クロロエタンの rasH2 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：0929

TABLES

TABLES

TABLES (CONTINUED)

TABLES (CONTINUED)

TABLE A

CONCENTRATIONS OF CHLOROETHANE IN THE INHALATION CHAMBER OF rasH2 MICE IN THE 26-WEEK CARCINOGENICITY STUDY

TABLE A CONCENTRATIONS OF CHLOROETHANE IN THE INHALATION CHAMBER OF rasH2 MICE IN THE 26-WEEK CARCINOGENICITY

Group Name	Concentration(ppm) Mean ± S.D.
Control	0.0 ± 0.0
2400 ppm	2403.3 ± 7.0
6000 ppm	5997.3 ± 24.4
15000 ppm	15014.9 ± 34.8

TABLE B1

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0929 SURVIVAL ANIMAL NUMBERS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	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	24/25	24/25	24/25	23/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	96.0	96.0	96.0	92.0

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	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
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
		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	24/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	96.0
15000 ppm	25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25	23/25
		92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0	92.0

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

TABLE B2

SURVIVAL ANIMAL NUMBERS : 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(%)

STUDY NO. : 0929 SURVIVAL ANIMAL NUMBERS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1 26
SEX : FEMALE

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	25/25	25/25	25/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	23/25	23/25
		100.0	100.0	100.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	92.0	92.0
6000 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
		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
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. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	1	1
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	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	1	1	1	1	1
EXTERNAL MASS	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	1	1	1	1	1	1	1	1	1
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
M.TAIL	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	1	1	1	1	1	1	1	1	1
	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	25	25	25	25	25	25	25	25	25	25	25	25	25
	6000 ppm	25	25	25	25	25	24	24	24	24	24	24	24	24	24
	15000 ppm	25	25	25	25	25	25	25	25	25	24	24	24	23	23

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	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	1
	15000 ppm	1	1	1	1	1	1	1	1	1	1	1	1
MORIBUND SACRIFICE	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	1	1	1	1	1	1	1	1	1	1	1	1
EXTERNAL MASS	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	1	1	1	1	1	1	1	1	1	1	1	1
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
M.TAIL	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	1	1	1	1	1	1	1	1	1	1	1	1
	15000 ppm	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
	2400 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	6000 ppm	24	24	24	24	24	24	24	24	24	24	24	23
	15000 ppm	23	23	23	23	23	23	23	23	23	23	23	23

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	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
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
TRAUMA	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
INTERNAL MASS	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
SWELLING	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
HEMORRHAGE	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	25	25	25	25	25	25	25	25	25	25	25	25	25
	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	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	0	0	0	0	0	0	0	0	0	0	0
	2400 ppm	0	0	0	0	0	0	0	0	0	0	1	1
	6000 ppm	0	0	0	1	1	1	1	1	1	1	1	1
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	1	1	1	1	1
	2400 ppm	0	0	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
PARALYTIC GAIT	Control	0	0	0	0	0	0	0	0	1	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
TRAUMA	Control	0	0	0	0	0	0	0	0	1	1	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
INTERNAL MASS	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	1	0	0	0	0	0	0	0	0	0
	15000 ppm	0	0	0	0	0	0	0	0	0	0	0	0
SWELLING	Control	0	0	0	0	0	0	0	0	1	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
HEMORRHAGE	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	1	0	0	0	0	1	1
	15000 ppm	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
	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	1
NON REMARKABLE	Control	25	25	25	25	25	25	25	24	23	23	24	24
	2400 ppm	25	25	24	24	24	24	24	24	24	24	23	23
	6000 ppm	25	25	24	24	24	23	24	24	24	24	23	23
	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. : 0929 MEAN BODY WEIGHTS AND SURVIVAL
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25.0 (25)	25/25	25.0 (25)	100	25/25	25.0 (25)	100	25/25	25.0 (25)	100	25/25
1-7	25.9 (25)	25/25	25.8 (25)	100	25/25	25.5 (25)	98	25/25	26.3 (25)	102	25/25
2-7	26.3 (25)	25/25	26.4 (25)	100	25/25	26.4 (25)	100	25/25	26.9 (25)	102	25/25
3-7	27.1 (25)	25/25	26.8 (25)	99	25/25	26.8 (25)	99	25/25	27.5 (25)	101	25/25
4-7	27.4 (25)	25/25	27.4 (25)	100	25/25	27.0 (25)	99	25/25	27.6 (25)	101	25/25
5-7	27.9 (25)	25/25	27.9 (25)	100	25/25	27.6 (25)	99	25/25	28.1 (25)	101	25/25
6-7	28.4 (25)	25/25	28.1 (25)	99	25/25	27.7 (25)	98	25/25	28.2 (25)	99	25/25
7-7	29.0 (25)	25/25	28.7 (25)	99	25/25	28.2 (25)	97	25/25	28.7 (25)	99	25/25
8-7	29.2 (25)	25/25	29.0 (25)	99	25/25	28.7 (25)	98	25/25	28.9 (25)	99	25/25
9-7	29.6 (25)	25/25	29.1 (25)	98	25/25	29.1 (25)	98	25/25	29.6 (25)	100	25/25
10-7	29.9 (25)	25/25	29.6 (25)	99	25/25	29.2 (25)	98	25/25	29.4 (24)	98	24/25
11-7	29.9 (25)	25/25	29.9 (25)	100	25/25	29.6 (25)	99	25/25	30.0 (24)	100	24/25
12-7	30.5 (25)	25/25	29.8 (25)	98	25/25	29.5 (25)	97	25/25	30.1 (24)	99	24/25
13-7	30.7 (25)	25/25	30.1 (25)	98	25/25	29.9 (25)	97	25/25	30.2 (23)	98	23/25
14-7	30.5 (25)	25/25	30.4 (25)	100	25/25	30.0 (25)	98	25/25	30.3 (23)	99	23/25
15-7	31.1 (25)	25/25	30.5 (25)	98	25/25	30.4 (25)	98	25/25	30.4 (23)	98	23/25
16-7	31.1 (25)	25/25	30.6 (25)	98	25/25	30.5 (25)	98	25/25	30.7 (23)	99	23/25
17-7	31.5 (25)	25/25	31.3 (25)	99	25/25	30.6 (25)	97	25/25	30.5 (23)	97	23/25
18-7	31.5 (25)	25/25	31.3 (25)	99	25/25	30.5 (25)	97	25/25	30.8 (23)	98	23/25
19-7	31.9 (25)	25/25	31.0 (25)	97	25/25	31.2 (25)	98	25/25	31.1 (23)	97	23/25
20-7	31.6 (25)	25/25	31.3 (25)	99	25/25	31.0 (25)	98	25/25	30.8 (23)	97	23/25
21-7	32.0 (25)	25/25	31.9 (25)	100	25/25	31.2 (25)	98	25/25	31.2 (23)	98	23/25
22-7	31.9 (25)	25/25	32.1 (25)	101	25/25	31.3 (25)	98	25/25	31.4 (23)	98	23/25
23-7	32.1 (25)	25/25	31.8 (25)	99	25/25	31.5 (25)	98	25/25	31.2 (23)	97	23/25
24-7	31.9 (25)	25/25	31.9 (25)	100	25/25	31.3 (25)	98	25/25	31.4 (23)	98	23/25
25-7	32.2 (25)	25/25	32.4 (25)	101	25/25	31.9 (25)	99	25/25	31.6 (23)	98	23/25
26-7	32.7 (25)	25/25	32.7 (25)	100	25/25	32.1 (24)	98	24/25	31.8 (23)	97	23/25

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

TABLE D2

BODY WEIGHT CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0929 MEAN BODY WEIGHTS AND SURVIVAL
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	20.1 (25)	25/25	20.1 (25)	100	25/25	20.1 (25)	100	25/25	20.1 (25)	100	25/25
1-7	21.0 (25)	25/25	20.7 (25)	99	25/25	21.1 (25)	100	25/25	21.3 (25)	101	25/25
2-7	20.9 (25)	25/25	20.9 (25)	100	25/25	21.2 (25)	101	25/25	21.4 (25)	102	25/25
3-7	20.9 (25)	25/25	21.1 (25)	101	25/25	21.2 (25)	101	25/25	21.6 (25)	103	25/25
4-7	21.1 (25)	25/25	21.2 (25)	100	25/25	21.4 (25)	101	25/25	21.6 (25)	102	25/25
5-7	21.7 (25)	25/25	21.7 (25)	100	25/25	21.8 (25)	100	25/25	22.2 (25)	102	25/25
6-7	22.0 (25)	25/25	21.8 (25)	99	25/25	21.9 (25)	100	25/25	22.4 (25)	102	25/25
7-7	22.2 (25)	25/25	22.1 (25)	100	25/25	22.2 (25)	100	25/25	22.6 (25)	102	25/25
8-7	22.6 (25)	25/25	22.6 (25)	100	25/25	23.0 (25)	102	25/25	23.2 (25)	103	25/25
9-7	22.9 (25)	25/25	22.6 (25)	99	25/25	22.7 (25)	99	25/25	23.3 (25)	102	25/25
10-7	22.7 (25)	25/25	23.0 (25)	101	25/25	23.1 (25)	102	25/25	24.1 (25)	106	25/25
11-7	23.3 (25)	25/25	23.3 (25)	100	25/25	23.2 (25)	100	25/25	24.0 (25)	103	25/25
12-7	23.2 (25)	25/25	23.4 (25)	101	25/25	23.3 (25)	100	25/25	24.0 (25)	103	25/25
13-7	23.7 (25)	25/25	23.2 (25)	98	25/25	23.5 (25)	99	25/25	24.4 (25)	103	25/25
14-7	24.3 (25)	25/25	24.0 (25)	99	25/25	23.4 (25)	96	25/25	24.4 (25)	100	25/25
15-7	24.0 (25)	25/25	23.3 (25)	97	25/25	23.8 (25)	99	25/25	24.2 (25)	101	25/25
16-7	24.1 (25)	25/25	23.8 (25)	99	25/25	24.1 (25)	100	25/25	24.5 (25)	102	25/25
17-7	24.9 (25)	25/25	24.2 (24)	97	24/25	23.6 (25)	95	25/25	24.7 (25)	99	25/25
18-7	24.5 (25)	25/25	24.1 (24)	98	24/25	24.0 (24)	98	24/25	24.6 (25)	100	25/25
19-7	24.5 (25)	25/25	24.0 (24)	98	24/25	24.7 (24)	101	24/25	24.9 (25)	102	25/25
20-7	24.5 (25)	25/25	24.6 (24)	100	24/25	24.3 (24)	99	24/25	24.9 (25)	102	25/25
21-7	24.5 (25)	25/25	24.1 (24)	98	24/25	24.3 (24)	99	24/25	24.7 (25)	101	25/25
22-7	24.9 (24)	24/25	24.7 (24)	99	24/25	24.3 (24)	98	24/25	25.3 (25)	102	25/25
23-7	24.7 (24)	24/25	24.1 (24)	98	24/25	23.8 (24)	96	24/25	24.7 (25)	100	25/25
24-7	24.8 (24)	24/25	24.5 (24)	99	24/25	24.2 (24)	98	24/25	24.4 (25)	98	25/25
25-7	24.6 (24)	24/25	24.5 (23)	100	23/25	24.6 (24)	100	24/25	24.9 (25)	101	25/25
26-7	25.3 (24)	24/25	25.6 (23)	101	23/25	25.0 (24)	99	24/25	25.4 (25)	100	25/25

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

TABLE D3

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25.0± 1.3	25.9± 1.4	26.3± 1.5	27.1± 1.7	27.4± 1.5	27.9± 1.5	28.4± 1.6
2400 ppm	25.0± 1.3	25.8± 1.3	26.4± 1.4	26.8± 1.5	27.4± 1.7	27.9± 1.5	28.1± 1.6
6000 ppm	25.0± 1.3	25.5± 1.3	26.4± 1.5	26.8± 1.5	27.0± 1.7	27.6± 1.7	27.7± 1.7
15000 ppm	25.0± 1.3	26.3± 1.2	26.9± 1.4	27.5± 1.6	27.6± 1.4	28.1± 1.6	28.2± 1.7

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	29.0± 1.6	29.2± 1.5	29.6± 1.5	29.9± 1.8	29.9± 1.6	30.5± 1.7	30.7± 2.0
2400 ppm	28.7± 1.6	29.0± 1.7	29.1± 1.9	29.6± 1.8	29.9± 2.1	29.8± 2.1	30.1± 2.4
6000 ppm	28.2± 1.8	28.7± 2.1	29.1± 1.9	29.2± 2.0	29.6± 2.0	29.5± 2.0	29.9± 2.3
15000 ppm	28.7± 1.8	28.9± 1.6	29.6± 1.8	29.4± 1.9	30.0± 1.8	30.1± 1.8	30.2± 2.0

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	30.5± 1.8	31.1± 2.0	31.1± 2.0	31.5± 2.1	31.5± 2.2	31.9± 1.8	31.6± 1.9
2400 ppm	30.4± 2.4	30.5± 2.7	30.6± 2.7	31.3± 2.8	31.3± 2.9	31.0± 2.7	31.3± 2.8
6000 ppm	30.0± 2.4	30.4± 2.5	30.5± 2.3	30.6± 2.5	30.5± 2.5	31.2± 2.6	31.0± 2.5
15000 ppm	30.3± 2.0	30.4± 2.0	30.7± 1.8	30.5± 2.0	30.8± 2.1	31.1± 2.1	30.8± 2.2

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	32.0± 2.0	31.9± 1.9	32.1± 1.8	31.9± 1.8	32.2± 1.7	32.7± 1.8
2400 ppm	31.9± 3.1	32.1± 3.2	31.8± 3.3	31.9± 3.3	32.4± 3.3	32.7± 3.4
6000 ppm	31.2± 2.7	31.3± 2.5	31.5± 2.6	31.3± 2.5	31.9± 2.8	32.1± 2.6
15000 ppm	31.2± 1.9	31.4± 2.1	31.2± 2.1	31.4± 2.0	31.6± 2.1	31.8± 2.1

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

TABLE D4

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	20.1± 1.1	21.0± 1.1	20.9± 1.4	20.9± 1.2	21.1± 1.1	21.7± 1.0	22.0± 1.1
2400 ppm	20.1± 1.1	20.7± 1.2	20.9± 1.3	21.1± 1.1	21.2± 1.1	21.7± 1.2	21.8± 0.8
6000 ppm	20.1± 1.1	21.1± 1.1	21.2± 1.1	21.2± 1.0	21.4± 1.0	21.8± 0.7	21.9± 0.9
15000 ppm	20.1± 1.1	21.3± 1.2	21.4± 1.1	21.6± 0.9	21.6± 1.0	22.2± 1.0	22.4± 1.0

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	22.2± 0.7	22.6± 1.2	22.9± 1.2	22.7± 1.1	23.3± 1.0	23.2± 1.0	23.7± 1.3
2400 ppm	22.1± 1.4	22.6± 1.4	22.6± 1.4	23.0± 1.3	23.3± 1.4	23.4± 1.4	23.2± 1.4
6000 ppm	22.2± 1.1	23.0± 1.2	22.7± 1.0	23.1± 1.2	23.2± 0.9	23.3± 1.2	23.5± 1.2
15000 ppm	22.6± 0.9	23.2± 0.9	23.3± 1.3	24.1± 1.3**	24.0± 1.0*	24.0± 1.2	24.4± 1.1

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	24.3± 1.7	24.0± 1.1	24.1± 1.4	24.9± 1.6	24.5± 1.6	24.5± 1.6	24.5± 1.7
2400 ppm	24.0± 1.7	23.3± 1.2	23.8± 1.6	24.2± 1.7	24.1± 1.2	24.0± 1.1	24.6± 1.7
6000 ppm	23.4± 1.1	23.8± 1.4	24.1± 1.6	23.6± 1.1**	24.0± 1.4	24.7± 1.4	24.3± 1.6
15000 ppm	24.4± 0.9	24.2± 1.1	24.5± 1.3	24.7± 1.2	24.6± 1.3	24.9± 1.7	24.9± 1.7

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	24.5± 1.7	24.9± 1.8	24.7± 1.7	24.8± 1.4	24.6± 1.4	25.3± 1.2
2400 ppm	24.1± 1.5	24.7± 1.7	24.1± 1.3	24.5± 2.0	24.5± 1.6	25.6± 1.9
6000 ppm	24.3± 1.2	24.3± 1.3	23.8± 1.3	24.2± 1.7	24.6± 1.2	25.0± 1.3
15000 ppm	24.7± 2.1	25.3± 2.2	24.7± 2.2	24.4± 2.6	24.9± 2.8	25.4± 3.0

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

TABLE E1

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0929 MEAN FOOD CONSUMPTION(FC) AND SURVIVAL
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

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	4.0 (25)	25/25	4.1 (25)	103	25/25	4.2 (25)	105	25/25	4.5 (25)	113	25/25
2-7	3.7 (25)	25/25	3.9 (25)	105	25/25	3.9 (25)	105	25/25	4.0 (25)	108	25/25
3-7	4.0 (25)	25/25	4.0 (25)	100	25/25	4.0 (25)	100	25/25	4.4 (25)	110	25/25
4-7	4.0 (25)	25/25	4.0 (25)	100	25/25	3.9 (25)	98	25/25	4.2 (25)	105	25/25
5-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.1 (25)	100	25/25	4.5 (25)	110	25/25
6-7	4.2 (25)	25/25	4.1 (25)	98	25/25	4.1 (25)	98	25/25	4.5 (25)	107	25/25
7-7	4.2 (25)	25/25	4.2 (25)	100	25/25	4.1 (25)	98	25/25	4.5 (25)	107	25/25
8-7	4.3 (25)	25/25	4.2 (25)	98	25/25	4.2 (25)	98	25/25	4.5 (25)	105	25/25
9-7	4.3 (25)	25/25	4.2 (25)	98	25/25	4.1 (25)	95	25/25	4.6 (25)	107	25/25
10-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.1 (25)	95	25/25	4.7 (24)	109	24/25
11-7	4.2 (25)	25/25	4.3 (25)	102	25/25	4.2 (25)	100	25/25	4.7 (24)	112	24/25
12-7	4.3 (25)	25/25	4.2 (25)	98	25/25	4.2 (25)	98	25/25	4.6 (24)	107	24/25
13-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.2 (25)	98	25/25	4.6 (23)	107	23/25
14-7	4.2 (25)	25/25	4.2 (25)	100	25/25	4.2 (25)	100	25/25	4.7 (23)	112	23/25
15-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.3 (25)	100	25/25	4.7 (23)	109	23/25
16-7	4.2 (25)	25/25	4.3 (25)	102	25/25	4.2 (25)	100	25/25	4.6 (23)	110	23/25
17-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.2 (25)	98	25/25	4.6 (23)	107	23/25
18-7	4.2 (25)	25/25	4.2 (25)	100	25/25	4.2 (25)	100	25/25	4.7 (23)	112	23/25
19-7	4.3 (25)	25/25	4.0 (25)	93	25/25	4.2 (25)	98	25/25	4.7 (23)	109	23/25
20-7	4.1 (25)	25/25	4.3 (25)	105	25/25	4.2 (25)	102	25/25	4.7 (23)	115	23/25
21-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.3 (25)	100	25/25	4.8 (23)	112	23/25
22-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.1 (25)	100	25/25	4.6 (23)	112	23/25
23-7	4.4 (25)	25/25	4.4 (25)	100	25/25	4.4 (25)	100	25/25	5.2 (23)	118	23/25
24-7	4.4 (25)	25/25	4.4 (25)	100	25/25	4.4 (25)	100	25/25	5.1 (23)	116	23/25
25-7	4.5 (25)	25/25	4.5 (25)	100	25/25	4.5 (25)	100	25/25	5.0 (23)	111	23/25
26-7	4.5 (25)	25/25	4.5 (25)	100	25/25	4.5 (24)	100	24/25	4.9 (23)	109	23/25

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

TABLE E2

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0929 MEAN FOOD CONSUMPTION(FC) AND SURVIVAL
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

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	3.6 (25)	25/25	3.5 (25)	97	25/25	3.8 (25)	106	25/25	3.9 (25)	108	25/25
2-7	3.1 (25)	25/25	3.1 (25)	100	25/25	3.0 (25)	97	25/25	3.4 (25)	110	25/25
3-7	3.3 (25)	25/25	3.3 (25)	100	25/25	3.3 (25)	100	25/25	3.7 (25)	112	25/25
4-7	3.5 (25)	25/25	3.4 (25)	97	25/25	3.4 (25)	97	25/25	3.8 (25)	109	25/25
5-7	3.7 (25)	25/25	3.6 (25)	97	25/25	3.5 (25)	95	25/25	4.0 (25)	108	25/25
6-7	3.8 (25)	25/25	3.7 (25)	97	25/25	3.6 (25)	95	25/25	4.1 (25)	108	25/25
7-7	3.8 (25)	25/25	3.7 (25)	97	25/25	3.7 (25)	97	25/25	4.2 (25)	111	25/25
8-7	3.9 (25)	25/25	3.8 (25)	97	25/25	3.8 (25)	97	25/25	4.1 (25)	105	25/25
9-7	4.0 (25)	25/25	3.7 (25)	93	25/25	3.7 (25)	93	25/25	4.2 (25)	105	25/25
10-7	3.9 (25)	25/25	3.8 (25)	97	25/25	3.8 (25)	97	25/25	4.4 (25)	113	25/25
11-7	3.9 (25)	25/25	3.7 (25)	95	25/25	3.8 (25)	97	25/25	4.4 (25)	113	25/25
12-7	4.0 (25)	25/25	3.8 (25)	95	25/25	3.8 (25)	95	25/25	4.3 (25)	108	25/25
13-7	4.1 (25)	25/25	3.7 (25)	90	25/25	3.7 (25)	90	25/25	4.4 (25)	107	25/25
14-7	4.0 (25)	25/25	3.9 (25)	98	25/25	3.8 (25)	95	25/25	4.4 (25)	110	25/25
15-7	4.0 (25)	25/25	3.7 (25)	93	25/25	3.8 (25)	95	25/25	4.3 (25)	108	25/25
16-7	4.0 (25)	25/25	3.9 (25)	98	25/25	3.9 (25)	98	25/25	4.4 (25)	110	25/25
17-7	4.1 (25)	25/25	3.8 (24)	93	24/25	3.7 (25)	90	25/25	4.4 (25)	107	25/25
18-7	4.0 (25)	25/25	3.7 (24)	93	24/25	3.9 (24)	98	24/25	4.3 (25)	108	25/25
19-7	3.9 (25)	25/25	3.7 (24)	95	24/25	4.0 (24)	103	24/25	4.4 (25)	113	25/25
20-7	4.0 (25)	25/25	4.0 (24)	100	24/25	3.8 (24)	95	24/25	4.5 (25)	113	25/25
21-7	4.0 (25)	25/25	3.7 (24)	93	24/25	3.8 (24)	95	24/25	4.5 (25)	113	25/25
22-7	4.0 (24)	24/25	3.8 (24)	95	24/25	3.8 (24)	95	24/25	4.5 (25)	113	25/25
23-7	4.1 (24)	24/25	3.9 (24)	95	24/25	4.2 (24)	102	24/25	5.0 (25)	122	25/25
24-7	4.3 (24)	24/25	4.2 (24)	98	24/25	4.2 (24)	98	24/25	5.0 (25)	116	25/25
25-7	4.3 (24)	24/25	4.1 (23)	95	23/25	4.1 (24)	95	24/25	4.9 (25)	114	25/25
26-7	4.5 (24)	24/25	4.2 (23)	93	23/25	4.2 (24)	93	24/25	4.8 (25)	107	25/25

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

TABLE E3

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	4.0± 0.3	3.7± 0.3	4.0± 0.3	4.0± 0.3	4.1± 0.3	4.2± 0.3	4.2± 0.3
2400 ppm	4.1± 0.3	3.9± 0.3	4.0± 0.3	4.0± 0.3	4.1± 0.3	4.1± 0.3	4.2± 0.3
6000 ppm	4.2± 0.4	3.9± 0.3	4.0± 0.3	3.9± 0.3	4.1± 0.4	4.1± 0.3	4.1± 0.4
15000 ppm	4.5± 0.3**	4.0± 0.4**	4.4± 0.4**	4.2± 0.4	4.5± 0.4**	4.5± 0.5	4.5± 0.4*

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	4.3± 0.3	4.3± 0.3	4.3± 0.3	4.2± 0.4	4.3± 0.3	4.3± 0.4	4.2± 0.4
2400 ppm	4.2± 0.3	4.2± 0.4	4.3± 0.3	4.3± 0.3	4.2± 0.3	4.3± 0.3	4.2± 0.3
6000 ppm	4.2± 0.4	4.1± 0.3	4.1± 0.4	4.2± 0.4	4.2± 0.4	4.2± 0.4	4.2± 0.4
15000 ppm	4.5± 0.5	4.6± 0.5*	4.7± 0.5*	4.7± 0.5**	4.6± 0.5	4.6± 0.5*	4.7± 0.4**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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± 0.3	4.2± 0.3	4.3± 0.4	4.2± 0.3	4.3± 0.3	4.1± 0.4	4.3± 0.4
2400 ppm	4.3± 0.3	4.3± 0.3	4.3± 0.3	4.2± 0.3	4.0± 0.3	4.3± 0.3	4.3± 0.4
6000 ppm	4.3± 0.4	4.2± 0.3	4.2± 0.5	4.2± 0.4	4.2± 0.4	4.2± 0.5	4.3± 0.5
15000 ppm	4.7± 0.6**	4.6± 0.4**	4.6± 0.4*	4.7± 0.5**	4.7± 0.4**	4.7± 0.6**	4.8± 0.6**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	4.1± 0.3	4.4± 0.4	4.4± 0.4	4.5± 0.3	4.5± 0.4
2400 ppm	4.1± 0.3	4.4± 0.3	4.4± 0.3	4.5± 0.4	4.5± 0.3
6000 ppm	4.1± 0.4	4.4± 0.4	4.4± 0.5	4.5± 0.5	4.5± 0.5
15000 ppm	4.6± 0.5**	5.2± 0.6**	5.1± 0.6**	5.0± 0.5**	4.9± 0.5**

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

TABLE E4

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	3.6± 0.4	3.1± 0.4	3.3± 0.2	3.5± 0.4	3.7± 0.2	3.8± 0.3	3.8± 0.3
2400 ppm	3.5± 0.4	3.1± 0.3	3.3± 0.3	3.4± 0.2	3.6± 0.3	3.7± 0.4	3.7± 0.5
6000 ppm	3.8± 0.2	3.0± 0.3	3.3± 0.3	3.4± 0.3	3.5± 0.4	3.6± 0.3	3.7± 0.3
15000 ppm	3.9± 0.4*	3.4± 0.4**	3.7± 0.4**	3.8± 0.4**	4.0± 0.3**	4.1± 0.4*	4.2± 0.4**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	3.9± 0.3	4.0± 0.3	3.9± 0.4	3.9± 0.3	4.0± 0.4	4.1± 0.5	4.0± 0.5
2400 ppm	3.8± 0.4	3.7± 0.3*	3.8± 0.4	3.7± 0.3	3.8± 0.4	3.7± 0.4*	3.9± 0.4
6000 ppm	3.8± 0.4	3.7± 0.4*	3.8± 0.4	3.8± 0.3	3.8± 0.4	3.7± 0.4*	3.8± 0.4
15000 ppm	4.1± 0.4	4.2± 0.4*	4.4± 0.4**	4.4± 0.5**	4.3± 0.5*	4.4± 0.5*	4.4± 0.5*

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	4.0± 0.4	4.0± 0.4	4.1± 0.4	4.0± 0.5	3.9± 0.4	4.0± 0.5	4.0± 0.5
2400 ppm	3.7± 0.4*	3.9± 0.3	3.8± 0.5	3.7± 0.4	3.7± 0.3	4.0± 0.5	3.7± 0.4
6000 ppm	3.8± 0.4	3.9± 0.4	3.7± 0.4**	3.9± 0.4	4.0± 0.5	3.8± 0.5	3.8± 0.4
15000 ppm	4.3± 0.5**	4.4± 0.4**	4.4± 0.4*	4.3± 0.5*	4.4± 0.5**	4.5± 0.4**	4.5± 0.4**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	4.0± 0.6	4.1± 0.5	4.3± 0.5	4.3± 0.5	4.5± 0.5
2400 ppm	3.8± 0.5	3.9± 0.5	4.2± 0.5	4.1± 0.5	4.2± 0.5
6000 ppm	3.8± 0.5	4.2± 0.4	4.2± 0.6	4.1± 0.6	4.2± 0.5
15000 ppm	4.5± 0.4**	5.0± 0.6**	5.0± 0.7**	4.9± 0.5**	4.8± 0.5

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

TABLE F1

URINALYSIS : MALE

STUDY NO. : 0929 URINALYSIS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	0	0	0	0	2	0	23		0	8	16	1	0	0		25	0	0	0	0	0		1	14	9	1	0	0		25	0	0	0	0	
2400 ppm	11	0	0	0	1	0	1	9		0	8	2	1	0	0	*	11	0	0	0	0	0		3	4	3	1	0	0		11	0	0	0	0	
6000 ppm	19	0	0	2	1	0	3	13	*	0	7	12	0	0	0		19	0	0	0	0	0		4	10	4	1	0	0		19	0	0	0	0	
15000 ppm	21	0	0	3	2	2	5	9	**	0	9	11	1	0	0		21	0	0	0	0	0		6	6	7	1	1	0		21	0	0	0	0	

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

STUDY NO. : 0929 URINALYSIS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

Group Name	NO. of Animals	Urobilinogen ±	+	2+	3+	4+	CHI
Control	25	25	0	0	0	0	
2400 ppm	11	11	0	0	0	0	
6000 ppm	19	19	0	0	0	0	
15000 ppm	21	21	0	0	0	0	

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

TABLE F2

URINALYSIS : FEMALE

STUDY NO. : 0929 URINALYSIS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	21	0	0	2	2	4	10	3		2	18	1	0	0	0		21	0	0	0	0	0		0	17	4	0	0	0		21	0	0	0	0	
2400 ppm	21	0	0	3	1	5	9	3		3	11	6	1	0	0		21	0	0	0	0	0		1	14	6	0	0	0		21	0	0	0	0	
6000 ppm	22	0	0	4	4	0	8	6		4	12	6	0	0	0		22	0	0	0	0	0		1	15	5	1	0	0		21	0	0	0	0	1
15000 ppm	23	0	1	1	2	2	7	10		8	11	4	0	0	0	*	23	0	0	0	0	0		3	17	1	2	0	0		23	0	0	0	0	

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

STUDY NO. : 0929 URINALYSIS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

Group Name	NO. of Animals	Urobilinogen ±	＋	2+	3+	4+	CHI
Control	21	21	0	0	0	0	
2400 ppm	21	21	0	0	0	0	
6000 ppm	22	22	0	0	0	0	
15000 ppm	23	23	0	0	0	0	

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

TABLE G1

HEMATOLOGY : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	11.22±	0.30	16.5±	0.4	46.4±	1.1	41.4±	0.6	14.7±	0.2	35.5±	0.3	1358±	77
2400 ppm	24	11.21±	0.32	16.6±	0.4	46.6±	1.2	41.5±	0.4	14.8±	0.2*	35.7±	0.2*	1368±	100
6000 ppm	24	10.72±	1.31*	15.9±	2.0	44.8±	5.0	41.8±	0.9	14.8±	0.2	35.4±	1.0	1343±	98
15000 ppm	23	10.75±	0.75**	16.0±	1.6	45.8±	3.2	42.6±	0.5**	14.9±	0.7**	34.9±	1.7	1384±	248

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

Group Name	NO. of Animals	RETICULOCYTE %
Control	25	3.5± 0.2
2400 ppm	24	3.6± 0.2
6000 ppm	24	4.4± 3.1**
15000 ppm	23	5.1± 5.6**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	1.70±	0.70	28.6±	6.7	67.6±	6.1	1.7±	0.7	2.0±	1.6	0.0±	0.0
2400 ppm	24	1.75±	0.91	28.8±	7.3	68.0±	6.9	1.7±	0.8	1.5±	1.2	0.0±	0.1
6000 ppm	24	1.50±	0.76	30.4±	9.6	66.1±	8.8	1.6±	0.5	1.9±	1.9	0.0±	0.0
15000 ppm	23	1.76±	1.19	27.4±	10.9	68.8±	9.9	1.7±	0.8	2.1±	2.0	0.0±	0.0

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

TABLE G2

HEMATOLOGY : FEMALE

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	10.95± 0.29	16.6± 0.5	45.8± 1.1	41.9± 0.6	15.1± 0.2	36.2± 0.5	1210± 86
2400 ppm	23	11.05± 0.28	16.7± 0.4	46.3± 1.2	41.9± 0.6	15.1± 0.2	36.0± 0.4	1189± 100
6000 ppm	24	10.83± 0.33	16.3± 0.7	45.4± 1.4	41.9± 0.5	15.0± 0.3	35.8± 0.6*	1173± 118
15000 ppm	24	10.67± 0.30**	16.3± 0.5	45.7± 1.2	42.8± 0.8**	15.3± 0.3	35.7± 0.6**	1155± 92

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

Group Name	NO. of Animals	RETICULOCYTE %
Control	24	3.8± 0.8
2400 ppm	23	4.0± 0.7
6000 ppm	24	4.2± 1.4
15000 ppm	24	4.3± 0.9

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	24	1.87±	1.31	32.7±	12.2	63.8±	11.5	2.4±	1.1	1.1±	0.9	0.0±	0.0
2400 ppm	23	2.14±	1.45	28.6±	9.7	66.7±	9.0	2.5±	0.9	2.2±	1.3**	0.0±	0.0
6000 ppm	24	2.04±	1.21	33.8±	14.2	62.1±	13.3	2.4±	1.1	1.7±	1.3	0.0±	0.0
15000 ppm	24	2.22±	1.38	26.0±	12.9	68.9±	11.9	2.3±	0.9	2.8±	1.6**	0.0±	0.0

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

TABLE H1

BIOCHEMISTRY : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	5.2±	0.2	2.9±	0.1	1.3±	0.1	0.10±	0.01	213±	24	84±	7	55±	15
2400 ppm	24	5.2±	0.2	3.0±	0.1	1.3±	0.1	0.09±	0.01	209±	24	82±	7	48±	14
6000 ppm	24	5.1±	0.3	2.9±	0.2	1.3±	0.1	0.10±	0.02	223±	23	83±	13	48±	17
15000 ppm	23	5.2±	0.2	2.9±	0.2	1.3±	0.1	0.11±	0.02*	232±	35*	91±	7**	48±	15

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	175± 15	58± 10	21± 5	267± 49	206± 14	0.2± 0.2	98± 62
2400 ppm	24	169± 13	56± 8	20± 6	249± 48	216± 15	0.2± 0.2	84± 65
6000 ppm	24	169± 25	52± 13	19± 5	225± 30**	217± 25	0.2± 0.2	65± 18*
15000 ppm	23	183± 13	54± 17	20± 10	255± 92	211± 27	0.1± 0.2	85± 126**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
MEASURE. TIME : 1
SEX : MALE 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	25	22.8±	3.1	150±	2	3.6±	0.3	117±	2	8.4±	0.2	5.7±	0.9
2400 ppm	24	24.3±	4.9	150±	1	3.7±	0.3	117±	2	8.4±	0.2	5.6±	0.7
6000 ppm	24	23.2±	4.4	149±	2	3.8±	0.3	117±	3	8.4±	0.2	5.6±	0.8
15000 ppm	23	22.2±	5.2	149±	1	3.9±	0.4*	117±	2	8.6±	0.3**	5.6±	1.0

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

TABLE H2

BIOCHEMISTRY : FEMALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
MEASURE. TIME : 1
SEX : FEMALE 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±	0.2	3.1±	0.1	1.5±	0.1	0.09±	0.01	177±	29	65±	7	38±	14
2400 ppm	23	5.2±	0.2	3.1±	0.1	1.4±	0.1	0.09±	0.01	191±	34	64±	7	43±	17
6000 ppm	24	5.2±	0.3	3.1±	0.2	1.5±	0.2	0.09±	0.01	197±	22*	66±	12	37±	12
15000 ppm	24	5.1±	0.2	3.1±	0.1	1.5±	0.1	0.10±	0.02	221±	23**	68±	7	42±	14

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
MEASURE. TIME : 1
SEX : FEMALE 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	132± 15	90± 34	25± 7	232± 59	349± 48	0.1± 0.1	95± 31
2400 ppm	23	132± 14	69± 18*	21± 5	214± 36	324± 45	0.1± 0.1	83± 36
6000 ppm	24	134± 23	87± 49	22± 9	246± 90	337± 60	0.2± 0.2	92± 32
15000 ppm	24	132± 15	60± 19**	21± 8	195± 40	298± 30**	0.2± 0.2	62± 18**

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	18.0± 3.5	149± 2	3.2± 0.2	118± 1	8.7± 0.2	5.6± 0.9
2400 ppm	23	17.1± 2.6	149± 2	3.2± 0.3	117± 1	8.7± 0.3	5.4± 0.8
6000 ppm	24	17.3± 3.2	149± 2	3.3± 0.3	118± 2	8.7± 0.2	5.4± 0.9
15000 ppm	24	16.2± 4.1	149± 1	3.4± 0.3**	117± 2	8.8± 0.2	4.7± 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 (%)
lung	white zone		1 (4)	2 (8)	2 (8)	7 (28)
	nodule		2 (8)	2 (8)	1 (4)	7 (28)
spleen	enlarged		0 (0)	0 (0)	0 (0)	2 (8)
	black zone		4 (16)	2 (8)	0 (0)	5 (20)
	nodule		1 (4)	0 (0)	1 (4)	0 (0)
stomach	forestomach:white zone		0 (0)	2 (8)	4 (16)	9 (36)
	glandular stomach:nodule		0 (0)	1 (4)	0 (0)	0 (0)
small intes	red zone		0 (0)	0 (0)	1 (4)	0 (0)
	nodule		0 (0)	0 (0)	0 (0)	1 (4)
	dilated		0 (0)	0 (0)	1 (4)	0 (0)
large intes	nodule		0 (0)	1 (4)	0 (0)	0 (0)
liver	white zone		2 (8)	1 (4)	0 (0)	0 (0)
	red zone		0 (0)	0 (0)	1 (4)	0 (0)
kidney	white zone		0 (0)	0 (0)	1 (4)	0 (0)
urin bladd	urine:marked retention		0 (0)	0 (0)	0 (0)	1 (4)
testis	absence		1 (4)	0 (0)	0 (0)	0 (0)
Harder gl	nodule		1 (4)	0 (0)	0 (0)	0 (0)
abdominal c	hemorrhage		0 (0)	0 (0)	0 (0)	1 (4)
thoracic ca	hemorrhage		0 (0)	0 (0)	0 (0)	1 (4)
other	tail:nodule		0 (0)	0 (0)	1 (4)	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 (%)
subcutis	red zone		0 (0)	0 (0)	0 (0)	1 (4)
	hemorrhage		1 (4)	0 (0)	0 (0)	0 (0)
	edema		1 (4)	1 (4)	0 (0)	0 (0)
lung	white zone		0 (0)	4 (16)	2 (8)	5 (20)
	nodule		3 (12)	5 (20)	6 (24)	5 (20)
spleen	enlarged		1 (4)	1 (4)	0 (0)	0 (0)
	black zone		1 (4)	2 (8)	5 (20)	3 (12)
	nodule		0 (0)	1 (4)	1 (4)	3 (12)
oral cavity	nodule		0 (0)	0 (0)	0 (0)	1 (4)
stomach	forestomach:white zone		3 (12)	1 (4)	4 (16)	19 (76)
	glandular stomach:red zone		0 (0)	1 (4)	0 (0)	0 (0)
large intes	nodule		0 (0)	0 (0)	1 (4)	0 (0)
liver	red zone		0 (0)	0 (0)	1 (4)	0 (0)
uterus	red zone		1 (4)	0 (0)	0 (0)	1 (4)
muscle	red		0 (0)	1 (4)	0 (0)	0 (0)
	black		1 (4)	0 (0)	0 (0)	0 (0)
abdominal c	hemorrhage		0 (0)	1 (4)	1 (4)	0 (0)
thoracic ca	pleural fluid		0 (0)	2 (8)	0 (0)	0 (0)
whole body	anemic		0 (0)	1 (4)	0 (0)	0 (0)

TABLE J1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	28.6± 1.7	0.016± 0.003	0.252± 0.037	0.185± 0.014	0.167± 0.015	0.584± 0.054
2400 ppm	25	28.6± 3.3	0.016± 0.003	0.252± 0.040	0.187± 0.017	0.170± 0.009	0.587± 0.034
6000 ppm	24	28.2± 2.4	0.015± 0.003	0.240± 0.041	0.185± 0.019	0.167± 0.015	0.592± 0.054
15000 ppm	23	28.1± 2.0	0.014± 0.003	0.265± 0.042	0.188± 0.015	0.193± 0.085	0.630± 0.048**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1
SEX : MALE
UNIT: g

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

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	25	0.075± 0.019	1.234± 0.085	0.470± 0.015
2400 ppm	25	0.071± 0.010	1.249± 0.090	0.474± 0.013
6000 ppm	24	0.076± 0.025	1.240± 0.113	0.476± 0.018
15000 ppm	23	0.087± 0.053	1.289± 0.108	0.478± 0.010

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

TABLE J2

ORGAN WEIGHT, ABSOLUTE : FEMALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	24	21.4± 1.2	0.016± 0.003	0.039± 0.008	0.146± 0.010	0.164± 0.011	0.425± 0.028
2400 ppm	23	21.7± 1.6	0.016± 0.003	0.036± 0.008	0.141± 0.009	0.169± 0.028	0.411± 0.025
6000 ppm	24	21.2± 1.2	0.015± 0.002	0.036± 0.006	0.149± 0.037	0.161± 0.011	0.408± 0.028
15000 ppm	25	22.0± 2.4*	0.016± 0.003	0.038± 0.011	0.145± 0.016	0.209± 0.199	0.421± 0.046

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

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

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	24	0.086± 0.010	1.054± 0.082	0.489± 0.016
2400 ppm	23	0.090± 0.014	1.094± 0.114	0.496± 0.019
6000 ppm	24	0.088± 0.021	1.056± 0.084	0.485± 0.017
15000 ppm	25	0.094± 0.019*	1.097± 0.144	0.484± 0.023

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

TABLE K1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	25	28.6± 1.7	0.055± 0.010	0.885± 0.128	0.648± 0.025	0.586± 0.043	2.045± 0.163
2400 ppm	25	28.6± 3.3	0.055± 0.011	0.889± 0.166	0.659± 0.072	0.598± 0.062	2.065± 0.155
6000 ppm	24	28.2± 2.4	0.054± 0.011	0.852± 0.144	0.655± 0.057	0.593± 0.051	2.100± 0.128
15000 ppm	23	28.1± 2.0	0.051± 0.008	0.944± 0.155	0.669± 0.034	0.691± 0.326	2.243± 0.096**

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1
SEX : MALE
UNIT: %

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

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	25	0.261± 0.058	4.321± 0.199	1.651± 0.094
2400 ppm	25	0.248± 0.036	4.383± 0.239	1.674± 0.161
6000 ppm	24	0.273± 0.099	4.397± 0.266	1.694± 0.114
15000 ppm	23	0.310± 0.200	4.586± 0.207**	1.707± 0.115

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

TABLE K2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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	24	21.4± 1.2	0.073± 0.012	0.182± 0.033	0.683± 0.039	0.770± 0.058	1.990± 0.120
2400 ppm	23	21.7± 1.6	0.072± 0.011	0.167± 0.031	0.649± 0.048	0.781± 0.126	1.896± 0.122*
6000 ppm	24	21.2± 1.2	0.070± 0.010	0.168± 0.030	0.700± 0.150	0.761± 0.044	1.925± 0.132
15000 ppm	25	22.0± 2.4*	0.074± 0.011	0.169± 0.041	0.660± 0.037	1.029± 1.338	1.913± 0.089

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

STUDY NO. : 0929
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

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

Group Name	NO. of Animals	SPLEEN	LIVER	BRAIN
Control	24	0.402± 0.045	4.932± 0.234	2.296± 0.135
2400 ppm	23	0.413± 0.049	5.031± 0.258	2.290± 0.156
6000 ppm	24	0.411± 0.088	4.978± 0.257	2.292± 0.122
15000 ppm	25	0.425± 0.067	4.969± 0.270	2.219± 0.214**

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
{Integumentary system/appandage}						
skin/app			<25>	<25>	<25>	<25>
	squamous cell papilloma		0 (0%)	0 (0%)	1 (4%)	0 (0%)
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	bronchiolar-alveolar adenoma		4 (16%)	6 (24%)	3 (12%)	10 (40%)
	bronchiolar-alveolar carcinoma		1 (4%)	1 (4%)	0 (0%)	5 (20%)
{Hematopoietic system}						
spleen			<25>	<25>	<25>	<25>
	hemangioma		0 (0%)	0 (0%)	1 (4%)	1 (4%)
	malignant lymphoma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
	hemangiosarcoma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
{Digestive system}						
small intes			<25>	<25>	<25>	<25>
	hemangiosarcoma		0 (0%)	0 (0%)	1 (4%)	1 (4%)
large intes			<25>	<25>	<25>	<25>
	hemangiosarcoma		0 (0%)	1 (4%)	0 (0%)	0 (0%)
liver			<25>	<25>	<25>	<25>
	hepatocellular adenoma		2 (8%)	1 (4%)	0 (0%)	0 (0%)
{Urinary system}						
urethra			<25>	<25>	<25>	<25>
	transitional cell carcinoma		0 (0%)	0 (0%)	0 (0%)	1 (4%)

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

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of animals on Study	25	25	25	25
{Special sense organs/appendage}						
Harder gl			<25>	<25>	<25>	<25>
	adenoma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
	adenocarcinoma		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 L2

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : FEMALE

Organ	Findings	Group Name / No. of animals on Study	Control 25	2400 ppm 25	6000 ppm 25	15000 ppm 25
{Integumentary system/appandage}						
subcutis			<25>	<25>	<25>	<25>
	hemangiosarcoma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	bronchiolar-alveolar adenoma		3 (12%)	6 (24%)	6 (24%)	6 (24%)
	bronchiolar-alveolar carcinoma		3 (12%)	2 (8%)	2 (8%)	3 (12%)
{Hematopoietic system}						
thymus			<25>	<25>	<25>	<25>
	malignant lymphoma		0 (0%)	1 (4%)	0 (0%)	0 (0%)
spleen			<25>	<25>	<25>	<25>
	hemangioma		0 (0%)	0 (0%)	1 (4%)	0 (0%)
	hemangiosarcoma		0 (0%)	1 (4%)	1 (4%)	3 (12%)
{Digestive system}						
oral cavity			<25>	<25>	<25>	<25>
	squamous cell papilloma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
stomach			<25>	<25>	<25>	<25>
	squamous cell papilloma		2 (8%)	1 (4%)	0 (0%)	0 (0%)
	hemangioma		0 (0%)	1 (4%)	0 (0%)	0 (0%)
large intes			<25>	<25>	<25>	<25>
	hemangiosarcoma		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

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of animals on Study	25	25	25	25
{Urinary system}						
urethra			<25>	<25>	<25>	<25>
	transitional cell papilloma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
{Reproductive system}						
uterus			<25>	<25>	<25>	<25>
	stromal polyp		0 (0%)	1 (4%)	0 (0%)	1 (4%)
{Special sense organs/appendage}						
Harder gl			<25>	<25>	<25>	<25>
	adenoma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
	adenocarcinoma		1 (4%)	1 (4%)	0 (0%)	0 (0%)
{Musculoskeletal system}						
muscle			<25>	<25>	<25>	<25>
	hemangiosarcoma		1 (4%)	1 (4%)	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 : lung				
TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	4/25(16.0)	6/25(24.0)	3/25(12.0)	10/25(40.0)
Adjusted rates(b)	16.00	24.00	12.50	43.48
Terminal rates(c)	4/25(16.0)	6/25(24.0)	3/24(12.5)	10/23(43.5)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.0157*			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.0460*			
Fisher Exact test(e)		P = 0.3626	P = 0.5000	P = 0.0568
SITE : lung				
TUMOR : bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	1/25(4.0)	0/25(0.0)	5/25(20.0)
Adjusted rates(b)	4.00	4.00	0.00	21.74
Terminal rates(c)	1/25(4.0)	1/25(4.0)	0/24(0.0)	5/23(21.7)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.0067**			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.0122*			
Fisher Exact test(e)		P = 0.7551	P = 0.5000	P = 0.0947
SITE : liver				
TUMOR : hepatocellular adenoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	1/25(4.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.00	4.00	0.00	0.00
Terminal rates(c)	2/25(8.0)	1/25(4.0)	0/24(0.0)	0/23(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9575			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.1191			
Fisher Exact test(e)		P = 0.5000	P = 0.2449	P = 0.2449

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : ALL SITE				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	1/25(4.0)	2/25(8.0)	2/25(8.0)
Adjusted rates(b)	4.00	4.00	4.17	8.70
Terminal rates(c)	1/25(4.0)	1/25(4.0)	1/24(4.2)	2/23(8.7)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.3741			
Prevalence method(d)	P = 0.2086			
Combined analysis(d)	P = 0.2204			
Cochran-Armitage test(e)	P = 0.4919			
Fisher Exact test(e)		P = 0.7551	P = 0.5000	P = 0.5000
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma,bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	4/25(16.0)	7/25(28.0)	3/25(12.0)	13/25(52.0)
Adjusted rates(b)	16.00	28.00	12.50	56.52
Terminal rates(c)	4/25(16.0)	7/25(28.0)	3/24(12.5)	13/23(56.5)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.0012**			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.0045**			
Fisher Exact test(e)		P = 0.2481	P = 0.5000	P = 0.0078**

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : Harderian gland				
TUMOR : adenoma,adenocarcinoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	0/25(0.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.00	0.00	0.00	0.00
Terminal rates(c)	2/25(8.0)	0/25(0.0)	0/24(0.0)	0/23(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9643			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.1425			
Fisher Exact test(e)		P = 0.2449	P = 0.2449	P = 0.2449

(a): Number of tumor-bearing 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 rates.
? : 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)	3/25(12.0)	6/25(24.0)	6/25(24.0)	6/25(24.0)
Adjusted rates(b)	12.50	26.09	24.00	24.00
Terminal rates(c)	3/24(12.5)	6/23(26.1)	5/24(20.8)	6/25(24.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.2351			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.4495			
Fisher Exact test(e)		P = 0.2317	P = 0.2317	P = 0.2317
SITE : lung				
TUMOR : bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	3/25(12.0)	2/25(8.0)	2/25(8.0)	3/25(12.0)
Adjusted rates(b)	12.50	8.70	8.33	12.00
Terminal rates(c)	3/24(12.5)	2/23(8.7)	2/24(8.3)	3/25(12.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.4430			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.8469			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.6664
SITE : spleen				
TUMOR : hemangiosarcoma				
Tumor rate				
Overall rates(a)	0/25(0.0)	1/25(4.0)	1/25(4.0)	3/25(12.0)
Adjusted rates(b)	0.00	4.35	0.00	12.00
Terminal rates(c)	0/24(0.0)	1/23(4.3)	0/24(0.0)	3/25(12.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.3909			
Prevalence method(d)	P = 0.0261*			
Combined analysis(d)	P = 0.0400*			
Cochran-Armitage test(e)	P = 0.0518			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.1173

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : stomach				
TUMOR : squamous cell papilloma				
Tumor rate				
Overall rates(a)	2/25(8.0)	1/25(4.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.33	4.35	0.00	0.00
Terminal rates(c)	2/24(8.3)	1/23(4.3)	0/24(0.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9626			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.1191			
Fisher Exact test(e)		P = 0.5000	P = 0.2449	P = 0.2449
SITE : ALL SITE				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	3/25(12.0)	3/25(12.0)	4/25(16.0)
Adjusted rates(b)	0.00	8.70	8.33	16.00
Terminal rates(c)	0/24(0.0)	2/23(8.7)	2/24(8.3)	4/25(16.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.8073			
Prevalence method(d)	P = 0.0383*			
Combined analysis(d)	P = 0.1474			
Cochran-Armitage test(e)	P = 0.2422			
Fisher Exact test(e)		P = 0.3046	P = 0.3046	P = 0.1743
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma,bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	6/25(24.0)	7/25(28.0)	8/25(32.0)	7/25(28.0)
Adjusted rates(b)	25.00	30.43	32.00	28.00
Terminal rates(c)	6/24(25.0)	7/23(30.4)	7/24(29.2)	7/25(28.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.4188			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.8146			
Fisher Exact test(e)		P = 0.5000	P = 0.3768	P = 0.5000

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
SITE : spleen				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	0/25(0.0)	1/25(4.0)	2/25(8.0)	3/25(12.0)
Adjusted rates(b)	0.00	4.35	4.17	12.00
Terminal rates(c)	0/24(0.0)	1/23(4.3)	1/24(4.2)	3/25(12.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.3909			
Prevalence method(d)	P = 0.0392*			
Combined analysis(d)	P = 0.0525			
Cochran-Armitage test(e)	P = 0.0725			
Fisher Exact test(e)		P = 0.5000	P = 0.2449	P = 0.1173

(a): Number of tumor-bearing 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 rates.
? : 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

HISTOPATHOLOGICAL FINDINGS : METASTASIS OF TUMOR (SUMMARY)
ALL ANIMALS (0- 27W)

Organ	Findings	Group Name	Control	2400 ppm	6000 ppm	15000 ppm
		No. of Animals on Study	25	25	25	25
{Hematopoietic system}						
thymus			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1

< 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	Control	2400 ppm	6000 ppm	15000 ppm
		No. of Animals on Study	25	25	25	25
{Integumentary system/appandage}						
subcutis			<25>	<25>	<25>	<25>
	metastasis:muscle tumor		1	0	0	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+ (%)
{Respiratory system}																		
nasal cavit			<25>				<25>				<25>				<25>			
	proliferation:gland		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)
	eosinophilic change:olfactory epithelium		2 (8)	0 (0)	0 (0)	0 (0)	7 (28)	0 (0)	0 (0)	0 (0)	2 (8)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	eosinophilic change:respiratory epithelium		1 (4)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	inflammation:respiratory epithelium		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)	0 (0)
	respiratory metaplasia:gland		14 (56)	0 (0)	0 (0)	0 (0)	14 (56)	0 (0)	0 (0)	0 (0)	11 (44)	0 (0)	0 (0)	0 (0)	13 (52)	0 (0)	0 (0)	0 (0)
	hyperplasia:respiratory epithelium		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)	1 (4)	0 (0)	0 (0)
nasopharynx			<25>				<25>				<25>				<25>			
	eosinophilic change		2 (8)	0 (0)	0 (0)	0 (0)	6 (24)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
lung			<25>				<25>				<25>				<25>			
	hemorrhage		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)	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+ (%)
{Respiratory system}																		
lung			<25>				<25>				<25>				<25>			
	granulation		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)
	inflammation: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)
	bronchiolar-alveolar cell hyperplasia		1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)
	atelectasis		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)
	vacuolic change:bronchial epithelium		0 (0)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	3 (12)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)
	eosinophilic crystalline pneumonia		0 (0)	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)
{Hematopoietic system}																		
thymus			<25>				<25>				<25>				<25>			
	hyperplasia:epithelium		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)	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 / 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+ (%)
{Hematopoietic system}																		
spleen			<25>				<25>				<25>				<25>			
	deposit of melanin		4	0	0	0	2	0	0	0	0	0	0	0	4	0	0	0
			(16)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
	fibrosis		3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
			(12)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	extramedullary hematopoiesis		0	0	0	0	0	0	0	0	1	0	0	0	0	0	1	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(4)	(0)
{Digestive system}																		
salivary gl			<25>				<25>				<25>				<25>			
	atrophy		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)
stomach			<25>				<25>				<25>				<25>			
	erosion:forestomach		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)
	ulcer:forestomach		0	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(8)	(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 / 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+ (%)
{Digestive system}																		
stomach		No. of Animals on Study	25				25				25				25			
	hyperplasia:forestomach	<25>	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	2 (8)	0 (0)	2 (8)	5 (20)	3 (12)	0 (0) **
	erosion:glandular stomach		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)	1 (4)	0 (0)	0 (0)
	hyperplasia:glandular stomach		0 (0)	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)
	dysplasia:glandular stomach		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)
liver		<25>					<25>				<25>				<25>			
	angiectasis		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)	0 (0)
	necrosis:focal		0 (0)	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)
	clear cell focus		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)
	basophilic 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)

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
{Digestive system}						
gall bladd			<25>	<25>	<25>	<25>
	hyperplasia	1+	0 (0)	0 (0)	0 (0)	0 (0)
		2+	0 (0)	0 (0)	0 (0)	1 (4)
		3+	0 (0)	0 (0)	0 (0)	0 (0)
		4+	0 (0)	0 (0)	0 (0)	0 (0)
{Urinary system}						
kidney			<25>	<25>	<25>	<25>
	hyaline cast	1+	0 (0)	0 (0)	0 (0)	1 (4)
		2+	0 (0)	0 (0)	0 (0)	0 (0)
		3+	0 (0)	0 (0)	0 (0)	0 (0)
		4+	0 (0)	0 (0)	0 (0)	0 (0)
	regeneration:renal tubule	1+	2 (8)	1 (4)	0 (0)	0 (0)
		2+	0 (0)	0 (0)	0 (0)	0 (0)
		3+	0 (0)	0 (0)	0 (0)	0 (0)
		4+	0 (0)	0 (0)	0 (0)	0 (0)
urin bladd			<25>	<25>	<25>	<25>
	simple hyperplasia:transitional epithelium	1+	0 (0)	0 (0)	0 (0)	1 (4)
		2+	0 (0)	0 (0)	0 (0)	0 (0)
		3+	0 (0)	0 (0)	0 (0)	0 (0)
		4+	0 (0)	0 (0)	0 (0)	0 (0)
{Endocrine system}						
pituitary			<25>	<25>	<25>	<25>
	cyst	1+	0 (0)	2 (8)	1 (4)	1 (4)
		2+	0 (0)	0 (0)	0 (0)	0 (0)
		3+	0 (0)	0 (0)	0 (0)	0 (0)
		4+	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+ (%)
{Endocrine system}																		
pituitary		<25>																
	hyperplasia:intermediate lobe		0 (0)	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)
parathyroid		<25>					<25>				<24>				<25>			
	ultimobranchial body remanet		2 (8)	0 (0)	0 (0)	0 (0)	4 (16)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)
adrenal		<25>					<25>				<25>				<25>			
	hemorrhage		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)	1 (4)	0 (0)	0 (0)
	spindle-cell hyperplasia		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)
{Reproductive system}																		
testis		<25>					<25>				<25>				<25>			
	aplasia		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)
	degeneration:seminiferous epithelium		2 (8)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	2 (8)	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 / 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+ (%)
{Reproductive system}																		
testis			<25>				<25>				<25>				<25>			
	tubular atrophy		0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)
	dilatation:rete testis		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)	1 (4)	0 (0)	0 (0)
epididymis			<25>				<25>				<25>				<25>			
	deformity		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)
	spermatogenic granuloma		2 (8)	1 (4)	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)
	decreased:sperma		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)
	debris of spermatic elements		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)
prostate			<25>				<25>				<25>				<25>			
	inflammation		0 (0)	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+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Musculoskeletal system}																		
bone			<25>				<25>				<25>				<25>			
	necrosis:focal		1	0	0	0	1	0	0	0	1	0	0	0	2	0	0	0
			(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(8)	(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+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Respiratory system}																		
nasal cavit			<25>				<25>				<25>				<25>			
	eosinophilic change:olfactory epithelium		7	0	0	0	7	0	0	0	4	0	0	0	0	0	0	0 *
			(28)	(0)	(0)	(0)	(28)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	eosinophilic change:respiratory epithelium		7	0	0	0	8	0	0	0	2	0	0	0	3	0	0	0
			(28)	(0)	(0)	(0)	(32)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(12)	(0)	(0)	(0)
	respiratory metaplasia:olfactory epithelium		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)
	respiratory metaplasia:gland		14	0	0	0	16	0	0	0	18	0	0	0	19	0	0	0
			(56)	(0)	(0)	(0)	(64)	(0)	(0)	(0)	(72)	(0)	(0)	(0)	(76)	(0)	(0)	(0)
	hyperplasia:respiratory epithelium		0	0	0	0	1	0	0	0	1	0	0	0	1	1	0	0
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(4)	(0)	(0)
nasopharynx			<25>				<25>				<25>				<25>			
	eosinophilic change		12	0	0	0	11	0	0	0	5	0	0	0	2	0	0	0 **
			(48)	(0)	(0)	(0)	(44)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
lung			<25>				<25>				<25>				<25>			
	inflammation:focal		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)

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+ (%)
{Respiratory system}																		
lung			<25>				<25>				<25>				<25>			
	bronchiolar-alveolar cell hyperplasia		0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	vacuolic change:bronchial epithelium		0 (0)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)
	eosinophilic crystalline pneumonia		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)	1 (4)	0 (0)	0 (0)
{Hematopoietic system}																		
thymus			<25>				<25>				<25>				<25>			
	atrophy		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)
	hyperplasia:epithelium		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)
spleen			<25>				<25>				<25>				<25>			
	deposit of melanin		1 (4)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	4 (16)	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+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Hematopoietic system}																		
spleen			<25>				<25>				<25>				<25>			
	fibrosis		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)
	extramedullary hematopoiesis		0	0	1	0	0	0	1	0	1	0	0	0	0	0	0	0
			(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	endothelial cell hyperplasia		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)
{Digestive system}																		
salivary gl			<25>				<25>				<25>				<25>			
	atrophy		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)
stomach			<25>				<25>				<25>				<25>			
	ulcer:forestomach		0	0	0	0	1	0	0	0	0	0	0	0	5	1	0	0 *
			(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(20)	(4)	(0)	(0)
	hyperplasia:forestomach		2	0	0	0	0	0	0	0	4	2	0	0	9	7	3	0 **
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(16)	(8)	(0)	(0)	(36)	(28)	(12)	(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+ (%)
{Digestive system}																		
stomach			<25>				<25>				<25>				<25>			
	erosion: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)
	hyperplasia:glandular stomach		0 (0)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
large intes			<25>				<25>				<25>				<25>			
	ulcer		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>				<25>				<25>				<25>			
	angiectasis		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)	0 (0)
	inflammatory cell nest		1 (4)	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)	0 (0)
	extramedullary hematopoiesis		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)
	basophilic cell focus		0 (0)	0 (0)	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)

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>			
	angiectasis		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)
	hyaline cast		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)
	degeneration		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)	(0)	(0)	(0)
{Endocrine system}																		
pituitary			<25>				<25>				<25>				<25>			
	cyst		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)
parathyroid			<25>				<25>				<25>				<25>			
	ultimobranchial body remanet		1	0	0	0	2	0	0	0	1	0	0	0	0	0	0	0
			(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
adrenal			<25>				<25>				<25>				<25>			
	spindle-cell hyperplasia		14	0	0	0	12	0	0	0	15	0	0	0	15	0	0	0
			(56)	(0)	(0)	(0)	(48)	(0)	(0)	(0)	(60)	(0)	(0)	(0)	(60)	(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+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
{Reproductive system}																		
uterus			<25>				<25>				<25>				<25>			
	atrophy		1	0	0	0	1	0	0	0	1	0	0	0	3	0	0	0
			(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(12)	(0)	(0)	(0)
	cystic endometrial hyperplasia		5	0	0	0	1	0	0	0	4	0	0	0	4	0	0	0
			(20)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
	vascular anomality		1	0	0	0	1	0	0	0	0	0	0	0	1	0	0	0
			(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
vagina			<25>				<25>				<25>				<25>			
	mucinous degeneration		3	0	0	0	7	0	0	0	3	0	0	0	4	0	0	0
			(12)	(0)	(0)	(0)	(28)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
{Special sense organs/appendage}																		
Harder gl			<25>				<25>				<25>				<25>			
	atypical 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)
nasolacr d			<25>				<25>				<25>				<25>			
	hyperplasia:gland		0	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)

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>			
	necrosis:focal		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)

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

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : CByB6F1-Tg(HRAS)2Jic(tg/wt) MALE MICE

TABLE P1 HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : CByB6F1-Tg(HRAS)2Jic(tg/wt) MALE MICE

Organs Tumors	No. of animals examined	No. of animals bearing tumor	Incidence (%)	Range (%)
Lung	225			
Bronchiolar-alveolar adenoma (A)		13	5.8	0 － 12
Bronchiolar-alveolar carcinoma (B)		1	0.4	0 － 4
A and/or B		14	6.2	0 － 12

8 carcinogenicity studies examined in Japan Bioassay Research Center were used.
Study No. : 0878, 0886, 0887, 0900, 0905, 0912, 0919, 0933

TABLE P2

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : CByB6F1-Tg(HRAS)2Jic(tg/wt) FEMALE MICE

TABLE P2 HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : CByB6F1-Tg(HRAS)2Jic(tg/wt) FEMALE MICE

Organs Tumors	No. of animals examined	No. of animals bearing tumor	Incidence (%)	Range (%)
Spleen	224			
Hemangioma (A)		6	2.7	0 － 8
Hemangiosarcoma (B)		3	1.3	0 － 4
A and/or B		8	3.6	0 － 8
All organ	224			
Hemangioma (C)		13	5.8	4 － 8
Hemangiosarcoma (D)		10	4.5	0 － 8
C and/or D		22	9.8	4 － 16

8 carcinogenicity studies examined in Japan Bioassay Research Center were used.
Study No. : 0878*, 0886, 0887, 0901, 0905, 0913, 0919, 0934
* : 49 animals were used.

TABLE Q1

CAUSE OF DEATH : MALE

STUDY NO. : 0929 COUSE OF DEATH (SUMMARY)
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt) (0- 27W)
SEX : MALE

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
Number of Dead and Moribund Animal	0	0	1	2
tumor d:leukemia	0	0	0	1
tumor d:small intes	0	0	1	0
tumor d:urethra	0	0	0	1

TABLE Q2

CAUSE OF DEATH : FEMALE

Group Name	Control	2400 ppm	6000 ppm	15000 ppm
Number of Dead and Moribund Animal	1	2	1	0
tumor d:leukemia	0	1	0	0
tumor d:spleen	0	0	1	0
tumor d:muscle	1	1	0	0