

イソキサチオンのラットを用いた  
吸入による28日間毒性試験報告書

試験番号：0752

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**TABLE A**

**CONCENTRATIONS OF ISOXATHION  
IN THE INHALATION CHAMBER  
OF THE 28-DAY INHALATION STUDY**

CONCENTRATIONS OF ISOXATHION IN THE INHALATION  
CHAMBER OF THE 28-DAY INHALATION STUDY

Group Name		Weight Measurement by Balance				Analysis by LC			
		Control	1 mg/m <sup>3</sup>	3 mg/m <sup>3</sup>	10 mg/m <sup>3</sup>	Control	1 mg/m <sup>3</sup>	3 mg/m <sup>3</sup>	10 mg/m <sup>3</sup>
2009/10/21	1	0.0	1.1	3.1	10.5	0.0	1.0	2.9	10.0
	2	0.0	1.1	3.0	9.8				
	3	0.0	1.0	3.1	10.2				
2009/10/22	1	0.0	1.0	3.0	10.5				
	2	0.0	1.0	3.0	10.6				
	3	0.0	1.0	3.0	10.4				
2009/10/23	1	0.0	1.0	2.9	10.2				
	2	0.0	1.1	3.1	9.6				
	3	0.0	0.9	2.9	9.8				
2009/10/26	1	0.0	1.0	3.2	10.4				
	2	0.0	1.1	3.0	9.8				
	3	0.0	1.1	3.1	10.1				
2009/10/27	1	0.0	1.0	3.1	9.8				
	2	0.0	1.0	3.1	10.0				
	3	0.0	1.1	3.1	9.9				
2009/10/28	1	0.0	1.1	3.0	10.7	0.0	0.9	3.1	10.4
	2	0.0	1.0	3.0	10.8				
	3	0.0	0.9	3.0	10.4				
2009/10/29	1	0.0	0.9	3.1	9.9				
	2	0.0	1.0	3.0	10.1				
	3	0.0	1.0	3.1	9.8				
2009/10/30	1	0.0	1.0	3.1	9.8				
	2	0.0	1.0	3.0	9.7				
	3	0.0	0.9	3.0	9.8				
2009/11/2	1	0.0	1.0	3.0	10.5				
	2	0.0	1.0	3.0	9.6				
	3	0.0	1.0	3.1	9.8				
2009/11/3	1	0.0	1.0	3.0	9.6				
	2	0.0	1.0	2.9	10.0				
	3	0.0	1.0	3.0	9.8				
2009/11/4	1	0.0	1.0	3.0	10.3	0.0	1.0	3.1	10.4
	2	0.0	0.9	2.9	9.8				
	3	0.0	1.0	3.0	9.8				
2009/11/5	1	0.0	1.0	3.1	9.9				
	2	0.0	1.0	3.1	10.5				
	3	0.0	1.0	3.0	9.8				
2009/11/6	1	0.0	1.0	2.9	10.3				
	2	0.0	1.0	3.0	9.9				
	3	0.0	1.0	3.0	9.9				
2009/11/9	1	0.0	1.1	3.0	10.4				
	2	0.0	1.0	3.1	10.3				
	3	0.0	1.0	3.0	10.1				
2009/11/10	1	0.0	1.1	3.1	10.2				
	2	0.0	0.9	3.1	10.1				
	3	0.0	1.0	3.0	9.9				
2009/11/11	1	0.0	1.0	2.9	10.1	0.0	1.1	3.3	11.0
	2	0.0	1.0	3.1	10.2				
	3	0.0	1.0	3.0	10.0				
2009/11/12	1	0.0	0.9	2.9	9.9				
	2	0.0	1.0	2.9	9.9				
	3	0.0	1.0	3.1	9.8				
2009/11/13	1	0.0	1.0	3.0	10.1				
	2	0.0	1.0	3.0	10.2				
	3	0.0	1.0	3.0	10.0				
2009/11/16	1	0.0	1.0	3.1	10.2				
	2	0.0	1.0	3.1	10.0				
	3	0.0	1.0	3.1	10.0				
2009/11/17	1	0.0	1.0	3.0	9.8				
	2	0.0	1.0	3.0	10.0				
	3	0.0	1.1	3.0	10.1				
Mean		0.0	1.0	3.0	10.1	0.0	1.0	3.1	10.5
S.D.		0.0	0.1	0.1	0.3	0.0	0.1	0.1	0.6

**TABLE B1**

**SURVIVAL ANIMAL NUMBERS : MALE**

SURVIVAL ANIMAL NUMBERS

STUDY NO. : 0752  
 ANIMAL : RAT Cr:1:CD(SD) [Cr:j:CD(SD) IGS]  
 REPORT TYPE : A1 4  
 SEX : MALE

Group Name	Animals At start	Administration (Weeks)				
		0	1	2	3	4
Control	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
1mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
3mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
10mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0

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

**TABLE B2**

**SURVIVAL ANIMAL NUMBERS : FEMALE**



SURVIVAL ANIMAL NUMBERS

STUDY NO. : 0752  
 ANIMAL : RAT Cr-I:CD(SD) [Cr-I:CD(SD) IGS]  
 REPORT TYPE : AI 4  
 SEX : FEMALE

Group Name	Animals At start	Administration (Weeks)				
		0	1	2	3	4
Control	10	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0
1mg/m3	10	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0
3mg/m3	10	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0
10mg/m3	10	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0	10/10 100.0

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

**TABLE C1**

**CLINICAL OBSERVATION : MALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr-I:CD(SD) [Cr-J:CD(SD) IGS]  
 REPORT TYPE : A1 4

SEX : MALE

CLINICAL OBSERVATION (SUMMARY)

ALL ANIMALS

PAGE : 1

Clinical sign	Group Name	Administration Week-day			
		1-7	2-7	3-7	4-7
NON REMARKABLE	Control	10	10	10	10
	1mg/m3	10	10	10	10
	3mg/m3	10	10	10	10
	10mg/m3	10	10	10	10

(HAN190)

BATS 4

**TABLE C2**

**CLINICAL OBSERVATION : FEMALE**

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

STUDY NO. : 0752  
ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]  
REPORT TYPE : A1 4

SEX : FEMALE

PAGE : 2

Clinical sign	Group Name	Administration Week-day			
		1-7	2-7	3-7	4-7
NON REMARKABLE	Control	10	10	10	10
	1mg/m <sup>3</sup>	10	10	10	10
	3mg/m <sup>3</sup>	10	10	10	10
	10mg/m <sup>3</sup>	10	10	10	10

(HNV190)

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**TABLE D1**

**BODY WEIGHT CHANGES AND SURVIVAL ANIMAL**

**NUMBERS : MALE**

MEAN BODY WEIGHTS AND SURVIVAL

STUDY NO. : 0752  
 ANIMAL : RAT CrI:CD(SD) [Crj:CD(SD)IGS]  
 UNIT : E  
 REPORT TYPE : A1 4  
 SEX : MALE

Week-Day on Study	Control			1mg/m3			3mg/m3			10mg/m3		
	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>
0-0	305 (10)	10/10	100	305 (10)	10/10	100	305 (10)	10/10	100	305 (10)	10/10	100
1-7	351 (10)	10/10	97	342 (10)	10/10	100	352 (10)	10/10	99	349 (10)	10/10	99
2-7	387 (10)	10/10	97	374 (10)	10/10	101	389 (10)	10/10	99	385 (10)	10/10	99
3-7	423 (10)	10/10	96	404 (10)	10/10	99	418 (10)	10/10	98	416 (10)	10/10	98
4-7	450 (10)	10/10	95	426 (10)	10/10	95	442 (10)	10/10	98	440 (10)	10/10	98

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

(BI0040)

**TABLE D2**

**BODY WEIGHT CHANGES AND SURVIVAL ANIMAL**

**NUMBERS : FEMALE**



MEAN BODY WEIGHTS AND SURVIVAL

STUDY NO. : 0752  
 ANIMAL : RAT Crl:CD(SD) [Crl:CD(SD)IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : FEMALE

Week-Day on Study	Control			1mg/m3			3mg/m3			10mg/m3		
	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>	Av. Wt.	No. of Surviv.	% of cont. <10>
0-0	215 (10)	10/10	100	215 (10)	10/10	100	215 (10)	10/10	100	215 (10)	10/10	100
1-7	237 (10)	10/10	100	238 (10)	10/10	100	237 (10)	10/10	100	237 (10)	10/10	100
2-7	254 (10)	10/10	98	248 (10)	10/10	100	253 (10)	10/10	100	246 (10)	10/10	97
3-7	266 (10)	10/10	99	263 (10)	10/10	100	266 (10)	10/10	100	258 (10)	10/10	97
4-7	274 (10)	10/10	100	275 (10)	10/10	100	278 (10)	10/10	101	266 (10)	10/10	97

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

(B10040)

**TABLE D3**

**BODY WEIGHT CHANGES : MALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr:1:CD(SD) [Cr:j:CD(SD) IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : MALE

BODY WEIGHT CHANGES  
 ALL ANIMALS (SUMMARY)

Group Name	Administration week-day				
	0-0	1-7	2-7	3-7	4-7
Control	305 ± 10	351 ± 15	387 ± 15	423 ± 17	450 ± 19
1mg/m <sup>3</sup>	305 ± 10	342 ± 14	374 ± 20	404 ± 22	426 ± 22
3mg/m <sup>3</sup>	305 ± 10	352 ± 16	389 ± 18	418 ± 24	442 ± 28
10mg/m <sup>3</sup>	305 ± 10	349 ± 12	385 ± 17	416 ± 20	440 ± 20

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

**TABLE D4**

**BODY WEIGHT CHANGES : FEMALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr:I:CD(SD) [Cr:j:CD(SD)IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : FEMALE

BODY WEIGHT CHANGES  
 ALL ANIMALS (SUMMARY)

Group Name	Administration week-day				
	0-0	1-7	2-7	3-7	4-7
Control	215± 8	237± 14	254± 16	266± 17	274± 19
1mg/m3	215± 9	236± 15	248± 14	263± 18	275± 17
3mg/m3	215± 9	238± 11	253± 12	266± 11	278± 15
10mg/m3	215± 9	237± 13	246± 16	258± 17	266± 18

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

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**TABLE E1**

**FOOD CONSUMPTION CHANGES : MALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr:1:CD(SD) [Crj:CD(SD)IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)  
 ALL ANIMALS

Group Name	Administration week-day(effective)			
	1-7(7)	2-7(7)	3-7(7)	4-7(7)
Control	26.9± 2.3	26.4± 1.7	26.5± 1.7	26.6± 2.0
1mg/m3	24.9± 1.9	24.9± 2.4	25.0± 1.7	25.1± 1.8
3mg/m3	26.4± 2.1	26.2± 1.9	25.7± 2.4	26.0± 3.0
10mg/m3	26.7± 1.9	26.5± 1.5	26.1± 1.8	26.1± 1.4

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Dunnett

(HAN260)

**TABLE E2**

**FOOD CONSUMPTION CHANGES : FEMALE**



STUDY NO. : 0752

ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]

UNIT : g

REPORT TYPE : AI 4

SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)

ALL ANIMALS

Group Name	Administration week-day(effective)			
	1-7(7)	2-7(7)	3-7(7)	4-7(7)
Control	18.6± 1.5	18.7± 1.9	18.7± 1.7	18.2± 1.9
1mg/m3	18.7± 1.6	18.3± 3.0	19.2± 2.3	18.7± 1.8
3mg/m3	18.7± 1.8	19.1± 2.6	19.7± 1.4	18.8± 2.8
10mg/m3	18.4± 2.0	17.1± 1.8	18.5± 1.8	16.9± 1.3

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

Test of Dunnett

(HAN260)

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**TABLE F1**

**HEMATOLOGY : MALE**

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : MALE  
REPORT TYPE : AI

Group Name	NO. of Animals	RED BLOOD CELL 10 <sup>6</sup> /μl	HEMOGLOBIN g/dl	HEMATOCRIT %	MCV fl	MCH pg	MCHC g/dl	PLATELET 10 <sup>3</sup> /μl
Control	10	8.41 ± 0.26	15.6 ± 0.5	45.0 ± 1.4	53.6 ± 1.4	18.6 ± 0.5	34.8 ± 0.3	1170 ± 130
1mg/m3	10	8.62 ± 0.42	15.8 ± 0.5	45.7 ± 1.6	53.1 ± 1.6	18.4 ± 0.6	34.7 ± 0.4	1216 ± 97
3mg/m3	10	8.55 ± 0.34	15.6 ± 0.5	45.3 ± 1.3	53.0 ± 1.3	18.3 ± 0.6	34.5 ± 0.6	1246 ± 126
10mg/m3	10	8.60 ± 0.34	15.8 ± 0.5	45.7 ± 1.4	53.2 ± 1.9	18.3 ± 0.7	34.5 ± 0.5	1162 ± 110

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01

Test of Dunnett

(HCL070)

BAIS 4

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : MALE REPORT TYPE : A1

Group Name	NO. of Animals	RETICULOCYTE %
Control	10	2.1 ± 0.2
1mg/m <sup>3</sup>	10	2.0 ± 0.3
3mg/m <sup>3</sup>	10	2.0 ± 0.3
10mg/m <sup>3</sup>	10	1.9 ± 0.3

Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL070)

BATS 4

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5#)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
MEASURE, TIME : 1  
SEX : MALE  
REPORT TYPE : A1

Group Name	NO. of Animals	WBC 10 <sup>3</sup> /μl	Differential NEUTRO	WBC (%) LYMPHO	MONO	EOSINO	BASO	OTHER
Control	10	10.73± 3.20	16± 5	79± 5	3± 1	1± 0	0± 0	1± 0
1mg/m3	10	11.26± 2.12	18± 5	77± 6	3± 1	1± 0	0± 0	1± 0
3mg/m3	10	12.86± 4.17	15± 5	80± 5	2± 1	1± 1	0± 0	1± 0
10mg/m3	10	12.24± 4.40	18± 6	77± 6	2± 0	2± 1	0± 0	1± 1

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01    Test of Dunnett

**TABLE F2**

**HEMATOLOGY : FEMALE**

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr1:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

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Group Name	NO. of Animals	RED BLOOD CELL 10 <sup>6</sup> /μl	HEMOGLOBIN g/dl	HEMATOCRIT %	MCV fl	MCH pg	MCHC g/dl	PLATELET 10 <sup>9</sup> /μl
Control	10	8.11 ± 0.50	15.1 ± 0.7	42.9 ± 1.9	53.0 ± 2.1	18.6 ± 0.7	35.2 ± 0.4	1392 ± 91
1mg/m3	10	8.12 ± 0.31	15.2 ± 0.6	43.4 ± 1.7	53.4 ± 1.0	18.8 ± 0.5	35.1 ± 0.6	1360 ± 102
3mg/m3	10	8.02 ± 0.38	15.3 ± 0.6	43.1 ± 1.9	53.7 ± 1.6	19.1 ± 0.5	35.5 ± 0.4	1328 ± 132
10mg/m3	10	8.14 ± 0.39	15.3 ± 0.6	43.2 ± 1.7	53.1 ± 1.2	18.8 ± 0.6	35.5 ± 0.4	1298 ± 145

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

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(HCL070)

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HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr1:CD(SD) IGS]  
MEASURE TIME : 1  
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	RETICULOCYTE %
Control	10	2.3 ± 0.5
1mg/m3	10	2.2 ± 0.4
3mg/m3	10	2.3 ± 0.7
10mg/m3	10	1.9 ± 0.5

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

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(HCL070)

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HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	WBC 10 <sup>3</sup> /μl	NEUTRO	Differential WBC LYMPHO (%)	MONO	EOSINO	BASO	OTHER						
Control	10	6.69± 2.50	21±	8	74±	8	3±	1	2±	1	0±	0	1±	0
1mg/m3	10	9.11± 3.78	16±	7	79±	7	3±	1	1±	0	0±	0	1±	0
3mg/m3	10	8.83± 5.37	16±	7	79±	6	2±	0	2±	1	0±	0	1±	0
10mg/m3	10	8.26± 2.46	17±	6	78±	6	3±	1	2±	1	0±	0	1±	0

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL070)

BAIS 4

**TABLE G1**

**BIOCHEMISTRY : MALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD) IGS]  
MEASURE, TIME : 1  
SEX : MALE REPORT TYPE : A1

PAGE : 1

Group Name	NO. of Animals	TOTAL PROTEIN g/dl	ALBUMIN g/dl	A/G RATIO	T-BILIRUBIN mg/dl	GLUCOSE mg/dl	T-CHOLESTEROL mg/dl	TRIGLYCERIDE mg/dl
Control	10	6.1± 0.4	3.5± 0.2	1.3± 0.1	0.11± 0.01	161± 9	57± 9	34± 10
1mg/m3	10	6.2± 0.2	3.5± 0.1	1.3± 0.1	0.12± 0.01	161± 11	55± 8	30± 11
3mg/m3	10	6.0± 0.2	3.5± 0.1	1.4± 0.1	0.11± 0.01	168± 13	55± 9	32± 13
10mg/m3	10	6.2± 0.3	3.5± 0.2	1.3± 0.1	0.12± 0.01	159± 12	55± 11	34± 17

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

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(HCL074)

BALS 4

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : MALE  
REPORT TYPE : A1

Group Name	NO. of Animals	PHOSPHOLIPID mg/dl	AST IU/l	ALT IU/l	LDH IU/l	ALP IU/l	G-GTP IU/l	CK IU/l
Control	10	94±	63±	25±	120±	466±	1±	131±
1mg/m3	10	92±	68±	29±	116±	581±	1±	139±
3mg/m3	10	90±	68±	26±	100±	522±	2±	127±
10mg/m3	10	94±	70±	26±	109±	534±	1±	133±

Significant difference ; \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL074)

BALS 4

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : MALE  
REPORT TYPE : AI

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	10	13.3±	0.5±	143±	3.7±	107±	10.2±	7.3±
1mg/m3	10	13.4±	0.5±	143±	3.6±	106±	10.2±	7.2±
3mg/m3	10	13.2±	0.5±	143±	3.7±	107±	10.1±	7.2±
10mg/m3	10	13.2±	0.5±	143±	3.5±	106±	10.2±	7.2±

Test of Dunnett

\*\* : P ≤ 0.01

\* : P ≤ 0.05

Significant difference :

(HCL074)

BAIS 4

**TABLE G2**

**BIOCHEMISTRY : FEMALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : FEMALE REPORT TYPE : A1

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	10	6.7± 0.3	4.1± 0.1	1.6± 0.2	0.14± 0.02	153± 22	68± 10	18± 8
1mg/m3	10	6.7± 0.2	3.9± 0.1	1.4± 0.1	0.13± 0.01	158± 14	69± 14	16± 7
3mg/m3	10	6.5± 0.3	3.9± 0.2	1.5± 0.2	0.13± 0.01	150± 14	67± 14	16± 9
10mg/m3	10	6.7± 0.4	4.0± 0.3	1.5± 0.2	0.13± 0.01	142± 23	65± 15	13± 5

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL074)

BAIS 4

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

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Group Name	NO. of Animals	PHOSPHOLIPID mg/dl	AST IU/l	ALT IU/l	LDH IU/l	ALP IU/l	G-GTP IU/l	CK IU/l
Control	10	130 ± 17	62 ± 13	21 ± 4	111 ± 44	263 ± 105	2 ± 1	94 ± 24
1mg/m3	10	129 ± 26	69 ± 16	32 ± 16	108 ± 56	323 ± 74	1 ± 0	113 ± 38
3mg/m3	10	125 ± 25	58 ± 4	21 ± 4	102 ± 28	287 ± 64	2 ± 1	101 ± 16
10mg/m3	10	120 ± 22	58 ± 6	21 ± 6	104 ± 51	260 ± 53	1 ± 1	93 ± 22

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL074)

BAIS 4



BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : AI

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	10	14.8±	0.6±	141±	3.3±	107±	10.3±	5.2±
1mg/m3	10	15.0±	0.6±	141±	3.5±	107±	10.2±	5.7±
3mg/m3	10	15.0±	0.6±	141±	3.4±	107±	10.2±	5.8±
10mg/m3	10	16.1±	0.6±	141±	3.8±	107±	10.3±	5.7±

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01

Test of Dunnett

(HCL074)

BAIS 4

TABLE H1

BIOCHEMISTRY

(ACETYLCHOLINEESTERASE ACTIVITY)

: MALE

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT CrI:CD(SD) [Crj:CD(SD) IGS]  
MEASURE, TIME : 1  
SEX : MALE  
REPORT TYPE : A1

Group Name	NO. of Animals	CHE (BRAIN) IU/ℓ	CHE (ERYTHROCYTE) IU/ℓ	CHE (PLASMA) IU/ℓ
Control	10	2268 ± 156	2106 ± 235	470 ± 43
1mg/m3	10	2162 ± 240	1754 ± 321	395 ± 89
3mg/m3	10	2227 ± 95	1745 ± 370	415 ± 84
10mg/m3	10	2199 ± 153	871 ± 400**	265 ± 58**

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01    Test of Dunnett

**TABLE H2**

**BIOCHEMISTRY**

**(ACETYLCHOLINEESTERASE ACTIVITY)**

**: FEMALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

PAGE : 8

Group Name	No. of Animals	CHE (BRAIN) IU/ℓ	CHE (ERYTHROCYTE) IU/ℓ	CHE (PLASMA) IU/ℓ
Control	10	2136± 190	2392± 501	2501± 522
1mg/m3	10	2233± 114	2219± 500	2284± 554
3mg/m3	10	2131± 212	1898± 424	1908± 301**
10mg/m3	10	2216± 124	1299± 484**	867± 212**

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01    Test of Dunnett

(HCL074)

BAIS 4

TABLE I1

LYMPHOCYTE-SUBSET TEST: MALE

### LYMPHOCYTE - SUBSET TEST

Flow cytometric analysis - Group mean values of peripheral blood lymphocyte in male rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Peripheral blood lymphocyte (10 <sup>3</sup> cells/ $\mu$ L)				Th/Tc ratio
		Pan-T cell (CD3+)	Helper-T cell (CD4+)	Cytotoxic-T cell (CD8+)		
0	10	Mean	2.21	1.16	2.08	
		S.D.	0.59	0.50	0.67	
1	10	Mean	2.20	1.19	1.86	
		S.D.	0.43	0.22	0.24	
3	10	Mean	2.56	1.28	2.09	
		S.D.	1.02	0.54	0.39	
10	10	Mean	2.46	1.21	2.20	
		S.D.	0.70	0.49	0.59	

Th/Tc ratio: Helper-T cell/Cytotoxic-T cell ratio.

S.D.: Standard deviation.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test.

Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .

### Lymphocyte - Subset Test

Flow cytometric analysis - Group mean values of thymic lymphocyte in male rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Thymic lymphocyte (10 <sup>7</sup> cells/rat)					
		Immature cells			Mature cells		
		DN cell (CD4-CD8-)	DP cell (CD4+CD8+)		Helper-T cell (CD4+CD8-)	Cytotoxic-T cell (CD4-CD8+)	
0	5	Mean	1.3	61.6		6.3	2.7
		S.D.	0.1	12.5		1.3	0.6
1	5	Mean	1.9	73.1		7.5	3.7
		S.D.	0.6	27.2		1.7	1.1
3	5	Mean	2.4 *	81.7		6.9	3.3
		S.D.	0.6	22.4		1.2	0.3
10	5	Mean	2.2	75.7		7.8	3.3
		S.D.	0.5	11.5		3.0	1.3

S.D.: Standard deviation.

Immature cells: DN; Double negative, DP; Double positive.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test. Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .



### Lymphocyte - Subset Test

Flow cytometric analysis - Group mean values of splenic lymphocyte in male rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Splenic lymphocyte (10 <sup>7</sup> cells/rat)		
		Pan-T cell (CD3+)	Helper-T cell (CD4+CD8-)	Cytotoxic-T cell (CD4-CD8+)
0	5	Mean	8.1	5.5
		S.D.	2.5	2.2
1	5	Mean	6.5	3.9
		S.D.	1.7	1.0
3	5	Mean	8.6	5.0
		S.D.	1.9	0.9
10	5	Mean	8.7	5.0
		S.D.	2.2	1.6

S.D.: Standard deviation.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test.

Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .

**TABLE 12**

**LYMPHOCYTE-SUBSET TEST: FEMALE**

LYMPHOCYTE - SUBSET TEST

Flow cytometric analysis - Group mean values of peripheral blood lymphocyte in female rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Peripheral blood lymphocyte (10 <sup>3</sup> cells/ $\mu$ L)				Th/Tc ratio
		Pan-T cell (CD3+)	Helper-T cell (CD4+)	Cytotoxic-T cell (CD8+)		
0	10	Mean	2.72	1.81	0.84	2.32
		S.D.	1.29	0.83	0.46	0.58
1	10	Mean	3.35	2.23	1.02	2.68
		S.D.	1.42	0.89	0.53	1.66
3	10	Mean	3.13	2.02	1.02	1.94
		S.D.	1.48	1.06	0.42	0.48
10	10	Mean	3.34	2.21	1.05	2.16
		S.D.	1.03	0.72	0.34	0.47

Th/Tc ratio: Helper-T cell/Cytotoxic-T cell ratio.

S.D.: Standard deviation.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test.

Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .

**LYMPHOCYTE - SUBSET TEST**  
Flow cytometric analysis - Group mean values of thymic lymphocyte in female rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Thymic lymphocyte (10 <sup>7</sup> cells/rat)					
		Immature cells			Mature cells		
		DN cell (CD4-CD8-)	DP cell (CD4+CD8+)	Helper-T cell (CD4+CD8-)	Cytotoxic-T cell (CD4-CD8+)		
0	5	Mean	1.9	53.0	6.5	2.9	
		S.D.	0.4	12.2	2.0	0.9	
1	5	Mean	2.4	64.3	7.1	3.3	
		S.D.	0.9	21.9	2.3	1.3	
3	5	Mean	2.5	63.6	5.7	2.9	
		S.D.	1.0	11.7	0.8	0.6	
10	5	Mean	2.8	70.5	7.0	3.3	
		S.D.	0.8	15.7	1.2	0.9	

S.D.: Standard deviation.

Immature cells: DN; Double negative, DP; Double positive.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test.

Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .

### LYMPHOCYTE - SUBSET TEST

Flow cytometric analysis - Group mean values of splenic lymphocyte in female rats

Dose (mg/m <sup>3</sup> )	Number of animals examined	Splenic lymphocyte (10 <sup>7</sup> cells/rat)		
		Pan-T cell (CD3+)	Helper-T cell (CD4+CD8-)	Cytotoxic-T cell (CD4-CD8+)
0	5	Mean	5.7	3.6
		S.D.	3.9	3.2
1	5	Mean	6.2	4.3
		S.D.	2.9	1.9
3	5	Mean	4.2	3.5
		S.D.	0.7	0.7
10	5	Mean	6.1	4.7
		S.D.	3.2	2.8

S.D.: Standard deviation.

Data were statistically analyzed by Dunnett's test following one-way ANOVA or Dunnett-type test following Kruskal-Wallis test.

Significantly different from control: \*,  $p \leq 0.05$ ; \*\*,  $p \leq 0.01$ .

**TABLE J**

**GROSS FINDINGS : FEMALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)  
 ALL ANIMALS (0- 5W)

Organ	Findings	Group Name NO. of Animals	Control 10 (%)	1mg/m3 10 (%)	3mg/m3 10 (%)	10mg/m3 10 (%)
liver	herniation		1 ( 10)	0 ( 0)	0 ( 0)	0 ( 0)

(HPT080)

**TABLE K1**

**ORGAN WEIGHT, ABSOLUTE : MALE**



ORGAN WEIGHT: ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD) IGS]  
REPORT TYPE : A1  
SEX : MALE  
UNIT : g

Group Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	10	418 ± 18	0.446 ± 0.100	0.078 ± 0.015	2.905 ± 0.531	1.414 ± 0.165	1.339 ± 0.077
1mg/m3	10	395 ± 19	0.417 ± 0.082	0.072 ± 0.007	3.134 ± 0.274	1.361 ± 0.107	1.315 ± 0.091
3mg/m3	10	412 ± 27	0.454 ± 0.088	0.074 ± 0.007	3.185 ± 0.301	1.381 ± 0.113	1.328 ± 0.087
10mg/m3	10	407 ± 22	0.489 ± 0.113	0.071 ± 0.011	3.065 ± 0.255	1.401 ± 0.096	1.355 ± 0.042

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

Test of Dunnett

(HCL040)

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD)IGS]  
REPORT TYPE : A1  
SEX : MALE  
UNIT : g

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	2.883 ± 0.243	0.704 ± 0.097	11.157 ± 1.252	1.980 ± 0.079
1mg/m3	10	2.713 ± 0.216	0.657 ± 0.083	10.589 ± 0.582	1.962 ± 0.076
3mg/m3	10	2.866 ± 0.267	0.726 ± 0.089	10.994 ± 0.903	1.990 ± 0.067
10mg/m3	10	2.904 ± 0.232	0.691 ± 0.080	11.266 ± 0.900	1.975 ± 0.068

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Dunnett

(HCL040)

**TABLE K2**

**ORGAN WEIGHT, ABSOLUTE : FEMALE**

STUDY NO. : 0752  
 ORGAN WEIGHT: ABSOLUTE (SUMMARY)  
 SURVIVAL ANIMALS ( 5W)

ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE  
 UNIT : g

Group Name	No. of Animals	Body Weight	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	10	253 ± 18	0.374 ± 0.070	0.082 ± 0.010	0.140 ± 0.025	0.880 ± 0.057	1.039 ± 0.095
1mg/m3	10	251 ± 14	0.390 ± 0.087	0.083 ± 0.005	0.137 ± 0.019	0.905 ± 0.064	1.072 ± 0.089
3mg/m3	10	254 ± 12	0.419 ± 0.067	0.079 ± 0.013	0.145 ± 0.019	0.919 ± 0.075	1.078 ± 0.061
10mg/m3	10	247 ± 18	0.418 ± 0.048	0.077 ± 0.009	0.130 ± 0.036	0.918 ± 0.054	1.030 ± 0.045

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Dunnett

(HCL040)

ORGAN WEIGHT:ABSOLUTE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr:1:CD(SD) [Cr:j:CD(SD) IGS]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT : g

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	1.727 ± 0.126	0.478 ± 0.070	6.877 ± 0.530	1.867 ± 0.067
1mg/m3	10	1.792 ± 0.149	0.509 ± 0.084	6.934 ± 0.639	1.861 ± 0.054
3mg/m3	10	1.808 ± 0.163	0.480 ± 0.065	6.847 ± 0.498	1.851 ± 0.099
10mg/m3	10	1.804 ± 0.126	0.471 ± 0.068	6.635 ± 0.674	1.894 ± 0.090

Significant difference : \* : P ≤ 0.05    \*\* : P ≤ 0.01    Test of Dunnett

(HCL040)

**TABLE L1**

**ORGAN WEIGHT, RELATIVE : MALE**

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
REPORT TYPE : A1  
SEX : MALE  
UNIT : %

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	10	418 ± 18	0.107 ± 0.023	0.019 ± 0.004	0.695 ± 0.127	0.338 ± 0.030	0.321 ± 0.013
1mg/m3	10	395 ± 19	0.105 ± 0.017	0.018 ± 0.002	0.794 ± 0.069	0.344 ± 0.021	0.333 ± 0.024
3mg/m3	10	412 ± 27	0.110 ± 0.017	0.018 ± 0.002	0.779 ± 0.108	0.336 ± 0.024	0.323 ± 0.021
10mg/m3	10	407 ± 22	0.120 ± 0.029	0.017 ± 0.002	0.755 ± 0.075	0.345 ± 0.026	0.334 ± 0.018

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL042)

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD) IGS]  
REPORT TYPE : AI  
SEX : MALE  
UNIT : %

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	0.690 ± 0.045	0.168 ± 0.020	2.665 ± 0.203	0.474 ± 0.020
1mg/m3	10	0.687 ± 0.056	0.166 ± 0.020	2.682 ± 0.135	0.498 ± 0.038
3mg/m3	10	0.698 ± 0.077	0.176 ± 0.019	2.668 ± 0.087	0.484 ± 0.025
10mg/m3	10	0.714 ± 0.048	0.170 ± 0.017	2.766 ± 0.118	0.486 ± 0.028

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01 Test of Dunnett

(HCL042)



**TABLE L2**

**ORGAN WEIGHT, RELATIVE : FEMALE**

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT : %

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	10	253 ± 18	0.148 ± 0.028	0.033 ± 0.003	0.056 ± 0.012	0.349 ± 0.022	0.412 ± 0.042
1mg/m3	10	251 ± 14	0.155 ± 0.032	0.033 ± 0.003	0.054 ± 0.007	0.361 ± 0.027	0.427 ± 0.028
3mg/m3	10	254 ± 12	0.165 ± 0.024	0.031 ± 0.004	0.057 ± 0.007	0.363 ± 0.023	0.426 ± 0.034
10mg/m3	10	247 ± 18	0.170 ± 0.021	0.032 ± 0.004	0.053 ± 0.015	0.374 ± 0.032	0.419 ± 0.030

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL042)

ORGAN WEIGHT:RELATIVE (SUMMARY)  
SURVIVAL ANIMALS ( 5W)

STUDY NO. : 0752  
ANIMAL : RAT Cr:1:CD(SD) [Cr:J:CD(SD) IGS]  
REPORT TYPE : A1  
SEX : FEMALE  
UNIT : %

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	0.686± 0.062	0.189± 0.026	2.721± 0.113	0.741± 0.054
1mg/m3	10	0.713± 0.049	0.202± 0.024	2.755± 0.146	0.742± 0.045
3mg/m3	10	0.713± 0.062	0.189± 0.026	2.702± 0.174	0.731± 0.049
10mg/m3	10	0.733± 0.055	0.191± 0.022	2.687± 0.192	0.769± 0.040

Significant difference : \* : P ≤ 0.05 \*\* : P ≤ 0.01

Test of Dunnett

(HCL042)

**TABLE M1**

**HISTOPATHOLOGICAL FINDINGS :**  
**NON-NEOPLASTIC LESIONS : MALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
 REPORT TYPE : AI  
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0- 5W)

Organ	Findings	Control				1mg/m3				3mg/m3				10mg/m3			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
		Grade				Grade				Grade				Grade			
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
(Respiratory system)																	
larynx	inflammatory infiltration	0	0	0	0	1	0	0	0	<10>	<10>	<10>	<10>	0	0	0	0
		(0)	(0)	(0)	(0)	(10)	(0)	(0)	(0)					(0)	(0)	(0)	(0)
lung	accumulation of foamy cells	0	0	0	0	1	0	0	0	<10>	<10>	<10>	<10>	0	0	0	0
		(0)	(0)	(0)	(0)	(10)	(0)	(0)	(0)					(0)	(0)	(0)	(0)
{Urinary system}																	
kidney	retention cyst	1	0	0	0	0	0	0	0	<10>	<10>	<10>	<10>	0	0	0	0
		(10)	(0)	(0)	(0)	(0)	(0)	(0)	(0)					(0)	(0)	(0)	(0)
(Reproductive system)																	
testis	atrophy	1	0	0	0	0	0	0	0	<10>	<10>	<10>	<10>	0	0	0	0
		(10)	(0)	(0)	(0)	(0)	(0)	(0)	(0)					(0)	(0)	(0)	(0)

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

**TABLE M2**

**HISTOPATHOLOGICAL FINDINGS :**  
**NON-NEOPLASTIC LESIONS : FEMALE**

STUDY NO. : 0752  
 ANIMAL : RAT Cr1:CD(SD) [Cr:j:CD(SD) IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NON-NEOPLASTIC LESIONS (SUMMARY)  
 ALL ANIMALS (0- 5W)

Organ	Findings	Control				1mg/m3				3mg/m3				10mg/m3			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
(Digestive system)																	
liver	herniation					<10>											
		1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(10)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
(Endocrine system)																	
thyroid	ultramembranial body remanet					<10>											
		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

TABLE N

ISOXATHION AND HYDROXY ISOXAZOLE  
CONCENTRATIONS IN PLASMA OF RATS



## Isoxathion and Hydroxy isoxazole concentration in plasma of rats

Group Name	Sex	Animal Number	Isoxathion (ng/mL)	Hydroxy isoxazole (ng/mL)
Control	Male	1001	- 1)	- 2)
		1002	-	-
		1003	-	-
		1004	-	-
		1005	-	-
	Female	2001	-	-
		2002	-	-
		2003	-	-
		2004	-	-
		2005	-	-
1 mg/m <sup>3</sup>	Male	1101	-	-
		1102	-	-
		1103	-	-
		1104	-	-
		1105	-	-
	Female	2101	-	-
		2102	-	-
		2103	-	-
		2104	-	-
		2105	-	-
3 mg/m <sup>3</sup>	Male	1201	-	-
		1202	-	-
		1203	-	-
		1204	-	-
		1205	-	-
	Female	2201	-	-
		2202	-	-
		2203	-	-
		2204	-	-
		2205	-	-
10 mg/m <sup>3</sup>	Male	1301	5.61	-
		1302	3.92	-
		1303	3.96	-
		1304	4.06	-
		1305	4.18	-
	Female	2301	5.58	-
		2302	4.03	-
		2303	3.61	-
		2304	4.46	-
		2305	5.85	-

1) : Below limit of quantification. Limit of quantification of Isoxathion was 2ng/mL.  
 2) : Below limit of quantification. Limit of quantification of Hydroxy isoxazole was 50ng/mL.