クロロエタンの p53 KO マウスを用いた
吸入による中期発がん性試験報告書

試験番号：0922

TABLES

TABLES

TABLES (CONTINUED)

TABLES (CONTINUED)

TABLE A

CONCENTRATIONS OF CHLOROETHANE IN THE INHALATION CHAMBER OF p53 KO MICE IN THE 26-WEEK CARCINOGENICITY STUDY

TABLE A CONCENTRATIONS OF CHLOROETHANE IN THE INHALATION CHAMBER OF p53 KO MICE IN THE 26-WEEK CARCINOGENICITY

Group Name	Concentration(ppm) Mean ± S.D.
Control	0.0 ± 0.0
2400 ppm	2401.0 ± 8.2
6000 ppm	5990.1 ± 35.7
15000 ppm	15002.4 ± 50.6

TABLE B1

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0922 SURVIVAL ANIMAL NUMBERS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1 26
SEX : MALE

Group Name	Animals At start	Administration (Weeks) 0	1	2	3	4	5	6	7	8	9	10	11	12	13
Control	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
2400 ppm	25	25/25	25/25	25/25	25/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25
		100.0	100.0	100.0	100.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0
6000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
15000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0

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

STUDY NO. : 0922 SURVIVAL ANIMAL NUMBERS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1 26
SEX : MALE

Group Name	Animals At start	Administration (Weeks) 14	15	16	17	18	19	20	21	22	23	24	25	26
Control	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	24/25	24/25	24/25	24/25	24/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	96.0	96.0	96.0	96.0	96.0
2400 ppm	25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25
		96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0
6000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
15000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	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

STUDY NO. : 0922 SURVIVAL ANIMAL NUMBERS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1 26
SEX : FEMALE

Group Name	Animals At start	Administration (Weeks) 0	1	2	3	4	5	6	7	8	9	10	11	12	13
Control	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
2400 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
6000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0
15000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0

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

Group Name	Animals At start	Administration (Weeks) 14	15	16	17	18	19	20	21	22	23	24	25	26
Control	25	25/25	25/25	24/25	24/25	24/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25
		100.0	100.0	96.0	96.0	96.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0
2400 ppm	25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25
		96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0
6000 ppm	25	25/25	25/25	25/25	25/25	25/25	24/25	24/25	24/25	23/25	23/25	23/25	23/25	23/25
		100.0	100.0	100.0	100.0	100.0	96.0	96.0	96.0	92.0	92.0	92.0	92.0	92.0
15000 ppm	25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25	25/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	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. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1 26

CLINICAL OBSERVATION (SUMMARY)
ALL ANIMALS

SEX : MALE

Clinical sign	Group Name	Administration Week-day 1-7	2-7	3-7	4-7	5-7	6-7	7-7	8-7	9-7	10-7	11-7	12-7	13-7	14-7
		1	1	1	1	1	1	1	1	1	1	1	1	1	1
DEATH	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	1	1	1	1	1	1	1	1	1	1	1
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ROUGH FUR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	1	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
MALOCCLUSION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	1	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EROSION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
IRREGULAR BREATHING	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
NON REMARKABLE	Control	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	2400 ppm	25	24	24	24	24	24	24	24	24	24	24	24	24	24
	6000 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	15000 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25

Clinical sign	Group Name	Administration Week-day 15-7 1	16-7 1	17-7 1	18-7 1	19-7 1	20-7 1	21-7 1	22-7 1	23-7 1	24-7 1	25-7 1	26-7 1
DEATH	Control	0	0	0	0	0	0	0	1	1	1	1	1
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	1	1	1	1	1	1	1	1	1	1	1	1
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
ROUGH FUR	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
MALOCCLUSION	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
EROSION	Control	1	1	1	1	1	1	1	2	2	2	2	2
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
IRREGULAR BREATHING	Control	0	0	1	1	1	1	1	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
NON REMARKABLE	Control	24	24	23	23	23	23	23	22	22	22	22	22
	2400 ppm	24	24	24	24	24	24	24	24	24	24	24	24
	6000 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	15000 ppm	25	25	25	25	25	25	25	25	25	25	25	25

TABLE C2

CLINICAL OBSERVATION : FEMALE

Clinical sign	Group Name	Administration Week-day 1-7 1	2-7 1	3-7 1	4-7 1	5-7 1	6-7 1	7-7 1	8-7 1	9-7 1	10-7 1	11-7 1	12-7 1	13-7 1	14-7 1
DEATH	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
PARALYTIC GAIT	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EXTERNAL MASS	Control	0	0	0	0	0	0	0	0	0	0	0	0	1	1
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M.HINDLIMB	Control	0	0	0	0	0	0	0	0	0	0	0	0	1	1
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M.GENITALIA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANEMIA	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EROSION	Control	0	0	0	0	0	0	0	0	0	1	1	1	1	1
	2400 ppm	0	0	0	0	0	0	0	1	1	1	1	1	1	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	1	1
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
IRREGULAR BREATHING	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Clinical sign	Group Name	Administration Week-day 15-7	16-7	17-7	18-7	19-7	20-7	21-7	22-7	23-7	24-7	25-7	26-7
		1	1	1	1	1	1	1	1	1	1	1	1
DEATH	Control	0	1	1	1	2	2	2	2	2	2	2	2
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	1	1	1	1	1	1	1	1	1	1	1	1
	6000 ppm	0	0	0	0	1	1	1	2	2	2	2	2
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
PARALYTIC GAIT	Control	1	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
EXTERNAL MASS	Control	1	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	1	1	1	1	1	1	1	1	1	1	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
M.HINDLIMB	Control	1	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
M.GENITALIA	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	1	1	1	1	1	1	1	1	1	1	0
	6000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
ANEMIA	Control	0	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	1	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
EROSION	Control	1	1	1	1	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	1	1	1	1	0	0	0	0	1	1	1	1
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	1
IRREGULAR BREATHING	Control	1	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	6000 ppm	0	0	0	1	0	0	1	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0

Clinical sign	Group Name	Administration Week-day 1-7	2-7	3-7	4-7	5-7	6-7	7-7	8-7	9-7	10-7	11-7	12-7	13-7	14-7
		1	1	1	1	1	1	1	1	1	1	1	1	1	1
NON REMARKABLE	Control	25	25	25	25	25	25	25	25	25	24	24	24	23	23
	2400 ppm	25	25	25	25	25	25	25	24	24	24	24	24	24	24
	6000 ppm	25	25	25	25	25	25	25	25	25	25	25	25	24	24
	15000 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25

Clinical sign	Group Name	Administration Week-day 15-7	16-7	17-7	18-7	19-7	20-7	21-7	22-7	23-7	24-7	25-7	26-7
		1	1	1	1	1	1	1	1	1	1	1	1
NON REMARKABLE	Control	23	23	23	23	23	23	23	23	23	23	23	23
	2400 ppm	24	23	23	23	23	23	23	23	23	23	23	24
	6000 ppm	24	24	24	24	24	24	23	23	22	22	22	22
	15000 ppm	25	25	25	25	25	25	25	25	25	25	25	24

TABLE D1

BODY WEIGHT CHANGES AND

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0922 MEAN BODY WEIGHTS AND SURVIVAL
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

Week-Day on Study	Control Av.Wt.	Control No.of Surviv. <25>	2400 ppm Av.Wt.	2400 ppm % of cont. <25>	2400 ppm No.of Surviv.	6000 ppm Av.Wt.	6000 ppm % of cont. <25>	6000 ppm No.of Surviv.	15000 ppm Av.Wt.	15000 ppm % of cont. <25>	15000 ppm No.of Surviv.
0-0	23.9 (25)	25/25	24.0 (25)	100	25/25	23.9 (25)	100	25/25	24.0 (25)	100	25/25
1-7	25.4 (25)	25/25	25.3 (25)	100	25/25	25.5 (25)	100	25/25	25.5 (25)	100	25/25
2-7	26.8 (25)	25/25	25.9 (25)	97	25/25	26.4 (25)	99	25/25	26.1 (25)	97	25/25
3-7	27.5 (25)	25/25	26.7 (25)	97	25/25	27.3 (25)	99	25/25	27.1 (25)	99	25/25
4-7	28.6 (25)	25/25	28.2 (24)	99	24/25	28.2 (25)	99	25/25	27.6 (25)	97	25/25
5-7	29.3 (25)	25/25	29.1 (24)	99	24/25	29.0 (25)	99	25/25	28.1 (25)	96	25/25
6-7	29.6 (25)	25/25	29.4 (24)	99	24/25	28.9 (25)	98	25/25	28.2 (25)	95	25/25
7-7	30.4 (25)	25/25	30.3 (24)	100	24/25	29.8 (25)	98	25/25	28.8 (25)	95	25/25
8-7	30.9 (25)	25/25	30.9 (24)	100	24/25	30.5 (25)	99	25/25	29.6 (25)	96	25/25
9-7	31.4 (25)	25/25	31.2 (24)	99	24/25	30.7 (25)	98	25/25	29.5 (25)	94	25/25
10-7	31.5 (25)	25/25	31.4 (24)	100	24/25	31.0 (25)	98	25/25	29.9 (25)	95	25/25
11-7	31.9 (25)	25/25	31.9 (24)	100	24/25	31.1 (25)	97	25/25	29.9 (25)	94	25/25
12-7	32.5 (25)	25/25	32.3 (24)	99	24/25	31.7 (25)	98	25/25	30.5 (25)	94	25/25
13-7	33.0 (25)	25/25	33.2 (24)	101	24/25	31.8 (25)	96	25/25	31.0 (25)	94	25/25
14-7	33.5 (25)	25/25	33.7 (24)	101	24/25	33.0 (25)	99	25/25	31.5 (25)	94	25/25
15-7	33.6 (25)	25/25	34.0 (24)	101	24/25	33.2 (25)	99	25/25	31.5 (25)	94	25/25
16-7	33.9 (25)	25/25	34.3 (24)	101	24/25	33.3 (25)	98	25/25	31.8 (25)	94	25/25
17-7	34.3 (25)	25/25	34.6 (24)	101	24/25	33.1 (25)	97	25/25	31.4 (25)	92	25/25
18-7	34.7 (25)	25/25	35.4 (24)	102	24/25	34.0 (25)	98	25/25	32.1 (25)	93	25/25
19-7	35.1 (25)	25/25	35.9 (24)	102	24/25	34.6 (25)	99	25/25	32.7 (25)	93	25/25
20-7	35.7 (25)	25/25	36.1 (24)	101	24/25	34.9 (25)	98	25/25	33.0 (25)	92	25/25
21-7	35.7 (25)	25/25	36.8 (24)	103	24/25	34.9 (25)	98	25/25	33.1 (25)	93	25/25
22-7	36.6 (24)	24/25	36.9 (24)	101	24/25	35.6 (25)	97	25/25	33.4 (25)	91	25/25
23-7	36.5 (24)	24/25	36.9 (24)	101	24/25	35.5 (25)	97	25/25	33.2 (25)	91	25/25
24-7	36.8 (24)	24/25	37.2 (24)	101	24/25	35.8 (25)	97	25/25	34.0 (25)	92	25/25
25-7	37.2 (24)	24/25	37.4 (24)	101	24/25	36.4 (25)	98	25/25	34.4 (25)	92	25/25
26-7	38.0 (24)	24/25	37.7 (24)	99	24/25	36.8 (25)	97	25/25	34.9 (25)	92	25/25

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

TABLE D2

BODY WEIGHT CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0922 MEAN BODY WEIGHTS AND SURVIVAL
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

Week-Day on Study	Control Av.Wt.	Control No.of Surviv. <25>	2400 ppm Av.Wt.	2400 ppm % of cont. <25>	2400 ppm No.of Surviv.	6000 ppm Av.Wt.	6000 ppm % of cont. <25>	6000 ppm No.of Surviv.	15000 ppm Av.Wt.	15000 ppm % of cont. <25>	15000 ppm No.of Surviv.
0-0	20.0 (25)	25/25	20.0 (25)	100	25/25	20.0 (25)	100	25/25	20.0 (25)	100	25/25
1-7	21.4 (25)	25/25	21.4 (25)	100	25/25	21.7 (25)	101	25/25	21.6 (25)	101	25/25
2-7	21.8 (25)	25/25	21.8 (25)	100	25/25	22.1 (25)	101	25/25	21.4 (25)	98	25/25
3-7	21.9 (25)	25/25	22.1 (25)	101	25/25	22.0 (25)	100	25/25	21.6 (25)	99	25/25
4-7	22.9 (25)	25/25	22.7 (25)	99	25/25	22.4 (25)	98	25/25	22.2 (25)	97	25/25
5-7	23.2 (25)	25/25	23.3 (25)	100	25/25	22.9 (25)	99	25/25	22.5 (25)	97	25/25
6-7	23.2 (25)	25/25	23.2 (25)	100	25/25	22.9 (25)	99	25/25	22.6 (25)	97	25/25
7-7	24.1 (25)	25/25	23.8 (25)	99	25/25	23.8 (25)	99	25/25	23.0 (25)	95	25/25
8-7	24.5 (25)	25/25	23.9 (25)	98	25/25	24.0 (25)	98	25/25	23.3 (25)	95	25/25
9-7	24.5 (25)	25/25	24.5 (25)	100	25/25	23.9 (25)	98	25/25	23.2 (25)	95	25/25
10-7	24.3 (25)	25/25	24.3 (25)	100	25/25	23.9 (25)	98	25/25	23.5 (25)	97	25/25
11-7	24.6 (25)	25/25	24.3 (25)	99	25/25	23.7 (25)	96	25/25	22.9 (25)	93	25/25
12-7	24.6 (25)	25/25	24.3 (25)	99	25/25	23.9 (25)	97	25/25	23.4 (25)	95	25/25
13-7	25.1 (25)	25/25	24.6 (25)	98	25/25	24.2 (25)	96	25/25	24.2 (25)	96	25/25
14-7	25.6 (25)	25/25	25.0 (24)	98	24/25	24.9 (25)	97	25/25	24.1 (25)	94	25/25
15-7	25.4 (25)	25/25	25.2 (24)	99	24/25	25.0 (25)	98	25/25	24.1 (25)	95	25/25
16-7	25.9 (24)	24/25	25.3 (24)	98	24/25	24.7 (25)	95	25/25	24.2 (25)	93	25/25
17-7	25.6 (24)	24/25	25.5 (24)	100	24/25	24.7 (25)	96	25/25	23.9 (25)	93	25/25
18-7	26.3 (24)	24/25	25.4 (24)	97	24/25	24.8 (25)	94	25/25	24.2 (25)	92	25/25
19-7	25.9 (23)	23/25	26.2 (24)	101	24/25	25.5 (24)	98	24/25	24.7 (25)	95	25/25
20-7	26.1 (23)	23/25	26.0 (24)	100	24/25	25.4 (24)	97	24/25	24.5 (25)	94	25/25
21-7	26.2 (23)	23/25	26.3 (24)	100	24/25	25.3 (24)	97	24/25	24.9 (25)	95	25/25
22-7	26.3 (23)	23/25	26.2 (24)	100	24/25	25.8 (23)	98	23/25	24.9 (25)	95	25/25
23-7	26.3 (23)	23/25	26.4 (24)	100	24/25	25.3 (23)	96	23/25	24.6 (25)	94	25/25
24-7	26.7 (23)	23/25	26.6 (24)	100	24/25	26.1 (23)	98	23/25	25.2 (25)	94	25/25
25-7	26.9 (23)	23/25	26.7 (24)	99	24/25	26.0 (23)	97	23/25	25.2 (25)	94	25/25
26-7	27.0 (23)	23/25	26.8 (24)	99	24/25	26.2 (23)	97	23/25	25.6 (25)	95	25/25

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

TABLE D3

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 0-0	1-7	2-7	3-7	4-7	5-7	6-7
Control	23.9± 1.4	25.4± 1.7	26.8± 1.5	27.5± 1.5	28.6± 1.7	29.3± 1.9	29.6± 1.9
2400 ppm	24.0± 1.4	25.3± 1.1	25.9± 2.6	26.7± 2.6	28.2± 1.5	29.1± 1.5	29.4± 1.8
6000 ppm	23.9± 1.4	25.5± 1.5	26.4± 1.3	27.3± 1.4	28.2± 1.4	29.0± 1.7	28.9± 1.6
15000 ppm	24.0± 1.4	25.5± 1.3	26.1± 1.1	27.1± 1.2	27.6± 1.2	28.1± 1.3*	28.2± 1.2**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 7-7	8-7	9-7	10-7	11-7	12-7	13-7
Control	30.4± 1.9	30.9± 1.9	31.4± 2.0	31.5± 2.1	31.9± 2.3	32.5± 2.5	33.0± 2.8
2400 ppm	30.3± 1.8	30.9± 2.1	31.2± 2.2	31.4± 2.2	31.9± 2.2	32.3± 2.4	33.2± 2.5
6000 ppm	29.8± 1.8	30.5± 1.8	30.7± 2.0	31.0± 2.0	31.1± 2.1	31.7± 2.0	31.8± 2.1
15000 ppm	28.8± 1.3**	29.6± 1.6*	29.5± 1.5**	29.9± 1.3**	29.9± 1.3**	30.5± 1.5**	31.0± 1.5*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 14-7	15-7	16-7	17-7	18-7	19-7	20-7
Control	33.5± 3.0	33.6± 3.2	33.9± 3.4	34.3± 3.8	34.7± 3.8	35.1± 3.9	35.7± 4.3
2400 ppm	33.7± 3.0	34.0± 2.9	34.3± 3.1	34.6± 3.4	35.4± 3.8	35.9± 3.8	36.1± 3.9
6000 ppm	33.0± 2.1	33.2± 2.2	33.3± 2.4	33.1± 2.5	34.0± 2.5	34.6± 2.5	34.9± 2.7
15000 ppm	31.5± 1.5*	31.5± 1.4*	31.8± 1.6*	31.4± 1.5**	32.1± 1.6**	32.7± 1.9**	33.0± 2.2*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 21-7	22-7	23-7	24-7	25-7	26-7
Control	35.7± 4.5	36.6± 3.7	36.5± 3.8	36.8± 4.0	37.2± 3.8	38.0± 4.0
2400 ppm	36.8± 3.6	36.9± 3.8	36.9± 3.9	37.2± 4.2	37.4± 4.1	37.7± 4.3
6000 ppm	34.9± 2.6	35.6± 2.6	35.5± 2.7	35.8± 2.9	36.4± 2.7	36.8± 2.9
15000 ppm	33.1± 1.9**	33.4± 2.2**	33.2± 2.1**	34.0± 2.1*	34.4± 2.2*	34.9± 2.3**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE D4

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 0-0	1-7	2-7	3-7	4-7	5-7	6-7
Control	20.0± 1.6	21.4± 1.7	21.8± 1.3	21.9± 1.2	22.9± 1.2	23.2± 1.4	23.2± 1.4
2400 ppm	20.0± 1.6	21.4± 1.7	21.8± 1.7	22.1± 1.8	22.7± 1.8	23.3± 1.8	23.2± 1.8
6000 ppm	20.0± 1.6	21.7± 1.6	22.1± 1.6	22.0± 1.4	22.4± 1.6	22.9± 1.7	22.9± 1.6
15000 ppm	20.0± 1.6	21.6± 1.7	21.4± 1.6	21.6± 1.6	22.2± 1.6	22.5± 1.7	22.6± 1.5

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 7-7	8-7	9-7	10-7	11-7	12-7	13-7
Control	24.1± 1.4	24.5± 1.2	24.5± 1.3	24.3± 1.5	24.6± 1.4	24.6± 1.9	25.1± 2.1
2400 ppm	23.8± 1.7	23.9± 1.8	24.5± 1.6	24.3± 1.4	24.3± 1.7	24.3± 1.7	24.6± 1.4
6000 ppm	23.8± 1.5	24.0± 1.4	23.9± 1.4	23.9± 1.4	23.7± 1.5	23.9± 1.4	24.2± 1.4
15000 ppm	23.0± 1.9	23.3± 1.7	23.2± 1.6**	23.5± 1.7	22.9± 1.6**	23.4± 1.8	24.2± 1.5

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 14-7	15-7	16-7	17-7	18-7	19-7	20-7
Control	25.6± 2.1	25.4± 2.2	25.9± 2.3	25.6± 2.2	26.3± 2.1	25.9± 1.6	26.1± 1.5
2400 ppm	25.0± 1.5	25.2± 1.3	25.3± 1.7	25.5± 1.7	25.4± 1.7	26.2± 1.5	26.0± 1.5
6000 ppm	24.9± 1.6	25.0± 1.5	24.7± 1.5	24.7± 1.5	24.8± 1.8*	25.5± 1.9	25.4± 1.5
15000 ppm	24.1± 1.4**	24.1± 1.7*	24.2± 1.6**	23.9± 1.7**	24.2± 1.5**	24.7± 1.6*	24.5± 1.5**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day 21-7	22-7	23-7	24-7	25-7	26-7
Control	26.2± 1.5	26.3± 1.8	26.3± 1.4	26.7± 1.8	26.9± 1.5	27.0± 1.7
2400 ppm	26.3± 2.0	26.2± 1.4	26.4± 1.8	26.6± 1.6	26.7± 1.7	26.8± 1.5
6000 ppm	25.3± 1.7	25.8± 1.9	25.3± 1.5	26.1± 1.9	26.0± 2.1	26.2± 1.8
15000 ppm	24.9± 1.5*	24.9± 1.8*	24.6± 1.8**	25.2± 1.6**	25.2± 1.6**	25.6± 1.7*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE E1

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

MEAN FOOD CONSUMPTION(FC) AND SURVIVAL

Week-Day on Study	Control Av.FC.	Control No.of Surviv. <25>	2400 ppm Av.FC.	2400 ppm % of cont. <25>	2400 ppm No.of Surviv.	6000 ppm Av.FC.	6000 ppm % of cont. <25>	6000 ppm No.of Surviv.	15000 ppm Av.FC.	15000 ppm % of cont. <25>	15000 ppm No.of Surviv.
1-7	3.9 (25)	25/25	3.9 (25)	100	25/25	4.1 (25)	105	25/25	4.1 (25)	105	25/25
2-7	3.8 (25)	25/25	3.7 (25)	97	25/25	3.9 (25)	103	25/25	4.0 (25)	105	25/25
3-7	3.9 (25)	25/25	3.9 (25)	100	25/25	4.0 (25)	103	25/25	4.1 (25)	105	25/25
4-7	4.1 (25)	25/25	4.0 (24)	98	24/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25
5-7	4.1 (25)	25/25	4.1 (24)	100	24/25	4.2 (25)	102	25/25	4.2 (25)	102	25/25
6-7	4.1 (25)	25/25	4.2 (24)	102	24/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25
7-7	4.1 (25)	25/25	4.1 (24)	100	24/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25
8-7	4.1 (25)	25/25	4.1 (24)	100	24/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25
9-7	4.1 (25)	25/25	4.2 (24)	102	24/25	4.1 (25)	100	25/25	4.3 (25)	105	25/25
10-7	4.1 (25)	25/25	4.1 (24)	100	24/25	4.2 (25)	102	25/25	4.4 (25)	107	25/25
11-7	4.3 (25)	25/25	4.3 (24)	100	24/25	4.2 (25)	98	25/25	4.5 (25)	105	25/25
12-7	4.3 (25)	25/25	4.3 (24)	100	24/25	4.5 (25)	105	25/25	4.5 (25)	105	25/25
13-7	4.4 (25)	25/25	4.3 (24)	98	24/25	4.2 (25)	95	25/25	4.5 (25)	102	25/25
14-7	4.4 (25)	25/25	4.4 (24)	100	24/25	4.4 (25)	100	25/25	4.6 (25)	105	25/25
15-7	4.3 (25)	25/25	4.4 (24)	102	24/25	4.4 (25)	102	25/25	4.5 (25)	105	25/25
16-7	4.4 (25)	25/25	4.4 (24)	100	24/25	4.4 (25)	100	25/25	4.6 (25)	105	25/25
17-7	4.5 (25)	25/25	4.5 (24)	100	24/25	4.5 (25)	100	25/25	4.7 (25)	104	25/25
18-7	4.5 (25)	25/25	4.5 (24)	100	24/25	4.6 (25)	102	25/25	4.7 (25)	104	25/25
19-7	4.5 (25)	25/25	4.4 (24)	98	24/25	4.5 (25)	100	25/25	4.7 (25)	104	25/25
20-7	4.8 (25)	25/25	4.5 (24)	94	24/25	4.6 (25)	96	25/25	5.0 (25)	104	25/25
21-7	4.7 (25)	25/25	4.6 (24)	98	24/25	4.6 (25)	98	25/25	4.9 (25)	104	25/25
22-7	4.7 (24)	24/25	4.5 (24)	96	24/25	4.7 (25)	100	25/25	4.9 (25)	104	25/25
23-7	4.6 (24)	24/25	4.5 (24)	98	24/25	4.7 (25)	102	25/25	4.8 (25)	104	25/25
24-7	4.7 (24)	24/25	4.5 (24)	96	24/25	4.7 (25)	100	25/25	4.9 (25)	104	25/25
25-7	4.7 (24)	24/25	4.4 (24)	94	24/25	4.5 (25)	96	25/25	4.7 (25)	100	25/25
26-7	4.7 (24)	24/25	4.5 (24)	96	24/25	4.5 (25)	96	25/25	4.9 (25)	104	25/25

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

TABLE E2

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

MEAN FOOD CONSUMPTION(FC) AND SURVIVAL

Week-Day on Study	Control Av.FC.	Control No.of Surviv. <25>	2400 ppm Av.FC.	2400 ppm % of cont. <25>	2400 ppm No.of Surviv.	6000 ppm Av.FC.	6000 ppm % of cont. <25>	6000 ppm No.of Surviv.	15000 ppm Av.FC.	15000 ppm % of cont. <25>	15000 ppm No.of Surviv.
1-7	3.8 (25)	25/25	3.8 (25)	100	25/25	3.9 (25)	103	25/25	3.9 (25)	103	25/25
2-7	3.4 (25)	25/25	3.5 (25)	103	25/25	3.6 (25)	106	25/25	3.5 (25)	103	25/25
3-7	3.4 (25)	25/25	3.6 (25)	106	25/25	3.4 (25)	100	25/25	3.6 (25)	106	25/25
4-7	3.8 (25)	25/25	3.6 (25)	95	25/25	3.4 (25)	89	25/25	4.0 (25)	105	25/25
5-7	3.9 (25)	25/25	3.8 (25)	97	25/25	3.7 (25)	95	25/25	3.9 (25)	100	25/25
6-7	4.0 (25)	25/25	3.7 (25)	93	25/25	3.7 (25)	93	25/25	4.0 (25)	100	25/25
7-7	4.0 (25)	25/25	3.8 (25)	95	25/25	3.9 (25)	98	25/25	4.0 (25)	100	25/25
8-7	4.1 (25)	25/25	3.8 (25)	93	25/25	3.8 (25)	93	25/25	4.1 (25)	100	25/25
9-7	4.1 (25)	25/25	4.1 (25)	100	25/25	3.9 (25)	95	25/25	4.1 (25)	100	25/25
10-7	4.1 (25)	25/25	4.0 (25)	98	25/25	3.8 (25)	93	25/25	4.3 (25)	105	25/25
11-7	4.2 (25)	25/25	4.0 (25)	95	25/25	3.8 (25)	90	25/25	4.2 (25)	100	25/25
12-7	4.4 (25)	25/25	4.0 (25)	91	25/25	4.0 (25)	91	25/25	4.5 (25)	102	25/25
13-7	4.3 (25)	25/25	4.1 (25)	95	25/25	4.0 (25)	93	25/25	4.7 (25)	109	25/25
14-7	4.4 (25)	25/25	4.2 (24)	95	24/25	4.2 (25)	95	25/25	4.6 (25)	105	25/25
15-7	4.4 (25)	25/25	4.2 (24)	95	24/25	4.2 (25)	95	25/25	4.7 (25)	107	25/25
16-7	4.4 (24)	24/25	4.1 (24)	93	24/25	4.1 (25)	93	25/25	4.8 (25)	109	25/25
17-7	4.5 (24)	24/25	4.3 (24)	96	24/25	4.4 (25)	98	25/25	5.0 (25)	111	25/25
18-7	4.5 (24)	24/25	4.1 (24)	91	24/25	4.2 (25)	93	25/25	4.9 (25)	109	25/25
19-7	4.3 (23)	23/25	4.2 (24)	98	24/25	4.2 (24)	98	24/25	4.8 (25)	112	25/25
20-7	4.3 (23)	23/25	4.3 (24)	100	24/25	4.1 (24)	95	24/25	5.3 (25)	123	25/25
21-7	4.4 (23)	23/25	4.4 (24)	100	24/25	4.2 (24)	95	24/25	5.2 (25)	118	25/25
22-7	4.3 (23)	23/25	4.4 (24)	102	24/25	4.3 (23)	100	23/25	5.2 (25)	121	25/25
23-7	4.3 (23)	23/25	4.4 (24)	102	24/25	4.2 (23)	98	23/25	5.2 (25)	121	25/25
24-7	4.4 (23)	23/25	4.4 (24)	100	24/25	4.4 (23)	100	23/25	5.3 (25)	120	25/25
25-7	4.3 (23)	23/25	4.2 (24)	98	24/25	4.1 (23)	95	23/25	4.9 (25)	114	25/25
26-7	4.3 (23)	23/25	4.3 (24)	100	24/25	4.2 (23)	98	23/25	5.1 (25)	119	25/25

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

TABLE E3

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
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)	5-7(7)	6-7(7)	7-7(7)
Control	3.9± 0.2	3.8± 0.3	3.9± 0.3	4.1± 0.3	4.1± 0.3	4.1± 0.3	4.1± 0.3
2400 ppm	3.9± 0.3	3.7± 0.8	3.9± 0.6	4.0± 0.4	4.1± 0.3	4.2± 0.3	4.1± 0.3
6000 ppm	4.1± 0.2*	3.9± 0.3	4.0± 0.3	4.1± 0.3	4.2± 0.2	4.1± 0.3	4.1± 0.2
15000 ppm	4.1± 0.3*	4.0± 0.3*	4.1± 0.3*	4.2± 0.2	4.2± 0.2	4.2± 0.3	4.2± 0.3

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 8-7(7)	9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	14-7(7)
Control	4.1± 0.3	4.1± 0.3	4.1± 0.3	4.3± 0.3	4.3± 0.4	4.4± 0.4	4.4± 0.4
2400 ppm	4.1± 0.3	4.2± 0.3	4.1± 0.3	4.3± 0.3	4.3± 0.3	4.3± 0.4	4.4± 0.4
6000 ppm	4.1± 0.2	4.1± 0.2	4.2± 0.2	4.2± 0.2	4.5± 0.2	4.2± 0.3	4.4± 0.2
15000 ppm	4.2± 0.3	4.3± 0.3	4.4± 0.3**	4.5± 0.4	4.5± 0.4	4.5± 0.4	4.6± 0.5

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 15-7(7)	16-7(7)	17-7(7)	18-7(7)	19-7(7)	20-7(7)	21-7(7)
Control	4.3± 0.5	4.4± 0.5	4.5± 0.6	4.5± 0.5	4.5± 0.5	4.8± 0.7	4.7± 0.5
2400 ppm	4.4± 0.4	4.4± 0.4	4.5± 0.5	4.5± 0.5	4.4± 0.4	4.5± 0.4	4.6± 0.4
6000 ppm	4.4± 0.3	4.4± 0.3	4.5± 0.3	4.6± 0.3	4.5± 0.4	4.6± 0.4	4.6± 0.3
15000 ppm	4.5± 0.4	4.6± 0.4	4.7± 0.4	4.7± 0.4	4.7± 0.3	5.0± 0.5	4.9± 0.4*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 22-7(7)	23-7(7)	24-7(7)	25-7(7)	26-7(7)
Control	4.7± 0.5	4.6± 0.5	4.7± 0.5	4.7± 0.5	4.7± 0.5
2400 ppm	4.5± 0.5	4.5± 0.5	4.5± 0.4	4.4± 0.5	4.5± 0.5
6000 ppm	4.7± 0.3	4.7± 0.3	4.7± 0.4	4.5± 0.3	4.5± 0.2
15000 ppm	4.9± 0.3	4.8± 0.4	4.9± 0.4	4.7± 0.4	4.9± 0.5

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE E4

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
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)	5-7(7)	6-7(7)	7-7(7)
Control	3.8± 0.3	3.4± 0.3	3.4± 0.3	3.8± 0.4	3.9± 0.4	4.0± 0.4	4.0± 0.4
2400 ppm	3.8± 0.3	3.5± 0.3	3.6± 0.3	3.6± 0.4	3.8± 0.4	3.7± 0.5	3.8± 0.4
6000 ppm	3.9± 0.3	3.6± 0.3*	3.4± 0.3	3.4± 0.4*	3.7± 0.3	3.7± 0.4	3.9± 0.3
15000 ppm	3.9± 0.4	3.5± 0.3	3.6± 0.3	4.0± 0.4	3.9± 0.4	4.0± 0.4	4.0± 0.4

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 8-7(7)	9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	14-7(7)
Control	4.1± 0.3	4.1± 0.4	4.1± 0.6	4.2± 0.7	4.4± 1.0	4.3± 0.9	4.4± 0.7
2400 ppm	3.8± 0.3*	4.1± 0.4	4.0± 0.5	4.0± 0.5	4.0± 0.6	4.1± 0.7	4.2± 0.4
6000 ppm	3.8± 0.4*	3.9± 0.3*	3.8± 0.3	3.8± 0.3	4.0± 0.3	4.0± 0.4	4.2± 0.4
15000 ppm	4.1± 0.5	4.1± 0.5	4.3± 0.5*	4.2± 0.4	4.5± 0.5	4.7± 0.5**	4.6± 0.5

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 15-7(7)	16-7(7)	17-7(7)	18-7(7)	19-7(7)	20-7(7)	21-7(7)
Control	4.4± 0.8	4.4± 1.0	4.5± 1.0	4.5± 1.1	4.3± 0.4	4.3± 0.4	4.4± 0.4
2400 ppm	4.2± 0.4	4.1± 0.3	4.3± 0.4	4.1± 0.4	4.2± 0.4	4.3± 0.4	4.4± 0.4
6000 ppm	4.2± 0.4	4.1± 0.4	4.4± 0.7	4.2± 0.6	4.2± 0.5	4.1± 0.6	4.2± 0.5
15000 ppm	4.7± 0.6**	4.8± 0.5**	5.0± 0.6**	4.9± 0.8**	4.8± 0.7**	5.3± 0.8**	5.2± 0.8**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
ALL ANIMALS

Group Name	Administration week-day(effective) 22-7(7)	23-7(7)	24-7(7)	25-7(7)	26-7(7)
Control	4.3± 0.4	4.3± 0.3	4.4± 0.5	4.3± 0.4	4.3± 0.5
2400 ppm	4.4± 0.5	4.4± 0.3	4.4± 0.4	4.2± 0.4	4.3± 0.3
6000 ppm	4.3± 0.5	4.2± 0.4	4.4± 0.5	4.1± 0.4	4.2± 0.5
15000 ppm	5.2± 0.6**	5.2± 0.7**	5.3± 0.7**	4.9± 0.6**	5.1± 0.7**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE F1

URINALYSIS : MALE

STUDY NO. : 0922 URINALYSIS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

Group Name	NO. of Animals	pH 5.0	6.0	6.5	7.0	7.5	8.0	8.5	CHI	Protein −	±	+	2+	3+	4+	CHI	Glucose −	±	+	2+	3+	4+	CHI	Ketone body −	±	+	2+	3+	4+	CHI	Occult blood −	±	+	2+	3+	CHI
Control	22	0	4	0	4	1	5	8		3	7	10	2	0	0		22	0	0	0	0	0		6	8	6	2	0	0		22	0	0	0	0	
2400 ppm	24	0	2	2	0	0	7	13		1	5	12	6	0	0		24	0	0	0	0	0		6	4	12	2	0	0		24	0	0	0	0	
6000 ppm	23	0	1	4	0	5	5	8	*	1	6	15	0	1	0		23	0	0	0	0	0		7	8	7	1	0	0		23	0	0	0	0	
15000 ppm	25	0	3	3	2	2	10	5		2	15	7	1	0	0		25	0	0	0	0	0		9	10	5	0	1	0		25	0	0	0	0	

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of CHI SQUARE

STUDY NO. : 0922 URINALYSIS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

Group Name	NO. of Animals	Urobilinogen ±	＋	2+	3+	4+	CHI
Control	22	22	0	0	0	0	
2400 ppm	24	24	0	0	0	0	
6000 ppm	23	23	0	0	0	0	
15000 ppm	25	25	0	0	0	0	

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of CHI SQUARE

TABLE F2

URINALYSIS : FEMALE

STUDY NO. : 0922 URINALYSIS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	pH 5.0	6.0	6.5	7.0	7.5	8.0	8.5	CHI	Protein −	±	+	2+	3+	4+	CHI	Glucose −	±	+	2+	3+	4+	CHI	Ketone body −	±	+	2+	3+	4+	CHI	Occult blood −	±	+	2+	3+	CHI
Control	22	0	4	8	4	3	3	0		8	12	1	0	1	0		22	0	0	0	0	0		3	14	5	0	0	0		22	0	0	0	0	
2400 ppm	23	0	4	6	3	2	6	2		5	16	2	0	0	0		23	0	0	0	0	0		2	17	4	0	0	0		23	0	0	0	0	
6000 ppm	20	0	6	1	1	1	8	3	*	4	14	2	0	0	0		20	0	0	0	0	0		3	10	6	1	0	0		20	0	0	0	0	
15000 ppm	24	0	4	4	3	6	7	0		4	18	2	0	0	0		24	0	0	0	0	0		0	19	5	0	0	0		24	0	0	0	0	

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of CHI SQUARE

STUDY NO. : 0922 URINALYSIS
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	Urobilinogen ±	＋	2+	3+	4+	CHI
Control	22	22	0	0	0	0	
2400 ppm	23	23	0	0	0	0	
6000 ppm	20	20	0	0	0	0	
15000 ppm	24	24	0	0	0	0	

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of CHI SQUARE

TABLE G1

HEMATOLOGY : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	RED BLOOD CELL 10^6／μℓ	HEMOGLOBIN g／dℓ	HEMATOCRIT %	MCV f ℓ	MCH p g	MCHC g／dℓ	PLATELET 10^3／μℓ
Control	24	10.11± 0.81	14.4± 1.3	42.4± 3.4	41.9± 0.8	14.2± 0.4	33.9± 0.9	1453± 199
2400 ppm	24	10.27± 0.51	14.8± 0.4	43.4± 1.3	42.3± 1.2	14.4± 0.4	34.1± 0.5	1455± 158
6000 ppm	23	10.17± 0.44	14.7± 0.4	43.0± 1.3	42.3± 0.8	14.4± 0.3*	34.1± 0.4	1460± 189
15000 ppm	24	9.86± 0.35**	14.3± 0.4**	42.1± 1.3	42.7± 0.7*	14.5± 0.3**	34.1± 0.6	1371± 228

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

Group Name	NO. of Animals	RETICULOCYTE %	
Control	24	4.4±	2.8
2400 ppm	24	3.8±	0.5
6000 ppm	23	4.0±	0.7
15000 ppm	24	4.6±	0.6**

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	WBC $10^3/\mu\ell$		Differential WBC (%) NEUTRO		LYMPHO		MONO		EOSINO		BASO	
Control	24	2.98±	1.50	14.8±	9.3	80.8±	10.1	3.6±	1.7	0.9±	0.5	0.0±	0.0
2400 ppm	24	2.99±	1.64	12.5±	6.1	83.8±	6.1	2.7±	0.9	0.9±	0.6	0.0±	0.0
6000 ppm	23	2.92±	1.67	11.7±	3.7	84.6±	3.9	2.9±	0.7	0.8±	0.4	0.0±	0.0
15000 ppm	24	2.52±	1.49	9.7±	4.1	86.2±	4.3	3.2±	1.1	0.9±	0.6	0.0±	0.0

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE G2

HEMATOLOGY : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	RED BLOOD CELL 10^6／μℓ	HEMOGLOBIN g／dℓ	HEMATOCRIT %	MCV f ℓ	MCH p g	MCHC g／dℓ	PLATELET 10^3／μℓ
Control	23	10.26± 0.50	14.8± 0.6	43.0± 1.9	42.0± 1.0	14.4± 0.3	34.4± 0.4	1242± 193
2400 ppm	24	10.36± 0.42	15.0± 0.5	43.7± 1.6	42.2± 0.8	14.5± 0.4	34.4± 0.5	1177± 189
6000 ppm	22	10.41± 0.42	15.1± 0.5	43.7± 1.5	42.0± 0.7	14.5± 0.3	34.5± 0.6	1205± 234
15000 ppm	24	9.94± 0.92	14.7± 1.2	43.1± 2.7	43.5± 2.2**	14.8± 0.5**	34.1± 1.1	1091± 215

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

Group Name	NO. of Animals	RETICULOCYTE %
Control	23	4.3± 1.2
2400 ppm	24	3.9± 0.5
6000 ppm	22	3.8± 0.5
15000 ppm	24	5.7± 5.1

Significant difference : ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	WBC $10^3/\mu\ell$		Differential WBC (%) NEUTRO		LYMPHO		MONO		EOSINO		BASO	
Control	23	1.91±	0.90	15.4±	7.4	81.4±	7.4	2.8±	1.1	0.4±	0.5	0.0±	0.0
2400 ppm	24	1.90±	0.94	16.2±	5.8	79.7±	6.7	3.5±	2.8	0.6±	0.7	0.0±	0.0
6000 ppm	22	1.77±	0.98	13.8±	6.7	81.5±	6.8	4.0±	4.1	0.3±	0.5	0.4±	1.9
15000 ppm	24	1.67±	0.91	19.1±	11.7	77.1±	11.5	3.2±	1.2	0.6±	0.6	0.0±	0.1

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE H1

BIOCHEMISTRY : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (27W)

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	24	5.2± 0.3	2.9± 0.3	1.3± 0.2	0.11± 0.02	224± 50	77± 12	44± 11
2400 ppm	24	5.3± 0.3	3.1± 0.2	1.4± 0.1	0.11± 0.02	247± 59	81± 14	43± 14
6000 ppm	23	5.2± 0.2	3.0± 0.2	1.4± 0.1	0.11± 0.02	240± 37	78± 14	41± 9
15000 ppm	24	5.0± 0.2	2.9± 0.2	1.4± 0.1	0.13± 0.03*	255± 40	76± 15	36± 13

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	PHOSPHOLIPID mg／dℓ	AST U／L	ALT U／L	LDH U／L	ALP U／L	G-GTP U／L	CK U／L
Control	24	160± 24	65± 31	21± 11	207± 60	195± 40	0.3± 0.4	81± 46
2400 ppm	24	168± 29	55± 18	19± 5	199± 54	216± 31	0.4± 0.4	86± 75
6000 ppm	23	161± 24	58± 34	18± 10	191± 30	213± 41	0.5± 0.3	64± 29
15000 ppm	24	151± 27	58± 35	22± 24*	201± 54	228± 23**	0.4± 0.3	57± 26*

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

Group Name	NO. of Animals	UREANITROGEN mg／dℓ	SODIUM mEq／ℓ	POTASSIUM mEq／ℓ	CHLORIDE mEq／ℓ	CALCIUM mg／dℓ	INORGANIC PHOSPHRUS mg／dℓ
Control	24	30.0± 2.8	151± 2	4.0± 0.2	119± 2	9.0± 0.2	5.9± 1.1
2400 ppm	24	30.1± 5.3	151± 2	4.1± 0.2	119± 2	9.0± 0.3	5.5± 1.0
6000 ppm	23	28.4± 3.1	150± 2	4.1± 0.2	118± 2	8.9± 0.2	5.7± 1.0
15000 ppm	24	29.0± 5.6	149± 2	4.1± 0.2	118± 2	8.9± 0.2	5.7± 1.2

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

TABLE H2

BIOCHEMISTRY : FEMALE

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	23	5.2± 0.3	3.1± 0.2	1.5± 0.1	0.12± 0.04	173± 34	64± 21	27± 8
2400 ppm	23	5.2± 0.3	3.1± 0.2	1.5± 0.1	0.12± 0.03	183± 30	64± 14	29± 10
6000 ppm	22	5.3± 0.2	3.2± 0.2	1.5± 0.1	0.11± 0.03	189± 26	67± 12	29± 9
15000 ppm	24	5.1± 0.3	3.0± 0.3	1.5± 0.2	0.12± 0.03	191± 37	69± 17	27± 10

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

Group Name	NO. of Animals	PHOSPHOLIPID mg／dℓ	AST U／L	ALT U／L	LDH U／L	ALP U／L	G-GTP U／L	CK U／L
Control	23	134± 38	82± 36	24± 17	202± 48	374± 80	0.2± 0.2	107± 113
2400 ppm	23	138± 31	73± 25	19± 5	191± 48	364± 56	0.4± 0.3	101± 80
6000 ppm	22	145± 24	67± 19	22± 6	198± 67	386± 80	0.2± 0.3	116± 227*
15000 ppm	24	137± 29	73± 37	22± 17	202± 66	377± 127	0.3± 0.3	68± 26*

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (27W)

Group Name	NO. of Animals	UREANITROGEN mg／dℓ	SODIUM mEq／ℓ	POTASSIUM mEq／ℓ	CHLORIDE mEq／ℓ	CALCIUM mg／dℓ	INORGANIC PHOSPHRUS mg／dℓ
Control	23	24.2± 4.6	151± 2	3.6± 0.2	119± 2	8.9± 0.2	6.5± 1.1
2400 ppm	23	23.5± 4.1	151± 2	3.6± 0.3	119± 3	8.9± 0.2	6.4± 1.1
6000 ppm	22	23.8± 2.1	151± 2	3.6± 0.3	120± 2	9.0± 0.2	6.0± 1.2
15000 ppm	24	24.4± 4.8	151± 2	3.5± 0.3	120± 2	8.9± 0.2	6.5± 1.0

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

TABLE I1

GROSS FINDINGS : MALE

Organ	Findings	Group Name / NO. of Animals	Control 25	(%)	2400 ppm 25	(%)	6000 ppm 25	(%)	15000 ppm 25	(%)
skin/app	erosion		1	(4)	0	(0)	0	(0)	0	(0)
lymph node	enlarged		3	(12)	0	(0)	0	(0)	0	(0)
thymus	atrophic		0	(0)	1	(4)	0	(0)	0	(0)
spleen	enlarged		1	(4)	0	(0)	0	(0)	0	(0)
	black zone		2	(8)	2	(8)	2	(8)	0	(0)
stomach	forestomach:nodule		1	(4)	1	(4)	0	(0)	0	(0)
kidney	white zone		0	(0)	1	(4)	0	(0)	1	(4)
	hydronephrosis		0	(0)	1	(4)	0	(0)	0	(0)
urin bladd	urine:marked retention		1	(4)	0	(0)	0	(0)	1	(4)
thoracic ca	pleural fluid		1	(4)	0	(0)	0	(0)	0	(0)

TABLE I2

GROSS FINDINGS : FEMALE

Organ	Findings	Group Name / NO. of Animals	Control 25	(%)	2400 ppm 25	(%)	6000 ppm 25	(%)	15000 ppm 25	(%)
skin/app	erosion		1	(4)	1	(4)	2	(8)	1	(4)
lung	white zone		1	(4)	0	(0)	0	(0)	0	(0)
	red zone		0	(0)	0	(0)	1	(4)	0	(0)
	adhesion		0	(0)	1	(4)	0	(0)	0	(0)
lymph node	enlarged		0	(0)	0	(0)	1	(4)	0	(0)
thymus	enlarged		0	(0)	0	(0)	1	(4)	0	(0)
spleen	enlarged		2	(8)	0	(0)	3	(12)	0	(0)
	black zone		0	(0)	1	(4)	0	(0)	1	(4)
stomach	glandular stomach:erosion		0	(0)	1	(4)	1	(4)	1	(4)
kidney	enlarged		0	(0)	0	(0)	1	(4)	0	(0)
	nodule		0	(0)	1	(4)	0	(0)	0	(0)
	cyst		0	(0)	1	(4)	0	(0)	0	(0)
	hydronephrosis		0	(0)	1	(4)	0	(0)	1	(4)
ovary	enlarged		0	(0)	0	(0)	1	(4)	0	(0)
	cyst		0	(0)	1	(4)	0	(0)	0	(0)
thoracic ca	pleural fluid		0	(0)	0	(0)	1	(4)	0	(0)
other	hindlimb:nodule		1	(4)	0	(0)	0	(0)	0	(0)

TABLE J1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	Body Weight	ADRENALS	TESTES	HEART	LUNGS	KIDNEYS
Control	24	33.6± 4.0	0.009± 0.003	0.227± 0.017	0.178± 0.018	0.157± 0.015	0.465± 0.047
2400 ppm	24	33.8± 4.1	0.009± 0.002	0.230± 0.025	0.170± 0.015	0.153± 0.014	0.436± 0.036
6000 ppm	25	33.0± 2.6	0.009± 0.002	0.220± 0.022	0.174± 0.020	0.156± 0.013	0.456± 0.052
15000 ppm	25	31.2± 2.4	0.009± 0.002	0.218± 0.032	0.169± 0.019	0.157± 0.010	0.476± 0.054

Significant difference : ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	24	0.094± 0.059	1.400± 0.223	0.479± 0.016
2400 ppm	24	0.078± 0.010	1.399± 0.291	0.471± 0.017
6000 ppm	25	0.081± 0.012	1.281± 0.173	0.475± 0.020
15000 ppm	25	0.088± 0.015	1.232± 0.155**	0.478± 0.019

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE J2

ORGAN WEIGHT, ABSOLUTE : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	Body Weight	ADRENALS	OVARIES	HEART	LUNGS	KIDNEYS
Control	23	23.7± 1.6	0.013± 0.002	0.029± 0.004	0.146± 0.014	0.153± 0.015	0.362± 0.036
2400 ppm	24	23.5± 1.5	0.013± 0.002	0.033± 0.009	0.145± 0.015	0.149± 0.010	0.399± 0.198
6000 ppm	23	22.9± 1.5	0.012± 0.002	0.030± 0.007	0.138± 0.010	0.151± 0.014	0.340± 0.026
15000 ppm	25	22.3± 1.3**	0.012± 0.002	0.028± 0.007	0.141± 0.017	0.152± 0.015	0.377± 0.066

Significant difference : * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	23	0.095± 0.019	0.990± 0.114	0.484± 0.015
2400 ppm	24	0.088± 0.008	1.021± 0.153	0.484± 0.016
6000 ppm	23	0.099± 0.055	1.041± 0.124	0.479± 0.019
15000 ppm	25	0.101± 0.042	1.025± 0.145	0.479± 0.015

Significant difference ; ＊ : P ≦ 0.05 ＊＊ : P ≦ 0.01 Test of Dunnett

TABLE K1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	Body Weight (g)	ADRENALS	TESTES	HEART	LUNGS	KIDNEYS
Control	24	33.6± 4.0	0.026± 0.010	0.680± 0.069	0.540± 0.101	0.474± 0.070	1.405± 0.248
2400 ppm	24	33.8± 4.1	0.028± 0.008	0.688± 0.097	0.508± 0.057	0.458± 0.060	1.304± 0.162
6000 ppm	25	33.0± 2.6	0.029± 0.006	0.671± 0.083	0.530± 0.063	0.475± 0.038	1.384± 0.155
15000 ppm	25	31.2± 2.4	0.029± 0.007	0.698± 0.099	0.544± 0.074	0.505± 0.036*	1.530± 0.197

Significant difference : * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	24	0.292± 0.238	4.172± 0.537	1.448± 0.208
2400 ppm	24	0.233± 0.030	4.116± 0.522	1.411± 0.168
6000 ppm	25	0.247± 0.037	3.880± 0.417	1.447± 0.111
15000 ppm	25	0.281± 0.046**	3.945± 0.404	1.541± 0.139

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE K2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	Body Weight (g)	ADRENALS	OVARIES	HEART	LUNGS	KIDNEYS
Control	23	23.7± 1.6	0.053± 0.007	0.123± 0.019	0.616± 0.052	0.649± 0.063	1.529± 0.111
2400 ppm	24	23.5± 1.5	0.056± 0.008	0.142± 0.041	0.616± 0.054	0.636± 0.041	1.702± 0.861
6000 ppm	23	22.9± 1.5	0.054± 0.010	0.131± 0.026	0.603± 0.039	0.661± 0.048	1.491± 0.112
15000 ppm	25	22.3± 1.3**	0.056± 0.006	0.128± 0.029	0.635± 0.072	0.684± 0.051	1.690± 0.244*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

STUDY NO. : 0922
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (27W)

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	23	0.398± 0.064	4.202± 0.547	2.055± 0.145
2400 ppm	24	0.373± 0.039	4.345± 0.594	2.068± 0.139
6000 ppm	23	0.432± 0.230	4.557± 0.492	2.101± 0.129
15000 ppm	25	0.453± 0.203	4.605± 0.618	2.157± 0.105*

Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01 Test of Dunnett

TABLE L1

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : MALE

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of animals on Study	25	25	25	25
{Hematopoietic system}						
lymph node			<25>	<25>	<25>	<25>
	malignant lymphoma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
{Special sense organs/appendage}						
Harder gl			<25>	<25>	<25>	<25>
	hemangioma		0 (0%)	0 (0%)	1 (4%)	0 (0%)

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

TABLE L2

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : FEMALE

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of animals on Study	25	25	25	25
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	bronchiolar-alveolar adenoma		1 (4%)	1 (4%)	0 (0%)	0 (0%)
{Hematopoietic system}						
bone marrow			<25>	<25>	<25>	<25>
	hemangioma		0 (0%)	0 (0%)	1 (4%)	0 (0%)
thymus			<25>	<25>	<25>	<25>
	malignant lymphoma		0 (0%)	0 (0%)	1 (4%)	0 (0%)
{Musculoskeletal system}						
bone			<25>	<25>	<25>	<25>
	osteosarcoma		1 (4%)	0 (0%)	0 (0%)	0 (0%)

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

TABLE M1

NEOPLASTIC LESIONS-INCIDENCE AND

STATISTICAL ANALYSIS : MALE

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : lymph node				
TUMOR : malignant lymphoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	0/25(0.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	0.00	0.00	0.00	0.00
Terminal rates(c)	0/24(0.0)	0/24(0.0)	0/25(0.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 1.0000 ?			
Prevalence method(d)	P = ------			
Combined analysis(d)	P = 1.0000 ?			
Cochran-Armitage test(e)	P = 0.3021			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.5000
SITE : Harderian gland				
TUMOR : hemangioma				
Tumor rate				
Overall rates(a)	0/25(0.0)	0/25(0.0)	1/25(4.0)	0/25(0.0)
Adjusted rates(b)	0.00	0.00	4.00	0.00
Terminal rates(c)	0/24(0.0)	0/24(0.0)	1/25(4.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.3948			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.9789			
Fisher Exact test(e)		P = N.C.	P = 0.5000	P = N.C.

(a): Number of tumor-beaning animals/number of animals examined at the site.
(b): Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.
(c): Observed tumor incidence at terminal kill.
(d): Beneath the control incidence are the P-values associated with the trend test.
Standard method : Death analysis
Prevalence method : Incidental tumor test
Combined analysis : Death analysis + Incidental tumor test
(e): The Cochran-Armitage and Fisher exact test compare directly the overall incidence retes.
? : The conditional probabilities of the largest and smallest possible out comes can not estimated or this P-value is beyond the estimated P-value.
------ : There is no data which should be statistical analysis.
Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01
N.C.:Statistical value cannot be calculated and was not significant.

TABLE M2

NEOPLASTIC LESIONS-INCIDENCE AND

STATISTICAL ANALYSIS : FEMALE

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	1/25(4.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	4.35	4.17	0.00	0.00
Terminal rates(c)	1/23(4.3)	1/24(4.2)	0/23(0.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.8854			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.2437			
Fisher Exact test(e)		P = 0.7551	P = 0.5000	P = 0.5000
SITE : bone marrow				
TUMOR : hemangioma				
Tumor rate				
Overall rates(a)	0/25(0.0)	0/25(0.0)	1/25(4.0)	0/25(0.0)
Adjusted rates(b)	0.00	0.00	4.35	0.00
Terminal rates(c)	0/23(0.0)	0/24(0.0)	1/23(4.3)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.3975			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.9789			
Fisher Exact test(e)		P = N.C.	P = 0.5000	P = N.C.
SITE : thymus				
TUMOR : malignant lymphoma				
Tumor rate				
Overall rates(a)	0/25(0.0)	0/25(0.0)	1/25(4.0)	0/25(0.0)
Adjusted rates(b)	0.00	0.00	0.00	0.00
Terminal rates(c)	0/23(0.0)	0/24(0.0)	0/23(0.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.3981			
Prevalence method(d)	P = ------			
Combined analysis(d)	P = 0.3981			
Cochran-Armitage test(e)	P = 0.9789			
Fisher Exact test(e)		P = N.C.	P = 0.5000	P = N.C.

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : bone				
TUMOR : osteosarcoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	0/25(0.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	0.00	0.00	0.00	0.00
Terminal rates(c)	0/23(0.0)	0/24(0.0)	0/23(0.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 1.0000 ?			
Prevalence method(d)	P = ------			
Combined analysis(d)	P = 1.0000 ?			
Cochran-Armitage test(e)	P = 0.3021			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.5000

(a): Number of tumor-beaning animals/number of animals examined at the site.
(b): Kaplan-Meier estimated tumor incidence at the end of the study after adjusting for intercurrent mortality.
(c): Observed tumor incidence at terminal kill.
(d): Beneath the control incidence are the P-values associated with the trend test.
Standard method : Death analysis
Prevalence method : Incidental tumor test
Combined analysis : Death analysis + Incidental tumor test
(e): The Cochran-Armitage and Fisher exact test compare directly the overall incidence retes.
? : The conditional probabilities of the largest and smallest possible out comes can not estimated or this P-value is beyond the estimated P-value.
------ : There is no data which should be statistical analysis.
Significant difference ; * : P ≦ 0.05 ** : P ≦ 0.01
N.C.:Statistical value cannot be calculated and was not significant.

TABLE N1

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR : MALE

Organ	Findings	Group Name / No. of Animals on Study	Control	2400 ppm	6000 ppm	15000 ppm
			25	25	25	25
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0
{Hematopoietic system}						
bone marrow			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0
spleen			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0
{Digestive system}						
liver			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0
{Urinary system}						
kidney			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0
{Nervous system}						
brain			<25>	<25>	<25>	<25>
	leukemic cell infiltration		1	0	0	0

< a > a : Number of animals examined at the site
b b : Number of animals with lesion

TABLE N2

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR : FEMALE

Organ	Findings	Group Name / No. of Animals on Study	Control	2400 ppm	6000 ppm	15000 ppm
			25	25	25	25
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
	metastasis:bone tumor		1	0	0	0
{Hematopoietic system}						
bone marrow			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
lymph node			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
spleen			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Circulatory system}						
heart			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Digestive system}						
liver			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Urinary system}						
kidney			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Endocrine system}						
pituitary			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0

< a > a : Number of animals examined at the site
b b : Number of animals with lesion

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of Animals on Study	25	25	25	25
{Reproductive system}						
ovary			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Nervous system}						
brain			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0
{Musculoskeletal system}						
bone			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	1	0

< a > a : Number of animals examined at the site
b b : Number of animals with lesion

TABLE O1

HISTOPATHOLOGICAL FINDINGS :

NON-NEOPLASTIC LESIONS : MALE

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Integumentary system/appandage}																		
skin/app				<25>				<25>				<25>				<25>		
	ulcer		0	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Respiratory system}																		
larynx				<25>				<25>				<25>				<25>		
	hyperplasia:squamous epithelium		0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
lung				<25>				<25>				<25>				<25>		
	hemorrhage		0	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	bronchiolar-alveolar cell hyperplasia		0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	vacuolic change:bronchial epithelium		0	0	0	0	1	0	0	0	8	1	0	0 **	3	8	0	0 **
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(32)	(4)	(0)	(0)	(12)	(32)	(0)	(0)
{Hematopoietic system}																		
bone marrow				<25>				<25>				<25>				<25>		
	increased hemotopoiesis:granulocyte/monocyte		0	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

Organ	Findings	Group Name / No. of Animals on Study / Grade	Control 25 1+ (%)	Control 25 2+ (%)	Control 25 3+ (%)	Control 25 4+ (%)	2400 ppm 25 1+ (%)	2400 ppm 25 2+ (%)	2400 ppm 25 3+ (%)	2400 ppm 25 4+ (%)	6000 ppm 25 1+ (%)	6000 ppm 25 2+ (%)	6000 ppm 25 3+ (%)	6000 ppm 25 4+ (%)	15000 ppm 25 1+ (%)	15000 ppm 25 2+ (%)	15000 ppm 25 3+ (%)	15000 ppm 25 4+ (%)
{Hematopoietic system}																		
lymph node		<25>																
	lymphadenitis		2 (8)	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)
thymus		<25>																
	atrophy		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	ultimobranchial body remanet		1 (4)	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)
spleen		<25>																
	deposit of melanin		2 (8)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	extramedullary hematopoiesis		1 (4)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)
{Circulatory system}																		
heart		<25>																
	endocardial hyperplasia		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)

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

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Circulatory system}																		
heart			<25>				<25>				<25>				<25>			
	valvulopathy		0	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Digestive system}																		
stomach			<25>				<25>				<25>				<25>			
	erosion:forestomach		0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	hyperplasia:forestomach		0	1	0	0	1	0	0	0	1	0	0	0	1	0	0	0
			(0)	(4)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	erosion:glandular stomach		0	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
small intes			<25>				<25>				<25>				<25>			
	hyperplasia		0	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
liver			<25>				<25>				<25>				<25>			
	fatty change:central		3	6	0	0	2	4	2	0	4	0	0	0 *	0	0	0	0 **
			(12)	(24)	(0)	(0)	(8)	(16)	(8)	(0)	(16)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

Organ	Findings	Group Name / No. of Animals on Study / Grade	Control 25 1+ (%)	Control 25 2+ (%)	Control 25 3+ (%)	Control 25 4+ (%)	2400 ppm 25 1+ (%)	2400 ppm 25 2+ (%)	2400 ppm 25 3+ (%)	2400 ppm 25 4+ (%)	6000 ppm 25 1+ (%)	6000 ppm 25 2+ (%)	6000 ppm 25 3+ (%)	6000 ppm 25 4+ (%)	15000 ppm 25 1+ (%)	15000 ppm 25 2+ (%)	15000 ppm 25 3+ (%)	15000 ppm 25 4+ (%)
{Digestive system}																		
liver		<25>																
	granulation		1 (4)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)
	mixed cell focus		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
pancreas		<25>																
	necrosis:focal		1 (4)	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)
{Urinary system}																		
kidney		<25>																
	hydronephrosis		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
ureter		<25>																
	hyperplasia:epithelium		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
urin bladd		<25>																
	dilatation		1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)

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

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Endocrine system}																		
adrenal			<25>				<25>				<25>				<25>			
	necrosis:focal		1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Reproductive system}																		
testis			<25>				<25>				<25>				<25>			
	tubular atrophy		0	0	0	0	1	0	0	0	1	1	0	0	4	2	0	0 *
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(4)	(0)	(0)	(16)	(8)	(0)	(0)
epididymis			<25>				<25>				<25>				<25>			
	granulomatous inflammation		0	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	spermatogenic granuloma		0	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)
{Musculoskeletal system}																		
bone			<25>				<25>				<25>				<25>			
	osteofibrosis		1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

TABLE O2

HISTOPATHOLOGICAL FINDINGS :

NON-NEOPLASTIC LESIONS : FEMALE

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Integumentary system/appandage}																		
skin/app			<25>				<25>				<25>				<25>			
	ulcer		0	1	0	0	0	1	0	0	0	2	0	0	1	0	0	0
			(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(8)	(0)	(0)	(4)	(0)	(0)	(0)
{Respiratory system}																		
lung			<25>				<25>				<25>				<25>			
	inflammatory infiltration		1	0	0	0	2	0	0	0	0	1	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)
	vacuolic change:bronchial epithelium		0	0	0	0	2	0	0	0	0	3	0	0	1	4	0	0
			(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(0)	(12)	(0)	(0)	(4)	(16)	(0)	(0)
	accumulation:macrophage		2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Hematopoietic system}																		
bone marrow			<25>				<25>				<25>				<25>			
	increased hemotopoiesis:granulocyte/monocyte		0	0	0	0	0	1	0	0	1	0	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(4)	(0)	(0)

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

Organ	Findings	Group Name / Grade	Control 1+ (%)	Control 2+ (%)	Control 3+ (%)	Control 4+ (%)	2400 ppm 1+ (%)	2400 ppm 2+ (%)	2400 ppm 3+ (%)	2400 ppm 4+ (%)	6000 ppm 1+ (%)	6000 ppm 2+ (%)	6000 ppm 3+ (%)	6000 ppm 4+ (%)	15000 ppm 1+ (%)	15000 ppm 2+ (%)	15000 ppm 3+ (%)	15000 ppm 4+ (%)
{Hematopoietic system}																		
lymph node		No. of Animals on Study	25				25				25				25			
	deposit of pigment	<25>	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	lymphadenitis	<25>	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
spleen		No. of Animals on Study	25				25				25				25			
	congestion	<25>	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	deposit of melanin	<25>	0	0	0	0	1	0	0	0	0	0	0	0	1	0	0	0
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	extramedullary hematopoiesis	<25>	4	2	0	0	7	1	0	0	3	1	0	0	7	1	0	0
			(16)	(8)	(0)	(0)	(28)	(4)	(0)	(0)	(12)	(4)	(0)	(0)	(28)	(4)	(0)	(0)
{Digestive system}																		
stomach		No. of Animals on Study	25				25				25				25			
	hyperplasia:forestomach	<25>	3	0	0	0	1	0	0	0	0	0	0	0	3	4	0	0
			(12)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(12)	(16)	(0)	(0)

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

Organ	Findings	Group Name / Grade	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
			1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)
{Digestive system}																		
stomach		<25>																
	erosion:glandular stomach		0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)
	ulcer:glandular stomach		1 (4)	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)
liver		<25>																
	necrosis:focal		1 (4)	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		2 (8)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	bile ductular proliferation		1 (4)	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)
gall bladd		<25>																
	eosinophilic change:focal		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
{Urinary system}																		
kidney		<25>																
	hydronephrosis		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)

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

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Urinary system}																		
kidney			<25>				<25>				<25>				<25>			
	regeneration:renal tubule		0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Reproductive system}																		
ovary			<25>				<25>				<25>				<25>			
	cystic extention:ovarian bursa		0	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
uterus			<25>				<25>				<25>				<25>			
	cystic endometrial hyperplasia		8	0	0	0	8	0	0	0	8	0	0	0	8	0	0	0
			(32)	(0)	(0)	(0)	(32)	(0)	(0)	(0)	(32)	(0)	(0)	(0)	(32)	(0)	(0)	(0)
{Nervous system}																		
spinal cord			<25>				<25>				<25>				<25>			
	ectopic tissue		1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
{Musculoskeletal system}																		
bone			<25>				<25>				<25>				<25>			
	osteosclerosis		0	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)

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

Organ	Findings	Group Name	Control				2400 ppm				6000 ppm				15000 ppm			
		No. of Animals on Study	25				25				25				25			
		Grade	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Musculoskeletal system}																		
bone			<25>				<25>				<25>				<25>			
	osteofibrosis		1	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)

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

TABLE P1

CAUSE OF DEATH : MALE

COUSE OF DEATH (SUMMARY)
(0- 27W)

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
Number of Dead and Moribund Animal	1	1	0	0
no microscop confirm	0	1	0	0
tumor d:leukemia	1	0	0	0

TABLE P2

CAUSE OF DEATH : FEMALE

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
Number of Dead and Moribund Animal	2	1	2	0
no microscop confirm	0	1	0	0
integumentary sy les	1	0	1	0
tumor d:leukemia	0	0	1	0
tumor d:bone	1	0	0	0