ヒドラジン一水加物のラット及びマウスを用いた経口投与によるがん原性予備試験(混水試験)報告書

APPENDIX

(A1-1~A12-4)

2週間試験:ラット/0261;マウス/0262

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RAT: FEMALE: SACRIFICED ANIMALS

(TWO-WEEK STUDIES)

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APPENDIX A 1-1

CLINICAL OBSERVATION (TWO-WEEK STUDY: SUMMARY)

RAT: MALE

STUDY NO.: 0261 ANIMAL: RAT F344 REPORT TYPE: A1 2 CLINICAL OBSERVATION (SUMMARY) ALL ANIMALS

SEX : MALE

Clinical sign	Group Name	Admini	stration We	eek-day			 	 		
		1-1	1-3	1-7	2-3	2-7				
		1	1	1	1	1				
							 ·	 		
EATII	Control	0	0	0	0	0				
Putn	50 ppm	0	0	0	0	0				
	100 ppill	0	0	0	0	0				
	100 ppm	0	-			-				
	200 ppm	v	0	0	0	0				
	400 ppm	0	0	0	0	1				
	Mqq 008	0	0	0	0	10			*	
UNCHBACK POSITION	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	0	0				
	400 ppm	Ö	0	Ö	Ö	4				
	800 ppm	Ö	Ö	Ö	ŏ	ō				
ILOERECTION	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	0	0				
	400 ppm	0	0	0	0	9				
	mqq 008	0	0	0	0	0				
DILED PERI GENITALIA	Control	0	0	0	0	0				
DOI GENTINEIN	50 ppm	Ö	Ö	0	0	0				
	100 ppm	Ô	Ő	0	0	Ö				
	200 ppm	0	Ô	0	0	0				
	400 ppm	0	0	0	0	5				
	800 ppm	0	0	0	2	0				
	WHA AND	v	v	v	۵	V				
IALL STOOL	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	0	0				
	400 ppm	0	0	9	10	10				
	mag 008	0	0	10	10	1				

STUDY NO.: 0261 ANIMAL : RAT F344 REPORT TYPE : A1 2

Group Name

Control

50 ppm 100 ppm 200 ppm

400 ppm

CLINICAL OBSERVATION (SUMMARY) ALL ANIMALS

2-3

1

9

Administration Week-day _

1-7

1

0

3

1-3

1

3

1-1

1

0

0

SEX : MALE

Clinical sign

OLIGO-STOOL

PAGE: 2

10 10 800 ppm 0 10 1 BAIS 2 (HAN190)

2-7

1

0 0

10

APPENDIX A 1-2

CLINICAL OBSERVATION (TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE

STUDY NO.: 0261 ANIMAL: RAT F344 REPORT TYPE: A1 2 CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX: FEMALE

linical sign	Group Name	Λdmini	stration We	eek-day			,		 	
		1-1	1-3	1-7	2-3	2-7				
		1	1	1	1	1			 	
EATH	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	0	0				
	400 ppm	0	0	0	3	6				
	mqq 008	0	0	1	9	-				
OCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0				
ACCOUNTAIN HOTEREAT DECK	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	0	0				
	400 ppm	0	0	1	0	1				
	mqq 008	0	0	1	0					
	MAC ANO	V	V	1	V	_				
UNCHBACK POSITION	Control	0	. 0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0 ·				
	200 ppm	0	0	0	0	0				
	400 ppm	0	0	2	1	3				
	mqq 008	0	0	3	1	-				
ASTING	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	Ö	0	0	0				
	400 ppm	0	ŏ	1	Ö	Ö				
	800 ppm	ő	Ő	î	Õ	_				
	200 PMII	v	v	•	·					
COILED	Control	0	0	0	0	0				
	50 ppm	0	0	0	0	. 0				
	100 ppm	0	0	0	0	0				
	200 ppm	0	0	0	1	1				
	400 ppm	0	0	0	1	1				
	800 ppm	0	0	0	0	_				

CLINICAL OBSERVATION (SUMMARY) ALL ANIMALS

STUDY NO.: 0261 ANIMAL: RAT F344 REPORT TYPE: A1 2

SEX : FEMALE

Clinical sign	Group Name	Admini	stration W	eek-day				
		1-1	1-3	1-7	2-3	2-7		
		1	1	1	1	1		
PILOERECTION	Control	0	0	0	0	0		
	50 ppm	0	0	0	0	0		
	100 ppm	0	0	0	0	0		
	200 ppm	0	0	0	0	0		
	400 ppm	0	0	0	0	4		
	mag 008	0	0	0	0	-		
coller per college			_					
SOILED PERI GENITALIA	Control	0	0	0	0	0		
	50 ppm	0	0	0	0	0		
	100 ppm	0	0	0	0	0		
	200 ppm	0	0	2	5	7		
	400 ppm	0	0	7	5	6		
	mqq 008	0	2	10	3	-		
NOSE HEMORRHAGIC DISCHA	Control	0	0	0	0	0		
	50 ppm	0	0	0	0	0		
	100 ppm	0	0	0	0	0		
	200 ppm	0	0	0	0	0		
	400 ppm	Ö	Ö	0	0	Ö		
	mqq 008	0	0	0	1	-		
CHAIL CTOOL	<i>a</i> , ,	•	0		•			
SHALL STOOL	Control	0	0	0 .	0	0		
	50 ppm	0	0	0	0	0		
	100 ppm	0	0	0	0	0		
	200 ppm	0	0	5	6	6		
	400 ppm	0	0	3	6	6		
	mqq 008	0	0	5	2	-		
OLIGO-STOOL	Control	0	0	0	0	0		
	50 ppm	0	0	0	0	0		
	100 ppm	0	0	Õ	ŏ	ő		
	200 ppm	ő	0	4	6	7		
	400 ppm	0	7	9	7	6		
	800 ppm	0	10	10	3	-		
	OAA PHIL	v	10	10	v	=		

(HAN190)

APPENDIX A 1-3

CLINICAL OBSERVATION (TWO-WEEK STUDY: SUMMARY)

MOSUE: MALE

ANIMAL : MOUSE BDF1 REPORT TYPE : A1 2

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

Clinical sign	Group Name	Adminis	stration W	eek-day		
•		1-1	1-3	1-7	2-3	2-7
		1	1	1	1	1
DEATH	Control	0	0	0	0	0
	50 ppm	0	0	0	0	0
	100 ppm	0	0	0	0	0
	200 ppm	0	0	0	0	0
	400 ppm	0	0	0	0	0
	Mqq 008	0	0	0	2	9
		•	v	·	2	ŭ
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0
	50 ppm	0	0	0	0	0
	100 ppm	0	0	0	0	0
	200 ppm	0	0	0	0	0
	400 ppm	0	0	0	0	0
	mqq 008	0	0	0	0	1
IIIIIIIII DOGUTA OU		_	_	_		_
HUNCHBACK POSITION	Control	0	0	0	0	0
	50 ppm	0	0	0	0	0
	100 ppm	0	0	0	0	0
	200 ppm	.0	0	0	0	0
	400 ppm	0	0	0	0	6
	mqq 008	0	0	2	5	1
PILOERECTION	Control	0	0	٥	0	0
FIBORNEOTION	Control 50 ppm	0	0	0	0	0
		0	0	0		
	100 ppm	v	0	0	0	0
	200 ppm	0	v	-	0	4
	400 ppm	0	0	3	4	10
	Mqq 008	0	0	8	6	1
IRREGULAR BREATHING	Control	0	0	0	0	0
	50 ppm	0	Õ	0	Ŏ	0
	100 ppm	Ô	0	0	0	Ŏ
	200 ppm	0	0	0	0	0
	400 ppm	0	0	0	0	0
	mag 008	0	0	0	0	1
	OAA PHIII	V	V	v	v	1

ANIMAL : MOUSE BDF1 REPORT TYPE: A1 2

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

PAGE: 2

1-1	Clinical sign	Group Name	Admini	stration W	eek-day _		
50 ppm						2-3	2-7
50 ppm			1	1	1	1	1
50 ppm							
100 ppm 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	ABNORMAL RESPIRATION	Control	0	0	0	0	0
200 ppm 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0		50 ppm	0	0	0	0	0
400 ppm 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0		100 ppm	0	0	0	0	0
800 ppm 0 0 0 0 1 GO-STOOL Control 0 0 0 0 0 0 50 ppm 0 0 0 0 0 100 ppm 0 0 0 0 0 200 ppm 0 0 0 0 0 400 ppm 0 0 0 0 0 10		200 ppm	0	0	0	0	0
GO-STOOL Control 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0		400 ppm	0	0	0	0	0
50 ppm 0 0 0 0 0 0 100 ppm 0 0 0 0 0 200 ppm 0 0 0 0 0 400 ppm 0 0 0 0 10		Mqq 008	0	0	0	0	1
100 ppm 0 0 0 0 0 0 0 200 ppm 0 0 0 0 0 0 0 400 ppm 0 0 0 0 10	OLIGO-STOOL	Control.	0	0	0	0	0
200 ppm 0 0 0 0 0 400 ppm 0 0 0 0 10		50 ppm	0	0	0	0	0
400 ppm 0 0 0 0 10		100 ppm	0	0	0	0	0
		200 ppm	0	0	0	0	0
800 ppm 0 2 5 5 1		400 ppm	0	0	0	0	10
		800 ppm	0	2	5	5	1
	00)						

(IIAN190)

BAIS 2

APPENDIX A 1-4

CLINICAL OBSERVATION (TWO-WEEK STUDY: SUMMARY)

MOSUE: FEMALE

STUDY NO. : 0262 ANIMAL : MOUSE BDF1
REPORT TYPE : A1 2

CLINICAL OBSERVATION (SUMMARY) ALL ANIMALS

SEX: FEMALE

Clinical sign	Group Name	Admini	stration W	eek-day			 		
		1-1	1-3	1-7	2-3	2-7			
		1	1	1	1	1			
DEATH	Control	0	0	0	0	0			
	50 ppm	0	0	0	0	0			
	100 ppm	0	0	0	0	0			
	200 ppm	0	0	0	0	0			
	400 ppm	0	0	0	0	1			
	mqq 008	0	0	0	3	10			
COMOTOR MOVEMENT DECR	Control	0	0	0	0	0			
	50 ppm	0	0	0	0	0		*	
	100 ppm	ő	0	0	0	0			
	200 ppm	0	0	0	0	0			
	400 ppm	0	0	0	0	0			
	mqq 008	0	0	1	0	0			
	ilidd ooo	V	V	ı	U	U			
UNCHBACK POSITION	Control	0	0	0	0	0			
	50 ppm	0	0	0	0	0			
	100 ppm	0	0	0	0	0			
	200 ppm	0	0	0	0	0			
	400 ppm	0	0	0	0	3			
	mqq 008	0	0	2	3	0			
LOERECTION	Control	0	0	0	0	0			
	50 ppm	0	0	0	0	0			
	100 ppm	0	0	0	0	Ö			
	200 ppm	0	ŏ	1	2	3			
	400 ppm	ō	ŏ.	5	9	9			
	800 ppm	0	0	9	7	0			
	OVU PPIII	v	v	Ü	•	v			
)LIGO-STOOL	Control	0	0	0	0	0			
	50 ppm	0	0	0	Ö	Ö			
	100 ppm	Ö	0	0	Ŏ	Ö			
	200 ppm	Ö	0	0	ő	Ö			
	400 ppm	Ö	1	6	6	10			
	mag 008	0	5	9	7	1			

APPENDIX A 2-1

BODY WEIGHT CHANGES (TWO-WEEK STUDY: SUMMARY)

RAT: MALE

ANIMAL : RAT F344

UNIT : g

REPORT TYPE : A1 2

SEX: MALE

BODY WEIGHT CHANGES (SUMMARY) ALL ANIMALS

PAGE: 1

oup Name	Administ	ration	n week-day								
	0-0		1-1		1-3	1-7		2-3		2-7	
Control	124±	5	128±	6	135± 7	150±	9	162±	10	178±	12
50 ppm	124±	4	125±	4	130± 5	144±	6	153±	8	166±	10*
100 ppm	124±	4	122±	4*	127± 6*	144±	6	150±	7	161±	7**
200 ppm	124±	4	120±	4**	123± 6**	126±	9**	129±	9**	137±	9**
400 ppm	124±	5	117±	4**	108± 5**	89±	5**	79±	6**	68±	5**
800 ppm	124±	4	115±	4**	102± 4**	82±	3**	70±	3**	_	
Significant differend	ce; *:P≦0.0)5	** : P ≤ 0.01			Test of Du	nnett				

(IIAN260)

BAIS 2

APPENDIX A 2-2

BODY WEIGHT CHANGES (TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE

ANIMAL : RAT F344

UNIT : g

REPORT TYPE : A1 2

SEX : FEMALE

BODY WEIGHT CHANGES ALL ANIMALS

(SUMMARY)

up Name	Admini	stration	n week-day									
	0-0		1-1		1-3		1-7		2-3		2-7	
Control	100±	4	100±	3	107土	4	117±	5	122±	5	129±	7
50 ppm	100±	4	96±	5*	101±	4*	108±	4 **	111±	4**	117±	4**
100 ppm	99±	4	96±	3*	94±	4**	104土	4**	104±	6**	110±	7**
200 ppm	99±	4	94±	4**	87±	4**	79±	6**	74±	8**	75±	9**
400 ppm	99±	4	92±	4**	83±	4**	65±	5**	62±	3**	57±	5**
800 ppm	100±	4	91±	4**	79±	4**	62±	4**	56±	0 ?	· –	
Significant differe	ence; *:P≦(0.05	**: P ≤ 0.0)1			Test of Du	nnett				

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(HAN260)

BAIS 2

APPENDIX A 2-3

BODY WEIGHT CHANGES (TWO-WEEK STUDY: SUMMARY)

MOSUE: MALE

ANIMAL : MOUSE BDF1

UNIT : g
REPORT TYPE : A1 2

SEX : MALE

BODY WEIGHT CHANGES ALL ANIMALS

(SUMMARY)

PAGE ; 1

owo Name	Λdministratio	n week-day					
	0-0	1-1	1-3	1-7	2-3	2-7	
Control	22.8± 0.9	22.3± 0.9	23.4± 0.8	24.2± 0.9	24.7± 1.0	25.2± 1.0	
50 ppm	22.7± 0.9	21.5± 0.8	21.8± 0.9**	22.7± 0.9*	23.3± 1.1*	24.1± 1.2	
100 ppm	22.8± 0.9	21.0± 0.8**	21.4± 0.8**	21.1± 1.5**	22.3± 0.8**	23.0± 0.9**	
200 ppm	22.7± 0.9	20.5± 0.8**	19.4± 0.8**	19.0± 0.9**	18.9± 1.0**	19.0± 1.3**	
400 ppm	22.7± 0.9	20.4± 0.9**	18.6± 1.0**	16.1± 1.3**	14.9土 1.4**	13.2± 1.1**	
MOC MOCH 008	22.7± 0.9	20.0± 0.9**	17.8± 1.0**	14.5± 1.1**	13.3± 0.7**	12.8± 0.0 ?	
Significant differer	nce; *:P≦0.05	** : P ≤ 0.01		Test of Dunnett			

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(HAN260)

BAIS 2

APPENDIX A 2-4

BODY WEIGHT CHANGES (TWO-WEEK STUDY: SUMMARY)

MOSUE: FEMALE

ANIMAL : MOUSE BDF1

UNIT : g
REPORT TYPE : A1 2
SEX : FEMALE

BODY WEIGHT CHANGES ALL ANIMALS

(SUMMARY)

oup Name	Administration	n week-day					
	0-0	1-1	1-3	1-7	2-3	2-7	
Control	18.6± 0.6	18.0± 0.4	18.7± 0.8	19.3± 0.6	19.3± 0.9	19.9± 0.7	
50 ppm	18.6± 0.6	16.9± 0.6**	17.0± 1.1	18.4± 1.0*	18.4± 0.8*	19.0± 0.8	
100 ppm	18.6± 0.6	16.6± 0.7**	16.1± 0.9	16.8± 1.1**	16.9± 1.0**	17.9± 1.0**	
200 ppm	18.6± 0.6	16.6± 0.4**	15.4生 0.4**	14.4± 0.6**	14.1± 0.6**	13.8± 0.9**	
400 ppm	18.6± 0.6	16.3± 0.4**	14.7± 0.5**	12.6± 0.7**	11.5± 0.6**	10.5生 0.4**	
800 ppm	18.6± 0.6	16.1± 0.6**	14.2± 0.7**	11.7± 0.7**	10.7± 0.4**	***	
Significant difference;	* : P ≤ 0.05	** : P ≤ 0.01		Test of Dunnett			
ANGCO)							

(IIAN260)

BAIS 2

APPENDIX A 3-1

WATER CONSUMPTION CHANGES: SUMMARY, RAT: MALE (TWO-WEEK STUDY)

ANIMAL : RAT F344

UNIT : g

REPORT TYPE : A1 2 SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY) ALL ANIMALS

PAGE: 1

Group Name	Administration			
	1-3(3)	1-7(4)	2-3(3)	2-7(4)
Control	17.44 2.0	17.3± 1.7	17 7 1 1 5	16 64 1 9
CONTRIBLE	17.4± 2.0	11.51 1.1	17.7± 1.5	16.6± 1.2
50 ppm	11.0± 1.0	11.7± 0.8	11.9± 1.0	11.4± 1.5**
100 ppm	9.7± 1.3*	12.3± 1.2	10.8± 0.7*	11.4± 1.2**
200 рра	8.4± 1.1**	9.5± 1.5**	9.5± 2.0**	9.6± 1.3**
400 ppm	3.9± 0.8**	2.0± 0.9**	2.4± 1.1**	2.1± 0.7**
800 ppm	1.8± 0.3**	1.3± 0.3**	1.1± 0.2**	-
Significant differe	nce; *:P ≤ 0.05	** : P ≤ 0.01		Test of Dunnett

(HAN260)

BAIS 2

APPENDIX A 3-2

WATER CONSUMPTION CHANGES: SUMMARY, RAT: FEMALE (TWO-WEEK STUDY)

ANIMAL : RAT F344

UNIT : g
REPORT TYPE : A1 2

SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY) ALL ANIMALS

PAGE: 2

Group Name	Administration	week-day(effective)		
	1-3(3)	1-7(4)	2-3(3)	2-7(4)
Control	14.6± 0.8	14.7± 1.1	14.8± 1.1	13.7± 2.2
50 ppm	9.0± 0.4	9.6± 1.0	9.1± 0.7**	8.8± 0.4
100 ppm	6.3± 1.1*	9.8± 2.0	7.7± 1.5**	8.2± 1.8**
200 ppm	4.0± 0.6**	4.4± 1.4**	4.9± 1.4**	5.6± 1.7**
400 ppm	2.4± 0.5**	1.4± 0.4**	3.7± 1,4**	3.1± 1.2**
800 pm	1.5士 0.2**	1.2± 0.1**	1.6± 0.0 ?	-
Significant differen	ce; $*:P \leq 0.05$	** : P ≤ 0.01		Test of Dunnett

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(HAN260)

BAIS 2

APPENDIX A 3-3

WATER CONSUMPTION CHANGES: SUMMARY, MOUSE: MALE (TWO-WEEK STUDY)

ANIMAL : MOUSE BDF1

UNIT : g

REPORT TYPE : A1 2

SEX : MALE

WATER CONSUMPTION CHANGES (SUMMARY)

PAGE: 1

ALL ANIMALS

up Name	Administration	n week-day(effective)			
	1-3(3)	1-7(4)	2-3(3)	2-7(4)	
Control	3.9± 0.5	4.0± 0.5	4.0± 0.5	3.8± 0.5	
50 ppm	1.9± 0.3	2.0± 0.2	1.8± 0.2	1.6± 0.2	
,					
100 ppm	1.6± 0.3	1.6± 0.6*	1.7± 0.6	1.6± 0.4*	
200 ppm	0.8± 0.2**	1.1± 0.1**	0.9± 0.2**	0.9± 0.3**	
400 ppm	0.5± 0.1**	0.6± 0.2**	0.4± 0.2**	0.4± 0.1**	
mag 008	0.3± 0.0**	0.3± 0.1**	0.3± 0.1**	0.3± 0.0 ?	
	$*: P \leq 0.05$	** : P ≤ 0.01		Test of Dunnett	

APPENDIX 3-4

WATER CONSUMPTION CHANGES: SUMMARY, MOUSE: FEMALE

(TWO-WEEK STUDY)

ANIMAL : MOUSE BDF1

UNIT : g

REPORT TYPE : A1 2

SEX : FEMALE

WATER CONSUMPTION CHANGES (SUMMARY)

ALL ANIMALS

Administration week-day(effective)_ Group Name 1-7(4) 2-3(3) 2-7(4) 1-3(3) Control 3.6± 0.4 4.0 ± 0.3 4.1± 0.3 4.1± 0.4 50 ppm 1.6± 0.5 2.0± 0.2 1.7 ± 0.3 1.7 ± 0.3 100 ppm 1.1± 0.3* 1.6± 0.2 1.3± 0.2* 1.6± 0.3* 200 ppm 0.7± 0.1** 0.8± 0.1** 0.8± 0.1** 0.8± 0.1** 400 ppm 0.4± 0.1** 0.5± 0.1** 0.4± 0.0** 0.5± 0.1** 800 ppm 0.3± 0.1** 0.3± 0.0** 0.3± 0.1** Significant difference; $*: P \leq 0.05$ ** : $P \le 0.01$ Test of Dunnett

(HAN260)

BAIS 2

APPENDIX A 4-1

FOOD CONSUMPTION CHANGES : SUMMARY, RAT : MALE
(TWO-WEEK STUDY)

ANIMAL : RAT F344

UNIT : g
REPORT TYPE : A1 2

SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)

ALL ANIMALS

Group Name	Administration	week		
	1	2		
Control	12.8± 1.0	13.5± 1.0		
50 ppm	11.3± 0.6	11.8± 1.1**		
100 ppm	11.2± 0.7	11.7± 0.7**		
200 ppm	9.4± 1.3**	9.7± 0.9**		
400 ppm	4.5± 0.4**	3.5± 0.5**		
800 ppm	3.5± 0.2**	-		
Significant diffe	erence; *: P ≤ 0.05	** : P ≤ 0.01	Test of Dunnett	
(HAN260)				BAIS

BAIS 3

APPENDIX A 4-2

FOOD CONSUMPTION CHANGES: SUMMARY, RAT: FEMALE

(TWO-WEEK STUDY)

STUDY NO.: 0261 ANIMAL : RAT F344

UNIT : g REPORT TYPE : A1 2 FOOD CONSUMPTION CHANGES (SUMMARY)

ALL ANIMALS

SEX : FEMALE

Group Name Administration week_ 1 2 Control 10.3± 0.7 10.1± 0.8 50 ppm 8.8± 0.4** 9.0± 0.3** 100 ppm 7.8± 0.4** 8.5± 0.7** 200 ppm 4.7± 0.6** 5.3± 0.8** 400 ppm 2.8± 0.4** 4.0± 0.6** 800 ppm 2.3± 0.3** Significant difference; $*: P \leq 0.05$ ** : $P \le 0.01$ Test of Dunnett (HAN260)

BAIS 3

PAGE: 2

APPENDIX A 4-3

FOOD CONSUMPTION CHANGES: SUMMARY, MOSUE: MALE

(TWO-WEEK STUDY)

STUDY NO.: 0262

ANIMAL : MOUSE BDF1

UNIT : g REPORT TYPE : A1 2 FOOD CONSUMPTION CHANGES (SUMMARY) ALL ANIMALS

SEX : MALE

(HAN260)

PAGE: 1 Group Name Administration week_ 2 1 Control 3.6 ± 0.2 3.7 ± 0.2 50 ppm 3.2生 0.2** 3.7 ± 0.2 100 ppm 2.9± 0.3** 3.7± 0.4 200 ppm 2.4生 0.1** 3.2± 0.2** 1.8± 0.2** 2.1士 0.2** 400 ppm 800 ppm 1.6± 0.2** 1.7生 0.0 ? Significant difference : $*: P \leq 0.05$ **: $P \le 0.01$ Test of Dunnett ?: Significant test is not applied, because No. of data in this group is less than 3.

APPENDIX A 4-4

FOOD CONSUMPTION CHANGES: SUMMARY, MOSUE: FEMALE
(TWO-WEEK STUDY)

STUDY NO.: 0262

ANIMAL : MOUSE BDF1

UNIT : g REPORT TYPE : A1 2

SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY) ALL ANIMALS

roup Name	Administration u	ueek		
	1	2		
Control	3.0± 0.2	3.4± 0.2		
50		0.1.10		
50 ppm	2.8± 0.3*	3.1生 0.2**		
100 ppm	2,5± 0,2**	3.1± 0.3*		
200 ppm	2.0± 0.2**	2.5± 0.3**		
400 ppm	1.5± 0.2**	2.0± 0.2**		
800 ppm	1.3 ± 0.2**	-		
Significant differe	ence; *:P≦0.05 **	*: P ≤ 0.01	Test of Dunnett	

(HAN260)

BAIS 3

PAGE: 2

APPENDIX A 5-1

CHEMICAL INTAKE CHANGES: SUMMARY, RAT: MALE

(TWO-WEEK STUDY)

STUDY NO. :0261
ANIMAL :RAT F344
UNIT :mg/kg/day
REPORT TYPE :A1 2
SEX :MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-	day(effective)	2-3(3)	2-7(4)	
Control	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	
50ppm	4.213 ± 0.292	4.046 ± 0.180	3.867 ± 0.161	3.429 ± 0.252	
100ppm	7.556 ± 0.711	8.572 ± 0.749	7.216 ± 0.388	7.111 ± 0.806	
200ррш	13.709 ± 1.427	15.135 ± 2.130	14.744 ± 3.165	13.970 ± 1.775	
400ppm	14.358 ± 2.579	8.611 ± 3.453	12.006 ± 5.000	12.045 ± 3.527	
800ppm	14.289 ± 1.701	12.355 ± 2.367	12.497 ± 1.842	-	

APPENDIX A 5-2

CHEMICAL INTAKE CHANGES: SUMMARY, RAT: FEMALE

(TWO-WEEK STUDY)

STUDY NO.

:0261

ANIMAL :RAT F344
UNIT :mg/kg/day
REPORT TYPE :A1 2 SEX :FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY) ALL ANIMALS

Group Name	Administration week-				
	1-3(3)	1-7(4)	2-3(3)	2-7(4)	
Control	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	
50ppm	4.452 ± 0.239	4.459 ± 0.520	4.118 ± 0.350	3.749 ± 0.232	
100ppm	6.693 ± 1.060	9.435 ± 1.942	7.395 ± 1.327	7.420 ± 1.360	
200ppm	9.111 ± 1.028	11.000 ± 2.982	13.342 ± 3.771	14.762 ± 3.802	
400ppm	11.442 ± 2.106	8.619 ± 2.149	23.967 ± 8.424	21.555 ± 6.572	
800ррш	15.219 ± 1.821	15.857 ± 2.083	22.857 ± 0.000		

APPENDIX A 5-3

CHEMICAL INTAKE CHANGES: SUMMARY, MOUSE: MALE (TWO-WEEK STUDY)

TUDY NO. :0262
ANIMAL :BDF1 MOUSE
UNIT :mg/kg/day
REPORT TYPE :A1 2
STX :MALE

CHEMICAL INTAKE CHANGES (SUMMARY)

ALL ANIMALS

Group Name	Administration week-	day(effective)			
	1-3(3)	1-7(4)	2-3(3)	2-7(4)	
Control	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	
50ррт	4.458 ± 0.786	4.480 ± 0.338	3.803 ± 0.384	3.382 ± 0.284	
.00ppm	7.376 ± 1.175	7.215 ± 2.542	7.704 ± 2.833	6.699 ± 1.278	
00ppm	7.950 ± 1.611	11.132 ± 1.139	9.385 ± 2.229	9.506 ± 1.976	
mqq00	10.716 ± 2.355	13.903 ± 4.046	11.748 ± 3.933	13.438 ± 3.953	
300ppm	12.637 ± 2.066	18.893 ± 3.163	18.914 ± 7.72	18.750 ± 0.000	

APPENDIX A 5-4

CHEMICAL INTAKE CHANGES: SUMMARY, MOUSE: FEMALE (TWO-WEEK STUDY)

STUDY NO. :0262

ANIMAL :BDF1 MOUSE
UNIT :mg/kg/day
REPORT TYPE :A1 2
SEX :FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY) ALL ANIMALS

Group Name	Administration week- 1-3(3)		0.0/0\	0.7/1)	
	1-3(3)	1-7(4)	2-3(3)	2-7(4)	
Control	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	0.000 ± 0.000	
50ppm	4.713 ± 1.181	5.483 ± 0.317	4.493 ± 0.651	4.557 ± 0.771	
100ppm	6.655 ± 1.547	9.270 ± 1.128	7.534 ± 0.981	8.889 ± 1.681	
200ppm	9.492 ± 1.149	11.649 ± 0.923	11.043 ± 1.514	11.733 ± 1.720	
400ppm	11.998 ± 3.094	15.217 ± 2.633	13.579 ± 1.355	18.226 ± 3.955	
800ppm	16.361 ± 4.120	19.235 ± 3.076	20.497 ± 6.278	_	

APPENDIX A 6-1

HEMATOLOGY (TWO-WEEK STUDY : SUMMARY)

RAT: MALE

STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1 HEMATOLOGY(1) (SUMMARY) SURVIVAL ANIMALS (2)

SEX : MALE

PAGE: 1

Group Name	NO. of Animals	RED BLC 1 O ⁶ /µ	OOD CELL	g∕dl HEMOGLO	BIN	HEMATOC %	CRIT	MCV f @		MCH pg		MCHC g∕dl		PLATELE 1 Ο³ / μ	
Control	5	7.52±	0.09	13.8±	0.2	40.7±	0.4	54,2±	0.6	18.4±	0.2	33.9±	0.3	691±	117
50 ppm	5	7.65±	0.35	14.0±	0.4	40.5±	1.8	52.9±	0.4	18.3±	0.4	34.6±	0.9	629±	51
100 ppm	5	7.76±	0.20	13.9±	0.4	40.8±	0.9	52.6±	0.4	17.9±	0.2	34.1±	0.3	632±	67
200 ppm	5	8.07±	0.30	14.2±	0.7	41.6±	1.7	51.5±	0.4**	17.6±	0.2**	34.2±	0.2	582±	164
400 ppm	5	11.80±	0.95**	21.5±	1.8**	66.8±	7.6*	56.5±	1.9	18.2±	0.2	32.3±	1.1	153±	76**
mqq 008	0	-		-		-		-		-		-		-	
Significan	t difference;	*: P ≦ ().05	**: P ≤ 0.0)1			Test of Dur	nnett						

(IICL070)

STUDY NO. : 0261 ANIMAL : RAT F344 HEMATOLOGY(1) (SUMMARY)
SURVIVAL ANIMALS (2)

REPORT TYPE : A1 SEX : MALE

PAGE: 2

Group Name	NO. of Animals	RETICUL	DCYTE	PROTIIROM sec	IBIN TIME	APTT sec					
Control	5	51±	6	13.3±	0.3	18.9±	1.6				
50 ppm	5	37±	9*	13.2±	0.4	19.7±	2.4				
100 ppm	5	34±	7**	13.2±	0.2	19.3±	0.6				
200 ppm	5	33±	10**	13.6±	0.6	19.8±	3.8		•		
400 ppm	5	9±	4**	15.5±	1.2	20.6±	2.5				
Mag 008	0			-		-					
Significan	t difference;	*: P ≤ 0	.05	**: P ≤ 0.01	1			Test of Dunnet			
(1101.070)		• • • • • • • • • • • • • • • • • • • •							 	· · · · · · · · · · · · · · · · · · ·	 DAIC

(IICL070)

STUDY NO.: 0261 ANIMAL: RAT F344 REPORT TYPE: A1 HEMATOLOGY(2) (SUMMARY)
SURVIVAL ANIMALS (2)

SEX : MALE

Group Name NO. of WBC Differential WBC (%) Animals 1 03/με N-BAND N-SEG EOSINO LYMPHO BASO NONO OTHERS Control 5 2.80± 0.92 0± $22\pm$ 0 7 $1\pm$ $0\pm$ $5\pm$ $72\pm$ 7 0 $0\pm$ 1 50 ppm 5 2.54± 0.67 $0\pm$ 1 $18\pm$ 6 $1\pm$ 1 0± 77± 0 $3\pm$ 7 $0\pm$ 1

100 ppm 5 2.74± 0.91 $0\pm$ 0 21土 7 0± $0\pm$ 0 2± 1 76± 6 0土 1 200 ppm 5 2.09± 0.83 0土 0 $19\pm$ 6 1土 1 0± 0 4土 2 $76 \pm$ 6 $1\pm$ 1 400 ppm 5 1.66± 0.27 $0\pm$ 41± 12** $1 \pm$ 1 0± 0 4± $53\pm$ 0± 11** 0 800 ppm 0

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

Test of Dufflett

(HCL071)

BAIS 2

PAGE: 1

APPENDIX A 6-2

HEMATOLOGY (TWO-WEEK STUDY : SUMMARY)

RAT: FEMALE

STUDY NO.: 0262
ANIMAL: MOUSE BDF1
REPORT TYPE: A1
SEX: FEMALE

HEMATOLOGY(1) (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 2

9.87± 10.33± 10.26±	0.38	15.1± 15.7± 15.3±	0.2	45.2± 46.8±	1.5	45.8±	0.5	15.3± 15.2±	0.6	33.5±	1.0	985± 1023±	77 77
							0.7	15.2±	0.5	33,5±	0.7	1023主	77
10.26±	0.28	15.3±	0.4	46 6±	1 4								
				40.04	1.4	45.4±	0.3	14.9±	0.1	32.8±	0.3	1020±	60
10.30±	0,80	15.7±	0.4	46.1±	3.5	44.7±	0.8	15.4±	1.0	34.3±	2.1	995±	99
10.94±	1.20	16.2±	1.8	49.9±	6.6	45.5±	1.1	14.8±	0.1	32.5±	0.7	1002±	164
-		-		~		-				-		-	
	10.94±	10.94± 1.20	10.94± 1.20 16.2±	10.94± 1.20 16.2± 1.8	10.94± 1.20 16.2± 1.8 49.9±	10.94± 1.20 16.2± 1.8 49.9± 6.6	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5±	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1 14.8±	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1 14.8± 0.1	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1 14.8± 0.1 32.5±	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1 14.8± 0.1 32.5± 0.7	10.94± 1.20 16.2± 1.8 49.9± 6.6 45.5± 1.1 14.8± 0.1 32.5± 0.7 1002±

(IICL070)

STUDY NO. : 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX: FEMALE

HEMATOLOGY(2) (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 2

roup Name	NO. of Animals	WBC 1 O³/	μl		ferential BAND	WBC (%		EOS	SINO	BAS	50	МОМ	10	LYì	IРНО	OT	THERS
Control	5	2.34±	1.01	0±	0	12±	3	2±	1	0±	0	1±	1	85±	3	0±	0
50 ppm	5	2.60±	0.94	1 ±	1	9±	3	1±	1	0±	0	1±	1	88±	3	0±	0
100 ppm	5	2.52±	0.94	0±	0	10±	2	2±	1	0±	0	0±	1	88±	3	0±	0
200 ppm	5	0.55±	0.47	1±	1	17士	8	1±	1	0±	0	2±	2	78±	7	0±	0
400 ppm	5	0.26±	0.18*	3±	2	45±	8	0±	0	0±	0	0±	0	51±	8**	0±	0
800 ppm	0	-		-		-		-		-		-		-			
	difference;	*: P ≦	0.05	**: P ≦	0.01			Test	of Dunne		-						

(HCL071)

APPENDIX A 6-3

HEMATOLOGY (TWO-WEEK STUDY : SUMMARY)

MOUSE: MALE

STUDY NO.: 0261
ANIMAL: RAT F344
REPORT TYPE: A1
SEX: FEMALE

HEMATOLOGY(1) (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 3

roup Name	NO. of Animals	RED BL	OOD CELL µ2	HEMOGLC g∕dl		HEMATOC %	CRIT	MCV f @		MCH pg		MCIIC g∕aશ		PLATELE 1 O³∕µ	
Control	5	8.12±	0.10	15.0±	0,2	43.5±	0.5	53.6±	0.5	18.5±	0.1	34.6±	0.3	666±	91
50 ppm	5	8.33±	0.15	15.1±	0.3	43.7±	1.1	52.4±	0.6	18.1±	0.1**	34.6±	0.1	508±	57*
100 ppm	5	8.40±	0.40	15.2±	0.7	43.7±	1.9	52.0±	0.3*	18.0±	0.2**	34.7±	0.4	565±	85
200 ppm	5	9.20±	0.51*	16.8±	1.1	48.2±	3.1	52.4±	0.5	18.2±	0.2*	34.8±	0.4	407±	75**
400 ppm	4	10.80±	0.97**	19.8±	1.7**	60.0±	7.2**	55.4±	1.8	18.4±	0.2	33.1±	1.1	288±	130**
800 ppm	0	-		-		-		-				-		_	

(IICL070)

STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1 SEX : FEMALE

HEMATOLOGY(1) (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 4

Group Name	NO. of Animals	RETICULO ‰	CYTE	PROTHRO sec	OMBIN TIME	APTT sec		
Control	5	30±	5	13.2±	0.2	18.1±	0.6	
50 ppm	5	27±	6	13.2±	0.1	18.3±	8.0	
100 ppm	5	28±	4	13.3±	0.2	17.9±	1,3	
200 ppm	5	9±	3**	13.5±	0.2	15.6±	8.0	
400 ppm	4	7±	2**	14.8±	1.0**	18.6±	4,2	
800 ppm	0	<u>-</u>		-		-		
Significan	t difference;	*: P ≤ 0	.05	**: P ≤ 0.0	01		Test of Dunnett	
IICL070)								BA

STUDY NO.: 0261
ANIMAL: RAT F344
REPORT TYPE: A1
SEX: FEMALE

HEMATOLOGY(2) (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 2

Group Name	NO. of Animals	WBC 1 O³∕µl			ferential AND	WBC (% N-S		EOS	SINO	BAS	0	MON	0	LY	ІРНО	OT	HERS
Control	5	2.62± 1.	30	0±	0	19±	5	2±	1	0±	0	2±	1	76±	5	0±	0
50 ppm	5	2.63± 0.	54	0±	0	18±	6	2±	1	0±	0	4±	1	76±	6	0±	0
100 ppm	5	2.86± 1.	19	0±	0	18土	6	1±	1	0±	0	3±	1	77±	6	1±	1
200 ppm	5	3.40± 1.	54	0±	0	26±	15	2±	1	0±	0	5±	2	66±	16	0±	0
400 ppm	4	2.28± 0.	54	0±	0	45±	8**	1±	1	0±	0	5±	2	50土	9**	0±	1
800 ppm	0	-		-				-		-		· •-		~		_	
Significan	nt difference :	*: P ≤ 0.	05 ×	**: P ≦	0.01	· · · · · · · · · · · · · · · · · · ·		Test	of Dunne	ett							

(IICL071)

APPENDIX A 6-4

HEMATOLOGY (TWO-WEEK STUDY : SUMMARY)

MOUSE: FEMALE

STUDY NO. : 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE HEMATOLOGY(1) (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 1

roup Name	NO. of Animals	RED BLO	OOD CELL	g∕dl BMOGLC	BIN	HEMATOC %	RIT	MCV f &		MCH pg		MCHC g∕dl		PLATELE 1 0 ³ /µ	
Control	5	10.63±	0.22	15.8±	0.4	49.0±	1.3	46.1±	0.3	14.8±	0.1	32.1±	0.4	1243土	45
50 ppm	5	10.77±	0.29	16.3±	0.5	49.7±	1.4	46.2±	0.7	15.1±	0.1	32.7±	0.5	1188±	67
100 ppm	5	10.84±	0.59	16.1±	0.8	50.0±	2.5	46.1±	0.6	14.8±	0.2	32.2±	0.3	1140±	44
200 ppm	5	11.36±	0.45	16.6±	0.7	50.5±	2.0	44.5±	0.7**	14.6±	0.2	33.0±	0.3	1099±	265
400 ppm	5	11.20±	0.95	16.8±	1.4	51.2±	5.1	45.6±	0.8	15.0±	0,8	33.0±	1.8	1045±	281
800 ppm	1	12.66±	0.00 ?	20.8±	0.0 ?	59.6±	0.0 ?	47.1±	0.0 ?	16.4±	0.0 ?	34.8±	0.0 ?	824±	0 ?

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL070)

STUDY NO. : 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE

HEMATOLOGY(2) (SUMMARY)
SURVIVAL ANIMALS (2)

Group Name	NO. of Animals	WBC 1 O ³	; - / με		ferential BAND		6) SEG	EOS	SINO	BAS	60	МОМ	10	LY	MPIIO	OTI	HERS
Control	5	2.15±	0.73	0±	0	9±	1	3±	2	0±	0	1±	1	87±	2	0±	0
50 ppm	5	2,23±	0.61	1±	1	9士	4	1±	2	0±	0	Ι±	1	88±	4	0±	0
100 ppm	5	2.08±	1.18	0±	1	10±	4	1±	1	0±	0	1±	1	88±	4	0±	0
200 ppm	5	1.85±	0.73	0±	1	17±	7	2±	1	0±	0	1±	1	80±	9	0±	0
400 ppm	5	0.29±	0.21**	4±	3	55±	22**	0±	0	0±	0	1±	1	40±	22*	0±	0
mqq 008	1	0.46±	0.00 ?	2±	0 ?	69±	0 ?	0±	0 ?	0±	0 ?	0±	0 ?	29±	0 ?	0±	0 ?
Significan	nt difference ;	* : P :	≤ 0.05	**:P≦	0.01			Test	of Dunnet	t							

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL071)

BAIS 2

PAGE: 1

APPENDIX A 7-1

BIOCHEMISTRY (TWO-WEEK STUDY : SUMMARY)

RAT: MALE

STUDY NO.: 0261 ANIMAL : RAT F344 REPORT TYPE : A1
SEX : MALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 1

oup Name	NO. of Animals	g∕dl TOTAL PI	ROTEIN	g∕dl ALBUMIN		A/G RAT	10	T-BILI mg/dl		GLUCOSE mg∕al		T-CHOLE mg∕dl	STEROL	PHOSPHO mg/dl	LIPID
Control	5	5.9±	0.1	3.3±	0.1	1.3±	0.1	0.19±	0.05	205±	16	60±	5	121±	8
50 ppm	5	5.9±	0.1	3.4±	0.0	1.4±	0.1	0.20±	0.07	195±	11	66±	3	128±	4
100 ppm	5	5.8±	0.2	3.4±	0.1	1.4±	0.1	0.18±	0.06	199±	10	59±	3	119±	7
200 ppm	5	6.0±	0.1	3.5±	0.1	1.5±	0.1	0.23±	0.08	190±	13	59±	4	115±	7
400 ppm	5	5.3±	1.0	3.0±	0.6	1.3±	0,3	0.89±	0.50*	105±	28**	63±	23	106±	30
mqq 008	0	-		-		, -		-		-		-		-	

(HCL074)

STUDY NO.: 0261
ANIMAL: RAT F344
REPORT TYPE: A1
SEX: MALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 2

TOUP Name	NO. of Animals	GOT I U / 4	e	GPT IU/e		LDII I U / 4	?	G−GTP IU∕ℓ		CPK IU/4	?	UREA N mg∕dl	ITROGEN	CREATIN mg∕dl	INE
Control	5	58±	6	19土	1	157±	13	0±	1	147±	43	14.2±	2.4	0.4±	0.1
50 ppm	5	54±	3	17±	1	176±	48	1±	1	142±	31	17.8±	2.9	0.4±	0.0
100 ppm	5	52±	4	16±	1	175±	32	0±	0	132±	5	17.9±	2.5	0.4±	0.0
200 ppm	5	57±	9	14±	1*	218±	64	0±	0	143±	32	17.5±	2.0	0.4±	0.0
400 ppm	5	565±	217	88±	34	923±	569**	1±	1	566±	452*	98.2±	66.7**	0.5±	0.2
800 ppm	0	-		-		-		-		-		-			

(IICL074)

STUDY NO.: 0261 ANIMAL: RAT F344 REPORT TYPE: A1 SEX: MALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 3

Group Name	NO. of Animals	SODIUM mEq/l	!	POTASSIU mEq/l		CHLORIDE mEq∕ℓ		CALCIUM mg/dl		INORGAN mg/dl	IC PHOSPHORUS
Control	5	142±	0	3.9±	0.2	106士	1	11.0±	0.5	8.3±	0.7
50 ppm	5	141±	1	4.0±	0.2	106生	1	10.9±	0.4	7.7±	0.7
100 ppm	5	141±	1	4.2±	0.1	105±	1	10.8±	0.2	7.6±	0.7
200 ppm	5	143±	2	4.8±	1.1	107±	1	10.2±	1.1	6.9±	0.6*
400 ppm	5	181±	14	4.8±	1.2	139±	12*	10.5±	0.4	11.6±	4.1
maja 008	. 0	-		-		-				-	
Significan	t difference;	*: P ≤ 0	.05	**: P ≤ 0.03	I			Test of Dunr	nett		

(HCL074)

APPENDIX A 7-2

BIOCHEMISTRY (TWO-WEEK STUDY : SUMMARY)

RAT: FEMALE

STUDY NO.: 0261
ANIMAL: RAT F344
REPORT TYPE: A1
SEX: FEMALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 4

roup Name	NO. of Animals	TOTAL PI g∕dl	ROTEIN	ALBUMIN g∕dશ		a∕G RAT	10	T-BILII mg/dl		GLUCOSE mg/dl		T-CHOLES	STEROL	PHOSPHOI mg/dl	JIPID
Control	5	5.7生	0.1	3.3±	0.0	1.4±	0.0	0.28±	0.04	199±	14	71±	3	127±	7
50 ppm	5	5.7±	0.1	3.4±	0.1	1.5±	0.1	0.34±	0.09	196±	10	74±	8	138±	9
100 ppm	5	5.8±	0.1	3.4±	0.1	1.5±	0.1	0.40±	0.12	183±	12	73±	3	133±	6
200 ppm	5	5.8±	0.3	3.5±	0.2	1.6±	0.1	0.49±	0.08	144±	20**	67±	8	127±	13
400 ppm	4	5.5±	0.4	3.1±	0.5	1.3±	0.3	0.78±	0.27**	92±	15**	92±	17	162±	16**
mqq 008	0	-		~		-		-		-		-		-	
Significan	t difference;	*: P ≤ 0	.05	** : P ≦ 0.0	1			Test of Du	nnett						

(IICL074)

STUDY NO. : 0261
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 5

oup Name	NO. of Animals	GOT IU/#	?	GPT IU∕ℓ		LDII IU/	e	G-GTP IU∕ℓ		CPK IU∕ℓ		UREA N mg∕dl	ITROGEN	CREATIN mg/dl	INE
Control	5	60±	2	19±	1	247±	84	1±	0	150±	12	17.1±	2.8	0.4±	0.1
50 ppm	5	62±	5	17±	1	252士	83	1±	0	140±	32	20.5±	2.3	0.4±	0.1
100 ppm	5	66±	5	17±	2	225±	58	1±	0	134±	32	19.7±	2.2	0.4±	0.1
200 ppm	5	112±	56*	21±	12	277±	70	2±	0	128±	27	25.6±	6.2	0.4±	0.1
400 ppm	4	223±	100**	29±	14	670±	260	3±	1	287±	99	53.4±	28.2**	0.3±	0.1**
Mqq 008	0	-		-		_		-		-		-		· -	

(HCL074)

STUDY NO. : 0261
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 6

Control	5	142±	1	3.9±	0.4	109±	2	10.3±	0.1	6.6±	0.8
50 ppm	5	142±	1	4.1±	0.2	108±	1	10.3±	0.1	7.0±	0.9
100 ppm	5	143±	1	4.4±	0.3*	109±	1	10.4±	0,3	7.1±	1.2
200 ppm	5	148±	1*	4.0±	0.1	112±	2	10.5±	0.4	6.3±	1.0
400 ppm	4	173±	17**	4.7±	0,3**	132±	13*	10.8±	0.4	9.7±	2,5**
mqq 008	0	_		-		-		-		-	

(HCL074)

APPENDIX A 7-3

BIOCHEMISTRY (TWO-WEEK STUDY : SUMMARY)

MOUSE: MALE

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 1

roup Name	NO. of Animals	TOTAL F	ROTEIN	g ∕aƙ ATBUMIN		A/G RAT	.10	T-BILI mg/dl		GLUCOSE mg/dl		T-CHOLES	STEROL	GOT IU/@	
Control	5	5.3±	0.2	3.0±	0.1	1.3±	0.0	0.36±	0.14	313±	15	101±	6	39±	11
50 ppm	5	5.0±	0.3	2.8±	0.1	1.3±	0.0	0.33±	0.09	325±	25	92±	7	41±	4
100 ppm	5	5.1±	0.2	2.9±	0.2	1.4±	0.1	0.30±	0.03	299±	18	96±	7	42±	6
200 ppm	5	5.4±	0.3	3.1±	0.2	1.3±	0.1	0.40±	0.13	264±	24*	108±	15	53±	10
400 ppm	5	5.6±	0.6	3.4±	0.4**	1.6±	0.1	0,49±	0.12	93±	38**	101±	14	251±	72**
mqq 008	1	5.4±	0.0 ?	3.3±	0.0 ?	1.6±	0.0 ?	0.54±	0.00 ?	144±	0 ?	99±	0 ?	522±	0 ?

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL074)

ANIMAL : MOUSE BDF1

REPORT TYPE : A1

SEX : MALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 2

Group Name	NO. of Animals	GPT IU/e		LDH IU/e		CPK I U/	?	UREA N mg∕d£	TROGEN	SODIUM mEq/l		POTASS:		CHLORIDE mEq∕ℓ	
Control	5	17±	5	212±	96	92±	49	23,4±	6.6	149±	2	5.5±	0.6	114±	7
50 ppm	5	16±	3	200±	42	66±	19	27,2±	4.2	149±	0	5.1±	0.4	117±	3
100 ppm	5	18±	3	223±	21	103±	25	29.4±	6.8	149±	2	5,3±	0.5	118±	3
200 ppm	5	20±	3	233±	87	92±	57	32.4±	7.9	153±	3	5.0±	0.8	119±	3
400 ppm	5	60±	13**	568±	99**	518±	174*	67.5±	4.9**	178±	7**	4.4±	0.4	133±	5**
800 ppm	1	150±	0 ?	1722±	0 ?	3306±	0 ?	95.4±	0.0 ?	189±	0 ?	3.0±	0.0 ?	150±	0 ?

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL074)

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

SEX : MALE

(IICL074)

PAGE: 3

oup Name	NO. of Animals	mg∕qr CVTCINW		INORGAN mg∕d2	C PHOSPHORUS		 	
Control	5	9.6±	0.4	8.8±	1.1			
50 ppm	5	9.3±	0.3	8.7±	1.8			
100 ppm	5	9.3±	0.2	8.8±	1.4			
200 ppm	5	9.7±	0.4	7.4±	1.2			
400 ppm	5	9.4±	0.3	9.1±	1.8			
mqq 008	1	10.8±	0.0 ?	9.0±	0.0 ?			
Significant	difference;	*: P ≤ 0	.05 **	*: P ≤ 0.0		Test of Dunnett		

BIOCHEMISTRY (TWO-WEEK STUDY : SUMMARY)

MOUSE: FEMALE

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : FEMALE BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 4

TOUP Name	NO. of Animals	TOTAL F g/dl		g∕dl ALBUMIN		A/G RAT	10	T-BILI mg/dl		GLUCOSE mg∕dl		T-CHOLES	STEROL	GOT IU/	?
Control	5	4.9±	0.2	3.0±	0.1	1.6±	0.1	0.29±	0.10	263±	32	82±	7	42±	7
50 ppm	5	4.7±	0.2	2.9±	0.1	1.6±	0.1	0.25±	0.03	270±	18	84±	6	44±	7
100 ppm	5	4.9±	0.4	2.9±	0.2	1.5±	0.0*	0.33±	0.09	266士	22	89±	2	46±	7
200 ppm	5	5.6±	0.2**	3.4±	0.2**	1.5±	0.1	0.42±	0.07	147±	57**	105±	6**	95±	33
400 ppm	5	5.7±	0.4**	3.5±	0.3**	1.6±	0.1	0.53±	0.20*	45±	32**	90±	17	278±	105**
mqq 008	0	-				-		-		-		-		-	

(IICL074)

ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 5

42 68± 18 41 68± 23 19 66± 17	22.8± 4.7 28.2± 4.0 30.8± 4.0	148± 1 149± 1 149± 1	$4.7\pm$ 0.7 $5.0\pm$ 0.2 $4.8\pm$ 0.5	122± 3 119± 4 119± 3
	30.8± 4.0	149± 1	4.8± 0.5	119± 3
110 108± 38	48.0± 8.0*	160± 3*	4.1± 0.6	125± 5
633** 1482± 1327*:	* 93.4± 25.1**	184士 11**	4.1± 0.1	146土 7**
-	-	-	-	-
	_			

(IICL074)

ANIMAL : MOUSE BDF1

REPORT TYPE : A1
SEX : FEMALE

BIOCHEMISTRY (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 6

iroup Name	NO. of Animals	CALCIUM mg∕dl		INORGAN mg∕dl	IC PHOSPHORUS					 	
Control	5	9.2±	0.2	7.8±	0.7						
50 ppm	5	9.1±	0.2	7.8±	0.9						
100 ppm	5	9.3±	0.2	7.3±	2.0						
200 ppm	5	9.5±	0.5	8.9±	1.6						
400 ppm	5	10.0±	0.8	11.3±	3.4						
mqq 008	0	-		-							
Significan	t difference ;	* : P ≤ (),05	**: P ≤ 0.0	1	 Test	of Dunnett	 		 	
(HCL074)						 		 	 	 	BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

RAT: MALE: DEAD AND MORIBUND ANIMALS

ANIMAL : RAT F344

REPORT TYPE : A1 SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

PAGE: 1

0rgan	Findings	Group Name NO. of Animals	Control 0 (%)	50 ppm 0 (%)	100 ppm 0 (%)	200 ppm 0 (%)
subcutis	dry		- (-)	- (-)	- (-)	- (-)
thymus	atrophic		- (-)	- (-)	- (-)	- (-)
gl stomach	hemorrhage		- (-)	- (-)	- ()	- (-)
adrenal	red		- (-)	- (-)	- (-)	- (-)
(HPT080)						BA

(HPT080)

ANIMAL : RAT F344 GROSS FINDINGS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1 SEX : MALE

400 ppm 1 (%) 800 ppm 10 (%) Group Name Organ___ Findings_ NO. of Animals subcutis 1 (100) 8 (80) dгу thymus 1 (100) 10 (100) atrophic hemorrhage 0 (0) 7 (70) gl stomach 0 (0) 3 (30) adrenal red (IIPT080) BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE: DEAD AND MORIBUND ANIMALS

ANIMAL : RAT F344

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1

; FEMALE

gan	Findings	Group Name NO. of Animals	Control 0 (%)	50 ppm 0 (%)	100 ppm 0 (%)	200 ppm 0 (%)
boutis	dry		- (-)	- (-)	- (-)	- (-)
mus	atrophic		- (-)	- (-)	- (-)	- (-)
	red zone		- (-)	- (-)	- (-)	- (-)
stomach	hemorrhage		- (-)	- (-)	- (-)	- (-)
enal	red		- (-)	- (-)	- (-)	- (-)

ANIMAL : RAT F344

REPORT TYPE : A1 SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

Group Name 400 ppm 800 ppm Findings_ 6 (%) 10 (%) Organ____ NO. of Animals dгу 5 (83) 9 (90) subcutis thymus 6 (100) 10 (100) atrophic 1 (17) 3 (30) red zone gl stomach hemorrhage 2 (33) 6 (60) adrena l red 2 (33) 7 (70)

(HPT080)

BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

RAT: MALE: SACRIFICED ANIMALS

ANIMAL : RAT F344

GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1

SEX : MALE

SHORT TODO INTIMADO (DIT)

PAGE: 1

Organ	Findings	Group Name NO. of Animals	Control 10 (%)	50 ppm 10 (%)	100 ppm 10 (%)	200 ppm 10 (%)
thymus	atrophic		0 (0)	0 (0)	0 (0)	0 (0)
l stomach	hemorrhage		0 (0)	0 (0)	0 (0)	0 (0)

.

ANIMAL : RAT F344

REPORT TYPE : A1 SEX : MALE GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

PAGE: 2

Organ	Findings	Group Name NO. of Animals	400 ppm 9 (%)	maq 008 (%)	
thymus	atrophic		9 (100)	- (-)	
gl stomach	hemorrhage		1 (11)	- (-)	
(HPT080)					RAIS 2

(HPT080)

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE: SACRIFICED ANIMALS

ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

Organ	Findings	Group Name NO, of Animals	Control 10 (%)	50 ppm 10 (%)	100 ppm 10 (%)	200 ppm 10 (%)
nymus	atrophic		0 (0)	0 (0)	0 (0)	6 (60)
	red zone		0 (0)	0 (0)	2 (20)	0 (0)
ver	herniation		1 (10)	0 (0)	0 (0)	0 (0)

STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1

GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

SEX : FEMALE

Organ	Findings	Group Name NO. of Animals	400 ppm 4 (%)	800 ppm 0 (%)	
thymus	atrophic		4 (100)	- (-)	
	red zone		0 (0)	- (-)	
iver	herniation		0 (0)	- (-)	

BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

MOUSE: MALE: DEAD AND MORIBUNDANIMALS

ANIMAL : MOUSE BDF1

GROSS FINDINGS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1 SEX : MALE

Organ	Findings	Group Name NO. of Animals	Cantral 0 (%)	50 ppm 0 (%)	100 ppm 0 (%)	200 ppm 0 (%)
ubcutis	dry		- (-)	- (-)	- (-)	- (-)
nymus	atrophic		- (-)	- (-)	- ()	- (-)
stomach	hemorrhage		- (-)	- (-)	- (-)	- (-)

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE

(IIPT080)

GROSS FINDINGS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

800 ppm Group Name 400 ppm 0 (%) Organ____ Findings_ NO. of Animals 9 (%) - (-) subcutis dry 5 (56) - (-) thymus atrophic 9 (100) - (-) gl stomach hemorrhage 2 (22)

BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

MOUSE: FEMALE: DEAD AND MORIBUNDANIMALS

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

Group Name 50 ppm 100 ppm 200 ppm Control Findings_ 0 (%) 0 (%) 0 (%) 0 (%) NO. of Animals Organ__ - (-) - (-) - (-) - (-) subcutis dгу - (-) - (-) - (-) - (-) thymus atrophic (HPT080) BAIS 2

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : FEMALE GROSS FINDINGS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

0rgan	Findings	Group Name NO. of Animals	400 ppm 1 (%)	800 ppm 10 (%)	
subcutis	dry		0 (0)	3 (30)	
thymus	atrophic		1 (100)	10 (100)	
(HPT080)			 		BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

MOUSE: MALE: SACRIFICED ANIMALS

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE

GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

)rgan	Findings	Group Name NO. of Animals	Control 10 (%)	50 ppm 10 (%)	100 ppm 10 (%)	200 ppm 10 (%)
<i>mus</i>	atrophic		0 (0)	0 (0)	0 (0)	0 (0)
een	black zone		0 (0)	0 (0)	0 (0)	1 (10)
stomach	hemorrhage		0 (0)	0 (0)	0 (0)	0 (0)

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

Organ	Findings	Group Name NO. of Animals	400 ppm 10 (%)	800 ppm 1 (%)	
thymus	atrophic		10 (100)	1 (100)	
spleen	black zone		0 (0)	0 (0)	
gl stomach	hemorrhage		1 (10)	0 (0)	
(HPT080)					BAIS 2

GROSS FINDINGS (TWO-WEEK STUDY: SUMMARY)

MOUSE: FEMALE: SACRIFICED ANIMALS

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : FEMALE GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

0rgan	Findings	Group Name NO. of Animals	Contral 10 (%)	50 ppm 10 (%)	100 ppm 10 (%)	200 ppm 10 (%)
NYMUS	atrophic		0 (0)	0 (0)	0 (0)	0 (0)
leen	black zone		1 (10)	0 (0)	0 (0)	0 (0)

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

GROSS FINDINGS (SUMMARY) SACRIFICED ANIMALS (2W)

SEX : FEMALE

Organ	Findings_	Group Name NO. of Animals	400 ppm 9 (%)	800 ppm 0 (%)	
thymus	atrophic		9 (100)	- (-)	
spleen	black zone		0 (0)	- (-)	
(IIPT080)					BAIS 2

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), ABSOLUTE

RAT: MALE

ANIMAL : RAT F344
REPORT TYPE : A1

SEX : MALE UNIT: g

ORGAN WEIGHT: ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 1

iroup Name	NO, of Animals	Body	Weight	ТНҮМ	JS	ADRE	ADRENALS		TESTES		HEART		LUNGS	
Control	5	177±	16	0.324±	0.054	0.043±	0.005	2.054±	0.233	0.649±	0.049	0.779±	0.022	
50 ppm	5	161±	12	0.328±	0.027	0.041±	0.009	1.644±	0.430	0.561±	0.057*	0.714±	0.030*	
100 ppm	5	164±	7	0.304±	0.028	0.046±	0.005	2.000±	0.253	0.587±	0.039	0.720±	0.027*	
200 ppm	5	137±	12**	0.181±	0.026**	0.048±	0.008	1.956±	0.273	0.486±	0.030**	0.646±	0.050**	
400 ppm	5	67±	6**	0.020±	0.009**	0.033±	0.004	0,610±	0.206**	0.289±	0.029**	0.521±	0.039**	
mag 008	0	-		-		-		-		-		-		
Significar	nt difference;	*: P ≦ 0.	05 **	: P ≤ 0.01			Tes	t of Dunnett				.,,		
(IICL040)														

(IICL040)

STUDY NO.: 0261 ANIMAL : RAT F344 REPORT TYPE : A1

ORGAN WEIGHT: ABSOLUTE (SUMMARY) SURVIVAL ANIMALS (2)

SEX : MALE UNIT: g

PAGE: 2

Graup Name	NO. of Animals	KID	KIDNEYS		SPLEEN		LIVER				
Contral	5	1.308±	0.086	0.411±	0.041	6.693±	0.678	1.723±	0.026		
Mqq 07	5	1.284±	0.103	0.369±	0.026	5.961±	0.707	1.699±	0,019		
100 ppm	5	1.399±	0.036	0.377±	0.031	6.407±	0.309	1.716±	0.048		
200 ppm	5	1.324±	0.087	0.319±	0.019**	5.502±	0.540**	1.675±	0.037		
400 ppm	5	0.899±	0.062**	0.087±	0.030**	2.428±	0.410**	1.573±	0.021**		
Mqq 008	0	-		•••		-		-			
Significar	nt difference;	*: P ≤ 0.	05 ** :	P ≤ 0.01			Tes	st of Dunnet		 	
(IICL040)										 	BAIS 2

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), ABSOLUTE

RAT: FEMALE

STUDY NO. : 0261 ANIMAL : RAT F344

REPORT TYPE : A1 SEX : FEMALE

UNIT: g

ORGAN WEIGHT: ABSOLUTE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 3

Control 50 ppm	5 5	126±	5	0.295±							HEART		LUNGS	
50 ppm				V-2001	0.023	0.051±	0.009	0.067±	0.008	0.461±	0.019	0.605±	0.024	
	J	116±	2	0.271±	0.025	0.040±	0.007*	0.054±	0.014	0.454±	0.028	0.567±	0.039	
100 ppm	5	107±	5**	0.239±	0.017**	0.042±	0.006	0.052±	0.006	0.394±	0.025**	0.573±	0.041	
200 ppm	5	75±	9**	0.051±	0.029**	0.032±	0.003**	0.044±	0.005**	0.312±	0.029**	0.490±	0.034**	
400 ppm	4	57±	5**	0.016±	0.007**	0.029±	0.004**	0.035±	0.010**	0.257±	0.036**	0.465±	0.048**	
800 ppm	0	-		_		-		-				-		
Significant	difference;	*: P ≤ 0.0	05 **	: P ≤ 0.01	 		Test	of Dunnett						

(IICL040)

REPORT TYPE : A1 SEX: FEMALE

UNIT: g

ORGAN WEIGHT: ABSOLUTE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 4

roup Name	NO. of Animals	KIDN	IEYS	SPL	EEN	LIV	ER	BRA		
Control	5	0.968±	0.052	0.296±	0.015	4.261±	0.367	1.580±	0.042	
50 ppm	5	1.073±	0.032*	0,280±	0.023	4.292±	0.190	1.595±	0.028	
100 ppm	5	1.072±	0.053	0.256±	0.011*	4.082±	0.397	1.584±	0.020	
200 ppm	5	0.893±	0.090	0.163±	0.028**	3.384±	0.396**	1.536±	0.021	
400 ppm	4	0.802±	0.072**	0.098±	0.030**	2.674±	0.501**	1.450±	0.060**	
mag 008	0	-		-		-		-		
Significan	t difference;	*: P ≤ 0.0)5 ** :	P ≤ 0.01			Tes	st of Dunnet		
ICL040)										

APPENDIX A 9-3

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), ABSOLUTE

MOUSE: MALE

STUDY NO.: 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE

SEX : MAL UNIT: g ORGAN WEIGHT: ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 1

roup Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	5	24.9± 0.9	0.063± 0.008	0.014± 0.000	0.187± 0.021	0.134± 0.012	0.144± 0.012
50 ppm	5	23.1± 0.7*	0.053± 0.007	0.016± 0.003	0.182± 0.013	0.127± 0.009	0.141± 0.013
100 ppm	5	23.4± 1.1	0.050± 0.010	0.016± 0.002	0.188± 0.024	0.126± 0.011	0.152± 0.016
200 ppm	5	19.4± 1.4**	0.032± 0.013**	0.013± 0.002	0.175± 0.013	0.113± 0.011*	0.134± 0.010
400 ppm	5	13.3± 0.6**	0.008± 0,005**	0.009± 0.001*	0.159± 0.024	0.081± 0.004**	0.125± 0.020
800 ppm	1	12.8± 0.0 ?	0.003± 0.000 ?	0.006± 0.000 ?	0.156± 0.000 ?	0.083± 0.000 ?	0.117± 0.000 ?

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL040)

STUDY NO. : 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1
SEX : MALE

UNIT: g

ORGAN WEIGHT: ABSOLUTE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 2

	NO. of Animals	KID	NEYS	SPL	EEN	LIV	ER	BRA	IN .
Control	5	0.368±	0.022	0.050±	0.004	1.252±	0.182	0.451±	0.019
50 ppm	5	0.377±	0.008	0.044±	0.002	1.164±	0.056	0.443±	0.015
100 ppm	5	0.385±	0.016	0.047±	0.008	1.185±	0.113	0,436±	0.014
200 ppm	5	0.354±	0.029	0.034±	0.010**	0.967±	0.081	0.443±	0.026
400 ppm	5	0.255±	0.021**	0.014±	0.007**	0.649±	0.034**	0.401±	0.012**
Mqq 008	1	0.224±	0.000 ?	0.007±	0.000 ?	0,651±	0.000 ?	0.381±	0.000 ?

(IICL040)

APPENDIX A 9-4

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), ABSOLUTE

MOUSE: FEMALE

STUDY NO.: 0262

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

REPORT TYPE: SEX: FEMALE UNIT: g ORGAN WEIGHT: ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 3

roup Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	5	20.0± 0.9	0.077± 0.008	0.012± 0.002	0.024± 0.009	0.118± 0.013	0.141± 0.013
50 ppm	5	19.1± 0.5	0.076± 0.006	0.012± 0.003	0.023± 0.005	0.105± 0.010	0.140± 0.019
100 ppm	5	18.3± 0.8**	0.066± 0.007	0.012± 0.002	0.023± 0.006	0.101± 0.011*	0.128± 0.019
200 ppm	5	13.9± 1.0**	0.024± 0.014**	0.009± 0.002*	0.009± 0.003**	0.087± 0.006**	0.127± 0.013
400 ppm	5	10.7± 0.5**	0.007± 0.004**	0.007± 0.001**	0.007± 0.004**	0.066± 0.006**	0.108± 0.010*
Mqq 008	0	-	-	-	-	-	-
Significar	nt difference;	*: P ≤ 0.05 **	: P ≤ 0.01	Test	of Dunnett		
IICL040)							

(IICL040)

STUDY NO. : 0262

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

REPORT TYPE: SEX: FEMALE UNIT: g ORGAN WEIGHT: ABSOLUTE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 4

roup Name	NO. of Animals	KID	NEYS	SPL	EEN	LIV	ER	BRA		
Contral	5	0.253±	0.012	0.054±	0.006	0.934±	0.119	0.440±	0.006	
50 ppm	5	0.280±	0.011**	0.049±	0.004	0.895±	0.038	0.443±	0,011	
100 ppm	5	0.273±	0.006	0.052±	0.008	0.898±	0.057	0.426±	0.015	
200 ppm	5	0.246±	0.018	0.021±	0.006**	0.640±	0.060**	0.426±	0.014	
400 ppm	5	0.200±	0.013**	0.009±	0.003**	0.474±	0.055**	0.390±	0.019**	
800 ppm	0	-		-		_		-		
Significan	t difference ;	*: P ≤ 0.	05 **	: P ≦ 0.01			Tes	st of Dunnet		
HCI 040)										

(IICL040)

APPENDIX A 10-1

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), RELATIVE

RAT: MALE

REPORT TYPE: A1
SEX: MALE
UNIT: %

ORGAN WEIGHT: RELATIVE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 1

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	5	177± 16	0.183± 0.024	0.025± 0.002	1.166± 0.138	0.369± 0.036	0.442± 0.031
50 ppm	5	161± 12	0.204± 0.020	0.026± 0.006	1.011± 0.206	0.348± 0.019	0.444± 0.018
100 ppm	5	164± 7	0.185± 0.014	0.028± 0.003	1.217± 0.142	0.357± 0.021	0.438± 0.015
200 ppm	5	137± 12**	0.131± 0.008**	0.035± 0.005*	1.423± 0.098	0.356± 0.012	0.474± 0.039
400 ppm	5	67± G**	0.030± 0.013**	0.049± 0.007**	0.898± 0.253	0.431± 0.014**	0.781± 0.074*
800 ppm	0	-		-	-	-	-
Significan	nt difference;	*: P ≤ 0.05 **	: P ≤ 0.01	Tes	t of Dunnett		

(IICL042)

STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1 ORGAN WEIGHT: RELATIVE (SUMMARY)
SURVIVAL ANIMALS (2)

SEX: MALE UNIT: %

PAGE: 2

roup Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN	
Control	5	0.742± 0.028	0.232± 0.006	3.783± 0.096	0.981± 0.092	
mqq 07	5	0.797± 0.014	0.229± 0.011	3.693± 0.258	1.058± 0.069	
100 ppm	5	0.851± 0.024**	0.229± 0.013	3.897± 0.083	1.045± 0.048	
200 ppm	5	0.968± 0.032**	0.234± 0.009	4.015± 0.097	1.230± 0.110**	
400 ppm	5	1.345± 0.064**	0.128± 0.037**	3.615± 0.457	2.360± 0.187**	
800 ppm	0	-	_	-	-	

(HCL042)

APPENDIX A 10-2

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), RELATIVE

RAT: FEMALE

REPORT TYPE: A1
SEX: FEMALE
UNIT: %

ORGAN WEIGHT: RELATIVE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 3

Group Name	NO. of Animals	Bady W	Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS	
Control	5	126±	5	0.235± 0.018	0.040± 0.005	0.053± 0.006	0.367± 0.012	0.483± 0.030	
50 ppm	5	116±	2	0.233± 0.020	0.035± 0.006	0.046± 0.013	0.390± 0.019	0.487± 0.036	
100 ppm	5	107±	5**	0.224± 0.017	0.039± 0.005	0.049± 0.004	0.370± 0.017	0.539± 0.056	
200 ppm	5	75±	9**	0.065± 0.028**	0.042± 0.004	0.058± 0.004	0.415± 0.025**	0.654± 0.052**	
400 ppm	4	57±	5**	0.028± 0.013**	0.050± 0.003*	0.061± 0.013	0.451± 0.033**	0.818± 0.082**	
800 ppm	0	-		-	-		-	-	
Significan	nt difference;	*: P ≤ 0.0)5 **	: P ≤ 0.01	Tes	t of Dunnett			

(IICL042)

STUDY NO.: 0261

ANIMAL : RAT F344
REPORT TYPE : A1

SEX : FEMALE UNIT: %

ORGAN WEIGHT: RELATIVE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 4

cup Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN	
Control	5	0.771± 0.033	0.236± 0.003	3.389± 0.195	1.260± 0.066	
50 ppm	5	0.922± 0.037**	0.241± 0.020	3.689± 0.199	1.370± 0.028	
100 ppm	5	1.006± 0.023**	0.240± 0.006	3.823± 0.202	1.489± 0.081	
200 ppm	5	1.188± 0.045**	0.216± 0.018	4.490± 0.144**	2.058± 0.214**	
400 ppm	4	1.408± 0.062**	0.170± 0.041*	4.667± 0.508**	2.558± 0.228**	
800 ppm	0	-	_	=	- -	

(IICL042)

APPENDIX A 10-3

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), RELATIVE

MOUSE: MALE

STUDY NO. : 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1 SEX : MALE UNIT: % ORGAN WEIGHT: RELATIVE (SUMMARY) SURVIVAL ANIMALS (2)

PAGE: 1

oup Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	5	24.9± 0.9	0.254± 0.035	0.057± 0.002	0.752± 0.076	0.538± 0.047	0.580± 0.065
50 ppm	5	23.1± 0.7*	0.231± 0.029	0.068± 0.015	0.786± 0.051	0.548± 0.031	0.609± 0.046
100 ppm	5	23.4± 1.1	0.212± 0.033	0.068± 0.007	0.807± 0.109	0.540± 0.041	0.649± 0.064
200 ppm	5	19.4生 1.4**	0.161± 0.056**	0.065± 0.010	0.908± 0.106	0.584± 0.066	0.688± 0.029
400 ppm	5	13.3± 0.6**	0.059± 0.041**	0.065± 0.011	1.188± 0.169**	0.605± 0.030	0.939± 0.156**
800 ppm	1	12.8± 0.0 ?	0.023± 0.000 ?	0.047± 0.000 ?	1.219± 0.000 ?	0.648± 0.000 ?	0.914± 0.000 ?

^{?:} Significant test is not applied, because No. of data in this group is less than 3.

(IICL042)

STUDY NO.: 0262

ANIMAL : MOUSE BDF1

REPORT TYPE : A1

SEX : MALE UNIT: %

ORGAN WEIGHT: RELATIVE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 2

oup Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN	
Control	5	1.476± 0.063	0.200± 0.010	5.012± 0.557	1.815± 0.087	
50 ppm	5	1.633± 0.034*	0.189± 0.005	5.034± 0.103	1.917± 0.095	
100 ppm	5	1.651± 0.095*	0.202± 0.029	5.069± 0.341	1.870± 0.101	
200 ppm	5	1.822± 0.098**	0.177± 0.044	4,978± 0.187	2.285± 0.076**	
400 ppm	5	1.910± 0.114**	0.106± 0.052*	4.866± 0.157	3.010± 0.061**	
mqq 008	1	1.750± 0.000 ?	0.055± 0.000 ?	5.086± 0.000 ?	2,977± 0,000 ?	

(HCL042)

APPENDIX A 10-4

ORGAN WEIGHT (TWO-WEEK STUDY : SUMMARY), RELATIVE

MOUSE: FEMALE

STUDY NO.: 0262

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

REPORT TYPE : SEX : FEMALE UNIT: % ORGAN WEIGHT: RELATIVE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 3

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS	
Control	5	20.0± 0.9	0.387± 0.043	0.062± 0.006	0,119± 0,047	0.588± 0.048	0.708± 0.088	
50 ppm	5	19.1± 0.5	0.398± 0.028	0.063± 0.014	0.120± 0.025	0.552± 0.065	0.734± 0.094	
100 ppm	5	18.3± 0.8**	0.358± 0.041	0.068± 0.011	0.127± 0.028	0.551± 0.052	0.699± 0.098	
200 ppm	5	13.9± 1.0**	0.167± 0.088**	0.063± 0.010	0.065± 0.021*	0,623± 0,025	0.911± 0.085**	
400 ppm	5	10.7± 0.5**	0.061± 0.034**	0.065± 0.010	0.069± 0.032	0.618± 0.074	1.009± 0.103**	
800 ppm	0	-	-	-	. -	-	_	
Significan	nt difference;	*: P ≤ 0.05 **	: P ≤ 0.01	Tes	st of Dunnett			

(HCL042)

STUDY NO. : 0262

ANIMAL : MOUSE BDF1
REPORT TYPE : A1

SEX : FEMALE UNIT: %

ORGAN WEIGHT: RELATIVE (SUMMARY)
SURVIVAL ANIMALS (2)

PAGE: 4

TOUP Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN	
Control	5	1.265± 0.074	0.268± 0.028	4.655± 0.392	2.203± 0.127	
50 ppm	5	1.468± 0.054*	0.255± 0.022	4.688± 0.146	2.322± 0.083	
100 ppm	5	1.492± 0.071*	0.285± 0.036	4.903± 0.268	2.327± 0.143	
200 ppm	5	1.774± 0.170**	0.151± 0.037**	4.588± 0.136	3.065± 0.172**	
400 ppm	5	1.870± 0.156**	0.082± 0.024**	4.400± 0.335	3.630± 0.161**	
800 ppm	0	_	_	-	-	

(IICL042)

APPENDIX A 11-1

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

RAT: MALE: DEAD AND MORIBUND ANIMALS

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY) DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1 SEX

: MALE

Organ	N	roup Name	50 ppm 0 1 2 3 4 (%) (%) (%) (%)	100 ppm 0 1 2 3 4 (%) (%) (%) (%)	200 ppm 0 1 2 3 4 (%) (%) (%) (%)
[Nematopoieti	c system]				
bone marrow	congestion	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)
	decreased hematopoiesis	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-)
thymus	atrophy	< 0> (-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)
spleen	atrophy	< 0> (-) (-) (-) (-)		(-) (-) (-) (-)	< 0> (-) (-) (-) (-)
(Endocrine s	vstem]				
adrenal	congestion	(-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)
[Reproductive	ə system]				
testis	atrophy	(-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)
Grade <a>> b (c)	1: Slight 2: Moderate 3: a: Number of animals examined at the sib: Number of animals with lesion c:b/a*100	: Marked 4 : Seuere te			

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1

(HPT150)

SEX : MALE

Organ	Findings	Group Name 400 ppm No. of Animals on Study 0 Grade 1 2 3 4 (%) (%) (%) (%)	800 ppm 2 2 1 2 3 4 (%) (%) (%)	
[Hematopoieti	c system]			
oone marrow	congestion	< 0> (-) (-) (-) (-)	<pre></pre>	
	decreased hematopoiesis	(-) (-) (-) (-)	0 2 0 0 (0) (100) (0) (0)	
thymus	atrophy	(-) (-) (-) (-)	<pre></pre>	
spleen	atrophy	< 0> (-) (-) (-) (-)	<pre></pre>	
(Endocrine sy	vstem]			
adrena l	congestion	(-) (-) (-) (-)	<pre></pre>	
[Reproductive	e system]			
testis	atrophy	< 0> (-) (-) (-) (-)	2 0 0 0 (100) (0) (0) (0)	
Grade <a> b (c)	1: Slight 2: Moderate a: Number of animals examined at b: Number of animals with lesion c: b/a * 100	3: Marked 4: Severe the site		

BAIS2

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (O- 2W)

REPORT TYPE : A1
SEX : MALE

0rgan		up Name of Animals on Study de(%)	Control 0 2 3 (%) (%)	<u>4</u> (%)	<u>1</u> (%)	50 ppm 0 2 3 (%) (%)	<u>4</u> (%)	1(%)	100 pp 0 2 (%)	3	<u>4</u> (%)	_1		3 4 (%)
[Reproductiv	pe system]													
epididymis			< 0>			< 0>			< 0	>			< 0>	
	debris of spermatic elements	(-)	(-) (-)	(-)	(-) (-) (-)	(-)	(-)	(-) (-) (- -)	(-) (-) (-) (-)
Grade (a > b (c)	1: Slight 2: Moderate 3: Ma a: Number of animals examined at the site b: Number of animals with lesion c: b/a * 100	arked 4 : Severe	9											
(IIPT150)						-								BA

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (O- 2W)

REPORT TYPE : A1 SEX : MALE

Organ	Findings	Group Name 400 pp No. of Animals on Study 0 Grade 1 2 (%) (%)	3 4 (%) (%)	800 ppm 2 1 2 3 4 (%) (%) (%) (%)	
[Reproductiv	e system]				
epididymis	debris of spermatic elements	< 0: (-) (<pre></pre>	
Grade <a>> b (c)	1: Slight 2: Moderate a: Number of animals examined at the b: Number of animals with lesion c: b/a*100	3 : Marked 4 : Severe site			
(IIPT150)					BAI

APPENDIX A 11-2

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE: DEAD AND MORIBUND ANIMALS

STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1

: FEMALE

SEX

(IIPT150)

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

PAGE: 5

100 ppm 200 ppm Group Name Control 50 ppm No. of Animals on Study 0 0 0 0 2 3 (%) (%) Findings [Respiratory system] lung hemorrhage (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) [Hematopoietic system] bone marrow congestion (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) decreased hematopoiesis (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) thymus atrophy (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) congestion (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) spleen atrophy (-) (-) (-) (-) Grade 1 : Slight 2 : Moderate 4 : Severe (a) a: Number of animals examined at the site b b: Number of animals with lesion c:b/a*100(c)

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

PAGE: 6

BAIS2

REPORT TYPE : A1

SEX : FEMALE

DEAD AND MORIBUND ANIMALS (0- 2W)

Group Name 400 ppm 800 ppm No. of Animals on Study 1 2 Grade 2 3 Findings [Respiratory system] lung < 1> < 2> 0 1 0 0 0 0 0 0 hemorrhage (0)(0)(0)(0) (0)(50)(0)(0) [Hematopoietic system] bone marrow < 1> < 2> 0 0 2 0 congestion 0 1 0 0 (0) (100) (0) (0) (0)(0)(100)(0) decreased hematopoiesis 0 1 0 0 0 2 0 0 (0) (100) (0) (0) (0) (100) (0) (0) thymus < 1> < 2> 0 0 1 0 0 0 2 0 atrophy (0)(0)(100)(0) (0)(0)(100)(0) congestion 0 0 0 0 0 1 0 0 (0) (0) (0) (0) (0)(50)(0)(0) spleen < 1> < 2> 1 0 0 0 0 2 0 0 atrophy (100) (0) (0) (0) (0)(100)(0)(0) Grade 1: Slight 3 : Marked 2 : Moderate 4 : Severe (a) a: Number of animals examined at the site b b: Number of animals with lesion (c) c : b / a * 100(IIPT150)

STUDY NO. : 0261

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

ANIMAL : RAT F344 REPORT TYPE: A1

: FEMALE SEX

DEAD AND MORIBUND ANIMALS (0- 2W)

PAGE: 7

Group Name 100 ppm 200 ppm Control 50 ppm No. of Animals on Study 0 Findings_ [Endocrine system] < 0> < 0> adrenal < 0> congestion (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) (-) 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(IIPT150) BAIS2 STUDY NO. : 0261 ANIMAL : RAT F344 REPORT TYPE : A1

SEX : FEMALE

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

PAGE: 8

DEAD AND MORIBUND ANIMALS (0- 2W)

mag 008 Group Name 400 ppm No. of Animals on Study 1 Organ____ Findings__ [Endocrine system] adrenal < 2> < 1> 1 1 0 0 congestion 0 1 0 0 (0) (100) (0) (0) (50) (50) (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 b (c) c:b/a*100(HPT150) BAIS2

APPENDIX A 11-3

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

RAT: MALE: SACRIFICED ANIMALS

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)

SACRIFICED ANIMALS (2W)

REPORT TYPE : A1 SEX : MALE

0rgan	Group Name No. of An Grade Findings	Control imals on Study 2 1 2 3 4 (%) (%) (%) (%)	50 ppm 2 1 2 3 4 (%) (%) (%) (%)	100 ppm 2 1 2 3 4 (%) (%) (%) (%)	200 ppm 2 1 2 3 4 (%) (%) (%) (%)
[Respirator)	v system]				
lung	hemorrhage	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	0 0 0 0 (0) (0) (0) (0)	<pre></pre>
[llematopoiet	ic system]				
bone marrow	congestion	<pre></pre>	<pre></pre>	(2> 0 0 0 0 (0) (0) (0) (0)	<pre></pre>
	decreased hematopoiesis	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)
thymus	atrophy	<pre></pre>	<pre></pre>	<pre></pre>	<pre></pre>
[Reproduction	pe system]				
testis	atrophy	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	0 0 0 0 (0) (0) (0) (0)	<pre></pre>
Grade <a> b	1: Slight 2: Moderate 3: Marked a: Number of animals examined at the site b: Number of animals with lesion c: b/a * 100	4 : Severe		-	

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

PAGE: 2

BAIS2

REPORT TYPE : A1

(IIPT150)

SEX : MALE

Group Name 400 ppm 800 ppm No. of Animals on Study 0 2 Findings [Respiratory system] lung < 2> hemorrhage 0 1 0 0 (0)(50)(0)(0) (-) (-) (-) [Nematopoietic system] bone marrow < 2> 0 2 0 0 congestion (-) (-) (-) (-) (0) (100) (0) (0) decreased hematopoiesis 0 2 0 0 (-) (-) (-) (-) (0) (100) (0) (0) thymus < 2> 0 0 2 0 atrophy (-) (-) (-) (0)(0)(100)(0) [Reproductive system] testis < 2> < 0> 2 0 0 0 atrophy (100) (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

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1 : MALE SEX

Organ	Findings	Group Name No. of Animals Grade	on Study _1	Control 2 2 3 (%) (%)	<u>4</u> (%)	<u>1</u> (%)		3 <u>4</u> %) (%)	1(%)	2 (%)	opm 2 3 (%)	<u>4</u> (%)	1 (%)	200		<u>4</u> (%)
[Reproductiv	e system]															
epididymis	debris of spermatic elements		(0) (< 2> 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)	< 2> 0) (0)	0 (0)
Grade <a> b (c)	1: Slight 2: Moderate a: Number of animals examined at th b: Number of animals with lesion c: b/a*100	3 : Marked e site	4: Severe													
(IIPT150)										· · · · · · · · · · · · · · · · · · ·						

STUDY NO. : 0261 ANIMAL

: RAT F344

REPORT TYPE : A1

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

SACRIFICED ANIMALS (2W)

SEX : MALE

PAGE: 4 Group Name 400 ppm 800 ppm No. of Animals on Study 0 Grade 3 Organ____ Findings [Reproductive system] epididymis < 2> < 0> debris of spermatic elements 1 0 0 0 (-) (-) (-) (-) (50) (0) (0) (0) Grade 1 : Slight 2 : Moderate 3 : Marked 4 : Severe a: Number of animals examined at the site (a) b b: Number of animals with lesion (c) c:b/a*100(IIPT150) BAIS2

APPENDIX A 11-4

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

RAT: FEMALE: SACRIFICED ANIMALS

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1

SEX : FEMALE

Organ	Findings	Group Name Control No. of Animals on Study 2 Grade 1 2 3 4 (%) (%) (%) (%)	50 ppm 2 1 2 3 4 (%) (%) (%) (%)	100 ppm 2 1 2 3 4 (%) (%) (%) (%)	200 ppm 2 1 2 3 4 (%) (%) (%) (%)
[Respiratory	system]				
lung	hemorrhage	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	<pre></pre>	<pre></pre>
[Hematopoiet	ic system]				
bone marrow	congestion	<pre></pre>	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	0 0 0 0 (0) (0) (0) (0)
	decreased hematopoiesis	0 0 0 0 0 (0) (0)	0 0 0 0 0 (0) (0)	0 0 0 0 0 0 (0) (0)	0 2 0 0 (0) (100) (0) (0)
thymus	atrophy	<pre></pre>	<pre></pre>	<pre></pre>	<pre></pre>
spleen	atrophy	0 0 0 0 (0) (0) (0) (0)	<pre></pre>	<pre></pre>	<pre></pre>
[Digestive s	system]				
liver	herniation	2> 1 0 0 0 (50) (0) (0) (0)	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	0 0 0 0 (0) (0) (0) (0)
Grade <a>> b (c)	1: Slight 2: Moderate a: Number of animals examined at the Number of animals with lesion c: b/a * 100	3: Marked 4: Severe me site			
(IIPT150)					ВЛ

STUDY NO. : 0261 ANIMAL : RAT F344 HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1 SEX

: FEMALE

PAGE: 6

Organ		Group Name 400 ppm No. of Animals on Study 2 Grade 1 2 3 4 (%) (%) (%) (%)	800 ppm 0 1 2 3 4 (%) (%) (%) (%)	
[Respiratory	system]			
lung	hemorrhage	0 1 0 0 (0) (50) (0) (0)	< 0> (-) (-) (-) (-)	
[Nematopoieti	ic system]			
bone marrow	congestion	2> 1 0 1 0 (50) (0) (50) (0)	(-) (-) (-) (-)	
	decreased hematapaiesis	0 2 0 0 (0) (100) (0) (0)	(-) (-) (-)	
thymus	atrophy	<pre></pre>	(-) (-) (-)	
spleen	atrophy	2> 1 0 0 0 (50) (0) (0) (0)	(-) (-) (-)	
[Digestive sy	ystem]			
liver	herniation	<pre></pre>	(-) (-) (-)	
Grade <a> b (c)	1: Slight 2: Moderate 3 a: Number of animals examined at the s b: Number of animals with lesion c: b/a * 100	: Marked 4 : Severe ite		

(IIPT150)

STUDY NO. : 0261 ANIMAL : RAT F344

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)

SACRIFICED ANIMALS (2W)

REPORT TYPE : A1
SEX : FEMALE

D	Tr. II	Group Name No. of Animals on Study Grade 1		3 4	50 ppm 2 1 2 3 4	100 ppm 2 1 2 3 4	200 ppm 2 1 2 3 4
)rgan	Findings	(%)	(%) (9	%) (%)	(%) (%) (%) (%)	(%) (%) (%)	(%) (%) (%) (%)
[Digestive :	system]						
liver	necrosis:middral	0 (0)	< 2> 0 (0) (0 0 0) (0)	<pre></pre>	<pre></pre>	<pre></pre>
(Endocrine :	system]						
pituitary	Rathke pouch	(0)		0 0 0) (0)	<pre></pre>	2> 1 0 0 0 (50) (0) (0) (0)	2> 1 0 0 0 (50) (0) (0) (0)
Grade (a) b	1: Slight 2: Moderate a: Number of animals examined at the b: Number of animals with lesion c:b/a*100	3 : Marked 4 : Severe site					

STUDY NO. : 0261

ANIMAL : RAT F344

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

SACRIFICED ANIMALS (2W)

REPORT TYPE : A1 SEX : FEMALE

Group Name 400 ppm 800 ppm No. of Animals on Study Grade Findings__ [Digestive system] liver < 2> necrosis:middral 1 0 0 0 (50) (0) (0) (0) (-) (-) (-) (-) [Endocrine system] pituitary 0 0 0 0 Rathke pouch (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) BAIS2

APPENDIX A 11-5

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

MOUSE: MALE: DEAD AND MORIBUNDANIMALS

STUDY NO. : 0262 ANIMAL : MOUSE BDF1 HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY) DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1

SEX : MALE

Organ	N	roup Name Control o. of Animals on Study 0 rade 1 2 3 4 (%) (%) (%) (%)	50 ppm 0 1 2 3 4 (%) (%) (%)	100 ppm 0 1 2 3 4 (%) (%) (%) (%)	200 ppm 0 1 2 3 4 (%) (%) (%) (%)
[Nematopoieti	c system]				
one marrow		< 0>	< 0>	< 0>	< 0>
	congestion	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)
	decreased hematopoiesis	(-) (-) (-) (-)	() () ()	(-) (-) (-) (-)	(-) (-) (-) (-)
:hymus		< 0>	< 0>	< 0>	< 0>
	atrophy	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)
pleen		< 0>	< 0>	< 0>	< 0>
	atrophy	(-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)
irade (a > b (c)	1: Slight 2: Moderate 3: a: Number of animals examined at the sit b: Number of animals with lesion c: b / a * 100	Marked 4: Seuere te			

STUDY NO. : 0262 ANIMAL : MOUSE BDF1 HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1 : MALE

(c)

(HPT150)

c : b / a * 100

Group Name 400 ppm mqq 008 No. of Animals on Study 0 3 (%) Findings_ [Hematopoietic system] bone marrow 0 3 0 0 congestion (-) (-) (-) (-) (0) (100) (0) (0) 0 3 0 0 decreased hematopoiesis (-) (-) (-) (-) (0) (100) (0) (0) < 0> < 2> thymus 0 0 2 0 atrophy (-) (-) (-) (-) (0)(0)(100)(0) < 0> < 3> spleen 0 3 0 0 atrophy (-) (-) (-) (-) (0)(100)(0)(0) 3 : Marked Grade 4 : Severe 1 : Slight 2 : Moderate <a>> a : Number of animals examined at the site b b: Number of animals with lesion

BAIS2

APPENDIX A 11-6

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

MOUSE: FEMALE: DEAD AND MORIBUNDANIMALS

STUDY NO. : 0262 ANIMAL : MOUSE BDF1

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0- 2W)

REPORT TYPE : A1 SEX

: FEMALE

0rgan	No	Dup Name Control . of Animals on Study 0 ade 1/2 3 4 (%) (%) (%) (%)	50 ppm 0 1 2 3 4 (%) (%) (%) (%)	100 ppm 0 1 2 3 4 (%) (%) (%) (%)	200 ppm 0 1 2 3 4 (%) (%) (%) (%)
[Nematopoietio	c system]				
bone marrow	congestion	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)	< 0> (-) (-) (-) (-)
	decreased hematopoiesis	 (-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-) (-)
thymus	atrophy	< 0> (-) (-) (-) (-)	·	(-) (-) (-) (-)	< 0> (-) (-) (-) (-)
pleen	atrophy	< 0> (-) (-) (-) (-)	(-) (-) (-) (-)	(-) (-) (-)	(-) (-) (-) (-)
irade (a> b (c)	1: Slight 2: Moderate 3: a: Number of animals examined at the site b: Number of animals with lesion c: b/a * 100	Marked 4: Severe			

STUDY NO. : 0262

: MOUSE BDF1

HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

PAGE: 4

DEAD AND MORIBUND ANIMALS (0- 2W)

ANIMAL REPORT TYPE : A1

SEX : FEMALE

Group Name 400 ppm mqq 008 No. of Animals on Study 2 0 Organ___ Findings [Hematopoietic system] bone marrow 1 1 0 0 (-) (-) (-) (-) congestion (50) (50) (0) (0) decreased hematopoiesis 1 1 0 0 (-) (-) (-) (-) (50) (50) (0) (0) thymus < 0> < 1> 0 0 1 0 atrophy (-) (-) (-) (-) (0) (100) (0) spleen < 0> < 2> 0 1 1 0 atrophy (-) (-) (-) (-) (0)(50)(50)(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(IIPT150) BAIS2

APPENDIX A 11-7

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

MOUSE: MALE: SACRIFICED ANIMALS

STUDY NO. : 0262 ANIMAL : MOUSE BDF1 HISTOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)

REPORT TYPE: A1

: MALE

SACRIFICED ANIMALS (2W)

SEX PAGE: 1 200 ppm Group Name Control 50 ppm 100 ppm 2 No. of Animals on Study 2 2 2 2 3 Grade 2 3 2 3 Findings_ (%) (%) (%) (%) (%) (%) (%) (%) (%) [Hematopoietic system] bone marrow < 2> < 2> < 2> < 2> congestion 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) decreased hematopoiesis 0 0 0 0 0 (0)(0)(0)(0) (0)(0)(0)(0) (0)(0)(0)(0) (0)(0)(0)(0) thymus 〈 2〉 〈 2〉 < 2> < 2> 2 0 0 0 atrophy 0 0 0 0 0 0 0 0 0 0 0 0 (0)(0)(0)(0) (0)(0)(0)(0) (0) (0) (0) (0) (100) (0) (0) (0) spleen < 2> < 2> < 2> < 2> 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 atrophy (0)(0)(0)(0) (0)(0)(0)(0) (0)(0)(0)(0) (0)(0)(0)(0) [Digestive system] liver 〈 2〉 < 2> < 2> necrosis:middral 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) granulation 0 0 0 0 0 0 0 0 0 0 0 1 0 0 (0)(0)(0)(0) (0)(0)(0)(0) (0)(50)(0)(0) (0)(0)(0)(0) Grade 3 : Marked 4 : Severe 1 : Slight 2 : Moderate (a) a: Number of animals examined at the site b b: Number of animals with lesion (c) c:b/a*100

(IIPT150)

BAIS2

STUDY NO. : 0262

HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
SACRIFICED ANIMALS (2W)

ANIMAL : MOUSE BDF1 REPORT TYPE : A1 SEX

: MALE

Organ	Group Nam No. of An Grade Findings	400 ppm imals on Study 2 1 2 3 4 (%) (%) (%) (%)	800 ppm 1 1 2 3 4 (%) (%) (%) (%)	
[Nematopoieti	ic system]			
bone marrow	congestion	2	(1) 0 1 0 0 (0) (100) (0) (0)	
	decreased hematopoiesis	0 2 0 0 (0) (100) (0) (0)	0 1 0 0 (0) (100) (0) (0) .	
thymus	atrophy	0 2 0 0 (0) (100) (0) (0)	0 1 0 0 (0) (100) (0) (0)	
spleen	atrophy	2 0 0 0 (100) (0) (0) (0)	0 1 0 0 (0) (100) (0) (0)	
[Digestive sy	ystem]			
liver	necrosis:middral	<pre></pre>	(0) (100) (0) (0)	
	granulation	0 0 0 0 0 (0) (0)	0 0 0 0 0 (0) (0)	
Grade <a>> b (c)	1: Slight 2: Moderate 3: Marked a: Number of animals examined at the site b: Number of animals with lesion c:b/a*100	4 : Severe		
(IIPT150)				ВЛІ

APPENDIX A 11-8

HISTOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS

(TWO-WEEK STUDY: SUMMARY)

MOUSE: FEMALE: SACRIFICED ANIMALS

STUDY NO. : 0262 ANIMAL : MOUSE BDF1 HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1

SEX

: FEMALE

Organ		p Name Control of Animals on Study 2 de 1 2 3 4 (%) (%) (%) (%)	50 ppm 2 1 2 3 4 (%) (%) (%) (%)	100 ppm 2 1 2 3 4 (%) (%) (%) (%)	200 ppm 2 1 2 3 4 (%) (%) (%) (%)
[Nematopoieti	c system]				
bone marrow	congestion	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	0 0 0 0 (0) (0) (0) (0)	<pre></pre>
	decreased hematopoiesis	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)
thymus	atrophy	<pre></pre>	<pre></pre>	<pre></pre>	2 0 0 0 (100) (0) (0) (0)
spleen	atrophy	<pre></pre>	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	<pre></pre>
[Digestive sy	vstem]				
liver	necrosis:focal	<pre></pre>	<pre></pre>	0 0 0 0 (0) (0) (0) (0)	2 0 0 0 (100) (0) (0) (0)
	granulation	1 0 0 0 0 (50) (50) (60)	1 0 0 0 0 (50) (50) (60)	1 0 0 0 (50) (0) (0) (0)	0 0 0 0 0 (0) (0)
Grade <a>> b (c)	1: Slight 2: Moderate 3: M a: Number of animals examined at the site b: Number of animals with lesion c: b/a * 100	arked 4: Severe			
(HPT150)					BAIS

STUDY NO. : 0262 ANIMAL : MOUSE BDF1 HISTOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY) SACRIFICED ANIMALS (2W)

REPORT TYPE : A1

SEX : FEMALE

: Al

Organ	No	roup Name 400 ppm b. of Animals on Study 2 rade 1 2 3 4 (%) (%) (%) (%)	800 ppm 0 1 2 3 4 (%) (%) (%) (%)	,	
[Nematopoieti	ic system]				
bone marrow	consestion	2	< 0> (-) (-) (-) (-)		
	decreased hematopoiesis	2 0 0 0 (100) (0) (0) (0)	(-) (-) (-)		
thymus	atrophy	0 2 0 0 (0) (100) (0) (0)	(-) (-) (-) (-)		
spleen	atrophy	<pre></pre>	< 0> (-) (-) (-) (-)		
[Digestive s	ystem]				
liver	necrosis:focal	2> 1 0 0 0 (50) (0) (0) (0)	< 0> (-) (-) (-) (-)		
	granulation	0 0 0 0 0 (0) (0)	(-) (-) (-) (-)		
Grade <a>> b (c)	1: Slight 2: Moderate 3: a: Number of animals examined at the sit b: Number of animals with lesion c: b / a * 100	Marked 4 : Severe e			
(IIPT150)					BAIS2

APPENDIX A 12-1 IDENTITY OF HYDRAZINE MONOHYDRATE (TWO-WEEK STUDIES)

IDENTITY OF HYDRAZINE MONOHYDRATE(TWO-WEEK STUDIES)

Test Substance Lot No. APF5206

1. Spectral data

(1) Mass Spectrometry

Instrument

: Hitachi M-80B

Ionization ·

: SIMS(Secondary Ionization Mass Spectrometry)

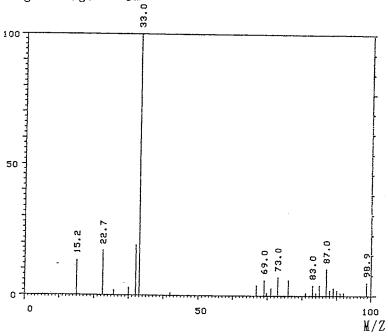
Matrix

: Glycerol

Primary Ion

: Xenone +

Accelerating Voltage : 8kv



Mass Spectrum of Test Substance

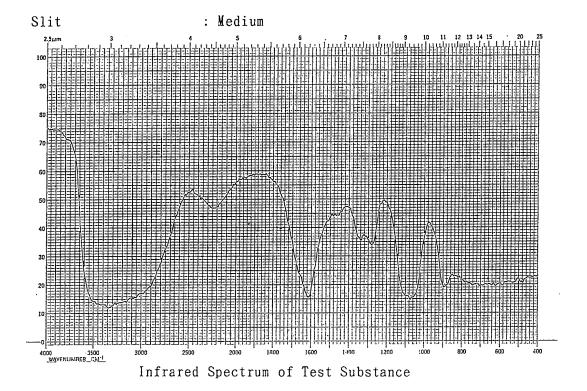
Result: The mass spectrum was consistent with calculated spectrum.

	Quasi Molecule Ion
Calculated Value	33. 0
(NH ₂ •NH ₂ +H ⁺) Determined Value	33.0

(2) Infrared Spectrometry

Instrument : Hitachi 270-30 Infrared Spectrometer

Cell : KBr Liquid Cell



Results: The infrared spectrum was consistent with literature spectrum.

<u>Determined Values</u>	<u>Literature Values*</u>
Wave Number(cm ⁻¹)	Wave Number(cm ⁻¹)
850~ 970	850~ 970
970~1220	970~1220
$1220 \sim 1400$	1220~1400
1500~1800	1500~1800
$2000 \sim 2500$	2000~2500
2500~3750	2500~3750
	(*Performed by WAKO PURE
	CHEMICAL INSUSTRIES, LTD.)

2. Conclusions: The test substance was identified as Hydrazine monohydrate, by the mass spectrum and infrared spectrum.

APPENDIX A 12-2 STABILITY OF HYDRAZINE MONOHYDRATE (TWO-WEEK STUDIES)

STABILITY OF HYDRAZINE MONOHYDRATE(TWO-WEEK STUDIES)

Test Substance Lot No. APF5206

1. Sample: This lot was used form 1994. 1.25 to 1994. 2.15. The test substance was stored in the dark at room temperature.

2. Infrared Spectrometry

Instrument

: Hitachi 270-30 Infrared Spectrometer

Cell

: KBr Liquid Cell

Slit

: Medium

Results: Infrared spectrum of the test substance agreed with before use and after use.

1994.01.11(date analyzed)	1994.02.25(date analyzed)
Wave Number(cm ⁻¹)	Wave Number(cm ⁻¹)
850~ 970	850~ 970
970~1220	$970 \sim 1220$
1220~1400	$1220 \sim 1400$
1500~1800	1500~1800
2000~2500	$2000 \sim 2500$
$2500 \sim 3750$	$2500 \sim 3750$

3. High Performance Liquid Chromatography

Instrument

: Shimadzu LC-10AD(Pump)

Esa Coulochem II (Detector)

Column

: TSK GEL ODS-80TM(4.6mm $\phi \times 15$ cm)

Column Temperature : 50°C

Flow Rate

: lml/min

Mobile Phase

: Water (pH. 7.0 Phosphate Buffer Powder +

0.2mol/l Sodium Perchlorate Monohydrate)

Oxidation Voltage : 600mV(Analytical Cell)

650mV(Guard Cell)

Detector

: ECD(Electrochemical Detector)

Injection Volume : $10 \mu 1$

Results: Chromatogram indicated one major peak analyzed at 1994.1.11 and one major peak analyzed at 1994.2.25. The new trace impurity peak in the test substance analyzed at 1994.2.25 was not detected.

Date	Retention Time(min)	AREA	
1994.01.11 (date analyzed)	2. 353	393562	
1994.02.25 (date analyzed)	2. 353	390976	

^{4.} Conclusions: The test substance was stable for about 6 weeks in the dark at room temperature.

APPENDIX A 12-3 CONCENTRATION HYDRAZINE MONOHYDRATE IN DRINKING WATER (TOW-WEEK STUDIES)

CONCENTRATION OF HYDRAZINE MONOHYDRATE IN DRINKING WATER (TWO-WEEK STUDIES)

(Rat)

		Target Concentra	tion(ppm)		
Date analyzed	50	100	200	400	800
1994.01.27	49.5(99.0)*	99.8(99.8)	204.6(102.3)	424.8(106.2)	799.4(99.9)
(Mouse)					
		Target Concentra	tion(ppm)		
Date analyzed	50	100	200	400	800
1994.02.10	50.1(100.2)	102.4(102.4)	212.0(106.0)	422.0(105.5)	846.4(105.8)

^{(*) %} of target concentration

Analytical method: The sample were analyzed by the HPLC.

: Shimadzu LC-10AD(Pump) Instrument

Esa Coulochem II (Detector)

: TSK GEL ODS-80TM(4.6mm $\phi \times 15$ cm)

Column Column Temperature: 50°C

Flow Rate : lml/min Mobile Phase

: Water(pll7. O Phosphate Buffer Powder +

0.2mol/1 Sodium Perchlorate Monohydrate)

Oxidation Voltage : 600mv(Analytical Cell)

650mv(Guard Cell)

Detector : ECD(Electrochemical Detector)

Injection Volume $: 10 \mu 1$

APPENDIX A 12-4 STABILITY OF HYDRAZINE MONOHYDRATE IN DRINKING WATER (TWO-WEEK STUDIES)

STABILITY OF HYDRAZINE MONOHYDRATE IN DRINKING WATER (TWO-WEEK STUDIES)

(Rat)

Date analyzed	Target Concentration(ppm)	
	50	800
1994.01.27(a)	49.5(99.0)*	799.4(99.9)
1994.02.01(b)	36.3(72.6)	707.3(88.4)

(Mouse)

	Target Concentration(ppm)	
Date analyzed	50	800
1994.02.10(a)	50.1(100.2)	846.4(105.8)
1994.02.15(b)	43.5(87.0)	716.3(89.5)

⁽a) Date of preparation

Analytical method: The sample were analyzed by the HPLC.

Instrument : Shimadzu LC-10AD(Pump)

Esa Coulochem II (Detector)

: TSK GEL ODS-80TM(4.6mm $\phi \times 15$ cm) Column

Column Temperature: 50°C

Flow Rate : lml/min Mobile Phase

Detector

: Water(pll7. O Phosphate Buffer Powder +

0. 2mol/1 Sodium Perchlorate Monohydrate)

Oxidation Voltage : 600mv(Analytical Cell)

650mv(Guard Cell)

: ECD(Electrochemical Detector)

Injection Volume $: 10 \mu 1$

⁽b) The stability of Hydrazine monohydrate in drinking water was established for 5 days when stored at room temperature.

^{(*) %} of target concentration