

トリクロルホ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：0726

# TABLES

## TABLES

TABLE A	CONCENTRATIONS OF TRICHLORFON IN THE INHALATION CHAMBER OF THE 28-DAY INHALATION STUDY
TABLE B 1	SURVIVAL ANIMAL NUMBERS: MALE
TABLE B 2	SURVIVAL ANIMAL NUMBERS: FEMALE
TABLE C 1	CLINICAL OBSERVATION: MALE
TABLE C 2	CLINICAL OBSERVATION: FEMALE
TABLE D 1	BODY WEIGHT CHANGES AND SURVIVAL ANIMAL NUMBERS : MALE
TABLE D 2	BODY WEIGHT CHANGES AND SURVIVAL ANIMAL NUMBERS : FEMALE
TABLE D 3	BODY WEIGHT CHANGES: MALE
TABLE D 4	BODY WEIGHT CHANGES: FEMALE
TABLE E 1	FOOD CONSUMPTION CHANGES: MALE
TABLE E 2	FOOD CONSUMPTION CHANGES: FEMALE
TABLE F 1	HEMATOLOGY: MALE
TABLE F 2	HEMATOLOGY: FEMALE
TABLE G 1	BIOCHEMISTRY: MALE
TABLE G 2	BIOCHEMISTRY: FEMALE
TABLE H 1	BIOCHEMISTRY (ACETYLCHOLINEESTERASE ACTIVITY) : MALE
TABLE H 2	BIOCHEMISTRY (ACETYLCHOLINEESTERASE ACTIVITY) : FEMALE

## TABLES (CONTINUED)

TABLE I 1	LYMPHOCYTE - SUBSET TEST: MALE
TABLE I 2	LYMPHOCYTE - SUBSET TEST: FEMALE
TABLE J	GROSS FINDINGS: FEMALE
TABLE K 1	ORGAN WEIGHT, ABSOLUTE: MALE
TABLE K 2	ORGAN WEIGHT, ABSOLUTE: FEMALE
TABLE L 1	ORGAN WEIGHT, RELATIVE: MALE
TABLE L 2	ORGAN WEIGHT, RELATIVE: FEMALE
TABLE M 1	HISTOPATHOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS : MALE
TABLE M 2	HISTOPATHOLOGICAL FINDINGS: NON-NEOPLASTIC LESIONS : FEMALE
TABLE N	DEP AND DDVP CONCENTRATIONS IN PLASMA OF RATS

TABLE A

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

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

Group Name		Weight Measurement by Balance				Analysis by LC-MS/MS			
		Control	10 mg/m <sup>3</sup>	30 mg/m <sup>3</sup>	100 mg/m <sup>3</sup>	Control	10 mg/m <sup>3</sup>	30 mg/m <sup>3</sup>	100 mg/m <sup>3</sup>
2008/11/5	1	0.0	10.0	33.0	97.1	0.0	10.3	33.5	108.1
	2	0.0	10.1	30.0	100.1				
	3	0.0	9.3	31.6	100.7				
2008/11/6	1	0.0	9.8	31.9	101.0				
	2	0.0	10.0	29.9	102.3				
	3	0.0	9.6	30.3	109.6				
2008/11/7	1	0.0	10.3	30.0	104.1				
	2	0.0	9.9	29.7	103.4				
	3	0.0	10.1	30.7	100.8				
2008/11/10	1	0.0	10.1	29.9	97.7				
	2	0.0	9.8	31.4	108.4				
	3	0.0	10.1	29.9	108.5				
2008/11/11	1	0.0	10.4	30.0	97.9				
	2	0.0	9.8	29.7	98.5				
	3	0.0	9.9	30.4	107.6				
2008/11/12	1	0.0	10.0	29.4	94.4	0.0	10.6	30.4	104.6
	2	0.0	10.3	29.2	104.7				
	3	0.0	10.0	31.0	100.2				
2008/11/13	1	0.0	10.0	30.2	101.9				
	2	0.0	10.2	29.6	99.0				
	3	0.0	9.9	29.4	100.4				
2008/11/14	1	0.0	9.8	29.0	104.3				
	2	0.0	9.8	29.2	103.5				
	3	0.0	9.8	32.1	103.9				
2008/11/17	1	0.0	9.9	30.9	105.4				
	2	0.0	9.7	30.7	99.0				
	3	0.0	10.3	29.7	98.1				
2008/11/18	1	0.0	10.0	30.5	98.8				
	2	0.0	9.8	30.3	100.3				
	3	0.0	10.2	29.7	98.9				
2008/11/19	1	0.0	9.1	29.6	94.1	0.0	9.6	29.5	98.4
	2	0.0	10.3	30.3	104.0				
	3	0.0	10.6	30.9	100.9				
2008/11/20	1	0.0	10.2	30.5	98.9				
	2	0.0	10.4	30.7	102.2				
	3	0.0	10.3	29.3	101.9				
2008/11/21	1	0.0	10.3	29.7	98.3				
	2	0.0	10.3	30.3	100.7				
	3	0.0	10.1	30.5	102.3				
2008/11/22	1	0.0	10.0	29.5	100.5				
	2	0.0	9.7	30.5	100.7				
	3	0.0	10.1	30.3	101.1				
2008/11/25	1	0.0	9.7	28.5	98.4				
	2	0.0	9.7	29.5	100.8				
	3	0.0	9.8	28.6	98.8				
2008/11/26	1	0.0	9.8	29.7	98.1	0.0	9.8	29.2	98.1
	2	0.0	10.0	30.8	100.0				
	3	0.0	10.3	32.5	103.0				
2008/11/27	1	0.0	10.5	31.3	99.0				
	2	0.0	9.9	31.0	101.5				
	3	0.0	9.9	31.7	103.7				
2008/11/28	1	0.0	10.0	29.9	97.7				
	2	0.0	10.2	30.3	102.2				
	3	0.0	9.8	30.7	98.6				
2008/12/1	1	0.0	10.0	29.1	97.7				
	2	0.0	10.0	29.6	99.4				
	3	0.0	10.2	28.9	100.5				
2008/12/2	1	0.0	10.0	29.0	98.5				
	2	0.0	10.1	29.5	100.9				
	3	0.0	10.1	30.3	102.3				
Mean		0.0	10.0	30.2	101.0	0.0	10.6	32.1	107.3
S.D.		0.0	0.3	0.9	3.1	0.0	0.6	1.8	6.0

**TABLE B1**

**SURVIVAL ANIMAL NUMBERS : MALE**

SURVIVAL ANIMAL NUMBERS

STUDY NO. : 0726  
 ANIMAL : RAT CrI:CD(SD) [Crj:CD(SD) IGS]  
 REPORT TYPE : AI 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
10mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
30mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
100mg/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. : 0726  
 ANIMAL : RAT Cr1:CD(SD) [Crj: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	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
30mg/m3	10	10/10	10/10	10/10	10/10	10/10
		100.0	100.0	100.0	100.0	100.0
100mg/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 C1**

**CLINICAL OBSERVATION : MALE**

STUDY NO. : 0726  
 ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
 REPORT TYPE : A1 4

SEX : MALE

CLINICAL OBSERVATION (SUMMARY)  
 ALL ANIMALS

Clinical sign	Group Name	Administration Week-day			
		1-7	2-7	3-7	4-7
NON REMARKABLE	Control	10	10	10	10
	10mg/m3	10	10	10	10
	30mg/m3	10	10	10	10
	100mg/m3	10	10	10	10

(HAN190)

**TABLE C2**

**CLINICAL OBSERVATION : FEMALE**

CLINICAL OBSERVATION (SUMMARY)  
ALL ANIMALS

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
REPORT TYPE : AI 4

SEX : FEMALE

Clinical sign	Group Name	Administration Week-day			
		1-7	2-7	3-7	4-7
NON REMARKABLE	Control	10	10	10	10
	10mg/m3	10	10	10	10
	30mg/m3	10	10	10	10
	100mg/m3	10	10	10	10

(HAN190)

**TABLE D1**

**BODY WEIGHT CHANGES AND SURVIVAL ANIMAL  
NUMBERS : MALE**

MEAN BODY WEIGHTS AND SURVIVAL

STUDY NO. : 0726  
 ANIMAL : RAT Cr:1:CD(SD) [Cr:1:CD(SD)IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : MALE

Week-Day on Study	Control				10mg/m <sup>3</sup>				30mg/m <sup>3</sup>				100mg/m <sup>3</sup>			
	Av. Wt.	No. of Surviv.	Av. Wt.	% of cont. <10>	No. of Surviv.	Av. Wt.	% of cont. <10>	No. of Surviv.	Av. Wt.	% of cont. <10>	No. of Surviv.	Av. Wt.	% of cont. <10>	No. of Surviv.	Av. Wt.	% of cont. <10>
0-0	298 (10)	10/10	298 (10)	100	10/10	298 (10)	100	10/10	298 (10)	100	10/10	298 (10)	100	10/10	298 (10)	100
1-7	395 (10)	10/10	329 (10)	98	10/10	330 (10)	99	10/10	331 (10)	99	10/10	363 (10)	98	10/10	398 (10)	98
2-7	372 (10)	10/10	362 (10)	97	10/10	365 (10)	98	10/10	399 (10)	98	10/10	426 (10)	98	10/10	428 (10)	98
3-7	406 (10)	10/10	396 (10)	98	10/10	399 (10)	98	10/10	426 (10)	98	10/10	426 (10)	98	10/10	428 (10)	98
4-7	436 (10)	10/10	426 (10)	98	10/10	426 (10)	98	10/10	426 (10)	98	10/10	426 (10)	98	10/10	428 (10)	98

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

(B10040)

**TABLE D2**

**BODY WEIGHT CHANGES AND SURVIVAL ANIMAL  
NUMBERS : FEMALE**



MEAN BODY WEIGHTS AND SURVIVAL

STUDY NO. : 0726  
 ANIMAL : RAT Cr:1:CD(SD) [Cr:1:CD(SD)IGS]  
 UNIT : g  
 REPORT TYPE : A1 4  
 SEX : FEMALE

Week-Day on Study	Control			10mg/m <sup>3</sup>			30mg/m <sup>3</sup>			100mg/m <sup>3</sup>		
	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	216 (10)	10/10	100	217 (10)	10/10	100	217 (10)	10/10	100	217 (10)	10/10	100
1-7	238 (10)	10/10	98	233 (10)	10/10	100	237 (10)	10/10	100	235 (10)	99	10/10
2-7	254 (10)	10/10	96	245 (10)	10/10	100	253 (10)	10/10	100	250 (10)	98	10/10
3-7	269 (10)	10/10	97	262 (10)	10/10	97	265 (10)	10/10	99	263 (10)	98	10/10
4-7	282 (10)	10/10	97	274 (10)	10/10	97	278 (10)	10/10	99	280 (10)	99	10/10

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

(BI0040)

**TABLE D3**

**BODY WEIGHT CHANGES : MALE**

STUDY NO. : 0726  
 ANIMAL : RAT Cr:1:CD(SD) [Cr:1:CD(SD)ICS]  
 UNIT : g  
 REPORT TYPE : AI 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	298 ± 14	335 ± 20	372 ± 24	406 ± 28	436 ± 29
10mg/m3	298 ± 14	329 ± 22	362 ± 29	396 ± 36	426 ± 37
30mg/m3	298 ± 13	330 ± 18	365 ± 23	399 ± 26	426 ± 30
100mg/m3	298 ± 15	331 ± 18	363 ± 20	398 ± 23	428 ± 28

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

Test of Dunnett

(HAN260)

**TABLE D4**

**BODY WEIGHT CHANGES : FEMALE**

STUDY NO. : 0726  
 ANIMAL : RAT Cr-1:CD(SD) [Cr-1: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	216 ± 12	238 ± 14	254 ± 20	269 ± 22	282 ± 24
10mg/m3	217 ± 11	233 ± 14	245 ± 12	262 ± 18	274 ± 22
30mg/m3	217 ± 11	237 ± 13	253 ± 19	265 ± 21	276 ± 18
100mg/m3	217 ± 11	235 ± 16	250 ± 20	263 ± 24	280 ± 28

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

Test of Dunnett

(HAN260)

BATS 4

**TABLE E1**

**FOOD CONSUMPTION CHANGES : MALE**

STUDY NO. : 0726  
 ANIMAL : RAT CrI:CD(SD) [CrI: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	25.1± 2.6	25.0± 2.2	25.5± 2.1	26.5± 3.1
10mg/m3	25.1± 2.6	24.1± 2.4	26.3± 2.8	26.7± 2.6
30mg/m3	25.7± 2.5	24.9± 2.4	26.1± 2.1	27.3± 2.1
100mg/m3	26.1± 1.7	25.1± 2.1	26.2± 2.1	27.3± 2.5

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

Test of Dunnett

(HAN260)

**TABLE E2**

**FOOD CONSUMPTION CHANGES : FEMALE**



STUDY NO. : 0726  
 ANIMAL : RAT CrI: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	19.0± 1.9	18.8± 2.6	19.0± 3.2	18.9± 3.3
10mg/m3	18.8± 1.9	19.1± 1.8	19.0± 2.3	18.8± 2.0
30mg/m3	20.4± 1.7	20.5± 2.0	19.1± 2.6	18.6± 3.1
100mg/m3	19.8± 1.6	19.3± 2.2	19.0± 2.2	19.4± 3.0

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

Test of Dunnett

(HAN260)

**TABLE F1**

**HEMATOLOGY : MALE**

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : MALE  
REPORT TYPE : A1

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	8.00 ± 0.50	15.4 ± 1.1	41.7 ± 3.1	52.0 ± 1.6	19.2 ± 0.7	37.0 ± 0.7	1220 ± 77
10mg/m <sup>3</sup>	10	8.25 ± 0.28	15.7 ± 0.6	42.3 ± 1.8	51.2 ± 2.2	19.0 ± 0.8	37.1 ± 0.3	1235 ± 175
30mg/m <sup>3</sup>	10	8.21 ± 0.31	15.4 ± 0.7	41.4 ± 1.5	50.4 ± 0.8	18.8 ± 0.3	37.3 ± 0.5	1256 ± 114
100mg/m <sup>3</sup>	10	8.10 ± 0.38	15.5 ± 0.5	41.0 ± 1.4	50.7 ± 1.4	19.1 ± 0.7	37.7 ± 0.5**	1262 ± 135

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

Test of Dunnett

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr: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± 0.6
10mg/m3	10	2.2± 0.3
30mg/m3	10	2.3± 0.4
100mg/m3	10	2.3± 0.5

Test of Dunnett

\*\* : P ≤ 0.01

\* : P ≤ 0.05

Significant difference :

(HCL070)

BATS 4

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : MALE REPORT TYPE : AI

Group Name	NO. of Animals	WBC 10 <sup>9</sup> /μl	Differential WBC (%)					BASO	OTHER
			NEUTRO	LYMPHO	MONO	EOSINO			
Control	10	13.88± 3.72	14± 4	80± 4	3± 4	1± 1	0± 0	1± 0	
10mg/m3	10	13.38± 3.01	17± 5	77± 4	3± 4	1± 1	0± 0	1± 1	
30mg/m3	10	13.77± 3.58	18± 4	75± 4**	4± 4	1± 1	0± 0	1± 0	
100mg/m3	10	13.36± 4.33	17± 3	78± 3	3± 3	2± 2	0± 0	1± 0	

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

**TABLE F2**

**HEMATOLOGY : FEMALE**

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE, TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

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	7.84± 0.18	15.4± 0.4	40.6± 0.9	51.9± 0.8	19.7± 0.4	38.0± 0.4	1266± 117
10mg/m3	10	7.74± 0.41	15.0± 0.7	39.6± 2.0	51.2± 1.5	19.4± 0.5	37.9± 0.5	1269± 122
30mg/m3	10	7.80± 0.20	15.2± 0.4	39.2± 1.1	50.3± 1.0*	19.4± 0.4	38.6± 0.6*	1344± 109
100mg/m3	10	7.79± 0.31	15.3± 0.6	39.5± 1.5	50.7± 1.3	19.6± 0.6	38.7± 0.7*	1235± 110

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

Test of Dunnett

(HCL070)

HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	RETICULOCYTE %
Control	10	1.8± 0.3
10mg/m3	10	1.9± 0.5
30mg/m3	10	2.0± 0.3
100mg/m3	10	2.0± 0.5

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

Test of Dunnett

(HCL070)



HEMATOLOGY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Crj: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	9.09± 1.75	15± 4	79± 4	3± 1	2± 1	0± 0	1± 0
10mg/m3	10	8.87± 5.23	16± 11	77± 14	4± 2	2± 1	0± 0	1± 1
30mg/m3	10	8.70± 2.07	19± 7	75± 7	4± 1	2± 1	0± 0	1± 0
100mg/m3	10	6.48± 4.24	16± 4	78± 5	3± 1	2± 1	0± 0	1± 0

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

Test of Dunnett

(HCL070)

BATS 4

**TABLE G1**

**BIOCHEMISTRY : MALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Crj:CD(SD) [Crj:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : MALE 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	6.1±	3.2±	1.1±	0.12±	169±	56±	41±
10mg/m3	10	6.0±	3.3±	1.2±	0.12±	176±	54±	37±
30mg/m3	10	6.0±	3.3±	1.2±	0.12±	169±	50±	39±
100mg/m3	10	6.1±	3.3±	1.1±	0.11±	174±	59±	38±

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

Test of Dunnett

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Crj: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± 17	64± 6	25± 3	83± 30	506± 109	1± 0	132± 27
10mg/m3	10	91± 13	66± 8	27± 5	101± 51	495± 74	1± 0	136± 27
30mg/m3	10	89± 15	67± 6	29± 8	81± 42	556± 151	1± 0	133± 31
100mg/m3	10	97± 16	62± 5	25± 2	86± 40	452± 56	1± 0	133± 23

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

Test of Dunnett

(HCL074)

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr:1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE, TIME : 1  
SEX : MALE  
REPORT TYPE : A1

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	13.5±	0.5±	142±	3.7±	107±	10.0±	7.1±
10mg/m3	10	13.3±	0.5±	143±	3.7±	107±	9.9±	7.2±
30mg/m3	10	13.0±	0.5±	144±	3.6±	108±	10.0±	7.4±
100mg/m3	10	12.7±	0.5±	143±	3.7±	107±	10.0±	7.2±

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

Test of Dunnett

(HCL074)

BATS 4

**TABLE G2**

**BIOCHEMISTRY : FEMALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr1:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

PAGE : 5

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	6.6±	3.9±	1.4±	0.14±	157±	75±	21±
10mg/m3	10	6.8±	3.9±	1.4±	0.15±	154±	77±	18±
30mg/m3	10	6.4±	3.7±	1.4±	0.13±	147±	68±	14±
100mg/m3	10	6.8±	4.0±	1.4±	0.15±	159±	77±	15±

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

Test of Dunnett

(HCL074)

BAS 4

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD)IGS]  
MEASURE, TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

PAGE : 6

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	143±	57±	23±	76±	239±	1±	94±
10mg/m3	10	140±	58±	21±	94±	241±	1±	105±
30mg/m3	10	122±	62±	21±	76±	300±	1±	110±
100mg/m3	10	140±	58±	20±	66±	257±	1±	100±

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

Test of Dunnett

(HCL074)

BALS 4



BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
MEASURE. TIME : 1  
SEX : FEMALE  
REPORT TYPE : A1

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	14.5±	0.6±	142±	3.5±	108±	10.1±	5.7±
10mg/m3	10	14.1±	0.6±	141±	3.4±	108±	10.2±	5.9±
30mg/m3	10	16.3±	0.7±	141±	3.4±	107±	10.0±	6.8±
100mg/m3	10	14.2±	0.6±	142±	3.3±	107±	10.1±	6.0±

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

Test of Dunnett

**TABLE H1**

**BIOCHEMISTRY**

**(ACETYLCHOLINEESTERASE ACTIVITY)**

**: MALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Cr-I:CD(SD) [Cr-j:CD(SD) IGS]  
MEASURE. TIME : 1  
SEX : MALE  
REPORT TYPE : A1

PAGE : 4

Group Name	NO. of Animals	CHE(BRAIN) IU/ℓ	CHE(ERYTHROCYTE) IU/ℓ	CHE(PLASMA) IU/ℓ
Control	10	2266 ± 126	2314 ± 189	501 ± 74
10mg/m3	10	2194 ± 125	2301 ± 285	413 ± 67*
30mg/m3	10	2121 ± 115*	1969 ± 275*	348 ± 41**
100mg/m3	10	1926 ± 72**	1736 ± 370**	376 ± 70**

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

(HCL074)

BATS 4

**TABLE H2**

**BIOCHEMISTRY**

**(ACETYLCHOLINEESTERASE ACTIVITY)**

**: FEMALE**

BIOCHEMISTRY (SUMMARY)  
ALL ANIMALS ( 5W)

STUDY NO. : 0726  
ANIMAL : RAT Crj: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	2227± 84	2626± 459	2399± 510
10mg/m3	10	2166± 86	2602± 412	2413± 887
30mg/m3	10	2066± 103**	2472± 455	1680± 439*
100mg/m3	10	1806± 78**	2145± 384	1394± 286**

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+CD8-)	Cytotoxic-T cell (CD4-CD8+)		
0	10	Mean	2.43	1.29	1.92	
		S.D.	0.66	0.38	0.29	
10	10	Mean	2.37	1.39	1.76	
		S.D.	0.72	0.36	0.64	
30	10	Mean	2.40	1.39	1.85	
		S.D.	0.54	0.48	0.46	
100	10	Mean	2.27	1.19	1.97	
		S.D.	0.89	0.51	0.49	

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.

## 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	1.6	75.6	7.7	4.1	
		S.D.	0.7	11.2	1.8	0.7	
10	5	Mean	1.4	91.5	9.3	6.0	
		S.D.	0.7	30.2	1.8	1.8	
30	5	Mean	1.4	76.4	7.7	5.3	
		S.D.	0.5	18.0	2.8	1.4	
100	5	Mean	1.6	107.1	9.4	5.7	
		S.D.	0.6	31.7	1.3	1.1	

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

S.D.: Standard deviation.

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



## 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	24.8	9.0	5.4
		S.D.	11.0	4.6	2.6
10	5	Mean	31.9	10.7	7.2
		S.D.	8.1	3.4	2.3
30	5	Mean	23.0	7.4	5.3
		S.D.	5.2	1.9	1.2
100	5	Mean	24.0	7.6	5.5
		S.D.	3.2	1.9	0.7

S.D.: Standard deviation.

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

**TABLE I2**

**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+CD8-)	Cytotoxic-T cell (CD4-CD8+)		
0	10	Mean	2.01	0.91	2.33	
		S.D.	0.34	0.28	0.61	
10	10	Mean	1.87	0.74	2.56	
		S.D.	0.79	0.25	0.63	
30	10	Mean	2.01	1.04	2.00	
		S.D.	0.56	0.36	0.33	
100	10	Mean	1.34	0.71	2.08	
		S.D.	0.69	0.48	0.51	

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.

## 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	1.0	58.5	6.1	4.0	
		S.D.	0.5	13.0	2.0	0.9	
10	5	Mean	1.7	94.2	8.8	6.4 *	
		S.D.	0.8	51.0	5.4	1.5	
30	5	Mean	1.3	67.8	7.8	5.0	
		S.D.	0.6	22.6	1.4	1.0	
100	5	Mean	1.1	57.5	6.5	4.3	
		S.D.	0.4	18.8	1.3	0.9	

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

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 ≤ 0.05; \*\*, p ≤ 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	16.6	5.3	3.3
		S.D.	2.8	1.0	0.9
10	5	Mean	15.7	4.9	2.9
		S.D.	2.8	0.6	0.9
30	5	Mean	13.6	3.9	3.0
		S.D.	3.0	0.9	1.2
100	5	Mean	18.0	5.1	4.1
		S.D.	4.3	2.0	1.2

S.D.: Standard deviation.

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

**TABLE J**

**GROSS FINDINGS : FEMALE**

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

STUDY NO. : 0726  
ANIMAL : RAT C-1:CD(SD) [Crj:CD(SD)IGS]  
REPORT TYPE : AI  
SEX : FEMALE

Organ	Findings	Group Name NO. of Animals	Control 10 (%)	10mg/m3 10 (%)	30mg/m3 10 (%)	100mg/m3 10 (%)
spleen	enlarged		0 ( 0)	1 ( 10)	0 ( 0)	0 ( 0)

(HPT080)

**TABLE K1**

**ORGAN WEIGHT, ABSOLUTE : MALE**



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

STUDY NO. : 0726  
ANIMAL : RAT Cr1:CD(SD) [Cr-j:CD(SD)IGS]  
REPORT TYPE : A1  
SEX : MALE  
UNIT : g

Group Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	10	409 ± 27	0.522 ± 0.068	0.072 ± 0.010	2.979 ± 0.292	1.415 ± 0.096	1.356 ± 0.125
10mg/m3	10	396 ± 35	0.555 ± 0.107	0.071 ± 0.010	3.158 ± 0.199	1.353 ± 0.096	1.324 ± 0.121
30mg/m3	10	398 ± 29	0.472 ± 0.085	0.071 ± 0.007	3.117 ± 0.173	1.420 ± 0.130	1.360 ± 0.097
100mg/m3	10	397 ± 26	0.551 ± 0.074	0.077 ± 0.012	3.013 ± 0.176	1.311 ± 0.106	1.334 ± 0.137

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

Test of Dunnett

(HCL040)

STUDY NO. : 0726  
 ANIMAL : RAT Cr1:CD(SD) [Crj:CD(SD) IGS]  
 REPORT TYPE : AI  
 SEX : MALE  
 UNIT : g

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

Group Name	NO. of Animals	KIDNEYS	SPLERN	LIVER	BRAIN
Control	10	2.898 ± 0.296	0.768 ± 0.150	11.445 ± 1.644	1.983 ± 0.097
10mg/m3	10	2.739 ± 0.300	0.739 ± 0.138	11.142 ± 1.226	1.945 ± 0.075
30mg/m3	10	2.811 ± 0.196	0.721 ± 0.128	11.124 ± 1.305	1.977 ± 0.072
100mg/m3	10	2.826 ± 0.354	0.787 ± 0.068	11.861 ± 0.989	1.963 ± 0.078

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

Test of Dunnett

(HCL040)

**TABLE K2**

**ORGAN WEIGHT, ABSOLUTE : FEMALE**

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

STUDY NO. : 0726  
ANIMAL : RAT Cx1:CD (SD) [Cx-j:CD (SD) IGS]  
REPORT TYPE : AI  
SEX : FEMALE  
UNIT: g

Group Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	10	260 ± 21	0.425 ± 0.105	0.080 ± 0.007	0.134 ± 0.015	0.902 ± 0.088	1.062 ± 0.064
10mg/m3	10	251 ± 19	0.426 ± 0.099	0.080 ± 0.010	0.138 ± 0.016	0.895 ± 0.087	1.071 ± 0.077
30mg/m3	10	257 ± 18	0.434 ± 0.105	0.088 ± 0.011	0.145 ± 0.028	0.893 ± 0.082	1.090 ± 0.049
100mg/m3	10	256 ± 24	0.424 ± 0.108	0.080 ± 0.010	0.140 ± 0.011	0.922 ± 0.080	1.065 ± 0.060

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

Test of Dunnett

(HCL040)

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

STUDY NO. : 0726  
ANIMAL : RAT C-1:CD(SD) [C-j:CD(SD)IGS]  
REPORT TYPE : AI  
SEX : FEMALE  
UNIT: g

Group Name	NO. of Animals	KIDNEYS	SPLLEN	LIVER	BRAIN
Control	10	1.767± 0.147	0.477± 0.027	7.206± 0.964	1.865± 0.099
10mg/m3	10	1.782± 0.144	0.533± 0.232	7.320± 1.008	1.881± 0.069
30mg/m3	10	1.819± 0.153	0.538± 0.086	7.337± 0.735	1.878± 0.064
100mg/m3	10	1.760± 0.135	0.497± 0.062	7.508± 0.724	1.881± 0.066

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

Test of Dunnett

(HCL040)

**TABLE L1**

**ORGAN WEIGHT, RELATIVE : MALE**

STUDY NO. : 0726  
 ANIMAL : RAT Cr1:CD (SD) [Crj:CD (SD) IGS]  
 REPORT TYPE : A1  
 SEX : MALE  
 UNIT : %

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

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	10	409 ± 27	0.128 ± 0.016	0.017 ± 0.002	0.731 ± 0.076	0.347 ± 0.010	0.332 ± 0.018
10mg/m3	10	396 ± 35	0.140 ± 0.021	0.018 ± 0.002	0.808 ± 0.090	0.342 ± 0.015	0.335 ± 0.024
30mg/m3	10	398 ± 29	0.118 ± 0.018	0.018 ± 0.002	0.786 ± 0.070	0.357 ± 0.032	0.342 ± 0.017
100mg/m3	10	397 ± 26	0.139 ± 0.017	0.019 ± 0.002	0.760 ± 0.044	0.331 ± 0.028	0.336 ± 0.022

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

Test of Dunnett

(HCL042)

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

STUDY NO. : 0726  
ANIMAL : RAT C-1:CD(SD) [C-j:CD(SD) IGS]  
REPORT TYPE : AI  
SEX : MALE  
UNIT : %

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	0.710 ± 0.065	0.188 ± 0.037	2.790 ± 0.262	0.486 ± 0.029
10mg/m3	10	0.694 ± 0.071	0.186 ± 0.029	2.815 ± 0.217	0.494 ± 0.042
30mg/m3	10	0.709 ± 0.069	0.182 ± 0.032	2.788 ± 0.188	0.499 ± 0.035
100mg/m3	10	0.710 ± 0.058	0.199 ± 0.013	2.984 ± 0.131	0.496 ± 0.038

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

Test of Dunnett

(HCL042)



**TABLE L2**

**ORGAN WEIGHT, RELATIVE : FEMALE**

STUDY NO. : 0726  
 ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD)IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE  
 UNIT : %

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

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	10	260± 21	0.164± 0.040	0.031± 0.003	0.052± 0.007	0.348± 0.031	0.411± 0.037
10mg/m3	10	251± 19	0.171± 0.045	0.032± 0.004	0.055± 0.008	0.356± 0.024	0.427± 0.032
30mg/m3	10	257± 18	0.169± 0.037	0.034± 0.005	0.057± 0.012	0.348± 0.032	0.426± 0.044
100mg/m3	10	256± 24	0.164± 0.030	0.031± 0.005	0.055± 0.004	0.361± 0.026	0.417± 0.027

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

Test of Dunnett

STUDY NO. : 0726  
 ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD)IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE  
 UNIT : %

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

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	10	0.681 ± 0.048	0.185 ± 0.015	2.766 ± 0.190	0.720 ± 0.050
10mg/m3	10	0.711 ± 0.063	0.213 ± 0.095	2.908 ± 0.303	0.752 ± 0.054
30mg/m3	10	0.710 ± 0.063	0.211 ± 0.040	2.854 ± 0.187	0.735 ± 0.072
100mg/m3	10	0.690 ± 0.058	0.196 ± 0.034	2.933 ± 0.125	0.740 ± 0.075

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

(HCL042)

**TABLE M1**

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

STUDY NO. : 0726  
 ANIMAL : RAT Cr-I:CD(SD) [Cr-J:CV(SD)IGS]  
 REPORT TYPE : AI  
 SEX : MALE

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

Organ	Findings	Control				10mg/m3				30mg/m3				100mg/m3							
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4				
Grade	No. of Animals on Study	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)				
{Urinary system}																					
kidney	eosinophilic body	1	0	0	0	<10>				3	0	0	0	<10>				1	0	0	0
		(10)	(0)	(0)	(0)	(30)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(10)	(0)	(0)	(0)
	regeneration:proximal tubule	2	0	0	0	3	0	0	0	1	0	0	0	1	0	0	0	2	0	0	0
		(20)	(0)	(0)	(0)	(30)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(10)	(0)	(0)	(0)	(20)	(0)	(0)	(0)

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

**TABLE M2**

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

STUDY NO. : 0726  
 ANIMAL : RAT Cr-1:CD(SD) [Cr-j:CD(SD) IGS]  
 REPORT TYPE : AI  
 SEX : FEMALE

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

Organ	Findings	Control				10mg/m <sup>3</sup>				30mg/m <sup>3</sup>				100mg/m <sup>3</sup>			
		1 (%)	2 (%)	3 (%)	4 (%)	1 (%)	2 (%)	3 (%)	4 (%)	1 (%)	2 (%)	3 (%)	4 (%)	1 (%)	2 (%)	3 (%)	4 (%)
<b>(Hematopoietic system)</b>																	
spleen	necrosis:focal	0 (0)	0 (0)	0 (0)	0 (0)	<10>	1 (10)	0 (0)	0 (0)	<10>	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
<b>(Digestive system)</b>																	
liver	necrosis:focal	0 (0)	0 (0)	0 (0)	0 (0)	<10>	1 (10)	0 (0)	0 (0)	<10>	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
<b>(Urinary system)</b>																	
kidney	regeneration:proximal tubule	1 (10)	0 (0)	0 (0)	0 (0)	<10>	2 (20)	0 (0)	0 (0)	<10>	0 (0)	0 (0)	0 (0)	1 (10)	0 (0)	0 (0)	0 (0)
<b>(Endocrine system)</b>																	
thyroid	ultimobranchial body remanet	2 (20)	0 (0)	0 (0)	0 (0)	<10>	1 (10)	0 (0)	0 (0)	<10>	0 (0)	0 (0)	0 (0)	1 (10)	0 (0)	0 (0)	0 (0)

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

STUDY NO. : 0726  
 ANIMAL : RAT Cf-1:CD(SD) [C+j:CD(SD) IGS]  
 REPORT TYPE : A1  
 SEX : FEMALE

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

Organ	Findings	Group Name				10mg/m3				30mg/m3				100mg/m3							
		No. of Animals on Study				Control				10				10				10			
		1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
eye	keratitis	0	0	0	0	<10>				0	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 )

(Special sense organs/appendage)

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**

**DEP AND DDVP CONCENTRATIONS  
IN PLASMA OF RATS**

## DEP and DDVP\* concentrations in plasma of rats

Group Name	Sex	Animal Number	DEP (ng/mL)	DDVP (ng/mL)
Control	Male	1001	- 1)	- 2)
		1002	-	-
		1003	-	-
		1004	-	-
		1005	-	-
	Female	2001	-	-
		2002	-	-
		2003	-	-
		2004	-	-
		2005	-	-
10 mg/m <sup>3</sup>	Male	1101	-	-
		1102	-	-
		1103	-	-
		1104	-	-
		1105	-	-
	Female	2101	6.71	-
		2102	-	-
		2103	-	-
		2104	-	-
		2105	-	-
30 mg/m <sup>3</sup>	Male	1201	13.33	-
		1202	-	-
		1203	5.08	-
		1204	-	-
		1205	-	-
	Female	2201	12.43	-
		2202	5.69	-
		2203	6.02	-
		2204	12.22	-
		2205	-	-
100 mg/m <sup>3</sup>	Male	1301	27.81	-
		1302	9.78	-
		1303	14.29	-
		1304	5.77	-
		1305	9.45	-
	Female	2301	43.79	-
		2302	48.13	-
		2303	24.45	-
		2304	21.18	-
		2305	21.06	-

1) : Below the limit of quantification. Limit of quantification of DEP was 5ng/mL.

2) : Below the limit of quantification. Limit of quantification of DDVP was 2ng/mL.

\* : 0,0-Dimethyl-(1-hydroxy-2,2,2-trichloroethyl)-phosphonate and 2,2-dichlorovinyl dimethyl phosphate were abbreviated as DEP and DDVP, respectively.