アリルアルコールの 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：0926

TABLES

TABLES

TABLES (CONTINUED)

TABLES (CONTINUED)

TABLE A

CONCENTRATIONS OF ALLYL ALCOHOL

IN THE INHALATION CHAMBER

TABLE A CONCENTRATIONS OF ALLYL ALCOHOL IN THE INHALATION CHAMBER

Group Name	Concentration (ppm) Mean ± S.D.	
Control	0.00 ± 0.00	
0.15 ppm	0.15 ± 0.00	
0.5 ppm	0.50 ± 0.01	
1.5 ppm	1.49 ± 0.01	(0.99 ± 0.01) [1)]

1) This value is the measured concentration of allyl alcohol in the inhalation chamber for the first three exposures set at the target concentration of 1 ppm.

TABLE B1

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0926 SURVIVAL ANIMAL NUMBERS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1 27
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
0.15 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
0.5 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
1.5 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. : 0926 SURVIVAL ANIMAL NUMBERS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1 27
SEX : MALE

Group Name	Animals At start	Administration (Weeks) 14	15	16	17	18	19	20	21	22	23	24	25	26	27
Control	25	25/25	25/25	25/25	25/25	25/25	25/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25	24/25
		100.0	100.0	100.0	100.0	100.0	100.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0	96.0
0.15 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	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	100.0	96.0
0.5 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
1.5 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	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	96.0

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

TABLE B2

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0926 SURVIVAL ANIMAL NUMBERS
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
REPORT TYPE : A1 27
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
0.15 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
0.5 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
1.5 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	27
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
0.15 ppm	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	24/25	24/25
		100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	100.0	96.0	96.0	96.0
0.5 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
1.5 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(%)

TABLE C1

CLINICAL OBSERVATION : MALE

STUDY NO. : 0926
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
MORIBUND SACRIFICE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	1
PILOERECTION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	1	0
TORTICOLLIS	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 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
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	1	1	0
NON REMARKABLE	Control	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	0.15 ppm	24	24	24	24	24	24	24	24	24	24	24	24	24	24
	0.5 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	1.5 ppm	25	25	25	25	25	25	25	25	25	25	25	24	24	24

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
MORIBUND SACRIFICE	Control	0	0	0	0	0	1	1	1	1	1	1	1
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	1	1	1	1	1	1	1	1	1	1	1	1
PILOERECTION	Control	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
TORTICOLLIS	Control	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	1	1	1	1	1	1	1	1	1	1	1	1
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 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
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
NON REMARKABLE	Control	25	25	25	25	25	24	24	24	24	24	24	24
	0.15 ppm	24	24	24	24	24	24	24	24	24	24	24	24
	0.5 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	1.5 ppm	24	24	24	24	24	24	24	24	24	24	24	24

TABLE C2

CLINICAL OBSERVATION : FEMALE

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
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 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
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
CICATRIX	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
VAGINAL PROLAPSE	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 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
	0.15 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	0.5 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	1.5 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
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	1	1
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	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
	0.15 ppm	0	0	0	0	1	1	1	1	1	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
CICATRIX	Control	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	1	1	1
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
VAGINAL PROLAPSE	Control	0	0	0	0	0	0	0	0	0	0	0	0
	0.15 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	0.5 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	1.5 ppm	0	0	0	0	1	1	1	1	1	1	1	1
NON REMARKABLE	Control	25	25	25	25	25	25	25	25	25	25	25	25
	0.15 ppm	25	25	25	25	24	24	24	24	24	24	23	23
	0.5 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	1.5 ppm	25	25	25	25	24	24	24	24	24	24	24	24

TABLE D1

BODY WEIGHT CHANGES AND

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0926
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
UNIT : g
REPORT TYPE : A1 26
SEX : MALE

MEAN BODY WEIGHTS AND SURVIVAL

Week-Day on Study	Control Av.Wt.	Control No.of Surviv. <25>	0.15 ppm Av.Wt.	0.15 ppm % of cont. <25>	0.15 ppm No.of Surviv.	0.5 ppm Av.Wt.	0.5 ppm % of cont. <25>	0.5 ppm No.of Surviv.	1.5 ppm Av.Wt.	1.5 ppm % of cont. <25>	1.5 ppm No.of Surviv.
0-0	25.6 (25)	25/25	25.6 (25)	100	25/25	25.6 (25)	100	25/25	25.6 (25)	100	25/25
1-7	26.0 (25)	25/25	26.0 (25)	100	25/25	25.6 (25)	98	25/25	24.7 (25)	95	25/25
2-7	27.0 (25)	25/25	26.6 (25)	99	25/25	26.2 (25)	97	25/25	25.2 (25)	93	25/25
3-7	27.8 (25)	25/25	27.5 (25)	99	25/25	27.2 (25)	98	25/25	25.4 (25)	91	25/25
4-7	28.4 (25)	25/25	28.1 (25)	99	25/25	27.6 (25)	97	25/25	25.7 (25)	90	25/25
5-7	29.1 (25)	25/25	28.6 (25)	98	25/25	28.2 (25)	97	25/25	26.1 (25)	90	25/25
6-7	29.6 (25)	25/25	29.5 (25)	100	25/25	29.1 (25)	98	25/25	26.6 (25)	90	25/25
7-7	30.3 (25)	25/25	30.2 (25)	100	25/25	29.3 (25)	97	25/25	26.8 (25)	88	25/25
8-7	30.7 (25)	25/25	30.2 (25)	98	25/25	29.7 (25)	97	25/25	27.3 (25)	89	25/25
9-7	31.0 (25)	25/25	30.6 (25)	99	25/25	30.2 (25)	97	25/25	27.6 (25)	89	25/25
10-7	31.2 (25)	25/25	30.9 (25)	99	25/25	30.2 (25)	97	25/25	27.4 (25)	88	25/25
11-7	31.6 (25)	25/25	31.1 (25)	98	25/25	30.5 (25)	97	25/25	27.6 (25)	87	25/25
12-7	31.9 (25)	25/25	31.4 (25)	98	25/25	31.1 (25)	97	25/25	28.0 (25)	88	25/25
13-7	32.0 (25)	25/25	32.0 (25)	100	25/25	31.3 (25)	98	25/25	27.9 (25)	87	25/25
14-7	32.3 (25)	25/25	31.9 (25)	99	25/25	31.4 (25)	97	25/25	28.1 (24)	87	24/25
15-7	32.4 (25)	25/25	32.1 (25)	99	25/25	31.7 (25)	98	25/25	28.3 (24)	87	24/25
16-7	32.8 (25)	25/25	32.4 (25)	99	25/25	31.6 (25)	96	25/25	28.3 (24)	86	24/25
17-7	32.8 (25)	25/25	32.1 (25)	98	25/25	32.0 (25)	98	25/25	28.3 (24)	86	24/25
18-7	32.9 (25)	25/25	32.2 (25)	98	25/25	31.9 (25)	97	25/25	28.3 (24)	86	24/25
19-7	33.1 (25)	25/25	32.7 (25)	99	25/25	32.5 (25)	98	25/25	28.8 (24)	87	24/25
20-7	33.7 (24)	24/25	33.0 (25)	98	25/25	32.8 (25)	97	25/25	29.0 (24)	86	24/25
21-7	33.7 (24)	24/25	33.0 (25)	98	25/25	32.7 (25)	97	25/25	28.9 (24)	86	24/25
22-7	34.2 (24)	24/25	33.5 (25)	98	25/25	33.3 (25)	97	25/25	29.0 (24)	85	24/25
23-7	34.1 (24)	24/25	33.7 (25)	99	25/25	33.1 (25)	97	25/25	29.0 (24)	85	24/25
24-7	34.7 (24)	24/25	34.1 (25)	98	25/25	33.4 (25)	96	25/25	29.2 (24)	84	24/25
25-7	34.7 (24)	24/25	34.2 (25)	99	25/25	33.7 (25)	97	25/25	29.3 (24)	84	24/25
26-7	34.9 (24)	24/25	34.2 (25)	98	25/25	33.9 (25)	97	25/25	29.1 (24)	83	24/25

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

TABLE D2

BODY WEIGHT CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0926 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>	0.15 ppm Av.Wt.	0.15 ppm % of cont. <25>	0.15 ppm No.of Surviv.	0.5 ppm Av.Wt.	0.5 ppm % of cont. <25>	0.5 ppm No.of Surviv.	1.5 ppm Av.Wt.	1.5 ppm % of cont. <25>	1.5 ppm No.of Surviv.
0-0	20.2 (25)	25/25	20.2 (25)	100	25/25	20.2 (25)	100	25/25	20.2 (25)	100	25/25
1-7	20.5 (25)	25/25	20.4 (25)	100	25/25	20.2 (25)	99	25/25	19.8 (25)	97	25/25
2-7	20.6 (25)	25/25	20.5 (25)	100	25/25	20.6 (25)	100	25/25	20.0 (25)	97	25/25
3-7	21.3 (25)	25/25	21.2 (25)	100	25/25	20.8 (25)	98	25/25	20.1 (25)	94	25/25
4-7	21.7 (25)	25/25	21.8 (25)	100	25/25	21.6 (25)	100	25/25	20.4 (25)	94	25/25
5-7	22.8 (25)	25/25	22.8 (25)	100	25/25	22.6 (25)	99	25/25	21.1 (25)	93	25/25
6-7	23.4 (25)	25/25	23.3 (25)	100	25/25	23.2 (25)	99	25/25	21.5 (25)	92	25/25
7-7	23.5 (25)	25/25	23.3 (25)	99	25/25	23.2 (25)	99	25/25	22.2 (25)	94	25/25
8-7	23.6 (25)	25/25	23.3 (25)	99	25/25	23.4 (25)	99	25/25	22.0 (25)	93	25/25
9-7	23.6 (25)	25/25	24.0 (25)	102	25/25	23.5 (25)	100	25/25	22.7 (25)	96	25/25
10-7	23.6 (25)	25/25	23.8 (25)	101	25/25	23.4 (25)	99	25/25	22.6 (25)	96	25/25
11-7	24.0 (25)	25/25	24.1 (25)	100	25/25	23.7 (25)	99	25/25	23.0 (25)	96	25/25
12-7	24.0 (25)	25/25	24.1 (25)	100	25/25	24.5 (25)	102	25/25	23.3 (25)	97	25/25
13-7	24.2 (25)	25/25	24.5 (25)	101	25/25	24.5 (25)	101	25/25	23.3 (25)	96	25/25
14-7	24.5 (25)	25/25	24.9 (25)	102	25/25	24.4 (25)	100	25/25	23.3 (25)	95	25/25
15-7	24.5 (25)	25/25	24.7 (25)	101	25/25	24.9 (25)	102	25/25	23.6 (25)	96	25/25
16-7	24.8 (25)	25/25	24.9 (25)	100	25/25	24.9 (25)	100	25/25	23.7 (25)	96	25/25
17-7	24.8 (25)	25/25	25.2 (25)	102	25/25	24.8 (25)	100	25/25	23.9 (25)	96	25/25
18-7	24.6 (25)	25/25	24.9 (25)	101	25/25	24.4 (25)	99	25/25	23.8 (25)	97	25/25
19-7	24.9 (25)	25/25	25.5 (25)	102	25/25	25.2 (25)	101	25/25	24.3 (25)	98	25/25
20-7	25.2 (25)	25/25	25.5 (25)	101	25/25	25.0 (25)	99	25/25	24.2 (25)	96	25/25
21-7	25.7 (25)	25/25	25.3 (25)	98	25/25	25.6 (25)	100	25/25	24.1 (25)	94	25/25
22-7	25.6 (25)	25/25	25.8 (25)	101	25/25	25.3 (25)	99	25/25	24.5 (25)	96	25/25
23-7	26.1 (25)	25/25	25.7 (25)	98	25/25	25.5 (25)	98	25/25	24.4 (25)	93	25/25
24-7	25.6 (25)	25/25	25.6 (25)	100	25/25	25.9 (25)	101	25/25	24.3 (25)	95	25/25
25-7	25.8 (25)	25/25	26.5 (24)	103	24/25	25.5 (25)	99	25/25	24.3 (25)	94	25/25
26-7	26.3 (25)	25/25	26.8 (24)	102	24/25	26.0 (25)	99	25/25	24.5 (25)	93	25/25

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

TABLE D3

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0926
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	25.6± 1.3	26.0± 1.4	27.0± 1.3	27.8± 1.4	28.4± 1.5	29.1± 1.6	29.6± 1.6
0.15 ppm	25.6± 1.3	26.0± 1.4	26.6± 1.4	27.5± 1.4	28.1± 1.8	28.6± 1.8	29.5± 1.7
0.5 ppm	25.6± 1.4	25.6± 1.3	26.2± 1.2	27.2± 1.2	27.6± 1.5	28.2± 1.4	29.1± 1.5
1.5 ppm	25.6± 1.3	24.7± 1.3**	25.2± 1.5**	25.4± 1.4**	25.7± 1.4**	26.1± 1.6**	26.6± 1.5**

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

STUDY NO. : 0926
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	30.3± 1.7	30.7± 1.8	31.0± 2.0	31.2± 2.0	31.6± 2.0	31.9± 2.2	32.0± 2.4
0.15 ppm	30.2± 1.9	30.2± 2.1	30.6± 2.0	30.9± 2.0	31.1± 2.0	31.4± 2.1	32.0± 2.3
0.5 ppm	29.3± 1.8	29.7± 1.8	30.2± 1.9	30.2± 2.2	30.5± 2.3	31.1± 2.6	31.3± 2.5
1.5 ppm	26.8± 1.5**	27.3± 1.5**	27.6± 1.4**	27.4± 1.5**	27.6± 1.6**	28.0± 1.7**	27.9± 1.8**

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

STUDY NO. : 0926
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	32.3± 2.7	32.4± 2.7	32.8± 2.8	32.8± 2.8	32.9± 2.9	33.1± 3.0	33.7± 3.0
0.15 ppm	31.9± 2.4	32.1± 2.3	32.4± 2.3	32.1± 2.1	32.2± 2.5	32.7± 2.5	33.0± 2.7
0.5 ppm	31.4± 2.6	31.7± 2.6	31.6± 2.8	32.0± 2.9	31.9± 3.1	32.5± 3.2	32.8± 3.3
1.5 ppm	28.1± 1.5**	28.3± 1.5**	28.3± 1.5**	28.3± 1.7**	28.3± 1.8**	28.8± 1.9**	29.0± 1.9**

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

STUDY NO. : 0926
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	33.7± 3.0	34.2± 3.4	34.1± 3.4	34.7± 3.5	34.7± 3.3	34.9± 3.1
0.15 ppm	33.0± 2.7	33.5± 2.5	33.7± 2.8	34.1± 3.1	34.2± 2.9	34.2± 2.8
0.5 ppm	32.7± 3.4	33.3± 3.4	33.1± 3.2	33.4± 3.3	33.7± 3.4	33.9± 3.1
1.5 ppm	28.9± 1.9**	29.0± 1.8**	29.0± 2.0**	29.2± 2.0**	29.3± 2.0**	29.1± 2.0**

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

TABLE D4

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0926
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	20.2± 1.2	20.5± 1.2	20.6± 1.1	21.3± 1.2	21.7± 1.1	22.8± 1.2	23.4± 1.1
0.15 ppm	20.2± 1.2	20.4± 1.1	20.5± 1.0	21.2± 1.5	21.8± 1.2	22.8± 1.3	23.3± 0.9
0.5 ppm	20.2± 1.2	20.2± 1.1	20.6± 1.1	20.8± 1.0	21.6± 1.4	22.6± 1.3	23.2± 1.4
1.5 ppm	20.2± 1.2	19.8± 1.4	20.0± 1.4	20.1± 1.3**	20.4± 1.1**	21.1± 1.3**	21.5± 1.3**

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

STUDY NO. : 0926
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	23.5± 1.2	23.6± 1.2	23.6± 1.2	23.6± 1.1	24.0± 1.0	24.0± 1.1	24.2± 1.5
0.15 ppm	23.3± 1.3	23.3± 1.2	24.0± 1.6	23.8± 1.4	24.1± 1.0	24.1± 1.4	24.5± 1.5
0.5 ppm	23.2± 1.6	23.4± 1.5	23.5± 1.1	23.4± 1.3	23.7± 1.1	24.5± 1.5	24.5± 1.7
1.5 ppm	22.2± 1.6**	22.0± 1.4**	22.7± 1.6	22.6± 1.4*	23.0± 1.4*	23.3± 1.4	23.3± 1.6

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

STUDY NO. : 0926
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	24.5± 1.7	24.5± 1.2	24.8± 1.6	24.8± 1.6	24.6± 1.6	24.9± 1.5	25.2± 1.8
0.15 ppm	24.9± 1.5	24.7± 1.6	24.9± 1.7	25.2± 1.7	24.9± 1.4	25.5± 1.9	25.5± 2.4
0.5 ppm	24.4± 1.5	24.9± 1.4	24.9± 1.6	24.8± 1.9	24.4± 1.4	25.2± 1.5	25.0± 1.9
1.5 ppm	23.3± 1.8*	23.6± 1.8	23.7± 1.9	23.9± 1.9	23.8± 2.1	24.3± 1.8	24.2± 1.8

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

STUDY NO. : 0926
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	25.7± 1.6	25.6± 1.7	26.1± 2.1	25.6± 1.5	25.8± 1.8	26.3± 2.0
0.15 ppm	25.3± 1.7	25.8± 1.8	25.7± 1.6	25.6± 1.3	26.5± 2.4	26.8± 2.7
0.5 ppm	25.6± 2.3	25.3± 2.0	25.5± 1.6	25.9± 1.7	25.5± 1.6	26.0± 1.5
1.5 ppm	24.1± 1.9**	24.5± 2.0	24.4± 2.2**	24.3± 1.9*	24.3± 2.1*	24.5± 2.3*

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

TABLE E1

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0926
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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>	0.15 ppm Av.FC.	0.15 ppm % of cont. <25>	0.15 ppm No.of Surviv.	0.5 ppm Av.FC.	0.5 ppm % of cont. <25>	0.5 ppm No.of Surviv.	1.5 ppm Av.FC.	1.5 ppm % of cont. <25>	1.5 ppm No.of Surviv.
1-7	4.0 (25)	25/25	4.0 (25)	100	25/25	3.9 (25)	98	25/25	3.7 (25)	93	25/25
2-7	4.1 (25)	25/25	4.2 (25)	102	25/25	4.0 (25)	98	25/25	3.8 (25)	93	25/25
3-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.1 (25)	95	25/25	3.8 (25)	88	25/25
4-7	4.4 (25)	25/25	4.5 (25)	102	25/25	4.3 (25)	98	25/25	3.9 (25)	89	25/25
5-7	4.7 (25)	25/25	4.6 (25)	98	25/25	4.4 (25)	94	25/25	4.0 (25)	85	25/25
6-7	4.7 (25)	25/25	4.6 (25)	98	25/25	4.5 (25)	96	25/25	4.0 (25)	85	25/25
7-7	4.7 (25)	25/25	4.6 (25)	98	25/25	4.5 (25)	96	25/25	4.1 (25)	87	25/25
8-7	4.7 (25)	25/25	4.6 (25)	98	25/25	4.5 (25)	96	25/25	4.1 (25)	87	25/25
9-7	4.7 (25)	25/25	4.7 (25)	100	25/25	4.6 (25)	98	25/25	4.2 (25)	89	25/25
10-7	4.7 (25)	25/25	4.7 (25)	100	25/25	4.5 (25)	96	25/25	4.1 (25)	87	25/25
11-7	4.8 (25)	25/25	4.7 (25)	98	25/25	4.6 (25)	96	25/25	4.1 (25)	85	25/25
12-7	5.0 (25)	25/25	4.9 (25)	98	25/25	4.8 (25)	96	25/25	4.4 (25)	88	25/25
13-7	4.8 (25)	25/25	4.8 (25)	100	25/25	4.7 (25)	98	25/25	4.2 (25)	88	25/25
14-7	4.9 (25)	25/25	4.7 (25)	96	25/25	4.6 (25)	94	25/25	4.2 (24)	86	24/25
15-7	4.8 (25)	25/25	4.8 (25)	100	25/25	4.6 (25)	96	25/25	4.2 (24)	88	24/25
16-7	4.8 (25)	25/25	4.7 (25)	98	25/25	4.6 (25)	96	25/25	4.2 (24)	88	24/25
17-7	4.8 (25)	25/25	4.7 (25)	98	25/25	4.8 (25)	100	25/25	4.3 (24)	90	24/25
18-7	4.8 (25)	25/25	4.5 (25)	94	25/25	4.6 (25)	96	25/25	4.1 (24)	85	24/25
19-7	4.7 (25)	25/25	4.6 (25)	98	25/25	4.6 (25)	98	25/25	4.1 (24)	87	24/25
20-7	4.8 (24)	24/25	4.6 (25)	96	25/25	4.5 (25)	94	25/25	4.2 (24)	88	24/25
21-7	4.8 (24)	24/25	4.5 (25)	94	25/25	4.6 (25)	96	25/25	4.1 (24)	85	24/25
22-7	4.9 (24)	24/25	4.6 (25)	94	25/25	4.6 (25)	94	25/25	4.1 (24)	84	24/25
23-7	4.8 (24)	24/25	4.7 (25)	98	25/25	4.5 (25)	94	25/25	4.2 (24)	88	24/25
24-7	5.0 (24)	24/25	4.7 (25)	94	25/25	4.7 (25)	94	25/25	4.2 (24)	84	24/25
25-7	4.9 (24)	24/25	4.7 (25)	96	25/25	4.6 (25)	94	25/25	4.1 (24)	84	24/25
26-7	4.9 (24)	24/25	4.6 (25)	94	25/25	4.7 (25)	96	25/25	4.2 (24)	86	24/25

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

TABLE E2

FOOD CONSUMPTION CHANGES AND

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0926
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt)
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>	0.15 ppm Av.FC.	0.15 ppm % of cont. <25>	0.15 ppm No.of Surviv.	0.5 ppm Av.FC.	0.5 ppm % of cont. <25>	0.5 ppm No.of Surviv.	1.5 ppm Av.FC.	1.5 ppm % of cont. <25>	1.5 ppm No.of Surviv.
1-7	3.3 (25)	25/25	3.4 (25)	103	25/25	3.4 (25)	103	25/25	3.2 (25)	97	25/25
2-7	3.5 (25)	25/25	3.5 (25)	100	25/25	3.5 (25)	100	25/25	3.2 (25)	91	25/25
3-7	3.7 (25)	25/25	3.9 (25)	105	25/25	3.8 (25)	103	25/25	3.4 (25)	92	25/25
4-7	4.1 (25)	25/25	4.3 (25)	105	25/25	4.3 (25)	105	25/25	3.7 (25)	90	25/25
5-7	4.7 (25)	25/25	4.7 (25)	100	25/25	4.5 (25)	96	25/25	3.9 (25)	83	25/25
6-7	4.4 (25)	25/25	4.6 (25)	105	25/25	4.5 (25)	102	25/25	3.9 (25)	89	25/25
7-7	4.4 (25)	25/25	4.9 (25)	111	25/25	4.7 (25)	107	25/25	4.1 (25)	93	25/25
8-7	4.4 (25)	25/25	4.5 (25)	102	25/25	4.6 (25)	105	25/25	4.1 (25)	93	25/25
9-7	4.5 (25)	25/25	4.9 (25)	109	25/25	4.7 (25)	104	25/25	4.1 (25)	91	25/25
10-7	4.4 (25)	25/25	4.6 (25)	105	25/25	4.7 (25)	107	25/25	4.1 (25)	93	25/25
11-7	4.6 (25)	25/25	4.8 (25)	104	25/25	4.8 (25)	104	25/25	4.3 (25)	93	25/25
12-7	4.9 (25)	25/25	4.8 (25)	98	25/25	5.1 (25)	104	25/25	4.4 (25)	90	25/25
13-7	4.7 (25)	25/25	4.8 (25)	102	25/25	5.0 (25)	106	25/25	4.2 (25)	89	25/25
14-7	4.7 (25)	25/25	4.8 (25)	102	25/25	4.7 (25)	100	25/25	4.1 (25)	87	25/25
15-7	4.7 (25)	25/25	4.8 (25)	102	25/25	4.9 (25)	104	25/25	4.1 (25)	87	25/25
16-7	4.6 (25)	25/25	4.7 (25)	102	25/25	4.9 (25)	107	25/25	4.1 (25)	89	25/25
17-7	4.7 (25)	25/25	4.7 (25)	100	25/25	4.9 (25)	104	25/25	4.2 (25)	89	25/25
18-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.7 (25)	109	25/25	3.9 (25)	91	25/25
19-7	4.4 (25)	25/25	4.6 (25)	105	25/25	4.8 (25)	109	25/25	4.0 (25)	91	25/25
20-7	4.4 (25)	25/25	4.3 (25)	98	25/25	4.5 (25)	102	25/25	3.9 (25)	89	25/25
21-7	4.6 (25)	25/25	4.4 (25)	96	25/25	4.9 (25)	107	25/25	4.0 (25)	87	25/25
22-7	4.4 (25)	25/25	4.5 (25)	102	25/25	4.7 (25)	107	25/25	4.1 (25)	93	25/25
23-7	4.7 (25)	25/25	4.4 (25)	94	25/25	4.7 (25)	100	25/25	4.0 (25)	85	25/25
24-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.6 (25)	107	25/25	4.0 (25)	93	25/25
25-7	4.6 (25)	25/25	4.6 (24)	100	24/25	4.5 (25)	98	25/25	4.1 (25)	89	25/25
26-7	4.6 (25)	25/25	4.5 (24)	98	24/25	4.6 (25)	100	25/25	4.0 (25)	87	25/25

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

TABLE E3

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0926
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	4.1± 0.2	4.3± 0.3	4.4± 0.3	4.7± 0.3	4.7± 0.3	4.7± 0.4
0.15 ppm	4.0± 0.3	4.2± 0.2	4.3± 0.3	4.5± 0.4	4.6± 0.4	4.6± 0.4	4.6± 0.5
0.5 ppm	3.9± 0.3	4.0± 0.2	4.1± 0.3	4.3± 0.3	4.4± 0.4*	4.5± 0.4	4.5± 0.4
1.5 ppm	3.7± 0.3**	3.8± 0.3**	3.8± 0.3**	3.9± 0.4**	4.0± 0.4**	4.0± 0.3**	4.1± 0.3**

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

STUDY NO. : 0926
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	4.7± 0.3	4.7± 0.3	4.7± 0.3	4.8± 0.4	5.0± 0.4	4.8± 0.4	4.9± 0.4
0.15 ppm	4.6± 0.5	4.7± 0.5	4.7± 0.6	4.7± 0.6	4.9± 0.7	4.8± 0.6	4.7± 0.6
0.5 ppm	4.5± 0.4	4.6± 0.4	4.5± 0.4	4.6± 0.5	4.8± 0.5	4.7± 0.4	4.6± 0.4
1.5 ppm	4.1± 0.4**	4.2± 0.4**	4.1± 0.4**	4.1± 0.4**	4.4± 0.5**	4.2± 0.5**	4.2± 0.4**

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

STUDY NO. : 0926
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	4.8± 0.4	4.8± 0.5	4.8± 0.4	4.8± 0.4	4.7± 0.4	4.8± 0.4	4.8± 0.5
0.15 ppm	4.8± 0.6	4.7± 0.6	4.7± 0.5	4.5± 0.6	4.6± 0.5	4.6± 0.5	4.5± 0.5
0.5 ppm	4.6± 0.4	4.6± 0.5	4.8± 0.4	4.6± 0.5	4.6± 0.4	4.5± 0.4	4.6± 0.5
1.5 ppm	4.2± 0.4**	4.2± 0.4**	4.3± 0.4**	4.1± 0.4**	4.1± 0.4**	4.2± 0.4**	4.1± 0.4**

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

STUDY NO. : 0926
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	4.9± 0.5	4.8± 0.4	5.0± 0.4	4.9± 0.4	4.9± 0.4
0.15 ppm	4.6± 0.5	4.7± 0.5	4.7± 0.6	4.7± 0.5	4.6± 0.4
0.5 ppm	4.6± 0.4	4.5± 0.4	4.7± 0.4*	4.6± 0.4	4.7± 0.4
1.5 ppm	4.1± 0.4**	4.2± 0.5**	4.2± 0.4**	4.1± 0.4**	4.2± 0.4**

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

TABLE E4

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0926
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	3.3± 0.5	3.5± 0.3	3.7± 0.4	4.1± 0.5	4.7± 1.3	4.4± 0.4	4.4± 0.4
0.15 ppm	3.4± 0.3	3.5± 0.3	3.9± 0.6	4.3± 1.0	4.7± 1.0	4.6± 0.6	4.9± 1.1
0.5 ppm	3.4± 0.3	3.5± 0.3	3.8± 0.4	4.3± 0.4	4.5± 0.4	4.5± 0.4	4.7± 0.6*
1.5 ppm	3.2± 0.4	3.2± 0.4**	3.4± 0.3*	3.7± 0.4**	3.9± 0.4**	3.9± 0.4**	4.1± 0.7*

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

STUDY NO. : 0926
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	4.4± 0.4	4.5± 0.6	4.4± 0.4	4.6± 0.6	4.9± 1.2	4.7± 0.6	4.7± 0.6
0.15 ppm	4.5± 0.5	4.9± 0.9	4.6± 0.4	4.8± 0.5	4.8± 0.6	4.8± 0.6	4.8± 0.4
0.5 ppm	4.6± 0.5	4.7± 0.4	4.7± 0.5	4.8± 0.5	5.1± 0.7	5.0± 0.9	4.7± 0.6
1.5 ppm	4.1± 0.7	4.1± 0.4*	4.1± 0.4	4.3± 0.5*	4.4± 0.5	4.2± 0.5**	4.1± 0.5**

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

STUDY NO. : 0926
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	4.7± 0.5	4.6± 0.5	4.7± 0.6	4.3± 0.5	4.4± 0.5	4.4± 0.5	4.6± 0.5
0.15 ppm	4.8± 0.6	4.7± 0.4	4.7± 0.4	4.3± 0.3	4.6± 0.5	4.3± 0.6	4.4± 0.6
0.5 ppm	4.9± 0.6	4.9± 0.7	4.9± 0.7	4.7± 0.5**	4.8± 1.1	4.5± 0.5	4.9± 1.1
1.5 ppm	4.1± 0.5**	4.1± 0.6*	4.2± 0.4**	3.9± 0.5*	4.0± 0.5*	3.9± 0.4**	4.0± 0.4**

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

STUDY NO. : 0926
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	4.4± 0.6	4.7± 0.9	4.3± 0.4	4.6± 0.4	4.6± 0.5
0.15 ppm	4.5± 0.4	4.4± 0.4	4.3± 0.4	4.6± 0.5	4.5± 0.6
0.5 ppm	4.7± 0.8	4.7± 0.7	4.6± 0.5*	4.5± 0.6	4.6± 0.5
1.5 ppm	4.1± 0.4	4.0± 0.5**	4.0± 0.4*	4.1± 0.5**	4.0± 0.5**

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

TABLE F1

URINALYSIS : MALE

STUDY NO. : 0926 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	21	0	0	0	0	2	3	16		0	9	8	4	0	0		21	0	0	0	0	0		1	12	6	0	2	0		21	0	0	0	0	
0.15 ppm	24	0	0	0	0	2	1	21		2	8	8	6	0	0		24	0	0	0	0	0		4	8	7	5	0	0	*	24	0	0	0	0	
0.5 ppm	21	0	0	0	0	2	5	14		0	2	12	6	1	0		21	0	0	0	0	0		1	9	4	5	2	0		21	0	0	0	0	
1.5 ppm	20	0	1	0	0	2	3	14		1	7	11	1	0	0		20	0	0	0	0	0		0	14	5	1	0	0		20	0	0	0	0	

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

STUDY NO. : 0926 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	21	21	0	0	0	0	
0.15 ppm	24	24	0	0	0	0	
0.5 ppm	21	21	0	0	0	0	
1.5 ppm	20	20	0	0	0	0	

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

TABLE F2

URINALYSIS : FEMALE

STUDY NO. : 0926 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	1	4	2	7	5	2		5	15	1	0	0	0		21	0	0	0	0	0		1	15	4	1	0	0		21	0	0	0	0	
0.15 ppm	21	0	0	1	4	5	9	2		12	9	0	0	0	0		21	0	0	0	0	0		4	16	1	0	0	0		21	0	0	0	0	
0.5 ppm	20	0	0	1	2	5	10	2		5	14	1	0	0	0		20	0	0	0	0	0		3	17	0	0	0	0		20	0	0	0	0	
1.5 ppm	23	0	0	1	1	3	11	7		11	11	1	0	0	0		23	0	0	0	0	0		6	16	1	0	0	0		23	0	0	0	0	

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

Group Name	NO. of Animals	Urobilinogen ±	＋	2+	3+	4+	CHI
Control	21	21	0	0	0	0	
0.15 ppm	21	21	0	0	0	0	
0.5 ppm	20	20	0	0	0	0	
1.5 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. : 0926
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	23	11.09± 0.34	16.4± 0.5	46.0± 1.4	41.5± 0.5	14.8± 0.2	35.7± 0.6	1439± 120
0.15 ppm	23	11.03± 0.29	16.4± 0.4	45.9± 1.3	41.6± 0.5	14.9± 0.3	35.8± 0.5	1395± 160
0.5 ppm	24	11.22± 0.26	16.6± 0.3	46.8± 1.2	41.7± 0.7	14.8± 0.2	35.5± 0.5	1438± 75
1.5 ppm	24	11.24± 0.32	16.7± 0.4*	46.9± 1.5*	41.8± 0.6	14.9± 0.3	35.6± 0.5	1371± 103

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

Group Name	NO. of Animals	RETICULOCYTE %
Control	23	3.0± 0.3
0.15 ppm	23	3.0± 0.1
0.5 ppm	24	3.0± 0.2
1.5 ppm	24	3.0± 0.2

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

STUDY NO. : 0926
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	23	1.67±	0.50	37.0±	9.8	59.5±	9.9	1.9±	0.8	1.6±	1.5	0.0±	0.0
0.15 ppm	23	1.75±	1.12	35.6±	11.0	60.9±	10.5	2.1±	0.9	1.4±	1.4	0.0±	0.0
0.5 ppm	24	1.61±	0.79	34.3±	6.7	62.0±	6.4	1.8±	0.7	2.0±	1.7	0.0±	0.1
1.5 ppm	24	1.22±	0.60**	37.1±	8.8	60.1±	8.2	1.7±	0.9	1.1±	1.4	0.0±	0.0

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

TABLE G2

HEMATOLOGY : FEMALE

STUDY NO. : 0926
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	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	10.79± 0.87	16.3± 1.0	44.8± 2.9	41.6± 1.3	15.2± 0.6	36.4± 0.7	1317± 123
0.15 ppm	24	10.84± 0.38	16.5± 0.5	45.3± 1.7	41.8± 0.6	15.2± 0.3	36.4± 0.6	1280± 91
0.5 ppm	25	10.79± 0.27	16.3± 0.4	45.3± 1.1	41.9± 0.7	15.1± 0.3	36.1± 0.6	1245± 158
1.5 ppm	24	10.98± 0.41	16.5± 0.5	45.7± 1.8	41.6± 0.5	15.1± 0.2	36.2± 0.8	1262± 126

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

Group Name	NO. of Animals	RETICULOCYTE %
Control	25	3.0± 0.9
0.15 ppm	24	2.9± 0.7
0.5 ppm	25	3.2± 0.7
1.5 ppm	24	3.2± 1.1

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

STUDY NO. : 0926
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	25	2.79±	1.88	39.1±	15.1	57.6±	13.9	2.2±	1.3	1.1±	1.4	0.0±	0.1
0.15 ppm	24	2.15±	1.61	35.4±	12.9	61.3±	12.0	2.3±	0.9	1.0±	1.1	0.0±	0.0
0.5 ppm	25	2.05±	1.12	39.8±	11.4	56.6±	10.4	2.3±	1.0	1.2±	1.4	0.0±	0.0
1.5 ppm	24	2.13±	1.23	39.4±	13.1	57.3±	12.4	2.2±	1.0	1.1±	0.9	0.0±	0.0

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

TABLE H1

BIOCHEMISTRY : MALE

STUDY NO. : 0926
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	24	5.2±	0.2	2.9±	0.1	1.3±	0.1	0.10±	0.01	230±	23	80±	10	56±	22
0.15 ppm	24	5.2±	0.2	2.9±	0.1	1.3±	0.1	0.10±	0.01	225±	28	80±	9	56±	17
0.5 ppm	25	5.0±	0.2*	2.8±	0.1	1.3±	0.1	0.10±	0.01	221±	24	75±	9	57±	17
1.5 ppm	24	5.1±	0.1	2.9±	0.1	1.3±	0.1	0.11±	0.01**	201±	17**	63±	8**	38±	12**

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

STUDY NO. : 0926
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	24	166± 19	66± 18	25± 12	296± 62	198± 14	0.2± 0.3	89± 46
0.15 ppm	24	167± 18	76± 24	30± 16	312± 73	199± 22	0.3± 0.4	94± 33
0.5 ppm	25	159± 17	61± 17	23± 14	282± 70	194± 16	0.3± 0.3	93± 39
1.5 ppm	24	135± 18**	70± 13	20± 4	293± 72	213± 25**	0.2± 0.2	138± 156

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

STUDY NO. : 0926
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	24	24.1±	6.3	149±	1	3.5±	0.3	118±	1	8.6±	0.2	5.3±	0.7
0.15 ppm	24	23.4±	4.8	149±	1	3.6±	0.2	118±	2	8.5±	0.1	5.6±	0.8
0.5 ppm	25	24.3±	5.3	149±	1	3.7±	0.3	118±	2	8.6±	0.3	5.4±	0.8
1.5 ppm	24	22.9±	3.6	149±	1	3.8±	0.3**	118±	2	8.6±	0.1	5.3±	0.9

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

TABLE H2

BIOCHEMISTRY : FEMALE

STUDY NO. : 0926
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	25	5.2±	0.2	3.1±	0.1	1.5±	0.1	0.10±	0.02	207±	25	67±	9	47±	16
0.15 ppm	24	5.3±	0.2	3.1±	0.1	1.5±	0.1	0.09±	0.01	204±	25	64±	7	41±	17
0.5 ppm	24	5.3±	0.2	3.1±	0.1	1.4±	0.1	0.10±	0.01	206±	24	66±	8	42±	14
1.5 ppm	25	5.3±	0.3	3.1±	0.3	1.4±	0.1	0.11±	0.05	197±	40	65±	19	37±	13

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

STUDY NO. : 0926
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	25	138± 17	117± 89	40± 46	279± 163	322± 35	0.2± 0.2	124± 82
0.15 ppm	24	131± 16	96± 48	29± 20	256± 71	322± 35	0.2± 0.2	93± 27
0.5 ppm	24	136± 17	114± 43	32± 16	291± 156	332± 46	0.3± 0.3	115± 52
1.5 ppm	25	128± 16	90± 35	26± 14	266± 132	340± 39	0.3± 0.4	155± 223

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

STUDY NO. : 0926
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	UREANITROGEN mg／dℓ	SODIUM mEq／ℓ	POTASSIUM mEq／ℓ	CHLORIDE mEq／ℓ	CALCIUM mg／dℓ	INORGANIC PHOSPHRUS mg／dℓ
Control	25	18.7± 3.0	149± 2	3.2± 0.3	118± 2	8.9± 0.2	5.3± 1.0
0.15 ppm	24	17.7± 2.6	149± 2	3.1± 0.3	118± 2	8.8± 0.2	5.4± 0.8
0.5 ppm	24	18.5± 2.5	149± 2	3.2± 0.2	119± 2	8.9± 0.2	5.4± 0.8
1.5 ppm	25	21.6± 12.8	149± 2	3.5± 0.6*	118± 3	8.9± 0.3	5.3± 1.3

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 (%)	0.15 ppm 25 (%)	0.5 ppm 25 (%)	1.5 ppm 25 (%)
lung	red		0 (0)	0 (0)	0 (0)	1 (4)
	white zone		2 (8)	1 (4)	2 (8)	1 (4)
	nodule		1 (4)	2 (8)	4 (16)	0 (0)
lymph node	enlarged		0 (0)	0 (0)	0 (0)	1 (4)
thymus	enlarged		0 (0)	0 (0)	0 (0)	1 (4)
spleen	enlarged		0 (0)	0 (0)	0 (0)	1 (4)
	black zone		3 (12)	3 (12)	8 (32)	4 (16)
	nodule		1 (4)	1 (4)	0 (0)	0 (0)
salivary gl	nodule		1 (4)	0 (0)	0 (0)	0 (0)
stomach	forestomach:nodule		0 (0)	0 (0)	0 (0)	2 (8)
	forestomach:thick		1 (4)	0 (0)	1 (4)	0 (0)
	glandular stomach:thick		2 (8)	1 (4)	5 (20)	0 (0)
liver	white zone		1 (4)	0 (0)	3 (12)	2 (8)
	red zone		0 (0)	0 (0)	1 (4)	0 (0)
	nodule		0 (0)	0 (0)	1 (4)	0 (0)
kidney	white zone		1 (4)	0 (0)	0 (0)	0 (0)
testis	small		1 (4)	0 (0)	0 (0)	0 (0)
	red zone		0 (0)	1 (4)	0 (0)	0 (0)
pleura	red zone		1 (4)	0 (0)	0 (0)	0 (0)
abdominal c	hemorrhage		0 (0)	1 (4)	0 (0)	0 (0)
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	(%)	0.15 ppm 25	(%)	0.5 ppm 25	(%)	1.5 ppm 25	(%)
skin/app	nodule		0	(0)	1	(4)	0	(0)	0	(0)
lung	white zone		0	(0)	0	(0)	3	(12)	0	(0)
	nodule		2	(8)	1	(4)	1	(4)	3	(12)
spleen	enlarged		1	(4)	0	(0)	0	(0)	0	(0)
	white zone		1	(4)	0	(0)	0	(0)	0	(0)
	black zone		5	(20)	4	(16)	3	(12)	0	(0)
	nodule		2	(8)	1	(4)	2	(8)	0	(0)
stomach	forestomach:nodule		2	(8)	2	(8)	0	(0)	1	(4)
	forestomach:thick		1	(4)	1	(4)	1	(4)	0	(0)
	glandular stomach:thick		8	(32)	5	(20)	8	(32)	3	(12)
liver	black zone		1	(4)	0	(0)	0	(0)	0	(0)
	nodule		1	(4)	0	(0)	0	(0)	0	(0)
ovary	enlarged		0	(0)	1	(4)	1	(4)	0	(0)
uterus	nodule		1	(4)	0	(0)	0	(0)	0	(0)
vagina	nodule		0	(0)	0	(0)	0	(0)	1	(4)
thoracic ca	pleural fluid		0	(0)	1	(4)	0	(0)	1	(4)

TABLE J1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0926
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	24	29.8± 3.3	0.014± 0.004	0.224± 0.037	0.197± 0.018	0.167± 0.017	0.606± 0.046
0.15 ppm	24	29.5± 2.7	0.014± 0.002	0.242± 0.053	0.197± 0.019	0.166± 0.017	0.604± 0.057
0.5 ppm	25	29.6± 2.9	0.013± 0.002	0.237± 0.045	0.190± 0.018	0.171± 0.013	0.586± 0.040
1.5 ppm	24	25.4± 1.9**	0.013± 0.004	0.236± 0.036	0.169± 0.017**	0.160± 0.018	0.547± 0.055**

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

STUDY NO. : 0926
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	24	0.072± 0.027	1.351± 0.126	0.473± 0.013
0.15 ppm	24	0.071± 0.012	1.333± 0.125	0.476± 0.014
0.5 ppm	25	0.068± 0.009	1.293± 0.096	0.476± 0.014
1.5 ppm	24	0.062± 0.012	1.135± 0.106**	0.469± 0.020

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

TABLE J2

ORGAN WEIGHT, ABSOLUTE : FEMALE

STUDY NO. : 0926
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	25	22.1± 1.9	0.016± 0.002	0.026± 0.005	0.149± 0.014	0.163± 0.022	0.427± 0.031
0.15 ppm	24	22.2± 1.6	0.015± 0.002	0.030± 0.015	0.148± 0.012	0.161± 0.013	0.420± 0.037
0.5 ppm	25	21.7± 1.7	0.015± 0.002	0.027± 0.005	0.152± 0.017	0.165± 0.017	0.427± 0.037
1.5 ppm	25	20.9± 2.3	0.015± 0.003	0.026± 0.006	0.137± 0.014*	0.189± 0.138	0.395± 0.038**

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

STUDY NO. : 0926
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	25	0.107± 0.095	1.134± 0.155	0.499± 0.015
0.15 ppm	24	0.091± 0.017	1.105± 0.095	0.494± 0.015
0.5 ppm	25	0.091± 0.035	1.098± 0.112	0.493± 0.012
1.5 ppm	25	0.078± 0.018	0.996± 0.106**	0.486± 0.014**

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

TABLE K1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0926
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	24	29.8± 3.3	0.049± 0.016	0.758± 0.130	0.667± 0.061	0.569± 0.084	2.047± 0.176
0.15 ppm	24	29.5± 2.7	0.046± 0.007	0.829± 0.213	0.669± 0.050	0.564± 0.044	2.046± 0.112
0.5 ppm	25	29.6± 2.9	0.044± 0.008	0.806± 0.165	0.643± 0.042	0.583± 0.054	1.988± 0.131
1.5 ppm	24	25.4± 1.9**	0.051± 0.015	0.934± 0.151**	0.668± 0.052	0.634± 0.064**	2.158± 0.166*

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

STUDY NO. : 0926
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	24	0.245± 0.101	4.549± 0.255	1.606± 0.168
0.15 ppm	24	0.241± 0.035	4.514± 0.186	1.623± 0.128
0.5 ppm	25	0.230± 0.032	4.384± 0.211	1.624± 0.152
1.5 ppm	24	0.244± 0.043	4.473± 0.217	1.856± 0.130**

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

TABLE K2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0926
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	25	22.1± 1.9	0.073± 0.010	0.120± 0.017	0.672± 0.038	0.739± 0.079	1.933± 0.113
0.15 ppm	24	22.2± 1.6	0.067± 0.008	0.135± 0.076	0.668± 0.038	0.726± 0.046	1.891± 0.095
0.5 ppm	25	21.7± 1.7	0.071± 0.011	0.124± 0.021	0.703± 0.065	0.764± 0.080	1.968± 0.122
1.5 ppm	25	20.9± 2.3	0.071± 0.012	0.126± 0.028	0.660± 0.056	0.937± 0.833	1.898± 0.141

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

STUDY NO. : 0926
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	25	0.467± 0.334	5.114± 0.320	2.270± 0.166
0.15 ppm	24	0.408± 0.065	4.970± 0.202	2.232± 0.124
0.5 ppm	25	0.417± 0.142	5.054± 0.209	2.284± 0.155
1.5 ppm	25	0.374± 0.077	4.777± 0.276**	2.346± 0.199

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

TABLE L1

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : MALE

Organ	Findings	Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
		No. of animals on Study	25	25	25	25
{Respiratory system}						
lung			<25>	<25>	<25>	<25>
	bronchiolar-alveolar adenoma		2 (8%)	6 (24%)	6 (24%)	3 (12%)
	hemangioma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
	bronchiolar-alveolar carcinoma		0 (0%)	1 (4%)	5 (20%)	0 (0%)
{Hematopoietic system}						
lymph node			<25>	<25>	<25>	<25>
	malignant lymphoma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
spleen			<25>	<25>	<25>	<25>
	hemangioma		2 (8%)	1 (4%)	0 (0%)	0 (0%)
	hemangiosarcoma		0 (0%)	1 (4%)	0 (0%)	0 (0%)
{Digestive system}						
salivary gl			<25>	<25>	<25>	<25>
	hemangioma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
liver			<25>	<25>	<25>	<25>
	hepatocellular adenoma		1 (4%)	0 (0%)	5 (20%)	1 (4%)
{Special sense organs/appendage}						
Harder gl			<25>	<25>	<25>	<25>
	adenoma		0 (0%)	1 (4%)	0 (0%)	0 (0%)
{Musculoskeletal system}						
muscle			<25>	<25>	<25>	<25>
	hemangioma		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

Organ	Findings	Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
		No. of animals on Study	25	25	25	25
{Body cavities}						
pleura			<25>	<25>	<25>	<25>
	hemangiosarcoma		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	0.15 ppm 25	0.5 ppm 25	1.5 ppm 25
{Integumentary system/appandage}						
skin/app			<25>	<25>	<25>	<25>
	squamous cell papilloma		0 (0%)	1 (4%)	0 (0%)	1 (4%)
{Respiratory system}						
nasal cavit			<25>	<25>	<25>	<25>
	papilloma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
lung			<25>	<25>	<25>	<25>
	bronchiolar-alveolar adenoma		1 (4%)	0 (0%)	4 (16%)	1 (4%)
	bronchiolar-alveolar carcinoma		1 (4%)	0 (0%)	1 (4%)	2 (8%)
{Hematopoietic system}						
spleen			<25>	<25>	<25>	<25>
	hemangioma		3 (12%)	3 (12%)	3 (12%)	0 (0%)
	hemangiosarcoma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
{Digestive system}						
stomach			<25>	<25>	<25>	<25>
	squamous cell papilloma		1 (4%)	1 (4%)	0 (0%)	1 (4%)
liver			<25>	<25>	<25>	<25>
	hemangioma		1 (4%)	0 (0%)	0 (0%)	0 (0%)
{Reproductive system}						
uterus			<25>	<25>	<25>	<25>
	endometrial stromal sarcoma		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

Organ	Findings	Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
		No. of animals on Study	25	25	25	25
{Special sense organs/appendage}						
Harder gl			<25>	<25>	<25>	<25>
	adenoma		1 (4%)	1 (4%)	2 (8%)	0 (0%)
{Musculoskeletal system}						
muscle			<25>	<25>	<25>	<25>
	hemangioma		0 (0%)	0 (0%)	0 (0%)	1 (4%)
{Body cavities}						
pleura			<25>	<25>	<25>	<25>
	hemangiosarcoma		0 (0%)	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	0.15 ppm	0.5 ppm	1.5 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	6/25(24.0)	6/25(24.0)	3/25(12.0)
Adjusted rates(b)	8.33	24.00	24.00	12.50
Terminal rates(c)	2/24(8.3)	5/24(20.8)	6/25(24.0)	3/24(12.5)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.6085			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.7370			
Fisher Exact test(e)		P = 0.1234	P = 0.1234	P = 0.5000
SITE : lung				
TUMOR : bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	0/25(0.0)	1/25(4.0)	5/25(20.0)	0/25(0.0)
Adjusted rates(b)	0.00	4.17	20.00	0.00
Terminal rates(c)	0/24(0.0)	1/24(4.2)	5/25(20.0)	0/24(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.6351			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.6787			
Fisher Exact test(e)		P = 0.5000	P = 0.0251*	P = N.C.
SITE : spleen				
TUMOR : hemangioma				
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	4.17	0.00	0.00
Terminal rates(c)	2/24(8.3)	1/24(4.2)	0/25(0.0)	0/24(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9551			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.1425			
Fisher Exact test(e)		P = 0.5000	P = 0.2449	P = 0.2449

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
SITE : liver				
TUMOR : hepatocellular adenoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	0/25(0.0)	5/25(20.0)	1/25(4.0)
Adjusted rates(b)	4.17	0.00	20.00	4.17
Terminal rates(c)	1/24(4.2)	0/24(0.0)	5/25(20.0)	1/24(4.2)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.4176			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.8735			
Fisher Exact test(e)		P = 0.5000	P = 0.0947	P = 0.7551
SITE : ALL SITE				
TUMOR : hemangioma				
Tumor rate				
Overall rates(a)	3/25(12.0)	1/25(4.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.33	4.17	0.00	0.00
Terminal rates(c)	2/24(8.3)	1/24(4.2)	0/25(0.0)	0/24(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 1.0000 ?			
Prevalence method(d)	P = 0.9551			
Combined analysis(d)	P = 0.9813			
Cochran-Armitage test(e)	P = 0.0808			
Fisher Exact test(e)		P = 0.3046	P = 0.1173	P = 0.1173
SITE : ALL SITE				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	3/25(12.0)	2/25(8.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.33	4.17	0.00	0.00
Terminal rates(c)	2/24(8.3)	1/24(4.2)	0/25(0.0)	0/24(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.8745			
Prevalence method(d)	P = 0.9551			
Combined analysis(d)	P = 0.9856			
Cochran-Armitage test(e)	P = 0.0609			
Fisher Exact test(e)		P = 0.5000	P = 0.1173	P = 0.1173

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma,bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	7/25(28.0)	9/25(36.0)	3/25(12.0)
Adjusted rates(b)	8.33	28.00	36.00	12.50
Terminal rates(c)	2/24(8.3)	6/24(25.0)	9/25(36.0)	3/24(12.5)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.6783			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.6032			
Fisher Exact test(e)		P = 0.0692	P = 0.0187*	P = 0.5000
SITE : spleen				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	2/25(8.0)	0/25(0.0)	0/25(0.0)
Adjusted rates(b)	8.33	4.17	0.00	0.00
Terminal rates(c)	2/24(8.3)	1/24(4.2)	0/25(0.0)	0/24(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.5359			
Prevalence method(d)	P = 0.9551			
Combined analysis(d)	P = 0.9673			
Cochran-Armitage test(e)	P = 0.1063			
Fisher Exact test(e)		P = 0.6954	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	0.15 ppm	0.5 ppm	1.5 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	0/25(0.0)	4/25(16.0)	1/25(4.0)
Adjusted rates(b)	4.00	0.00	16.00	4.00
Terminal rates(c)	1/25(4.0)	0/24(0.0)	4/25(16.0)	1/25(4.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.4094			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.8430			
Fisher Exact test(e)		P = 0.5000	P = 0.1743	P = 0.7551
SITE : lung				
TUMOR : bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	0/25(0.0)	1/25(4.0)	2/25(8.0)
Adjusted rates(b)	4.00	0.00	4.00	8.00
Terminal rates(c)	1/25(4.0)	0/24(0.0)	1/25(4.0)	2/25(8.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.1368			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.2386			
Fisher Exact test(e)		P = 0.5000	P = 0.7551	P = 0.5000
SITE : spleen				
TUMOR : hemangioma				
Tumor rate				
Overall rates(a)	3/25(12.0)	3/25(12.0)	3/25(12.0)	0/25(0.0)
Adjusted rates(b)	12.00	12.50	12.00	0.00
Terminal rates(c)	3/25(12.0)	3/24(12.5)	3/25(12.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9681			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.0843			
Fisher Exact test(e)		P = 0.6664	P = 0.6664	P = 0.1173

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
SITE : Harderian gland				
TUMOR : adenoma				
Tumor rate				
Overall rates(a)	1/25(4.0)	1/25(4.0)	2/25(8.0)	0/25(0.0)
Adjusted rates(b)	4.00	4.17	8.00	0.00
Terminal rates(c)	1/25(4.0)	1/24(4.2)	2/25(8.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.7944			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.3827			
Fisher Exact test(e)		P = 0.7551	P = 0.5000	P = 0.5000
SITE : ALL SITE				
TUMOR : hemangioma				
Tumor rate				
Overall rates(a)	4/25(16.0)	3/25(12.0)	3/25(12.0)	1/25(4.0)
Adjusted rates(b)	16.00	12.50	12.00	4.00
Terminal rates(c)	4/25(16.0)	3/24(12.5)	3/25(12.0)	1/25(4.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9175			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.1782			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.1743
SITE : ALL SITE				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	4/25(16.0)	4/25(16.0)	3/25(12.0)	1/25(4.0)
Adjusted rates(b)	16.00	12.50	12.00	4.00
Terminal rates(c)	4/25(16.0)	3/24(12.5)	3/25(12.0)	1/25(4.0)
Statistical analysis				
Peto test				
Standard method(d)	P = 0.5339			
Prevalence method(d)	P = 0.9175			
Combined analysis(d)	P = 0.9389			
Cochran-Armitage test(e)	P = 0.1335			
Fisher Exact test(e)		P = 0.6490	P = 0.5000	P = 0.1743

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma,bronchiolar-alveolar carcinoma				
Tumor rate				
Overall rates(a)	2/25(8.0)	0/25(0.0)	5/25(20.0)	3/25(12.0)
Adjusted rates(b)	8.00	0.00	20.00	12.00
Terminal rates(c)	2/25(8.0)	0/24(0.0)	5/25(20.0)	3/25(12.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.1864			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.3541			
Fisher Exact test(e)		P = 0.2449	P = 0.2087	P = 0.5000
SITE : spleen				
TUMOR : hemangioma,hemangiosarcoma				
Tumor rate				
Overall rates(a)	4/25(16.0)	3/25(12.0)	3/25(12.0)	0/25(0.0)
Adjusted rates(b)	16.00	12.50	12.00	0.00
Terminal rates(c)	4/25(16.0)	3/24(12.5)	3/25(12.0)	0/25(0.0)
Statistical analysis				
Peto test				
Standard method(d)	P = ------			
Prevalence method(d)	P = 0.9835			
Combined analysis(d)	P = ------			
Cochran-Armitage test(e)	P = 0.0508			
Fisher Exact test(e)		P = 0.5000	P = 0.5000	P = 0.0549

(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

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER :

Jic:CB6F1-Tg rasH2@Jcl MALE MICE

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER： Jic:CB6F1-Tg rasH2@Jcl MALE MICE

Organs Tumors	No. of animals examined	No. of animals bearing tumor	Incidence (%)	Min.－Max. (%)
Lung	150			
Bronchiolar-alveolar adenoma (A)		9	6.0	0 － 12
Bronchiolar-alveolar carcinoma (B)		0	0	0
A+B		9	6.0	0 － 12
Liver				
Hepatocellular adenoma	150	1	0.7	0 － 4

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

TABLE N2

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : Jic:CB6F1-Tg rasH2@Jcl FEMALE MICE

HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER： Jic:CB6F1-Tg rasH2@Jcl FEMALE MICE

Organs Tumors	No. of animals examined	No. of animals bearing tumor	Incidence (%)	Min.－Max. (%)
Lung	150			
Bronchiolar-alveolar adenoma (A)		6	4.0	0 － 12
Bronchiolar-alveolar carcinoma (B)		4	2.7	0 － 8
A+B		10	6.7	0 － 16
Nasal cavity				
Papilloma	150	0	0	0

6 carcinogenicity studies examined in Japan Bioassay Research Center were used.
Study No.： 0886, 0887, 0901, 0905, 0913, 0919

TABLE O

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR : MALE

Organ	Findings	Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
		No. of Animals on Study	25	25	25	25
{Respiratory system}						
nasal cavit			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
lung			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
{Hematopoietic system}						
bone marrow			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
thymus			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
spleen			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
{Digestive system}						
liver			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
{Urinary system}						
kidney			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
urin bladd			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
{Endocrine system}						
adrenal			<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

Organ	Findings	Group Name / No. of Animals on Study	Control	0.15 ppm	0.5 ppm	1.5 ppm
{Reproductive system}						
epididymis			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
{Special sense organs/appendage}						
eye			<25>	<25>	<25>	<25>
	leukemic cell infiltration		0	0	0	1
Harder gl			<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 P1

HISTOPATHOLOGICAL FINDINGS :

NON-NEOPLASTIC LESIONS : MALE

Organ	Findings	Group Name / No. of Animals on Study / Grade	Control 25 1+ (%)	Control 25 2+ (%)	Control 25 3+ (%)	Control 25 4+ (%)	0.15 ppm 25 1+ (%)	0.15 ppm 25 2+ (%)	0.15 ppm 25 3+ (%)	0.15 ppm 25 4+ (%)	0.5 ppm 25 1+ (%)	0.5 ppm 25 2+ (%)	0.5 ppm 25 3+ (%)	0.5 ppm 25 4+ (%)	1.5 ppm 25 1+ (%)	1.5 ppm 25 2+ (%)	1.5 ppm 25 3+ (%)	1.5 ppm 25 4+ (%)
{Respiratory system}																		
nasal cavit		<25>					<25>				<25>				<25>			
	hyperkeratosis		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)
	eosinophilic change:olfactory epithelium		7 (28)	9 (36)	0 (0)	0 (0)	6 (24)	6 (24)	0 (0)	0 (0)	8 (32)	7 (28)	0 (0)	0 (0)	1 (4)	2 (8)	0 (0)	0 (0) **
	eosinophilic change:respiratory epithelium		11 (44)	5 (20)	0 (0)	0 (0)	10 (40)	5 (20)	0 (0)	0 (0)	10 (40)	4 (16)	0 (0)	0 (0)	8 (32)	0 (0)	0 (0)	0 (0) *
	respiratory metaplasia:olfactory epithelium		1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	8 (32)	1 (4)	0 (0)	0 (0) *
	respiratory metaplasia:gland		18 (72)	7 (28)	0 (0)	0 (0)	21 (84)	3 (12)	0 (0)	0 (0)	22 (88)	3 (12)	0 (0)	0 (0)	21 (84)	4 (16)	0 (0)	0 (0)
	transitional cell hyperplasia		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	14 (56)	11 (44)	0 (0)	0 (0) **	12 (48)	13 (52)	0 (0)	0 (0) **
	atrophy:olfactory 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)	7 (28)	0 (0)	0 (0)	0 (0) *
nasopharynx		<25>					<25>				<25>				<25>			
	eosinophilic change:respiratory epithelium		20 (80)	0 (0)	0 (0)	0 (0)	19 (76)	0 (0)	0 (0)	0 (0)	16 (64)	0 (0)	0 (0)	0 (0)	5 (20)	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				0.15 ppm				0.5 ppm				1.5 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+ (%)
{Respiratory system}																		
lung			<25>				<25>				<25>				<25>			
	eosinophilic crystalline pneumonia		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)
{Hematopoietic system}																		
thymus			<25>				<25>				<25>				<25>			
	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>				<25>				<25>				<25>			
	deposit of melanin		3 (12)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	8 (32)	0 (0)	0 (0)	0 (0)	4 (16)	0 (0)	0 (0)	0 (0)
	extramedullary hematopoiesis		3 (12)	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)
{Circulatory system}																		
heart			<25>				<25>				<25>				<25>			
	mineralization		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 / Grade	Control 1+ (%)	Control 2+ (%)	Control 3+ (%)	Control 4+ (%)	0.15 ppm 1+ (%)	0.15 ppm 2+ (%)	0.15 ppm 3+ (%)	0.15 ppm 4+ (%)	0.5 ppm 1+ (%)	0.5 ppm 2+ (%)	0.5 ppm 3+ (%)	0.5 ppm 4+ (%)	1.5 ppm 1+ (%)	1.5 ppm 2+ (%)	1.5 ppm 3+ (%)	1.5 ppm 4+ (%)
{Digestive system}																		
stomach		No. of Animals on Study	25				25				25				25			
	ulcer:forestomach		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)
	hyperplasia:forestomach		2 (8)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)
	erosion:glandular stomach		2 (8)	1 (4)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)
	ulcer:glandular stomach		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)
	hyperplasia:glandular stomach		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)
liver		No. of Animals on Study	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		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)
	fatty change:central		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)

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	Grade	Control	0.15 ppm	0.5 ppm	1.5 ppm
		No. of Animals on Study	25	25	25	25
{Digestive system}						
liver			<25>	<25>	<25>	<25>
	granulation	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)
	acidophilic cell focus	1+	0 (0)	0 (0)	1 (4)	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)
	basophilic cell focus	1+	2 (8)	0 (0)	1 (4)	0 (0)
		2+	1 (4)	1 (4)	0 (0)	0 (0)
		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>
	regeneration:proximal tubule	1+	0 (0)	0 (0)	0 (0)	2 (8)
		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>
	Rathke pouch	1+	0 (0)	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)

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+ (%)	0.15 ppm 1+ (%)	0.15 ppm 2+ (%)	0.15 ppm 3+ (%)	0.15 ppm 4+ (%)	0.5 ppm 1+ (%)	0.5 ppm 2+ (%)	0.5 ppm 3+ (%)	0.5 ppm 4+ (%)	1.5 ppm 1+ (%)	1.5 ppm 2+ (%)	1.5 ppm 3+ (%)	1.5 ppm 4+ (%)
{Endocrine system}																		
parathyroid		No. of Animals on Study	25				25				25				25			
	ultimobranchial body remanet	<25>	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)
{Reproductive system}																		
testis		No. of Animals on Study	25				25				25				25			
	tubular atrophy	<25>	1 (4)	1 (4)	0 (0)	0 (0)	2 (8)	1 (4)	1 (4)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
epididymis		No. of Animals on Study	25				25				25				25			
	inflammatory infiltration	<25>	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)
	spermatogenic granuloma	<25>	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)
	debris of spermatic elements	<25>	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)
{Special sense organs/appendage}																		
Harder gl		No. of Animals on Study	25				25				25				25			
	hyperplasia	<25>	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				0.15 ppm				0.5 ppm				1.5 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	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)

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 P2

HISTOPATHOLOGICAL FINDINGS :

NON-NEOPLASTIC LESIONS : FEMALE

Organ	Findings	Group Name	Control				0.15 ppm				0.5 ppm				1.5 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>			
	exudate		0	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)
	eosinophilic change:olfactory epithelium		13	11	0	0	15	9	0	0	13	6	0	0	11	5	0	0 *
			(52)	(44)	(0)	(0)	(60)	(36)	(0)	(0)	(52)	(24)	(0)	(0)	(44)	(20)	(0)	(0)
	eosinophilic change:respiratory epithelium		13	12	0	0	11	13	1	0	7	14	0	0	10	11	0	0
			(52)	(48)	(0)	(0)	(44)	(52)	(4)	(0)	(28)	(56)	(0)	(0)	(40)	(44)	(0)	(0)
	respiratory metaplasia:olfactory epithelium		0	0	0	0	0	0	0	0	0	0	0	0	5	1	0	0 *
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(20)	(4)	(0)	(0)
	respiratory metaplasia:gland		19	6	0	0	13	10	2	0	16	6	2	0	11	13	1	0
			(76)	(24)	(0)	(0)	(52)	(40)	(8)	(0)	(64)	(24)	(8)	(0)	(44)	(52)	(4)	(0)
	transitional cell hyperplasia		0	0	0	0	0	0	0	0	16	3	0	0 **	15	8	0	0 **
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(64)	(12)	(0)	(0)	(60)	(32)	(0)	(0)
	atrophy:olfactory epithelium		0	0	0	0	0	0	0	0	1	0	0	0	2	0	0	0
			(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
nasopharynx			<25>				<25>				<25>				<25>			
	eosinophilic change:respiratory epithelium		25	0	0	0	25	0	0	0	22	0	0	0	24	0	0	0
			(100)	(0)	(0)	(0)	(100)	(0)	(0)	(0)	(88)	(0)	(0)	(0)	(96)	(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				0.15 ppm				0.5 ppm				1.5 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>			
	hemorrhage		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)
	fibrosis:focal		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)
	bronchiolar-alveolar 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)
{Hematopoietic system}																		
spleen			<25>				<25>				<25>				<25>			
	deposit of melanin		6	0	0	0	4	0	0	0	5	0	0	0	1	0	0	0
			(24)	(0)	(0)	(0)	(16)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	extramedullary hematopoiesis		2	0	0	0	0	0	0	0	0	0	0	0	2	0	0	0
			(8)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
{Digestive system}																		
stomach			<25>				<25>				<25>				<25>			
	erosion:forestomach		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	Grade	Control 1+ (%)	Control 2+ (%)	Control 3+ (%)	Control 4+ (%)	0.15 ppm 1+ (%)	0.15 ppm 2+ (%)	0.15 ppm 3+ (%)	0.15 ppm 4+ (%)	0.5 ppm 1+ (%)	0.5 ppm 2+ (%)	0.5 ppm 3+ (%)	0.5 ppm 4+ (%)	1.5 ppm 1+ (%)	1.5 ppm 2+ (%)	1.5 ppm 3+ (%)	1.5 ppm 4+ (%)
		No. of Animals on Study	25				25				25				25			
{Digestive system}																		
stomach			<25>				<25>				<25>				<25>			
	hyperplasia:forestomach		4	0	0	0	5	0	0	0	1	0	0	0	1	0	0	0
			(16)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	erosion:glandular stomach		7	0	0	0	3	0	0	0	5	0	0	0	4	0	0	0
			(28)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(16)	(0)	(0)	(0)
	hyperplasia:glandular stomach		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)
liver			<25>				<25>				<25>				<25>			
	inflammatory cell nest		1	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
			(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	basophilic cell focus		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)
{Urinary system}																		
kidney			<25>				<25>				<25>				<25>			
	regeneration:proximal 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)

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 1+ (%)	Control 2+ (%)	Control 3+ (%)	Control 4+ (%)	0.15 ppm 1+ (%)	0.15 ppm 2+ (%)	0.15 ppm 3+ (%)	0.15 ppm 4+ (%)	0.5 ppm 1+ (%)	0.5 ppm 2+ (%)	0.5 ppm 3+ (%)	0.5 ppm 4+ (%)	1.5 ppm 1+ (%)	1.5 ppm 2+ (%)	1.5 ppm 3+ (%)	1.5 ppm 4+ (%)
{Endocrine system}																		
parathyroid		25	<25>				<25>				<25>				<25>			
	ultimobranchial body remanet		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)
{Reproductive system}																		
ovary		25	<25>				<25>				<25>				<25>			
	thrombus		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
uterus		25	<25>				<25>				<25>				<25>			
	cystic endometrial hyperplasia		11 (44)	0 (0)	0 (0)	0 (0)	14 (56)	0 (0)	0 (0)	0 (0)	12 (48)	0 (0)	0 (0)	0 (0)	10 (40)	0 (0)	0 (0)	0 (0)
{Special sense organs/appendage}																		
Harder gl		25	<25>				<25>				<25>				<25>			
	hyperplasia		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)
{Musculoskeletal system}																		
bone		25	<25>				<25>				<25>				<25>			
	necrosis:focal		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	1 (4)	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 Q1

CAUSE OF DEATH : MALE

STUDY NO. : 0926 COUSE OF DEATH (SUMMARY)
ANIMAL : CByB6F1-Tg(HRAS)2Jic (tg/wt) (0- 27W)
SEX : MALE

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
Number of Dead and Moribund Animal	1	1	0	1
tumor d:leukemia	0	0	0	1
tumor d:spleen	0	1	0	0
tumor d:thoracic cav	1	0	0	0

TABLE Q2

CAUSE OF DEATH : FEMALE

Group Name	Control	0.15 ppm	0.5 ppm	1.5 ppm
Number of Dead and Moribund Animal	0	1	0	0
tumor d:pleura	0	1	0	0