

1 - ブロモブタンのラットを用いた
吸入による2週間毒性試験報告書

試験番号 : 0480

APPENDICES

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OF 1-BROMOBUTANE

APPENDIX A 1

IDENTITY OF 1-BROMOBUTANE IN THE 2-WEEK INHALATION STUDY

IDENTITY OF 1-BROMOBUTANE IN THE 2-WEEK INHALATION STUDY

Test Substance : 1-Bromobutane (Wako Pure Chemical Industries, Ltd.)

Lot No. : ASQ0017

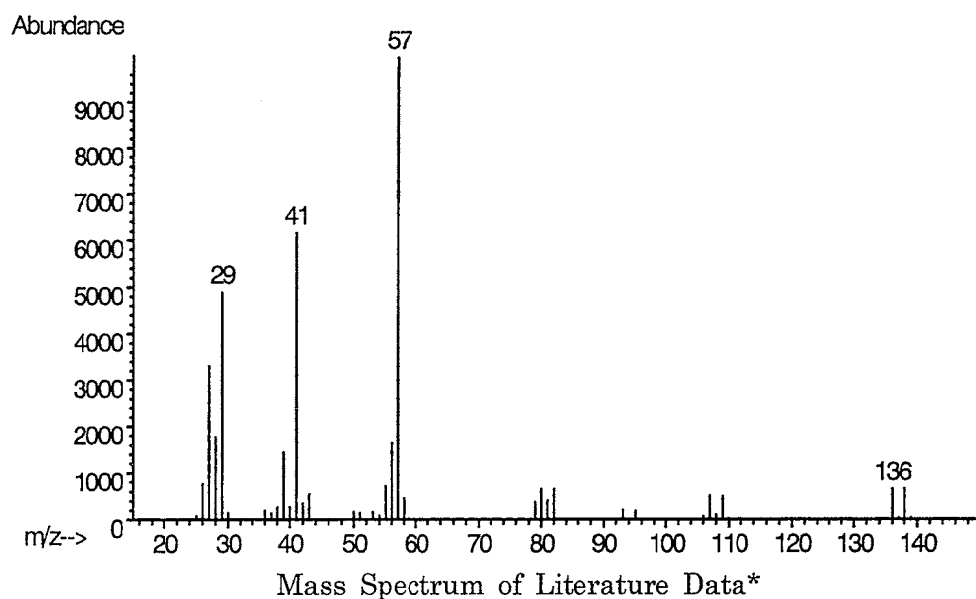
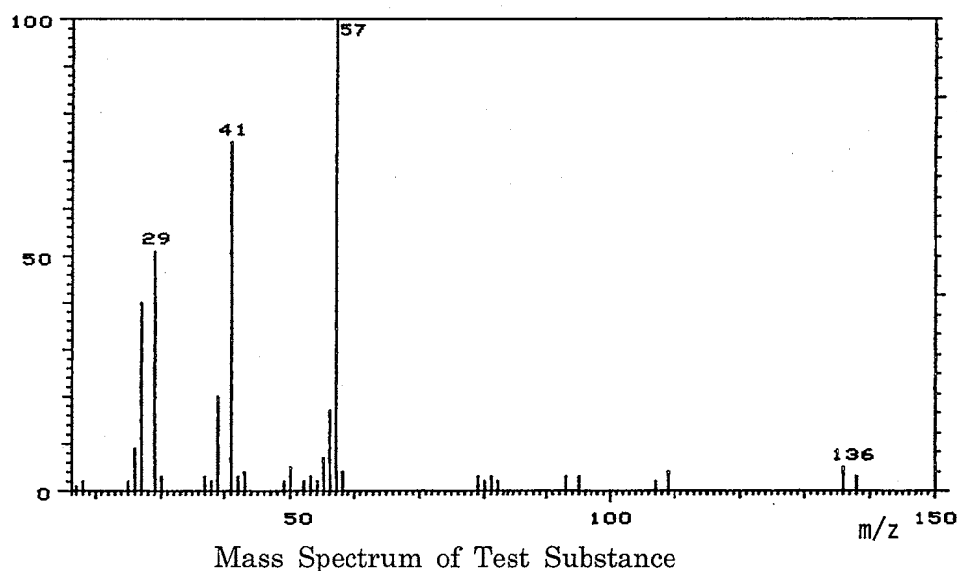
1. Spectral Data

Mass Spectrometry

Instrument : Hitachi M-80B Mass Spectrometer

Ionization : EI (Electron Ionization)

Ionization Voltage : 70eV



Result: The mass spectrum was consistent with literature spectrum.

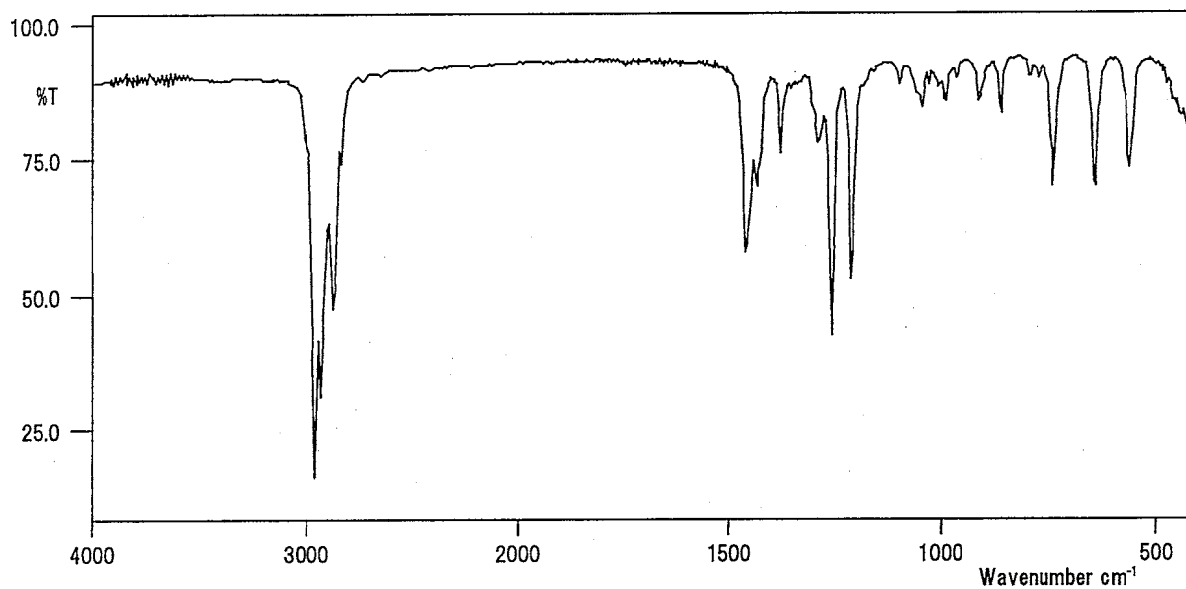
(*McLafferty FW, ed. 1994. Wiley Registry of Mass Spectral Data. 6th ed. New York, NY:John Wiley and Sons.)

Infrared Spectrometry

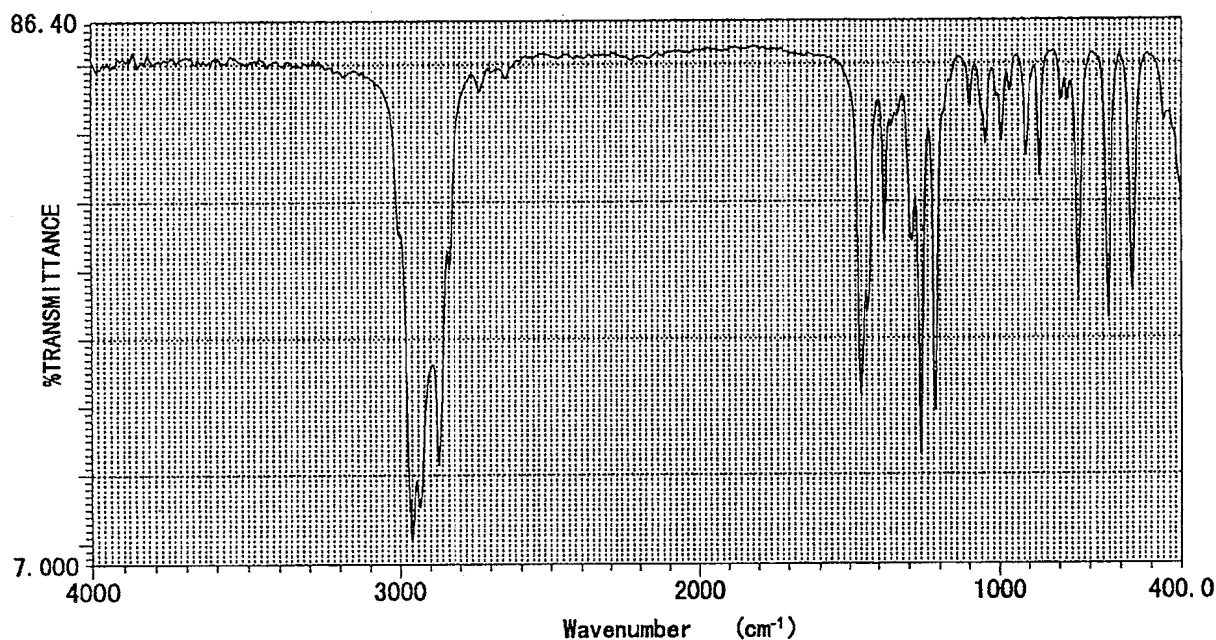
Instrument : Shimadzu FTIR-8200PC Infrared Spectrometer

Cell : KBr Liquid Cell

Resolution : 4 cm^{-1}



Infrared Spectrum of Test Substance



Infrared Spectrum of Literature Data*

Result: The infrared spectrum was consistent with literature spectrum.

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

2. Conclusion: The test substance was identified as 1-bromobutane by mass spectrum and infrared spectrum.

APPENDIX A 2

STABILITY OF 1-BROMOBUTANE IN THE 2-WEEK INHALATION STUDY

STABILITY OF 1-BROMOBUTANE IN THE 2-WEEK INHALATION STUDY

Test Substance : 1-Bromobutane (Wako Pure Chemical Industries, Ltd.)
Lot No. : ASQ0017
1. Sample : This lot was used from 2003.3.20 to 2003.4.2. Test substance was stored in a dark place at room temperature.

2. Gas Chromatography

Instrument : Hewlett Packard 5890A Gas Chromatograph
Column : Methyl Silicone (0.53 mm ϕ \times 60 m)
Column Temperature: 100° C
Flow Rate : 15 mL/min
Detector : FID (Flame Ionization Detector)
Injection Volume : 1 μ L

Date (date analyzed)	Peak No.	Retention Time (min)	Area (%)
2003.03.19	1	4.139	100
2003.04.25	1	4.127	100

Result: Gas chromatography indicated one major peak (peak No. 1) analyzed on 2003.3.19 and one major peak(peak No. 1) analyzed on 2003.4.25. No new trace impurity peak in the test substance analyzed on 2003.4.25 was detected.

3. Conclusion: The test substance was stable for about 1 month in a dark place at room temperature.

APPENDIX B 1

CONCENTRATION OF 1-BROMOBUTANE IN THE INHALATION CHAMBER OF THE 2-WEEK INHALATION STUDY

CONCENTRATION OF 1-BROMOBUTANE IN THE INHALATION
CHAMBER OF THE 2-WEEK INHALATION STUDY

Group Name	Concentration(ppm)
	Mean \pm S.D.
Control	0.0 \pm 0.0
500ppm	500.5 \pm 3.2
1000ppm	1003.0 \pm 5.3
2000ppm	2005.5 \pm 12.6
4000ppm	4007.0
8000ppm	7967.7

APPENDIX B 2

ENVIRONMENTAL CONDITIONS OF INHALATION CHAMBER IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

ENVIRONMENTAL CONDITIONS OF INHALATION CHAMBER IN THE 2-WEEK
INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Temperature (°C) Mean ± S.D.	Humidity (%) Mean ± S.D.	Ventilation Rate (L/min) Mean ± S.D.	Air Change (time/h) Mean
Control	22.4 ± 0.2	56.2 ± 0.6	212.7 ± 0.6	12.0
500ppm	22.3 ± 0.2	56.8 ± 0.7	212.6 ± 0.6	12.0
1000ppm	22.3 ± 0.2	56.0 ± 0.7	213.1 ± 0.5	12.1
2000ppm	21.7 ± 0.1	55.2 ± 1.0	213.0 ± 0.4	12.1
4000ppm	21.5 ± 0.4	56.6 ± 1.1	213.5 ± 0.6	12.1
8000ppm	22.1	52.1	213.2	12.1

APPENDIX C 1

CLINICAL OBSERVATION : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1 2

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

PAGE : 1

Clinical sign	Group Name	Administration Week-day					
		1-1	1-2	1-4	1-7	2-3	2-7
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	4	4	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
LATERAL	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	4	1	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
HUNCHBACK POSITION	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	0	3	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
WASTING	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	0	4	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
SOILED	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	1	1	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
PILORECTION	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	4	4	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1 2

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

PAGE : 2

Clinical sign	Group Name	Administration Week-day					
		1-1	1-2	1-4	1-7	2-3	2-7
IRREGULAR BREATHING	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	5	5	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
TACHYPNEA	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	1	1	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
BRADYPNEA	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	1	0	-	-
	4000ppm	0	-	-	-	-	-
	8000ppm	-	-	-	-	-	-

(HAN190)

BAIS 4

APPENDIX C 2

CLINICAL OBSERVATION : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1 2

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : FEMALE

PAGE : 3

Clinical sign	Group Name	Administration Week-day					
		1-1	1-2	1-4	1-7	2-3	2-7
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	0	0	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
PILOERECTION	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	0	0	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-
IRREGULAR BREATHING	Control	0	0	0	0	0	0
	500ppm	0	0	0	0	0	0
	1000ppm	0	0	0	0	0	0
	2000ppm	0	0	0	4	-	-
	4000ppm	5	-	-	-	-	-
	8000ppm	-	-	-	-	-	-

APPENDIX D 1

BODY WEIGHT CHANGES : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 UNIT : g
 REPORT TYPE : A1 2
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 1

Group Name	Administration		week-day									
	0-0		1-2		1-4		1-7		2-3		2-7	
Control	118±	5	125±	6	129±	8	137±	9	144±	10	156±	11
500ppm	119±	5	121±	5	126±	6	131±	5	140±	6	152±	6
1000ppm	119±	5	120±	4	123±	4	128±	3	133±	4	142±	5*
2000ppm	119±	5	115±	5	95±	10**	82±	23**	-		-	
4000ppm	119±	4	-		-		-		-		-	
8000ppm	119±	5	-		-		-		-		-	

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

Test of Dunnett

APPENDIX D 2

BODY WEIGHT CHANGES : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 UNIT : g
 REPORT TYPE : A1 2
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 2

Group Name	Administration		week-day									
	0-0		1-2		1-4		1-7		2-3		2-7	
Control	95±	3	99±	3	102±	4	106±	3	109±	4	115±	4
500ppm	95±	2	97±	3	98±	3	102±	4	105±	3	109±	3
1000ppm	95±	3	96±	3	97±	3	101±	3	105±	4	111±	3
2000ppm	95±	3	95±	4	92±	4**	94±	5**	-		-	
4000ppm	95±	3	-		-		-		-		-	
8000ppm	95±	3	-		-		-		-		-	

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

Test of Dunnett

(HAN260)

BAIS 4

APPENDIX E 1

FOOD CONSUMPTION CHANGES : SUMMARY, RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
UNIT : g
REPORT TYPE : A1 2
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 1

Group Name	Administration		week-day(effective)	
	1-7 (6)		2-7 (7)	
Control	13.8±	0.6	13.3±	1.6
500ppm	11.7±	0.6	13.7±	0.9
1000ppm	12.0±	0.4	14.0±	0.6
2000ppm	3.7±	4.2**	-	
4000ppm	-		-	
8000ppm	-		-	

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

Test of Dunnett

(HAN260)

BAIS 4

APPENDIX E 2

FOOD CONSUMPTION CHANGES : SUMMARY, RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
UNIT : g
REPORT TYPE : A1 2
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 2

Group Name	Administration week-day(effective)	
	1-7 (6)	2-7 (7)
Control	10.5± 0.5	10.0± 0.6
500ppm	10.0± 1.0	9.9± 0.3
1000ppm	10.1± 0.4	11.2± 0.5**
2000ppm	8.8± 0.9**	-
4000ppm	-	-
8000ppm	-	-

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

Test of Dunnett

APPENDIX F 1

HEMATOLOGY : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (3W)

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	5	8.89± 0.17	15.9± 0.4	45.5± 0.9	51.3± 0.4	17.8± 0.1	34.8± 0.3	805± 69
500ppm	5	8.97± 0.20	15.8± 0.3	46.2± 1.0	51.6± 0.2	17.7± 0.1	34.2± 0.2**	973± 55**
1000ppm	5	9.21± 0.15*	16.3± 0.3	47.7± 0.8**	51.7± 0.3	17.7± 0.1	34.2± 0.2**	936± 35**
2000ppm	0	-	-	-	-	-	-	-
4000ppm	0	-	-	-	-	-	-	-
8000ppm	0	-	-	-	-	-	-	-

Significant difference : * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
MEASURE. TIME : 1
SEX : MALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (3W)

PAGE : 2

Group Name	NO. of Animals	RETICULOCYTE %		PROTHROMBIN TIME s e c		APTT s e c	
Control	5	1.9±	0.3	12.7±	0.4	22.2±	0.4
500ppm	5	2.3±	0.3	12.7±	0.7	21.3±	1.5
1000ppm	5	2.2±	0.2	12.4±	0.4	19.5±	0.5**
2000ppm	0	-		-		-	
4000ppm	0	-		-		-	
8000ppm	0	-		-		-	

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (3W)

REPORT TYPE : A1

PAGE : 3

Group Name	NO. of Animals	WBC 1 O ³ /μl		Differential N-BAND		WBC (%) N-SBG		EOSINO		BASO		MONO		LYMPHO		OTHER	
Control	5	3.43±	1.02	0±	1	18±	6	1±	1	0±	0	2±	1	78±	6	0±	0
500ppm	5	5.69±	1.13*	1±	1	19±	2	0±	0	0±	0	2±	1	78±	3	0±	0
1000ppm	5	5.66±	1.12*	1±	1	18±	4	0±	1	0±	0	1±	1	79±	4	0±	0
2000ppm	0	-		-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-		-	

Significant difference : * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

APPENDIX F 2

HEMATOLOGY : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480

ANIMAL : RAT F344/DuCrj

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)

ALL ANIMALS (3W)

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	5	9.25±	0.13	16.8±	0.2	47.3±	0.5	51.2±	0.5	18.2±	0.2	35.5±	0.2	758±	84
500ppm	5	9.10±	0.12	16.6±	0.3	46.8±	0.7	51.4±	0.4	18.2±	0.1	35.4±	0.3	879±	84
1000ppm	5	9.29±	0.11	16.8±	0.3	47.9±	0.6	51.6±	0.1	18.1±	0.2	35.1±	0.3	829±	41
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

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

Test of Dunnett

(HCL070)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
MEASURE. TIME : 1
SEX : FEMALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (3W)

REPORT TYPE : A1

PAGE : 5

Group Name	NO. of Animals	RETICULOCYTE %		PROTHROMBIN TIME s e c		APTT s e c	
Control	5	1.4±	0.2	12.0±	0.4	18.9±	0.5
500ppm	5	1.6±	0.2	12.2±	0.2	18.3±	0.8
1000ppm	5	2.0±	0.1**	12.2±	0.5	18.1±	1.2
2000ppm	0	-		-		-	
4000ppm	0	-		-		-	
8000ppm	0	-		-		-	

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

Test of Dunnett

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : FEMALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (3W)

REPORT TYPE : A1

PAGE : 6

Group Name	NO. of Animals	WBC 1 G ³ /μℓ		Differential N-BAND		WBC (%) N-SEG		EOSINO		BASO		MONO		LYMPHO		OTHER	
Control	5	3.25±	0.46	0±	0	13±	4	1±	1	0±	0	2±	1	83±	4	0±	0
500ppm	5	4.28±	1.04	1±	1	16±	4	1±	1	0±	0	3±	1	80±	6	0±	0
1000ppm	5	4.61±	0.82	1±	1	16±	5	1±	1	0±	0	3±	1	79±	4	0±	0
2000ppm	0	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4000ppm	0	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
8000ppm	0	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

Significant difference : * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS 4

APPENDIX G 1

BIOCHEMISTRY : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480

ANIMAL : RAT F344/DuCrj

MEASURE. TIME : 1

SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (3W)

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	5	5.7±	0.2	3.4±	0.1	1.5±	0.1	0.12±	0.01	157±	12	49±	3	33±	11
500ppm	5	6.0±	0.1**	3.5±	0.1	1.5±	0.1	0.11±	0.01	146±	4	36±	4**	14±	6*
1000ppm	5	6.2±	0.1**	3.7±	0.2*	1.4±	0.1	0.14±	0.03	156±	6	32±	2**	13±	2*
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (3W)

REPORT TYPE : A1

PAGE : 2

Group Name	NO. of Animals	PHOSPHOLIPID mg/dl		GOT IU/l		GPT IU/l		LDH IU/l		ALP IU/l		G-GTP IU/l		CPK IU/l	
Control	5	99±	7	57±	9	32±	2	360±	73	677±	40	1±	1	255±	47
500ppm	5	77±	6**	66±	2*	28±	1	324±	51	569±	25**	2±	4	188±	16*
1000ppm	5	78±	2**	64±	4	29±	3	427±	96	517±	44**	1±	1	153±	27**
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (3W)

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	5	16.5±	1.0	0.5±	0.0	141±	1	4.1±	0.3	103±	1	10.1±	0.2	8.6±	0.7
500ppm	5	13.7±	1.4*	0.5±	0.1	141±	1	3.9±	0.4	105±	1**	10.3±	0.1*	8.2±	0.4
1000ppm	5	11.5±	1.6**	0.4±	0.1	139±	1*	4.2±	0.2	107±	0**	10.4±	0.1**	8.3±	0.3
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

APPENDIX G 2

BIOCHEMISTRY : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (3W)

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	5	5.7±	0.1	3.4±	0.1	1.5±	0.1	0.15±	0.01	138±	8	67±	5	14±	5
500ppm	5	5.8±	0.2	3.4±	0.1	1.4±	0.1	0.15±	0.01	134±	8	60±	4	15±	2
1000ppm	5	5.9±	0.2	3.5±	0.1	1.4±	0.1	0.15±	0.02	161±	3**	56±	4**	15±	4
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0480

ANIMAL : RAT F344/DuCrj

MEASURE. TIME : 1

SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)

ALL ANIMALS (3W)

PAGE : 5

Group Name	NO. of Animals	PHOSPHOLIPID mg/dl		GOT IU/l		GPT IU/l		LDH IU/l		ALP IU/l		G-GTP IU/l		CPK IU/l	
Control	5	122±	7	75±	2	32±	3	549±	154	571±	31	1±	0	258±	42
500ppm	5	117±	6	70±	5	24±	1**	553±	143	479±	26**	1±	0	214±	45
1000ppm	5	112±	7	69±	4*	27±	1**	507±	106	478±	31**	2±	1	158±	29**
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

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

Test of Dunnett

(HCL074)

BAIS 4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (3W)

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	5	18.3±	1.6	0.5±	0.0	139±	1	3.9±	0.4	105±	1	9.6±	0.2	7.2±	1.0
500ppm	5	14.0±	0.9**	0.5±	0.0	138±	1	4.1±	0.2	106±	1	9.8±	0.2	7.1±	0.5
1000ppm	5	11.2±	1.0**	0.5±	0.1	137±	1*	4.1±	0.3	107±	1**	10.1±	0.3*	7.4±	0.7
2000ppm	0	-		-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-		-	

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

Test of Dunnett

APPENDIX H 1

GROSS FINDINGS : SUMMARY,
RAT : MALE :
DEAD AND MORIBUND ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 1

Organ	Findings	Group Name NO. of Animals	Control		500ppm		1000ppm		2000ppm	
			0	(%)	0	(%)	0	(%)	5	(%)
lung	red zone		-	(-)	-	(-)	-	(-)	1	(20)
thymus	atrophic		-	(-)	-	(-)	-	(-)	4	(80)
	red		-	(-)	-	(-)	-	(-)	3	(60)
gl stomach	black zone		-	(-)	-	(-)	-	(-)	1	(20)
cecum	gas		-	(-)	-	(-)	-	(-)	1	(20)
urin bladd	urine:brown		-	(-)	-	(-)	-	(-)	2	(40)
thoracic ca	pleural fluid		-	(-)	-	(-)	-	(-)	0	(0)

(HPT080)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 2

Organ	Findings	Group Name NO. of Animals	4000ppm		8000ppm	
			5	(%)	5	(%)
lung	red zone		0	(0)	3	(60)
thymus	atrophic		0	(0)	0	(0)
	red		0	(0)	0	(0)
gl stomach	black zone		0	(0)	0	(0)
cecum	gas		0	(0)	0	(0)
urin bladd	urine:brown		0	(0)	0	(0)
thoracic ca	pleural fluid		5	(100)	0	(0)

(HPT080)

BAIS 4

APPENDIX H 2

GROSS FINDINGS : SUMMARY,
RAT : MALE :
SACRIFICED ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (3W)

PAGE : 1

Organ	Findings	Group Name NO. of Animals	Control		500ppm		1000ppm		2000ppm	
			5	(%)	5	(%)	5	(%)	0	(%)
liver	herniation		1	(20)	0	(0)	0	(0)	-	(-)

(HPT080)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (3W)

PAGE : 2

Organ	Findings	Group Name NO. of Animals	4000ppm 0 (%)	8000ppm 0 (%)
liver	herniation		- (-)	- (-)

(HPT080)

BAYS 4

APPENDIX H 3

GROSS FINDINGS : SUMMARY,
RAT : FEMALE :
DEAD AND MORIBUND ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 3

Organ	Findings	Group Name NO. of Animals	Control	500ppm	1000ppm	2000ppm
			0 (%)	0 (%)	0 (%)	5 (%)
lung	red zone		- (-)	- (-)	- (-)	0 (0)
thymus	atrophic		- (-)	- (-)	- (-)	1 (20)
	red zone		- (-)	- (-)	- (-)	0 (0)
liver	herniation		- (-)	- (-)	- (-)	1 (20)
urin bladd	red zone		- (-)	- (-)	- (-)	1 (20)
	urine: brown		- (-)	- (-)	- (-)	1 (20)
thoracic ca	pleural fluid		- (-)	- (-)	- (-)	0 (0)

(HPT080)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 4

Organ	Findings	Group Name NO. of Animals	4000ppm		8000ppm	
			5	(%)	5	(%)
lung	red zone		1	(20)	5	(100)
thymus	atrophic		0	(0)	0	(0)
	red zone		1	(20)	0	(0)
liver	herniation		1	(20)	0	(0)
urin bladd	red zone		0	(0)	0	(0)
	urine:brown		0	(0)	0	(0)
thoracic ca	pleural fluid		5	(100)	0	(0)

(HPT080)

BAIS 4

APPENDIX H 4

GROSS FINDINGS : SUMMARY,
RAT : FEMALE :
SACRIFICED ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (3W)

PAGE : 3

Organ	Findings	Group Name NO. of Animals	Control		500ppm		1000ppm		2000ppm	
			5	(%)	5	(%)	5	(%)	0	(%)
liver	herniation		2	(40)	1	(20)	2	(40)	-	(-)

(HPT080)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (3W)

PAGE : 4

Organ	Findings	Group Name		4000ppm		8000ppm	
		NO. of Animals		0	(%)	0	(%)
liver	herniation			-	(-)	-	(-)

(HPT080)

BAIS 4

APPENDIX I 1

ORGAN WEIGHT, ABSOLUTE : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE
 UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
 SURVIVAL ANIMALS (3W)

PAGE : 1

Group Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	5	140± 10	0.246± 0.035	0.039± 0.003	2.248± 0.246	0.578± 0.031	0.656± 0.037
500ppm	5	137± 5	0.238± 0.019	0.043± 0.005	2.316± 0.104	0.616± 0.020	0.697± 0.025
1000ppm	5	127± 4*	0.195± 0.034*	0.045± 0.007	2.190± 0.047	0.618± 0.019*	0.682± 0.023
2000ppm	0	-	-	-	-	-	-
4000ppm	0	-	-	-	-	-	-
8000ppm	0	-	-	-	-	-	-

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

Test of Dunnett

(HCL040)

BAIS 4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE
 UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
 SURVIVAL ANIMALS (3W)

PAGE : 2

Group Name	NO. of Animals	KIDNEYS		SPLEEN		LIVER		BRAIN	
Control	5	1.138±	0.085	0.305±	0.032	4.089±	0.299	1.699±	0.035
500ppm	5	1.296±	0.066**	0.301±	0.015	4.865±	0.235**	1.688±	0.027
1000ppm	5	1.314±	0.052**	0.261±	0.015*	5.177±	0.241**	1.606±	0.018**
2000ppm	0	-		-		-		-	
4000ppm	0	-		-		-		-	
8000ppm	0	-		-		-		-	

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

Test of Dunnett

(HCL040)

BAIS 4

APPENDIX I 2

ORGAN WEIGHT, ABSOLUTE : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE
 UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
 SURVIVAL ANIMALS (3W)

PAGE : 3

Group Name	NO. of Animals	Body Weight		THYMUS		ADRENALS		OVARIES		HEART		LUNGS	
Control	5	103±	4	0.262±	0.014	0.043±	0.002	0.068±	0.007	0.453±	0.020	0.575±	0.022
500ppm	5	98±	3	0.208±	0.011**	0.047±	0.004	0.069±	0.010	0.449±	0.006	0.583±	0.039
1000ppm	5	99±	4	0.166±	0.009**	0.054±	0.005**	0.069±	0.015	0.491±	0.019**	0.574±	0.041
2000ppm	0	-		-		-		-		-		-	
4000ppm	0	-		-		-		-		-		-	
8000ppm	0	-		-		-		-		-		-	

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

Test of Dunnett

(HCL040)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (3W)

PAGE : 4

Group Name	NO. of Animals	KIDNEYS		SPLEEN		LIVER		BRAIN	
Control	5	0.869±	0.041	0.251±	0.015	3.089±	0.181	1.579±	0.066
500ppm	5	0.974±	0.013**	0.239±	0.009	3.582±	0.133**	1.565±	0.016
1000ppm	5	1.028±	0.046**	0.213±	0.013**	4.242±	0.190**	1.529±	0.018
2000ppm	0	-		-		-		-	
4000ppm	0	-		-		-		-	
8000ppm	0	-		-		-		-	

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

Test of Dunnett

(HCL040)

BAIS 4

APPENDIX J 1

ORGAN WEIGHT, RELATIVE : SUMMARY,
RAT : MALE

(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (3W)

PAGE : 1

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	5	140± 10	0.175± 0.024	0.028± 0.003	1.598± 0.085	0.413± 0.019	0.469± 0.044
500ppm	5	137± 5	0.173± 0.016	0.031± 0.003	1.687± 0.088	0.448± 0.006**	0.508± 0.022
1000ppm	5	127± 4*	0.154± 0.028	0.035± 0.006*	1.731± 0.061	0.488± 0.018**	0.538± 0.014**
2000ppm	0	-	-	-	-	-	-
4000ppm	0	-	-	-	-	-	-
8000ppm	0	-	-	-	-	-	-

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (3W)

PAGE : 2

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	5	0.811± 0.023	0.217± 0.009	2.913± 0.077	1.217± 0.106
500ppm	5	0.943± 0.024**	0.219± 0.005	3.540± 0.085**	1.230± 0.057
1000ppm	5	1.038± 0.019**	0.206± 0.007	4.088± 0.081**	1.270± 0.045
2000ppm	0	-	-	-	-
4000ppm	0	-	-	-	-
8000ppm	0	-	-	-	-

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

Test of Dunnett

APPENDIX J 2

ORGAN WEIGHT, RELATIVE : SUMMARY,
RAT : FEMALE

(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE
 UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
 SURVIVAL ANIMALS (3W)

PAGE : 3

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	5	103± 4	0.253± 0.008	0.042± 0.002	0.066± 0.009	0.439± 0.030	0.557± 0.030
500ppm	5	98± 3	0.212± 0.016**	0.048± 0.003*	0.070± 0.011	0.457± 0.014	0.593± 0.033
1000ppm	5	99± 4	0.168± 0.012**	0.055± 0.006**	0.070± 0.014	0.496± 0.022**	0.580± 0.055
2000ppm	0	-	-	-	-	-	-
4000ppm	0	-	-	-	-	-	-
8000ppm	0	-	-	-	-	-	-

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

Test of Dunnett

(HCL042)

BAIS 4

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (3W)

PAGE : 4

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	5	0.841± 0.014	0.243± 0.008	2.987± 0.132	1.528± 0.041
500ppm	5	0.993± 0.032**	0.243± 0.011	3.647± 0.044	1.594± 0.037
1000ppm	5	1.036± 0.017**	0.215± 0.016**	4.276± 0.043**	1.543± 0.059
2000ppm	0	-	-	-	-
4000ppm	0	-	-	-	-
8000ppm	0	-	-	-	-

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

Test of Dunnett

(HCL042)

BAIS 4

APPENDIX K 1

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : SUMMARY,
RAT : MALE :
DEAD AND MORIBUND ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 1

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
nasal cavit			< 0>				< 0>				< 0>				< 5>			
	disarrangement:olfactory epithelium		-	-	-	-	-	-	-	-	-	-	-	-	4	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(80)	(0)	(0)	(0)
	atrophy:olfactory epithelium		-	-	-	-	-	-	-	-	-	-	-	-	1	1	3	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(20)	(60)	(0)
	necrosis:olfactory epithelium		-	-	-	-	-	-	-	-	-	-	-	-	0	4	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(80)	(0)	(0)
lung			< 0>				< 0>				< 0>				< 5>			
	congestion		-	-	-	-	-	-	-	-	-	-	-	-	0	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(0)	(0)
	hemorrhage		-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
	edema		-	-	-	-	-	-	-	-	-	-	-	-	0	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(0)	(0)
	edema:perivascular		-	-	-	-	-	-	-	-	-	-	-	-	4	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(80)	(20)	(0)	(0)
{Hematopoietic system}																		
bone marrow			< 0>				< 0>				< 0>				< 5>			
	congestion		-	-	-	-	-	-	-	-	-	-	-	-	0	4	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(80)	(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

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 2

Organ	Findings	Group Name	4000ppm				8000ppm				
		No. of Animals on Study	5				5				
		Grade	1	2	3	4	1	2	3	4	
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}											
nasal cavit			< 5>				< 5>				
	disarrangement:olfactory epithelium		0	0	0	0	0	0	0	0	
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	
	atrophy:olfactory epithelium		0	0	0	0	0	0	0	0	
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	
	necrosis:olfactory epithelium		0	0	5	0	0	0	0	0	
			(0)	(0)	(100)	(0)	(0)	(0)	(0)	(0)	
lung			< 5>				< 5>				
	congestion		0	0	0	0	2	3	0	0	
			(0)	(0)	(0)	(0)	(40)	(60)	(0)	(0)	
	hemorrhage		0	0	0	0	2	0	0	0	
			(0)	(0)	(0)	(0)	(40)	(0)	(0)	(0)	
	edema		0	4	1	0	5	0	0	0	
			(0)	(80)	(20)	(0)	(100)	(0)	(0)	(0)	
	edema:perivascular		0	5	0	0	0	5	0	0	
			(0)	(100)	(0)	(0)	(0)	(100)	(0)	(0)	

{Hematopoietic system}

bone marrow	congestion		< 5>					< 5>			
			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			

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3%)

PAGE : 3

Organ	Findings	Group Name No. of Animals on Study Grade	Control 0				500ppm 0				1000ppm 0				2000ppm 5			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																		
thymus	atrophy		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	1	0	4	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(0)	(80)	(0)	
	congestion		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	3	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(60)	(0)	(0)	(0)	
spleen	atrophy		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	1	4	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(80)	(0)	(0)	
{Circulatory system}																		
heart	hemorrhage		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	0	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)	
{Digestive system}																		
stomach	erosion:forestomach		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	1	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(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

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 4

		Group Name	4000ppm				8000ppm			
		No. of Animals on Study	5				5			
Organ	Findings	Grade	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}										
thymus			< 5>				< 5>			
	atrophy		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	congestion		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
spleen			< 5>				< 5>			
	atrophy		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Circulatory system}										
heart			< 5>				< 5>			
	hemorrhage		3	2	0	0	0	0	0	0
			(60)	(40)	(0)	(0)	(0)	(0)	(0)	(0)
{Digestive system}										
stomach			< 5>				< 5>			
	erosion:forestomach		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

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 5

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																		
liver			< 0>				< 0>				< 0>				< 5>			
	herniation	-	-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
	necrosis:central	-	-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
	{Endocrine system}																	
pituitary			< 0>				< 0>				< 0>				< 5>			
	Rathke pouch	-	-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
adrenal			< 0>				< 0>				< 0>				< 5>			
	hemorrhage	-	-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
{Reproductive system}																		
testis			< 0>				< 0>				< 0>				< 5>			
	germ cell necrosis	-	-	-	-	-	-	-	-	-	-	-	-	-	1	4	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(80)	(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																		

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 6

Organ	Findings	4000ppm				8000ppm			
		No. of Animals on Study				5			
		Grade				5			
		1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}									
liver	herniation	< 5>				< 5>			
		0	0	0	0	1	0	0	0
		(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)
	necrosis:central	1	0	4	0	0	0	0	0
		(20)	(0)	(80)	(0)	(0)	(0)	(0)	(0)
{Endocrine system}									
pituitary	Rathke pouch	< 5>				< 5>			
		1	0	0	0	0	0	0	0
		(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
adrenal	hemorrhage	< 5>				< 5>			
		0	1	0	0	0	0	0	0
		(0)	(20)	(0)	(0)	(0)	(0)	(0)	(0)
{Reproductive system}									
testis	germ cell necrosis	< 5>				< 5>			
		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

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 7

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
epididymis			< 0>				< 0>				< 0>				< 5>			
	decreased:sperma		-	-	-	-	-	-	-	-	-	-	-	-	0	1	4	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(80)	(0)
	debris of spermatic elements		-	-	-	-	-	-	-	-	-	-	-	-	2	3	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(40)	(60)	(0)	(0)
{Nervous system}																		
brain			< 0>				< 0>				< 0>				< 5>			
	degeneration:granular cell		-	-	-	-	-	-	-	-	-	-	-	-	0	2	3	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(40)	(60)	(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																	

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 8

Organ	Findings	Group Name No. of Animals on Study Grade	4000ppm				8000ppm			
			5				5			
			1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)

{Reproductive system}

epididymis	decreased:sperma	< 5>				< 5>			
		0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	debris of spermatic elements	< 5>				< 5>			
		0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

{Nervous system}

brain	degeneration:granular cell	< 5>				< 5>			
		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

(HPT150)

BAIS4

APPENDIX K 2

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : SUMMARY,
RAT : MALE :
SACRIFICED ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 SACRIFICED ANIMALS (3W)

PAGE : 1

Organ	Findings	Group Name No. of Animals on Study Grade	Control				500ppm				1000ppm				2000ppm			
			5				5				5				0			
			1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)

{Respiratory system}

nasal cavit			< 5>				< 5>				< 5>				< 0>			
	disarrangement:olfactory epithelium		0	0	0	0	4	0	0	0	5	0	0	0	-	-	-	-
			(0)	(0)	(0)	(0)	(80)	(0)	(0)	(0)	(100)	(0)	(0)	(0)	(-)	(-)	(-)	(-)
	atrophy:olfactory epithelium		0	0	0	0	0	0	0	0	1	0	0	0	-	-	-	-
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(-)	(-)	(-)	(-)

{Digestive system}

liver			< 5>				< 5>				< 5>				< 0>			
	herniation		1	0	0	0	1	0	0	0	0	0	0	0	-	-	-	-
			(20)	(0)	(0)	(0)	(20)	(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

(HPT150)

BAIS4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 SACRIFICED ANIMALS (3W)

PAGE : 2

		4000ppm				8000ppm			
		No. of Animals on Study				No. of Animals on Study			
		Grade				Grade			
Organ	Findings	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}									
nasal cavit		< 0>				< 0>			
	disarrangement:olfactory epithelium	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)
		< 0>				< 0>			
	atrophy:olfactory epithelium	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)
{Digestive system}									
liver		< 0>				< 0>			
	herniation	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)

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

(HPT150)

BAIS4

APPENDIX K 3

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : SUMMARY,
RAT : FEMALE :
DEAD AND MORIBUND ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 9

Organ	Findings	Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
nasal cavit	thrombus		< 0>				< 0>				< 0>				< 5>			
		-	-	-	-	-	-	-	-	-	-	-	-	0	0	1	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(20)	(0)	
	disarrangement:olfactory epithelium	-	-	-	-	-	-	-	-	-	-	-	-	3	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(60)	(0)	(0)	(0)	
	atrophy:olfactory epithelium	-	-	-	-	-	-	-	-	-	-	-	-	2	3	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(40)	(60)	(0)	(0)	
	necrosis:olfactory epithelium	-	-	-	-	-	-	-	-	-	-	-	-	5	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(100)	(0)	(0)	(0)	
lung	congestion		< 0>				< 0>				< 0>				< 5>			
		-	-	-	-	-	-	-	-	-	-	-	-	1	2	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(40)	(0)	(0)	
	hemorrhage	-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)	
	edema	-	-	-	-	-	-	-	-	-	-	-	-	3	2	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(60)	(40)	(0)	(0)	
	thrombus	-	-	-	-	-	-	-	-	-	-	-	-	0	1	0	0	
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(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																	

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 10

		4000ppm				8000ppm			
		No. of Animals on Study				No. of Animals on Study			
		Grade				Grade			
Organ	Findings	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}									
nasal cavit									
	thrombus	< 5>				< 5>			
		0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	disarrangement:olfactory epithelium	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	atrophy:olfactory epithelium	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	necrosis:olfactory epithelium	0	1	4	0	0	0	0	0
		(0)	(20)	(80)	(0)	(0)	(0)	(0)	(0)
lung									
	congestion	< 5>				< 5>			
		0	0	0	0	3	2	0	0
		(0)	(0)	(0)	(0)	(60)	(40)	(0)	(0)
	hemorrhage	0	0	0	0	0	1	0	0
		(0)	(0)	(0)	(0)	(0)	(20)	(0)	(0)
	edema	0	5	0	0	2	1	0	0
		(0)	(100)	(0)	(0)	(40)	(20)	(0)	(0)
	thrombus	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

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 11

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
lung	edema:perivascular		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	-	3	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(60)	(20)	(0)	(0)
{Hematopoietic system}																		
bone marrow	congestion		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	-	1	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(0)	(0)	(0)
thymus	atrophy		< 0>				< 0>				< 0>				< 4>			
			-	-	-	-	-	-	-	-	-	-	-	-	3	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(75)	(25)	(0)	(0)
	congestion		-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)
spleen	atrophy		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	-	4	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(80)	(20)	(0)	(0)
{Circulatory system}																		
heart	hemorrhage		< 0>				< 0>				< 0>				< 5>			
			-	-	-	-	-	-	-	-	-	-	-	-	0	1	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(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

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 12

		Group Name	4000ppm				8000ppm			
		No. of Animals on Study	5				5			
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>
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}										
lung			< 5>				< 5>			
	edema:perivascular		0	5	0	0	0	5	0	0
			(0)	(100)	(0)	(0)	(0)	(100)	(0)	(0)
{Hematopoietic system}										
bone marrow			< 5>				< 5>			
	congestion		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
thymus			< 5>				< 5>			
	atrophy		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	congestion		1	0	0	0	0	0	0	0
			(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
spleen			< 5>				< 5>			
	atrophy		0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Circulatory system}										
heart			< 5>				< 5>			
	hemorrhage		2	3	0	0	0	0	0	0
			(40)	(60)	(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

STUDY NO. : 0480
ANIMAL : RAT F344/DuCrj
REPORT TYPE : A1
SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 13

Organ	Findings	Group Name No. of Animals on Study				Control 0				500ppm 0				1000ppm 0				2000ppm 5			
		Grade																			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}																					
stomach	ulcer:forestomach	< 0>				< 0>				< 0>				< 5>				1	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(0)	(0)	(0)				
	erosion:glandular stomach																	1	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(0)	(0)	(0)				
liver	herniation	< 0>				< 0>				< 0>				< 5>				2	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(40)	(0)	(0)	(0)				
	necrosis:central																	1	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(20)	(0)	(0)	(0)				
	vacuolic change																	0	0	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)				
{Urinary system}																					
urin bladd	hemorrhage	< 0>				< 0>				< 0>				< 5>				0	1	0	0
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(20)	(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

(HPT150)

BAIS4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 14

		4000ppm				8000ppm			
		5				5			
		No. of Animals on Study							
		Grade							
Organ	Findings	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Digestive system}									
stomach		< 5>				< 5>			
	ulcer:forestomach	0	0	0	0	0	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)	(0)	(0)	(0)	(0)
liver		< 5>				< 5>			
	herniation	1	0	0	0	0	0	0	0
		(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	necrosis:central	1	0	0	0	0	0	0	0
		(20)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	vacuolic change	5	0	0	0	0	0	0	0
		(100)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Urinary system}									
urin bladd		< 5>				< 5>			
	hemorrhage	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

(HPT150)

BAIS4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 15

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	0				0				0				5			
		Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
Organ	Findings		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
<hr/>																		
{Endocrine system}																		
pituitary			< 0>				< 0>				< 0>				< 4>			
	Rathke pouch		-	-	-	-	-	-	-	-	-	-	-	-	1	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(25)	(0)	(0)	(0)	(0)
adrenal			< 0>				< 0>				< 0>				< 5>			
	hemorrhage		-	-	-	-	-	-	-	-	-	-	-	-	0	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(0)	(0)	(0)	(0)	(0)
 {Nervous system}																		
brain			< 0>				< 0>				< 0>				< 5>			
	degeneration:granular cell		-	-	-	-	-	-	-	-	-	-	-	-	5	0	0	0
			(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(100)	(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

(HPT150)

BAIS4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0- 3W)

PAGE : 16

		4000ppm				8000ppm			
		No. of Animals on Study				5			
		Grade				5			
Organ_____	Findings_____	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
<hr/>									
{Endocrine system}									
pituitary		< 5>				< 5>			
	Rathke pouch	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
adrenal		< 5>				< 5>			
	hemorrhage	0	1	2	0	0	0	0	0
		(0)	(20)	(40)	(0)	(0)	(0)	(0)	(0)
{Nervous system}									
brain		< 5>				< 5>			
	degeneration:granular cell	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

(HPT150)

BAIS4

APPENDIX K 4

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : SUMMARY,
RAT : FEMALE :
SACRIFICED ANIMALS
(2-WEEK STUDY)

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 SACRIFICED ANIMALS (3W)

PAGE : 3

		Group Name	Control				500ppm				1000ppm				2000ppm			
		No. of Animals on Study	5				5				5				0			
Organ	Findings	Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
nasal cavit			< 5>				< 5>				< 5>				< 0>			
	disarrangement:olfactory epithelium		0	0	0	0	5	0	0	0	5	0	0	0	-	-	-	-
			(0)	(0)	(0)	(0)	(100)	(0)	(0)	(0)	(100)	(0)	(0)	(0)	(-)	(-)	(-)	(-)
{Digestive system}																		
liver			< 5>				< 5>				< 5>				< 0>			
	herniation		2	0	0	0	1	0	0	0	2	0	0	0	-	-	-	-
			(40)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(40)	(0)	(0)	(0)	(-)	(-)	(-)	(-)
{Endocrine system}																		
pituitary			< 5>				< 5>				< 5>				< 0>			
	Rathke pouch		0	0	0	0	0	0	0	0	1	0	0	0	-	-	-	-
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(20)	(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

(HPT150)

BAIS4

STUDY NO. : 0480
 ANIMAL : RAT F344/DuCrj
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 SACRIFICED ANIMALS (3W)

PAGE : 4

Organ	Findings	Group Name				4000ppm				8000ppm			
		No. of Animals on Study				0				0			
		Grade				1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}													
nasal cavit		< 0>				< 0>				< 0>			
	disarrangement:olfactory epithelium	-	-	-	-	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)
{Digestive system}													
liver		< 0>				< 0>				< 0>			
	herniation	-	-	-	-	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)
{Endocrine system}													
pituitary		< 0>				< 0>				< 0>			
	Rathke pouch	-	-	-	-	-	-	-	-	-	-	-	-
		(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)	(-)

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

(HPT150)

BAIS4

APPENDIX L

METHODS, UNITS AND DECIMAL PLACE FOR HEMATOLOGY AND BIOCHEMISTRY IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

METHODS, UNITS AND DECIMAL PLACE FOR HEMATOLOGY AND BIOCHEMISTRY
IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

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
Prothrombin time	Quick one stage method ²⁾	sec	1
Activated partial thromboplastin time (APTT)	Ellagic acid activaterd method ²⁾	sec	1
White blood cell (WBC)	Light scattering method ¹⁾	$\times 10^3/\mu\text{L}$	2
Differential WBC	Pattern recognition method ³⁾ (Wright staining)	%	0
Biochemistry			
Total protein (TP)	Biuret method ⁴⁾	g/dL	1
Albumin (Alb)	BCG method ⁴⁾	g/dL	1
A/G ratio	Calculated as $\text{Alb} / (\text{TP} - \text{Alb})$ ⁴⁾	—	1
T-bilirubin	Alkaline azobilirubin method ⁴⁾	mg/dL	2
Glucose	GlcK·G-6-PDH method ⁴⁾	mg/dL	0
T-cholesterol	CE·COD·POD method ⁴⁾	mg/dL	0
Triglyceride	LPL·GK·GPO·POD method ⁴⁾	mg/dL	0
Phospholipid	PLD·ChOD·POD method ⁴⁾	mg/dL	0
Glutamic oxaloacetic transaminase (GOT)	JSCC method ⁴⁾	IU/L	0
Glutamic pyruvic transaminase (GPT)	JSCC method ⁴⁾	IU/L	0
Lactate dehydrogenase (LDH)	SFBC method ⁴⁾	IU/L	0
Alkaline phosphatase (ALP)	GSCC method ⁴⁾	IU/L	0
γ -Glutamyl transpeptidase (γ -GTP)	JSCC method ⁴⁾	IU/L	0
Creatine phosphokinase (CPK)	JSCC method ⁴⁾	IU/L	0
Urea nitrogen	Urease·GLDH method ⁴⁾	mg/dL	1
Creatinine	Jaffe method ⁴⁾	mg/dL	1
Sodium	Ion selective electrode method ⁴⁾	mEq/L	0
Potassium	Ion selective electrode method ⁴⁾	mEq/L	1
Chloride	Ion selective electrode method ⁴⁾	mEq/L	0
Calcium	OCPC method ⁴⁾	mg/dL	1
Inorganic phosphorus	PNP·XOD·POD method ⁴⁾	mg/dL	1

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

2) Automatic coagulometer (Sysmex CA-5000 : Sysmex Corporation)

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

4) Automatic analyzer (Hitachi 7070 : Hitachi, Ltd.)