDIX O 1

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS : MALE

STUDY No. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 1

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : subcutis TUMOR : fibroma				
Tumor rate				
Overall rates(a)	1/50(2.0)	8/50(16.0)	2/50(4.0)	3/50(6.0)
Adjusted rates(b)	2.27	15.91	4.65	5.56
Terminal rates(c)	1/44(2.3)	7/44(15.9)	1/42(2.4)	2/36(5.6)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.2228			
Prevalence method(d)	P = 0.5369			
Combined analysis(d)	P = 0.4100			
Cochran-Armitage test(e)	P = 0.9254			
Fisher Exact test(e)		P = 0.0154*	P = 0.5000	P = 0.3087
SITE : subcutis TUMOR : fibroma, fibrosarcoma				
Tumor rate				
Overall rates(a)	2/50(4.0)	8/50(16.0)	3/50(6.0)	5/50(10.0)
Adjusted rates(b)	2.27	15.91	6.98	8.33
Terminal rates(c)	1/44(2.3)	7/44(15.9)	2/42(4.8)	3/36(8.3)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.2293			
Prevalence method(d)	P = 0.3321			
Combined analysis(d)	P = 0.2295			
Cochran-Armitage test(e)	P = 0.6762			
Fisher Exact test(e)		P = 0.0458*	P = 0.5000	P = 0.2180
SITE : lung TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	6/50(12.0)	3/50(6.0)	2/50(4.0)	4/50(8.0)
Adjusted rates(b)	13.64	6.82	4.76	8.89
Terminal rates(c)	6/44(13.6)	3/44(6.8)	2/42(4.8)	3/36(8.3)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.6691			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.5552			
Fisher Exact test(e)		P = 0.2435	P = 0.1343	P = 0.3703

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 2

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : lung TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	7/50(14.0)	3/50(6.0)	2/50(4.0)	4/50(8.0)
Adjusted rates(b)	15.91	6.82	4.76	8.89
Terminal rates(c)	7/44(15.9)	3/44(6.8)	2/42(4.8)	3/36(8.3)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.7652			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.3782			
Fisher Exact test(e)		P = 0.1589	P = 0.0798	P = 0.2623
SITE : spleen TUMOR : mononuclear cell leukemia				
Tumor rate				
Overall rates(a)	5/50(10.0)	5/50(10.0)	6/50(12.0)	4/50(8.0)
Adjusted rates(b)	6.82	4.55	7.14	2.78
Terminal rates(c)	3/44(6.8)	2/44(4.5)	3/42(7.1)	1/36(2.8)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.3216			
Prevalence method(d)	P = 0.7340			
Combined analysis(d)	P = 0.5261			
Cochran-Armitage test(e)	P = 0.7499			
Fisher Exact test(e)		P = 0.6297	P = 0.5000	P = 0.5000
SITE : pancreas TUMOR : islet cell adenoma				
Tumor rate				
Overall rates(a)	3/50(6.0)	3/50(6.0)	2/50(4.0)	0/50(0.0)
Adjusted rates(b)	6.82	6.82	4.76	0.0
Terminal rates(c)	3/44(6.8)	3/44(6.8)	2/42(4.8)	0/36(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.9434			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.0877			
Fisher Exact test(e)		P = 0.6611	P = 0.5000	P = 0.1212

STUDY No. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 3

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : pituitary gland TUMOR : adenoma				
Tumor rate				
Overall rates(a)	6/50(12.0)	4/50(8.0)	7/50(14.0)	8/50(16.0)
Adjusted rates(b)	13.64	9.09	13.04	16.67
Terminal rates(c)	6/44(13.6)	4/44(9.1)	5/42(11.9)	6/36(16.7)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.0381*			
Prevalence method(d)	P = 0.3060			
Combined analysis(d)	P = 0.1240			
Cochran-Armitage test(e)	P = 0.3663			
Fisher Exact test(e)		P = 0.3703	P = 0.5000	P = 0.3871
SITE : thyroid TUMOR : C-cell adenoma				
Tumor rate				
Overall rates(a)	3/50(6.0)	6/50(12.0)	8/50(16.0)	6/50(12.0)
Adjusted rates(b)	6.82	13.64	19.05	16.67
Terminal rates(c)	3/44(6.8)	6/44(13.6)	8/42(19.0)	6/36(16.7)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.0999			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.3889			
Fisher Exact test(e)		P = 0.2435	P = 0.0999	P = 0.2435
SITE : thyroid TUMOR : C-cell adenoma, C-cell carcinoma				
Tumor rate				
Overall rates(a)	4/50(8.0)	7/50(14.0)	9/50(18.0)	6/50(12.0)
Adjusted rates(b)	9.09	15.91	19.05	16.67
Terminal rates(c)	4/44(9.1)	7/44(15.9)	8/42(19.0)	6/36(16.7)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.4016			
Prevalence method(d)	P = 0.1821			
Combined analysis(d)	P = 0.1784			
Cochran-Armitage test(e)	P = 0.6188			
Fisher Exact test(e)		P = 0.2623	P = 0.1168	P = 0.3703

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 4

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : adrenal gland TUMOR : pheochromocytoma				
Tumor rate				
Overall rates(a)	2/50(4.0)	2/50(4.0)	3/50(6.0)	3/50(6.0)
Adjusted rates(b)	4.55	4.55	6.67	6.38
Terminal rates(c)	2/44(4.5)	2/44(4.5)	2/42(4.8)	2/36(5.6)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.2914			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.5834			
Fisher Exact test(e)		P = 0.6913	P = 0.5000	P = 0.5000
SITE : adrenal gland TUMOR : pheochromocytoma, pheochromocytoma:malignant				
Tumor rate				
Overall rates(a)	2/50(4.0)	3/50(6.0)	4/50(8.0)	4/50(8.0)
Adjusted rates(b)	4.55	6.82	8.89	6.38
Terminal rates(c)	2/44(4.5)	3/44(6.8)	3/42(7.1)	2/36(5.6)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.1118			
Prevalence method(d)	P = 0.3306			
Combined analysis(d)	P = 0.2008			
Cochran-Armitage test(e)	P = 0.4098			
Fisher Exact test(e)		P = 0.5000	P = 0.3389	P = 0.3389
SITE : testis TUMOR : interstitial cell tumor				
Tumor rate				
Overall rates(a)	47/50(94.0)	49/50(98.0)	44/50(88.0)	48/50(96.0)
Adjusted rates(b)	100.00	100.00	97.73	100.00
Terminal rates(c)	44/44(100.0)	44/44(100.0)	41/42(97.6)	36/36(100.0)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.3712			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 1.0000			
Fisher Exact test(e)		P = 0.3087	P = 0.2435	P = 0.5000

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 5

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : mammary gland TUMOR : fibroadenoma				
Tumor rate				
Overall rates(a)	3/50(6.0)	0/50(0.0)	0/50(0.0)	2/50(4.0)
Adjusted rates(b)	6.82	0.0	0.0	2.78
Terminal rates(c)	3/44(6.8)	0/44(0.0)	0/42(0.0)	1/36(2.8)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.1093			
Prevalence method(d)	P = 0.8044			
Combined analysis(d)	P = 0.5183			
Cochran-Armitage test(e)	P = 0.8183			
Fisher Exact test(e)		P = 0.1212	P = 0.1212	P = 0.5000
SITE : mammary gland TUMOR : adenoma, fibroadenoma				
Tumor rate				
Overall rates(a)	3/50(6.0)	1/50(2.0)	0/50(0.0)	2/50(4.0)
Adjusted rates(b)	6.82	2.27	0.0	2.78
Terminal rates(c)	3/44(6.8)	1/44(2.3)	0/42(0.0)	1/36(2.8)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.1093			
Prevalence method(d)	P = 0.8376			
Combined analysis(d)	P = 0.5881			
Cochran-Armitage test(e)	P = 0.6742			
Fisher Exact test(e)		P = 0.3087	P = 0.1212	P = 0.5000

(HPT360A)

BAIS4

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 6

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : preputial/clitoral gland				
TUMOR : adenoma				
Tumor rate				
Overall rates(a)	3/50(6.0)	1/50(2.0)	1/50(2.0)	2/50(4.0)
Adjusted rates(b)	6.82	2.27	2.38	5.56
Terminal rates(c)	3/44(6.8)	1/44(2.3)	1/42(2.4)	2/36(5.6)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.5349			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.7450			
Fisher Exact test(e)		P = 0.3087	P = 0.3087	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 out comes can not 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.

APPENDIX O 2

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS : FEMALE

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 7

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : lung TUMOR : bronchiolar-alveolar adenoma, bronchiolar-alveolar 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	6.67	0.0	5.13
Terminal rates(c)	0/37(0.0)	3/45(6.7)	0/36(0.0)	2/39(5.1)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.2396			
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 : spleen TUMOR : mononuclear cell leukemia				
Tumor rate				
Overall rates(a)	8/50(16.0)	4/50(8.0)	9/50(18.0)	9/50(18.0)
Adjusted rates(b)	8.11	4.44	13.89	7.69
Terminal rates(c)	3/37(8.1)	2/45(4.4)	5/36(13.9)	3/39(7.7)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.2311			
Prevalence method(d)	P = 0.3862			
Combined analysis(d)	P = 0.2293			
Cochran-Armitage test(e)	P = 0.4615			
Fisher Exact test(e)		P = 0.1783	P = 0.5000	P = 0.5000
SITE : pituitary gland TUMOR : adenoma				
Tumor rate				
Overall rates(a)	11/50(22.0)	12/50(24.0)	10/50(20.0)	8/50(16.0)
Adjusted rates(b)	24.32	23.40	27.78	17.95
Terminal rates(c)	9/37(24.3)	10/45(22.2)	10/36(27.8)	7/39(17.9)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.8736			
Prevalence method(d)	P = 0.7252			
Combined analysis(d)	P = 0.8079			
Cochran-Armitage test(e)	P = 0.3587			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.3055

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 8

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : pituitary gland TUMOR : adenoma, adenocarcinoma				
Tumor rate				
Overall rates(a)	13/50(26.0)	12/50(24.0)	11/50(22.0)	9/50(18.0)
Adjusted rates(b)	27.03	23.40	27.78	20.51
Terminal rates(c)	10/37(27.0)	10/45(22.2)	10/36(27.8)	8/39(20.5)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.9090			
Prevalence method(d)	P = 0.7086			
Combined analysis(d)	P = 0.8315			
Cochran-Armitage test(e)	P = 0.3164			
Fisher Exact test(e)		P = 0.5000	P = 0.4076	P = 0.2348
SITE : thyroid TUMOR : C-cell adenoma				
Tumor rate				
Overall rates(a)	2/50(4.0)	2/50(4.0)	1/50(2.0)	4/50(8.0)
Adjusted rates(b)	4.88	4.44	2.78	10.26
Terminal rates(c)	1/37(2.7)	2/45(4.4)	1/36(2.8)	4/39(10.3)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.1524			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.3270			
Fisher Exact test(e)		P = 0.6913	P = 0.5000	P = 0.3389
SITE : thyroid TUMOR : C-cell adenoma, C-cell carcinoma				
Tumor rate				
Overall rates(a)	2/50(4.0)	2/50(4.0)	3/50(6.0)	4/50(8.0)
Adjusted rates(b)	4.88	4.44	8.33	10.26
Terminal rates(c)	1/37(2.7)	2/45(4.4)	3/36(8.3)	4/39(10.3)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.1464			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.3192			
Fisher Exact test(e)		P = 0.6913	P = 0.5000	P = 0.3389

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 9

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : uterus TUMOR : endometrial stromal polyp				
Tumor rate				
Overall rates(a)	11/50(22.0)	10/50(20.0)	12/50(24.0)	14/50(28.0)
Adjusted rates(b)	23.68	22.22	30.77	30.77
Terminal rates(c)	8/37(21.6)	10/45(22.2)	11/36(30.6)	12/39(30.8)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.9335 ?			
Prevalence method(d)	P = 0.1322			
Combined analysis(d)	P = 0.1818			
Cochran-Armitage test(e)	P = 0.3822			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.3224
SITE : uterus TUMOR : endometrial stromal sarcoma				
Tumor rate				
Overall rates(a)	4/50(8.0)	0/50(0.0)	2/50(4.0)	2/50(4.0)
Adjusted rates(b)	0.0	0.0	2.78	5.13
Terminal rates(c)	0/37(0.0)	0/45(0.0)	1/36(2.8)	2/39(5.1)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.9876			
Prevalence method(d)	P = 0.0367*			
Combined analysis(d)	P = 0.6632			
Cochran-Armitage test(e)	P = 0.6256			
Fisher Exact test(e)		P = 0.0587	P = 0.3389	P = 0.3389
SITE : mammary gland TUMOR : fibroadenoma				
Tumor rate				
Overall rates(a)	4/50(8.0)	8/50(16.0)	6/50(12.0)	6/50(12.0)
Adjusted rates(b)	9.09	15.22	16.67	15.38
Terminal rates(c)	3/37(8.1)	6/45(13.3)	6/36(16.7)	6/39(15.4)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.5714			
Prevalence method(d)	P = 0.3345			
Combined analysis(d)	P = 0.3764			
Cochran-Armitage test(e)	P = 0.7686			
Fisher Exact test(e)		P = 0.1783	P = 0.3703	P = 0.3703

STUDY No. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 SEX : FEMALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

PAGE : 10

Group Name	Control	25 ppm	50 ppm	100 ppm
SITE : mammary gland				
TUMOR : fibroadenoma, adenocarcinoma				
Tumor rate				
Overall rates(a)	4/50(8.0)	9/50(18.0)	8/50(16.0)	6/50(12.0)
Adjusted rates(b)	9.09	17.39	16.67	15.38
Terminal rates(c)	3/37(8.1)	7/45(15.6)	6/36(16.7)	6/39(15.4)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.5056			
Prevalence method(d)	P = 0.3769			
Combined analysis(d)	P = 0.3941			
Cochran-Armitage test(e)	P = 0.8066			
Fisher Exact test(e)		P = 0.1168	P = 0.1783	P = 0.3703

(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.

APPENDIX P 1

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR :

MALE

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 1

Organ	Findings	Group Name	Control	25 ppm	50 ppm	100 ppm
		No. of Animals on Study	50	50	50	50
{Respiratory system}						
nasal cavit			<50>	<50>	<50>	<50>
	leukemic cell infiltration		0	0	1	0
lung			<50>	<50>	<50>	<50>
	leukemic cell infiltration		3	2	4	3
	metastasis:subcutis tumor		0	0	0	1
	metastasis:skin/appendage tumor		1	0	0	0
{Hematopoietic system}						
bone marrow			<50>	<50>	<50>	<50>
	leukemic cell infiltration		1	1	1	1
lymph node			<50>	<50>	<50>	<50>
	leukemic cell infiltration		1	1	2	1
{Circulatory system}						
heart			<50>	<50>	<50>	<50>
	leukemic cell infiltration		0	0	0	1
	metastasis:skin/appendage tumor		1	0	0	0
{Digestive system}						
liver			<50>	<50>	<50>	<50>
	leukemic cell infiltration		2	2	6	2
pancreas			<50>	<50>	<50>	<50>
	leukemic cell infiltration		0	1	0	0
< a >	a : Number of animals examined at the site					
b	b : Number of animals with lesion					

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : MALE

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

PAGE : 2

Organ	Findings	Group Name No. of Animals on Study	Control 50	25 ppm 50	50 ppm 50	100 ppm 50
{Urinary system}						
kidney	leukemic cell infiltration		<50> 0	<50> 0	<50> 0	<50> 1
urin bladd	leukemic cell infiltration		<50> 0	<50> 1	<49> 0	<49> 0
{Endocrine system}						
adrenal	leukemic cell infiltration		<50> 0	<50> 0	<50> 1	<50> 0
{Nervous system}						
brain	leukemic cell infiltration		<50> 1	<50> 0	<50> 1	<50> 0
	metastasis:bone tumor		1	0	0	0
	metastasis:peripheral nerve tumor		1	0	0	0
spinal cord	leukemic cell infiltration		<50> 0	<50> 1	<50> 1	<50> 0
periph nerv	metastasis:bone tumor		<50> 1	<50> 0	<50> 0	<50> 0
{Body cavities}						
adipose	metastasis:kidney tumor		<50> 0	<50> 0	<50> 1	<50> 0
< a > a : Number of animals examined at the site						
b b : Number of animals with lesion						

(JPT150)

BAIS4

APPENDIX P 2

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR :

FEMALE

STUDY NO. : 0535
 ANIMAL : RAT F344/DuCr1j[F344/DuCrj]
 REPORT TYPE : A1
 SEX : FEMALE

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

PAGE : 3

Group Name No. of Animals on Study		Control 50	25 ppm 50	50 ppm 50	100 ppm 50
Organ	Findings				
{Integumentary system/appandage}					
subcutis	metastasis:uterus tumor	<50> 1	<50> 0	<50> 1	<50> 0
{Respiratory system}					
lung	leukemic cell infiltration	<50> 5	<50> 2	<50> 5	<50> 5
	metastasis:uterus tumor	1	0	2	0
	metastasis:bone tumor	0	0	1	0
{Hematopoietic system}					
bone marrow	leukemic cell infiltration	<50> 4	<50> 2	<50> 1	<50> 2
lymph node	leukemic cell infiltration	<50> 3	<50> 2	<50> 0	<50> 2
	metastasis:uterus tumor	0	0	1	0
	metastasis:thyroid tumor	0	0	1	0
{Digestive system}					
stomach	leukemic cell infiltration	<50> 1	<50> 0	<50> 0	<50> 0
large intes	metastasis:uterus tumor	<50> 1	<50> 0	<50> 1	<50> 0
liver	leukemic cell infiltration	<50> 7	<50> 2	<50> 6	<50> 9
< a > a : Number of animals examined at the site					
b : Number of animals with lesion					

STUDY NO. : 0535
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	25 ppm 50	50 ppm 50	100 ppm 50
Organ	Findings					
{Digestive system}						
liver	metastasis:uterus tumor		<50> 1	<50> 0	<50> 2	<50> 0
	metastasis:ovary tumor		0	1	0	0
pancreas	leukemic cell infiltration		<50> 3	<50> 0	<50> 1	<50> 0
	metastasis:uterus tumor		1	0	1	0
{Urinary system}						
kidney	leukemic cell infiltration		<50> 2	<50> 1	<50> 0	<50> 1
	metastasis:uterus tumor		0	0	1	0
urin bladd	leukemic cell infiltration		<50> 0	<50> 0	<50> 1	<50> 0
	metastasis:uterus tumor		1	0	1	0
{Endocrine system}						
pituitary	leukemic cell infiltration		<50> 0	<50> 1	<50> 0	<50> 1
	leukemic cell infiltration		<50> 0	<50> 1	<50> 1	<50> 0
{Reproductive system}						
ovary	metastasis:uterus tumor		<50> 1	<50> 0	<50> 0	<50> 0
< a >	a : Number of animals examined at the site					
b	b : Number of animals with lesion					

STUDY NO. : 0535
ANIMAL : RAT F344/DuCr1Cr1j[F344/DuCrj]
REPORT TYPE : A1
SEX : FEMALE

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

PAGE : 5

Organ_____	Findings_____	Group Name No. of Animals on Study	Control 50	25 ppm 50	50 ppm 50	100 ppm 50
{Reproductive system}						
uterus			<50>	<50>	<50>	<50>
	leukemic cell infiltration		1	0	1	1
{Nervous system}						
brain			<50>	<50>	<50>	<50>
	leukemic cell infiltration		1	0	2	1
	metastasis:pituitary tumor		2	0	1	1
spinal cord			<50>	<50>	<50>	<50>
	leukemic cell infiltration		1	0	1	1
{Body cavities}						
peritoneum			<50>	<50>	<50>	<50>
	metastasis:uterus tumor		1	0	0	1
	metastasis:ovary tumor		0	1	0	0
< a >	a : Number of animals examined at the site					
b	b : Number of animals with lesion					

(JPT150)

BAIS4

APPENDIX Q

METHODS, UNITS AND DECIMAL PLACE FOR
HEMATOLOGY AND BIOCHEMISTRY IN THE 2-YEAR
INHALATION STUDY OF PROPIONONITRILE

METHODS, UNITS AND DECIMAL PLACE FOR HEMATOLOGY AND BIOCHEMISTRY IN THE 2-YEAR INHALATION STUDY OF PROPIONONITRILE

Item	Method	Unit	Decimal place
Hematology			
Red blood cell (RBC)	Light scattering method ¹⁾	$\times 10^6/\mu\text{L}$	2
Hemoglobin(Hgb)	Cyanmethemoglobin method ¹⁾	g/dL	1
Hematocrit(Hct)	Calculated as $\text{RBC} \times \text{MCV} / 10$ ¹⁾	%	1
Mean corpuscular volume(MCV)	Light scattering method ¹⁾	fL	1
Mean corpuscular hemoglobin(MCH)	Calculated as $\text{Hgb} / \text{RBC} \times 10$ ¹⁾	pg	1
Mean corpuscular hemoglobin concentration (MCHC)	Calculated as $\text{Hgb} / \text{Hct} \times 100$ ¹⁾	g/dL	1
Platelet	Light scattering method ¹⁾	$\times 10^3/\mu\text{L}$	0
Reticulocyte	Light scattering method ¹⁾	%	1
White blood cell(WBC)	Light scattering method ¹⁾	$\times 10^3/\mu\text{L}$	2
Differential WBC	Pattern recognition method ²⁾ (Wright staining)	%	0
Biochemistry			
Total protein(TP)	Biuret method ³⁾	g/dL	1
Albumin (Alb)	BCG method ³⁾	g/dL	1
A/G ratio	Calculated as $\text{Alb} / (\text{TP} - \text{Alb})$ ³⁾	—	1
T-bilirubin	Alkaline azobilirubin method ³⁾	mg/dL	2
Glucose	GlcK·G-6-PDH method ³⁾	mg/dL	0
T-cholesterol	CE·COD·POD method ³⁾	mg/dL	0
Triglyceride	LPL·GK·GPO·POD method ³⁾	mg/dL	0
Phospholipid	PLD·ChOD·POD method ³⁾	mg/dL	0
Aspartate aminotransferase (AST)	JSCC method ³⁾	IU/L	0
Alanine aminotransferase (ALT)	JSCC method ³⁾	IU/L	0
Lactate dehydrogenase (LDH)	SFBC method ³⁾	IU/L	0
Alkaline phosphatase (ALP)	GSCC method ³⁾	IU/L	0
γ -Glutamyl transpeptidase (γ -GTP)	JSCC method ³⁾	IU/L	0
Creatine kinase (CK)	JSCC method ³⁾	IU/L	0
Urea nitrogen	Urease·GLDH method ³⁾	mg/dL	1
Creatinine	Jaffé method ³⁾	mg/dL	1
Sodium	Ion selective electrode method ³⁾	mEq/L	0
Potassium	Ion selective electrode method ³⁾	mEq/L	1
Chloride	Ion selective electrode method ³⁾	mEq/L	0
Calcium	OCPC method ³⁾	mg/dL	1
Inorganic phosphorus	PNP·XOD·POD method ³⁾	mg/dL	1

1) Automatic blood cell analyzer (ADVIA120 : Bayer Corporation)

2) Automatic blood cell differential analyzer (MICROX HEG-120NA : OMRON Corporation)

3) Automatic analyzer (Hitachi 7080 : Hitachi,Ltd.)