

二酸化窒素の p53KO マウスを用いた吸入による
中期がん原性試験報告書

試験番号 : 0904

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TABLE A

CONCENTRATIONS OF NITROGEN DIOXIDE
IN THE INHALATION CHAMBER
OF p53KO MICE IN THE 26-WEEK
CARCINOGENICITY STUDY

CONCENTRATIONS OF NITROGEN DIOXIDE IN THE INHALATION CHAMBER
OF p53KO MICE IN THE 26-WEEK CARCINOGENICITY STUDY

Group Name	Concentration(ppm) Mean \pm S.D.
Control	0.0 \pm 0.0
10 ppm	10.3 \pm 0.2
20 ppm	20.5 \pm 0.3
40 ppm	40.6 \pm 0.5

TABLE B1

SURVIVAL ANIMAL NUMBERS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

Group Name	Animals At start	Administration (Weeks)													
		0	1	2	3	4	5	6	7	8	9	10	11	12	13
Control	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
10 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
20 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
40 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0
		Number of survival/ Number of effective animals Survival rate(%)													

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26
 SEX : MALE

SURVIVAL ANIMAL NUMBERS

Group Name	Animals At start	Administration (Weeks)													
		14	15	16	17	18	19	20	21	22	23	24	25	26	
Control	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	
10 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	
20 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	
40 ppm	25	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	
		Number of survival/ Number of effective animals Survival rate(%)													

TABLE B2

SURVIVAL ANIMAL NUMBERS : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

Group Name	Animals At start	Administration (Weeks)													
		0	1	2	3	4	5	6	7	8	9	10	11	12	13
Control	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
10 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
20 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
40 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0
		Number of survival/ Number of effective animals Survival rate(%)													

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26
 SEX : FEMALE

SURVIVAL ANIMAL NUMBERS

Group Name	Animals At start	Administration (Weeks)													
		14	15	16	17	18	19	20	21	22	23	24	25	26	
Control	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	
10 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	24/25 96.0	
20 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	
40 ppm	25	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	25/25 100.0	
		Number of survival/ Number of effective animals Survival rate(%)													

TABLE C1

CLINICAL OBSERVATION : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

PAGE : 1

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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	1	1	1	1	1	1
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	1	0	0	0	0	0	0
ROUGH FUR	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ABNORMAL GROWTH OF TEETH	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	1	1	1	0	0	0	0	0	0
MALOCCLUSION	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	1	1	1	0	0	0	0	0	0
IRREGULAR BREATHING	Control	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	1	0	0	0	0	0	0
NON REMARKABLE	Control	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	10 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	20 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	40 ppm	25	25	25	25	25	24	24	24	24	24	24	24	24	24

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : MALE

PAGE : 2

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	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	1	1	1	1	1	1	1	1	1	1	1	1
LOCOMOTOR MOVEMENT DECR	Control	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0
ROUGH FUR	Control	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	1	1	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0
ABNORMAL GROWTH OF TEETH	Control	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0
MALOCCLUSION	Control	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	1	1	1	1	1	1	1	1	1
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	20 ppm	25	25	25	24	24	24	24	24	24	24	24	24
	40 ppm	24	24	24	24	24	24	24	24	24	24	24	24

TABLE C2

CLINICAL OBSERVATION : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : FEMALE

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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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	20 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	40 ppm	25	25	25	25	25	25	25	25	25	25	25	25	25	25

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1 26

CLINICAL OBSERVATION (SUMMARY)
 ALL ANIMALS

SEX : FEMALE

PAGE : 4

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
	10 ppm	0	0	0	0	0	0	1	1	1	1	1	1
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 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
	10 ppm	0	0	0	0	1	1	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0
EROSION	Control	1	1	1	1	1	1	2	1	1	1	1	1
	10 ppm	0	0	0	0	1	1	1	1	1	1	1	1
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	1	1	1
IRREGULAR BREATHING	Control	0	0	0	0	0	0	0	0	0	0	0	0
	10 ppm	0	0	0	0	1	1	0	0	0	0	0	0
	20 ppm	0	0	0	0	0	0	0	0	0	0	0	0
	40 ppm	0	0	0	0	0	0	0	0	0	0	0	0
NON REMARKABLE	Control	24	24	24	24	24	24	23	23	23	23	23	23
	10 ppm	25	25	25	25	23	23	23	23	23	23	23	23
	20 ppm	25	25	25	25	25	25	25	25	25	25	25	25
	40 ppm	25	25	25	25	25	25	25	25	25	24	24	24

TABLE D1

BODY WEIGHT CHANGES AND SURVIVAL ANIMAL

NUMBERS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

MEAN BODY WEIGHTS AND SURVIVAL

Week-Day on Study	Control		10 ppm		20 ppm		40 ppm				
	Av. Wt.	No. of Surviv. <25>	Av. Wt.	% of cont. <25>	No. of Surviv.	Av. Wt.	% of cont. <25>	No. of Surviv.	Av. Wt.	% of cont. <25>	No. of Surviv.
0-0	24.9 (25)	25/25	24.9 (25)	100	25/25	24.9 (25)	100	25/25	24.9 (25)	100	25/25
1-7	26.0 (25)	25/25	25.7 (25)	99	25/25	25.7 (25)	99	25/25	23.8 (25)	92	25/25
2-7	27.0 (25)	25/25	26.6 (25)	99	25/25	26.6 (25)	99	25/25	25.5 (25)	94	25/25
3-7	28.0 (25)	25/25	27.5 (25)	98	25/25	27.8 (25)	99	25/25	26.8 (25)	96	25/25
4-7	28.9 (25)	25/25	28.2 (25)	98	25/25	28.6 (25)	99	25/25	27.3 (25)	94	25/25
5-7	29.2 (25)	25/25	28.7 (25)	98	25/25	29.1 (25)	100	25/25	27.7 (25)	95	25/25
6-7	29.9 (25)	25/25	29.4 (25)	98	25/25	29.9 (25)	100	25/25	28.8 (25)	96	25/25
7-7	30.3 (25)	25/25	29.7 (25)	98	25/25	30.3 (25)	100	25/25	29.0 (25)	96	25/25
8-7	30.8 (25)	25/25	30.1 (25)	98	25/25	30.7 (25)	100	25/25	29.3 (25)	95	25/25
9-7	31.2 (25)	25/25	30.7 (25)	98	25/25	31.2 (25)	100	25/25	30.0 (24)	96	24/25
10-7	31.7 (25)	25/25	31.0 (25)	98	25/25	31.5 (25)	99	25/25	30.4 (24)	96	24/25
11-7	32.1 (25)	25/25	31.4 (25)	98	25/25	31.9 (25)	99	25/25	30.4 (24)	95	24/25
12-7	32.4 (25)	25/25	31.8 (25)	98	25/25	32.3 (25)	100	25/25	30.9 (24)	95	24/25
13-7	33.0 (25)	25/25	32.4 (25)	98	25/25	32.8 (25)	99	25/25	31.3 (24)	95	24/25
14-7	33.3 (25)	25/25	32.7 (25)	98	25/25	32.8 (25)	98	25/25	31.6 (24)	95	24/25
15-7	33.6 (25)	25/25	32.7 (25)	97	25/25	33.0 (25)	98	25/25	32.1 (24)	96	24/25
16-7	33.9 (25)	25/25	33.1 (25)	98	25/25	33.2 (25)	98	25/25	31.9 (24)	94	24/25
17-7	34.2 (25)	25/25	33.7 (25)	99	25/25	33.8 (25)	99	25/25	32.2 (24)	94	24/25
18-7	35.0 (25)	25/25	34.0 (25)	97	25/25	33.8 (25)	97	25/25	32.7 (24)	93	24/25
19-7	35.2 (25)	25/25	34.3 (25)	97	25/25	34.4 (25)	98	25/25	32.9 (24)	93	24/25
20-7	35.7 (25)	25/25	35.1 (25)	98	25/25	35.1 (25)	98	25/25	33.2 (24)	93	24/25
21-7	36.1 (25)	25/25	35.5 (25)	98	25/25	35.2 (25)	98	25/25	33.3 (24)	92	24/25
22-7	36.5 (25)	25/25	36.2 (25)	99	25/25	35.8 (25)	98	25/25	33.7 (24)	92	24/25
23-7	36.9 (25)	25/25	36.4 (25)	99	25/25	36.2 (25)	98	25/25	34.0 (24)	92	24/25
24-7	37.4 (25)	25/25	36.6 (25)	98	25/25	36.4 (25)	97	25/25	34.5 (24)	92	24/25
25-7	37.8 (25)	25/25	36.8 (25)	97	25/25	36.3 (25)	96	25/25	34.4 (24)	91	24/25
26-7	38.0 (25)	25/25	37.6 (25)	99	25/25	36.6 (25)	96	25/25	34.8 (24)	92	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. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

MEAN BODY WEIGHTS AND SURVIVAL

Week-Day on Study	Control		10 ppm		20 ppm		40 ppm				
	Av. Wt.	No. of Surviv. <25>	Av. Wt.	% of cont. <25>	No. of Surviv.	Av. Wt.	% of cont. <25>	No. of Surviv.	Av. Wt.	% of cont. <25>	No. of Surviv.
0-0	20.6 (25)	25/25	20.6 (25)	100	25/25	20.6 (25)	100	25/25	20.6 (25)	100	25/25
1-7	21.8 (25)	25/25	21.5 (25)	99	25/25	21.6 (25)	99	25/25	20.9 (25)	96	25/25
2-7	21.9 (25)	25/25	22.2 (25)	101	25/25	22.1 (25)	101	25/25	21.7 (25)	99	25/25
3-7	22.9 (25)	25/25	22.8 (25)	100	25/25	23.1 (25)	101	25/25	22.3 (25)	97	25/25
4-7	23.4 (25)	25/25	23.3 (25)	100	25/25	23.8 (25)	102	25/25	23.2 (25)	99	25/25
5-7	23.8 (25)	25/25	23.8 (25)	100	25/25	23.8 (25)	100	25/25	23.7 (25)	100	25/25
6-7	24.4 (25)	25/25	23.8 (25)	98	25/25	24.4 (25)	100	25/25	24.0 (25)	98	25/25
7-7	24.3 (25)	25/25	24.2 (25)	100	25/25	24.7 (25)	102	25/25	24.3 (25)	100	25/25
8-7	24.7 (25)	25/25	24.7 (25)	100	25/25	24.7 (25)	100	25/25	24.7 (25)	100	25/25
9-7	24.7 (25)	25/25	24.8 (25)	100	25/25	25.3 (25)	102	25/25	25.2 (25)	102	25/25
10-7	24.7 (25)	25/25	25.0 (25)	101	25/25	25.1 (25)	102	25/25	25.1 (25)	102	25/25
11-7	25.4 (25)	25/25	25.0 (25)	98	25/25	25.3 (25)	100	25/25	25.1 (25)	99	25/25
12-7	25.2 (25)	25/25	25.3 (25)	100	25/25	25.5 (25)	101	25/25	25.0 (25)	99	25/25
13-7	25.7 (25)	25/25	25.5 (25)	99	25/25	26.0 (25)	101	25/25	25.8 (25)	100	25/25
14-7	26.1 (25)	25/25	26.0 (25)	100	25/25	25.8 (25)	99	25/25	25.8 (25)	99	25/25
15-7	26.3 (25)	25/25	26.1 (25)	99	25/25	26.3 (25)	100	25/25	26.4 (25)	100	25/25
16-7	26.1 (25)	25/25	25.9 (25)	99	25/25	26.3 (25)	101	25/25	26.0 (25)	100	25/25
17-7	26.7 (25)	25/25	26.2 (25)	98	25/25	26.6 (25)	100	25/25	26.7 (25)	100	25/25
18-7	26.9 (25)	25/25	26.9 (25)	100	25/25	26.7 (25)	99	25/25	26.8 (25)	100	25/25
19-7	27.0 (25)	25/25	26.6 (25)	99	25/25	27.1 (25)	100	25/25	27.2 (25)	101	25/25
20-7	26.8 (25)	25/25	26.5 (25)	99	25/25	27.2 (25)	101	25/25	26.8 (25)	100	25/25
21-7	27.2 (25)	25/25	26.5 (24)	97	24/25	27.4 (25)	101	25/25	26.9 (25)	99	25/25
22-7	27.3 (24)	24/25	26.9 (24)	99	24/25	27.5 (25)	101	25/25	27.3 (25)	100	25/25
23-7	27.5 (24)	24/25	27.3 (24)	99	24/25	27.7 (25)	101	25/25	27.4 (25)	100	25/25
24-7	28.1 (24)	24/25	27.4 (24)	98	24/25	28.0 (25)	100	25/25	27.6 (25)	98	25/25
25-7	28.1 (24)	24/25	27.6 (24)	98	24/25	28.4 (25)	101	25/25	27.8 (25)	99	25/25
26-7	28.0 (24)	24/25	27.8 (24)	99	24/25	28.3 (25)	101	25/25	28.1 (25)	100	25/25

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

TABLE D3

BODY WEIGHT CHANGES : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day							
	0-0	1-7	2-7	3-7	4-7	5-7	6-7	
Control	24.9± 1.2	26.0± 1.2	27.0± 1.5	28.0± 1.5	28.9± 1.5	29.2± 1.4	29.9± 1.6	
10 ppm	24.9± 1.2	25.7± 1.1	26.6± 1.1	27.5± 1.2	28.2± 1.3	28.7± 1.3	29.4± 1.4	
20 ppm	24.9± 1.2	25.7± 1.3	26.6± 1.2	27.8± 1.2	28.6± 1.4	29.1± 1.4	29.9± 1.3	
40 ppm	24.9± 1.2	23.8± 1.7**	25.5± 1.9**	26.8± 2.2	27.3± 2.2**	27.7± 2.4	28.8± 2.9	

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration		week-day		9-7	10-7	11-7	12-7	13-7
	7-7	8-7	7-7	8-7					
Control	30.3± 1.5	30.8± 1.7	31.2± 1.7	31.7± 1.7	32.1± 1.8	32.4± 1.9	33.0± 2.1		
10 ppm	29.7± 1.3	30.1± 1.4	30.7± 1.4	31.0± 1.7	31.4± 1.7	31.8± 1.7	32.4± 1.8		
20 ppm	30.3± 1.5	30.7± 1.3	31.2± 1.2	31.5± 1.6	31.9± 1.6	32.3± 1.7	32.8± 1.7		
40 ppm	29.0± 3.0	29.3± 3.3	30.0± 1.8*	30.4± 2.1	30.4± 2.1**	30.9± 2.2*	31.3± 2.4*		

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day							
	14-7	15-7	16-7	17-7	18-7	19-7	20-7	
Control	33.3± 2.2	33.6± 2.0	33.9± 2.4	34.2± 2.5	35.0± 2.6	35.2± 2.5	35.7± 2.8	
10 ppm	32.7± 1.8	32.7± 2.1	33.1± 2.3	33.7± 2.4	34.0± 2.2	34.3± 2.4	35.1± 2.7	
20 ppm	32.8± 2.0	33.0± 1.7	33.2± 2.0	33.8± 2.3	33.8± 3.2	34.4± 3.1	35.1± 2.6	
40 ppm	31.6± 2.4*	32.1± 2.6	31.9± 2.6**	32.2± 2.9*	32.7± 2.7*	32.9± 2.8*	33.2± 2.8**	

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration		week-day		23-7		24-7		25-7		26-7	
	21-7		22-7									
Control	36.1±	2.8	36.5±	3.1	36.9±	3.1	37.4±	3.3	37.8±	3.4	38.0±	3.4
10 ppm	35.5±	2.9	36.2±	3.2	36.4±	3.1	36.6±	3.0	36.8±	3.1	37.6±	3.3
20 ppm	35.2±	2.8	35.8±	3.1	36.2±	3.1	36.4±	3.2	36.3±	3.6	36.6±	4.0
40 ppm	33.3±	2.8**	33.7±	3.4**	34.0±	3.3**	34.5±	3.5**	34.4±	3.5**	34.8±	3.5**

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

Test of Dunnett

TABLE D4

BODY WEIGHT CHANGES : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration		week-day											
	0-0		1-7	2-7	3-7	4-7	5-7	6-7						
Control	20.6±	1.7	21.8±	1.9	21.9±	1.4	22.9±	1.6	23.4±	1.4	23.8±	1.4	24.4±	1.5
10 ppm	20.6±	1.7	21.5±	1.7	22.2±	1.8	22.8±	2.0	23.3±	1.4	23.8±	1.6	23.8±	1.5
20 ppm	20.6±	1.6	21.6±	1.6	22.1±	1.4	23.1±	1.5	23.8±	1.5	23.8±	1.5	24.4±	1.2
40 ppm	20.6±	1.6	20.9±	1.6	21.7±	1.8	22.3±	1.8	23.2±	1.3	23.7±	1.4	24.0±	1.5

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration		week-day		9-7	10-7	11-7	12-7	13-7
	7-7	8-7	8-7	9-7					
Control	24.3± 1.4	24.7± 1.5	24.7± 1.5	24.7± 1.6	25.4± 1.7	25.2± 1.6	25.7± 1.7		
10 ppm	24.2± 1.6	24.7± 1.5	24.8± 1.5	25.0± 1.8	25.0± 1.7	25.3± 1.7	25.5± 1.4		
20 ppm	24.7± 1.5	24.7± 1.3	25.3± 1.6	25.1± 1.3	25.3± 1.3	25.5± 1.6	26.0± 1.3		
40 ppm	24.3± 1.7	24.7± 1.2	25.2± 1.6	25.1± 1.6	25.1± 1.4	25.0± 1.5	25.8± 1.5		

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day							
	14-7	15-7	16-7	17-7	18-7	19-7	20-7	
Control	26.1± 2.0	26.3± 1.9	26.1± 1.7	26.7± 2.0	26.9± 2.0	27.0± 1.9	26.8± 2.0	
10 ppm	26.0± 1.9	26.1± 1.7	25.9± 1.8	26.2± 2.0	26.9± 1.8	26.6± 1.7	26.5± 1.9	
20 ppm	25.8± 1.4	26.3± 1.7	26.3± 1.9	26.6± 1.5	26.7± 1.4	27.1± 1.6	27.2± 1.7	
40 ppm	25.8± 1.6	26.4± 1.6	26.0± 1.5	26.7± 1.6	26.8± 1.8	27.2± 1.7	26.8± 1.7	

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

BODY WEIGHT CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration		week-day		23-7	24-7	25-7	26-7
	21-7	22-7	21-7	22-7				
Control	27.2± 2.1	27.3± 2.0	27.5± 2.0	28.1± 2.0	28.1± 2.4	28.0± 2.3		
10 ppm	26.5± 2.2	26.9± 2.0	27.3± 1.9	27.4± 2.1	27.6± 2.1	27.8± 2.0		
20 ppm	27.4± 1.5	27.5± 1.8	27.7± 1.6	28.0± 1.9	28.4± 2.1	28.3± 1.9		
40 ppm	26.9± 1.7	27.3± 2.2	27.4± 2.3	27.6± 2.3	27.8± 2.2	28.1± 2.2		

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

Test of Dunnett

TABLE E1

FOOD CONSUMPTION CHANGES AND SURVIVAL ANIMAL
NUMBERS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

MEAN FOOD CONSUMPTION (FC) AND SURVIVAL

Week-Day on Study	Control		10 ppm			20 ppm			40 ppm		
	Av. FC.	No. of Surviv. <25>	Av. FC.	% of cont. <25>	No. of Surviv.	Av. FC.	% of cont. <25>	No. of Surviv.	Av. FC.	% of cont. <25>	No. of Surviv.
1-7	3.8 (25)	25/25	3.9 (25)	103	25/25	3.7 (25)	97	25/25	2.6 (25)	68	25/25
2-7	3.6 (25)	25/25	3.7 (25)	103	25/25	3.7 (25)	103	25/25	4.1 (25)	114	25/25
3-7	3.8 (25)	25/25	3.8 (25)	100	25/25	3.9 (25)	103	25/25	3.8 (25)	100	25/25
4-7	3.9 (25)	25/25	3.9 (25)	100	25/25	4.0 (25)	103	25/25	3.9 (25)	100	25/25
5-7	3.8 (25)	25/25	3.9 (25)	103	25/25	4.0 (25)	105	25/25	3.9 (25)	103	25/25
6-7	3.9 (25)	25/25	4.0 (25)	103	25/25	4.1 (25)	105	25/25	3.9 (25)	100	25/25
7-7	3.9 (25)	25/25	3.9 (25)	100	25/25	4.0 (25)	103	25/25	3.7 (25)	95	25/25
8-7	3.9 (25)	25/25	4.0 (25)	103	25/25	4.0 (25)	103	25/25	3.9 (25)	100	25/25
9-7	4.0 (25)	25/25	4.0 (25)	100	25/25	4.0 (25)	100	25/25	3.8 (25)	95	24/25
10-7	4.0 (25)	25/25	3.9 (25)	98	25/25	4.1 (25)	103	25/25	4.0 (24)	100	24/25
11-7	4.0 (25)	25/25	4.1 (25)	103	25/25	4.1 (25)	103	25/25	3.9 (24)	98	24/25
12-7	4.0 (25)	25/25	4.1 (25)	103	25/25	4.2 (25)	105	25/25	4.0 (24)	100	24/25
13-7	4.0 (25)	25/25	4.0 (25)	100	25/25	4.0 (25)	100	25/25	3.9 (24)	98	24/25
14-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.1 (25)	100	25/25	4.0 (24)	98	24/25
15-7	4.0 (25)	25/25	4.0 (25)	100	25/25	4.1 (25)	103	25/25	4.0 (24)	100	24/25
16-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25	4.0 (24)	98	24/25
17-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.1 (25)	100	25/25	3.9 (24)	95	24/25
18-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.1 (25)	100	25/25	4.0 (24)	98	24/25
19-7	4.1 (25)	25/25	4.1 (25)	100	25/25	4.2 (25)	102	25/25	4.0 (24)	98	24/25
20-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.3 (25)	100	25/25	4.2 (24)	98	24/25
21-7	4.4 (25)	25/25	4.3 (25)	98	25/25	4.3 (25)	98	25/25	4.0 (24)	91	24/25
22-7	4.4 (25)	25/25	4.4 (25)	100	25/25	4.4 (25)	100	25/25	4.2 (24)	95	24/25
23-7	4.3 (25)	25/25	4.3 (25)	100	25/25	4.4 (25)	102	25/25	4.2 (24)	98	24/25
24-7	4.4 (25)	25/25	4.4 (25)	100	25/25	4.5 (25)	102	25/25	4.3 (24)	98	24/25
25-7	4.4 (25)	25/25	4.4 (25)	100	25/25	4.3 (25)	98	25/25	4.2 (24)	95	24/25
26-7	4.4 (25)	25/25	4.5 (25)	102	25/25	4.4 (25)	100	25/25	4.4 (24)	100	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. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
UNIT : g
REPORT TYPE : A1 26
SEX : FEMALE

MEAN FOOD CONSUMPTION (FC) AND SURVIVAL

Week-Day on Study	Control		10 ppm			20 ppm			40 ppm		
	Av. FC.	No. of Surviv. <25>	Av. FC.	% of cont. <25>	No. of Surviv.	Av. FC.	% of cont. <25>	No. of Surviv.	Av. FC.	% of cont. <25>	No. of Surviv.
1-7	3.6 (25)	25/25	3.5 (25)	97	25/25	3.6 (25)	100	25/25	2.8 (25)	78	25/25
2-7	3.2 (25)	25/25	3.1 (25)	97	25/25	3.1 (25)	97	25/25	3.6 (25)	113	25/25
3-7	3.4 (25)	25/25	3.3 (25)	97	25/25	3.4 (25)	100	25/25	3.4 (25)	100	25/25
4-7	3.5 (25)	25/25	3.4 (25)	97	25/25	3.5 (25)	100	25/25	3.5 (25)	100	25/25
5-7	3.6 (25)	25/25	3.7 (25)	103	25/25	3.6 (25)	100	25/25	3.6 (25)	100	25/25
6-7	3.8 (25)	25/25	3.5 (25)	92	25/25	3.7 (25)	97	25/25	3.7 (25)	97	25/25
7-7	3.7 (25)	25/25	3.7 (25)	100	25/25	3.7 (25)	100	25/25	3.7 (25)	100	25/25
8-7	3.7 (25)	25/25	3.7 (25)	100	25/25	3.6 (25)	97	25/25	3.7 (25)	100	25/25
9-7	3.8 (25)	25/25	3.7 (25)	97	25/25	3.8 (25)	100	25/25	3.8 (25)	100	25/25
10-7	3.6 (25)	25/25	3.6 (25)	100	25/25	3.6 (25)	100	25/25	3.6 (25)	100	25/25
11-7	3.9 (25)	25/25	3.7 (25)	95	25/25	3.7 (25)	95	25/25	3.6 (25)	92	25/25
12-7	3.7 (25)	25/25	3.8 (25)	103	25/25	3.7 (25)	100	25/25	3.6 (25)	97	25/25
13-7	3.7 (25)	25/25	3.7 (25)	100	25/25	3.7 (25)	100	25/25	3.6 (25)	97	25/25
14-7	3.8 (25)	25/25	3.8 (25)	100	25/25	3.7 (25)	97	25/25	3.7 (25)	97	25/25
15-7	3.9 (25)	25/25	3.7 (25)	95	25/25	3.8 (25)	97	25/25	3.8 (25)	97	25/25
16-7	3.7 (25)	25/25	3.7 (25)	100	25/25	3.8 (25)	103	25/25	3.7 (25)	100	25/25
17-7	3.9 (25)	25/25	3.8 (25)	97	25/25	3.8 (25)	97	25/25	3.8 (25)	97	25/25
18-7	3.8 (25)	25/25	3.8 (25)	100	25/25	3.8 (25)	100	25/25	3.7 (25)	97	25/25
19-7	3.8 (25)	25/25	3.8 (25)	100	25/25	3.9 (25)	103	25/25	3.8 (25)	100	25/25
20-7	3.9 (25)	25/25	4.0 (25)	103	25/25	4.0 (25)	103	25/25	3.8 (25)	97	25/25
21-7	4.1 (25)	25/25	4.0 (25)	98	24/25	4.0 (25)	98	25/25	3.8 (25)	93	25/25
22-7	4.0 (25)	24/25	4.1 (24)	103	24/25	4.1 (25)	103	25/25	4.0 (25)	100	25/25
23-7	4.0 (24)	24/25	4.1 (24)	103	24/25	4.1 (25)	103	25/25	3.8 (25)	95	25/25
24-7	4.1 (24)	24/25	4.1 (24)	100	24/25	4.2 (25)	102	25/25	4.0 (25)	98	25/25
25-7	4.1 (24)	24/25	4.0 (24)	98	24/25	4.0 (25)	98	25/25	4.1 (25)	100	25/25
26-7	4.0 (24)	24/25	4.1 (24)	103	24/25	4.2 (25)	105	25/25	4.2 (25)	105	25/25

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

TABLE E3

FOOD CONSUMPTION CHANGES : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	1-7 (7)	2-7 (7)	3-7 (7)	4-7 (7)	5-7 (7)	6-7 (7)	7-7 (7)
Control	3.8± 0.2	3.6± 0.3	3.8± 0.3	3.9± 0.3	3.8± 0.2	3.9± 0.3	3.9± 0.2
10 ppm	3.9± 0.3	3.7± 0.3	3.8± 0.3	3.9± 0.3	3.9± 0.2	4.0± 0.3	3.9± 0.2
20 ppm	3.7± 0.3	3.7± 0.3	3.9± 0.3	4.0± 0.3	4.0± 0.3	4.1± 0.3	4.0± 0.3
40 ppm	2.6± 0.5**	4.1± 0.5**	3.8± 0.4	3.9± 0.4	3.9± 0.4	3.9± 0.6	3.7± 0.5

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	8-7(7)	9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	14-7(7)
Control	3.9± 0.3	4.0± 0.3	4.0± 0.3	4.0± 0.2	4.0± 0.2	4.0± 0.2	4.1± 0.3
10 ppm	4.0± 0.3	4.0± 0.3	3.9± 0.3	4.1± 0.3	4.1± 0.3	4.0± 0.3	4.1± 0.3
20 ppm	4.0± 0.3	4.0± 0.2	4.1± 0.3	4.1± 0.3	4.2± 0.3	4.0± 0.3	4.1± 0.3
40 ppm	3.9± 0.6	3.8± 0.6	4.0± 0.3	3.9± 0.3	4.0± 0.4	3.9± 0.3	4.0± 0.3

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	15-7(7)	16-7(7)	17-7(7)	18-7(7)	19-7(7)	20-7(7)	21-7(7)
Control	4.0± 0.3	4.1± 0.3	4.1± 0.2	4.1± 0.3	4.1± 0.3	4.3± 0.3	4.4± 0.3
10 ppm	4.0± 0.3	4.1± 0.3	4.1± 0.3	4.1± 0.3	4.1± 0.4	4.3± 0.4	4.3± 0.4
20 ppm	4.1± 0.3	4.2± 0.3	4.1± 0.3	4.1± 0.6	4.2± 0.3	4.3± 0.3	4.3± 0.3
40 ppm	4.0± 0.4	4.0± 0.3	3.9± 0.4	4.0± 0.3	4.0± 0.3	4.2± 0.4	4.0± 0.4**

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : MALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)				
	22-7 (7)	23-7 (7)	24-7 (7)	25-7 (7)	26-7 (7)
Control	4.4± 0.3	4.3± 0.3	4.4± 0.4	4.4± 0.3	4.4± 0.3
10 ppm	4.4± 0.4	4.3± 0.4	4.4± 0.4	4.4± 0.4	4.5± 0.4
20 ppm	4.4± 0.4	4.4± 0.4	4.5± 0.4	4.3± 0.5	4.4± 0.6
40 ppm	4.2± 0.4	4.2± 0.3	4.3± 0.4	4.2± 0.4	4.4± 0.4

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

TABLE E4

FOOD CONSUMPTION CHANGES : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	1-7 (7)	2-7 (7)	3-7 (7)	4-7 (7)	5-7 (7)	6-7 (7)	7-7 (7)
Control	3.6± 0.3	3.2± 0.3	3.4± 0.4	3.5± 0.3	3.6± 0.3	3.8± 0.2	3.7± 0.3
10 ppm	3.5± 0.3	3.1± 0.4	3.3± 0.4	3.4± 0.4	3.7± 0.3	3.5± 0.4	3.7± 0.2
20 ppm	3.6± 0.4	3.1± 0.3	3.4± 0.3	3.5± 0.4	3.6± 0.4	3.7± 0.4	3.7± 0.3
40 ppm	2.8± 0.4**	3.6± 0.4**	3.4± 0.3	3.5± 0.3	3.6± 0.3	3.7± 0.3	3.7± 0.3

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	8-7(7)	9-7(7)	10-7(7)	11-7(7)	12-7(7)	13-7(7)	14-7(7)
Control	3.7± 0.3	3.8± 0.2	3.6± 0.2	3.9± 0.3	3.7± 0.3	3.7± 0.4	3.8± 0.5
10 ppm	3.7± 0.3	3.7± 0.2	3.6± 0.3	3.7± 0.2**	3.8± 0.3	3.7± 0.3	3.8± 0.4
20 ppm	3.6± 0.2	3.8± 0.3	3.6± 0.2	3.7± 0.2*	3.7± 0.2	3.7± 0.2	3.7± 0.3
40 ppm	3.7± 0.3	3.8± 0.2	3.6± 0.2	3.6± 0.2**	3.6± 0.2	3.6± 0.2	3.7± 0.2

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)						
	15-7(7)	16-7(7)	17-7(7)	18-7(7)	19-7(7)	20-7(7)	21-7(7)
Control	3.9± 0.5	3.7± 0.5	3.9± 0.4	3.8± 0.5	3.8± 0.4	3.9± 0.7	4.1± 0.6
10 ppm	3.7± 0.3	3.7± 0.3	3.8± 0.3	3.8± 0.3	3.8± 0.4	4.0± 0.6	4.0± 0.6
20 ppm	3.8± 0.3	3.8± 0.3	3.8± 0.3	3.8± 0.2	3.9± 0.3	4.0± 0.3*	4.0± 0.3
40 ppm	3.8± 0.2	3.7± 0.3	3.8± 0.3	3.7± 0.2	3.8± 0.2	3.8± 0.2	3.8± 0.3

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 UNIT : g
 REPORT TYPE : A1 26
 SEX : FEMALE

FOOD CONSUMPTION CHANGES (SUMMARY)
 ALL ANIMALS

Group Name	Administration week-day(effective)				
	22-7(7)	23-7(7)	24-7(7)	25-7(7)	26-7(7)
Control	4.0± 0.8	4.0± 0.6	4.1± 0.6	4.1± 0.7	4.0± 0.5
10 ppm	4.1± 0.5	4.1± 0.5	4.1± 0.6	4.0± 0.4	4.1± 0.5
20 ppm	4.1± 0.3	4.1± 0.3*	4.2± 0.3	4.0± 0.3	4.2± 0.3
40 ppm	4.0± 0.3	3.8± 0.3	4.0± 0.5	4.1± 0.6	4.2± 0.7

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

Test of Dunnett

TABLE F1

URINALYSIS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 1

Group Name	NO. of Animals	pH_____							CHI	Protein_____					CHI	Glucose_____					CHI	Ketone body					CHI	Occult blood				CHI			
		5.0	6.0	6.5	7.0	7.5	8.0	8.5		-	±	+	2+	3+		4+	-	±	+	2+		3+	4+	-	±	+		2+	3+	4+	-		±	+	2+
Control	23	0	1	7	2	5	5	3		0	13	10	0	0	0		23	0	0	0	0	0		5	10	5	3	0	0		23	0	0	0	0
10 ppm	24	0	2	5	3	7	6	1		1	13	10	0	0	0		24	0	0	0	0	0		8	10	5	1	0	0		24	0	0	0	0
20 ppm	25	0	1	6	5	8	3	2		0	16	7	2	0	0		25	0	0	0	0	0		8	10	7	0	0	0		25	0	0	0	0
40 ppm	24	0	1	6	5	5	5	2		0	14	10	0	0	0		24	0	0	0	0	0		8	11	5	0	0	0		22	2	0	0	0

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

Test of CHI SQUARE

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 2

Group Name	NO. of Animals	Urobilinogen ± + 2+ 3+ 4+	CHI
Control	23	23 0 0 0 0	
10 ppm	24	24 0 0 0 0	
20 ppm	25	25 0 0 0 0	
40 ppm	24	24 0 0 0 0	

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

Test of CHI SQUARE

TABLE F2

URINALYSIS : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE TIME : 1
 SEX : FEMALE REPORT TYPE : A1

URINALYSIS

Group Name	NO. of Animals	pH							CHI	Protein					CHI	Glucose					CHI	Ketone body					CHI	Occult blood			CHI				
		5.0	6.0	6.5	7.0	7.5	8.0	8.5		-	±	+	2+	3+		4+	-	±	+	2+		3+	4+	-	±	+		2+	3+	4+		-	±	+	2+
Control	20	0	7	6	2	2	3	0		8	7	5	0	0	0		20	0	0	0	0	0		2	13	5	0	0	0		20	0	0	0	0
10 ppm	22	0	8	8	2	3	1	0		1	13	8	0	0	0	*	22	0	0	0	0	0		1	17	4	0	0	0		22	0	0	0	0
20 ppm	23	0	7	3	5	2	5	1		3	12	8	0	0	0		23	0	0	0	0	0		4	15	3	1	0	0		23	0	0	0	0
40 ppm	20	0	3	5	5	3	2	2		1	13	6	0	0	0	*	20	0	0	0	0	0		4	13	3	0	0	0		20	0	0	0	0

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

Test of CHI SQUARE

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE TIME : 1
SEX : FEMALE REPORT TYPE : A1

URINALYSIS

Group Name	NO. of Animals	Urobilinogen	CHI
		± + 2+ 3+ 4+	
Control	20	20 0 0 0 0	
10 ppm	22	22 0 0 0 0	
20 ppm	23	23 0 0 0 0	
40 ppm	20	20 0 0 0 0	

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

Test of CHI SQUARE

TABLE G1

HEMATOLOGY : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

PAGE : 1

Group Name	No. of Animals	RED BLOOD CELL 10 ⁹ /μℓ		HEMOGLOBIN g/dℓ		HEMATOCRIT %		MCV fℓ		MCH p g		MCHC g/dℓ		PLATELET 10 ⁹ /μℓ	
Control	25	10.73±	0.47	15.4±	0.6	46.6±	1.6	43.4±	1.0	14.4±	0.5	33.1±	0.4	1512±	234
10 ppm	25	10.75±	0.34	15.5±	0.4	46.7±	1.2	43.4±	0.9	14.4±	0.3	33.2±	0.4	1507±	207
20 ppm	25	10.92±	0.39	15.6±	0.4	47.1±	1.4	43.2±	1.0	14.3±	0.4	33.1±	0.5	1453±	236
40 ppm	24	10.76±	0.40	15.6±	0.4	46.9±	1.4	43.6±	0.9	14.5±	0.4	33.3±	0.4	1448±	213

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

Test of Dunnett

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

PAGE : 2

Group Name	NO. of Animals	RETICULOCYTE %	
Control	25	4.0±	0.5
10 ppm	25	3.8±	0.4
20 ppm	25	4.0±	0.5
40 ppm	24	4.1±	0.5

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

Test of Dunnett

(HCL070)

BAIS 6

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (27W)

Group Name	NO. of Animals	WBC		Differential		WBC (%)		MONO		EOSINO		BASO	
		$10^3/\mu\ell$		NEUTRO		LYMPHO							
Control	25	3.23±	1.92	12.7±	4.3	83.4±	4.6	3.0±	1.2	0.8±	0.5	0.0±	0.0
10 ppm	25	3.43±	2.10	12.2±	4.7	84.0±	4.9	3.1±	1.3	0.7±	0.6	0.0±	0.0
20 ppm	25	3.59±	2.07	13.9±	8.5	82.2±	8.7	3.0±	1.2	0.8±	0.7	0.0±	0.0
40 ppm	24	3.24±	2.07	16.7±	9.3	79.6±	9.1	2.7±	0.8	1.0±	1.0	0.0±	0.0

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

Test of Dunnett

TABLE G2

HEMATOLOGY : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : FEMALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

PAGE : 4

Group Name	No. of Animals	RED BLOOD CELL 10 ⁶ /μl		HEMOGLOBIN g/dl		HEMATOCRIT %		MCV fl		MCH pg		MCHC g/dl		PLATELET 10 ³ /μl	
Control	24	10.72±	0.54	15.8±	0.5	47.4±	2.0	44.2±	0.6	14.8±	0.4	33.4±	0.5	1287±	220
10 ppm	24	10.70±	0.37	15.9±	0.4	47.3±	1.4	44.3±	0.9	14.9±	0.4	33.6±	0.5	1290±	191
20 ppm	25	10.73±	0.50	15.9±	0.7	47.5±	1.9	44.3±	1.0	14.8±	0.5	33.4±	0.6	1260±	201
40 ppm	25	10.65±	0.52	15.8±	0.6	47.6±	1.8	44.8±	1.1	14.9±	0.4	33.2±	0.8	1191±	260

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

Test of Dunnett

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
MEASURE. TIME : 1
SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
ALL ANIMALS (27W)

PAGE : 5

Group Name	NO. of Animals	RETICULOCYTE %	
Control	24	4.0±	0.8
10 ppm	24	3.9±	0.5
20 ppm	25	4.2±	1.0
40 ppm	25	4.4±	1.8

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : FEMALE REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (27W)

Group Name	NO. of Animals	WBC		Differential		WBC (%)		MONO		EOSINO		BASO	
		$10^3/\mu\ell$		NEUTRO		LYMPHO							
Control	24	2.76±	1.87	16.2±	7.4	79.8±	6.8	3.3±	1.3	0.6±	0.6	0.0±	0.0
10 ppm	24	3.03±	1.97	18.6±	6.4	77.7±	6.4	3.3±	1.3	0.5±	0.5	0.0±	0.0
20 ppm	25	3.54±	2.22	16.8±	7.9	79.1±	7.7	3.4±	1.5	0.7±	0.7	0.0±	0.1
40 ppm	25	2.72±	2.19	16.5±	8.4	79.6±	7.8	3.1±	1.7	0.6±	0.5	0.2±	0.9

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

Test of Dunnett

TABLE H1

BIOCHEMISTRY : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

PAGE : 1

Group Name	NO. of Animals	TOTAL PROTEIN g/dl		ALBUMIN g/dl		A/G RATIO		T-BILIRUBIN mg/dl		GLUCOSE mg/dl		T-CHOLESTEROL mg/dl		TRIGLYCERIDE mg/dl	
Control	25	5.2±	0.2	2.8±	0.2	1.2±	0.1	0.07±	0.03	212±	38	73±	14	43±	12
10 ppm	25	5.3±	0.2	2.9±	0.1	1.2±	0.1	0.06±	0.02	226±	29	76±	13	38±	11
20 ppm	24	5.2±	0.3	2.8±	0.1	1.2±	0.1	0.08±	0.04	217±	38	67±	16	33±	12**
40 ppm	24	5.2±	0.2	2.8±	0.1	1.2±	0.1	0.08±	0.03	220±	31	65±	13	31±	8**

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

PAGE : 2

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	153±	26	76±	60	21±	12	211±	84	222±	28	0.3±	0.3	107±	93
10 ppm	25	158±	24	64±	23	22±	7	209±	55	226±	22	0.4±	0.4	144±	227
20 ppm	24	140±	32	80±	41	24±	9	215±	65	242±	34	0.5±	0.4	92±	62
40 ppm	24	142±	24	82±	49	23±	12	201±	58	235±	26	0.4±	0.3	103±	143

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : MALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

Group Name	NO. of Animals	UREA NITROGEN mg/dl		SODIUM mEq/l		POTASSIUM mEq/l		CHLORIDE mEq/l		CALCIUM mg/dl		INORGANIC PHOSPHORUS mg/dl	
Control	25	30.5±	1.6	151±	2	4.2±	0.3	119±	4	8.9±	0.2	5.8±	0.8
10 ppm	25	31.8±	3.1	152±	2	4.1±	0.2	120±	2	8.9±	0.1	5.6±	1.0
20 ppm	24	30.9±	3.7	151±	2	4.1±	0.2	119±	2	9.0±	0.2	5.6±	1.0
40 ppm	24	30.9±	2.9	151±	2	4.2±	0.2	119±	2	9.0±	0.2	5.6±	1.0

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

Test of Dunnett

TABLE H2

BIOCHEMISTRY : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

Group Name	NO. of Animals	TOTAL PROTEIN g/dl		ALBUMIN g/dl		A/G RATIO		T-BILIRUBIN mg/dl		GLUCOSE mg/dl		T-CHOLESTEROL mg/dl		TRIGLYCERIDE mg/dl	
Control	24	5.2±	0.2	2.9±	0.2	1.3±	0.1	0.07±	0.03	174±	30	63±	14	24±	10
10 ppm	24	5.2±	0.3	2.9±	0.2	1.3±	0.1	0.08±	0.03	177±	23	62±	11	23±	6
20 ppm	25	5.2±	0.3	2.9±	0.2	1.3±	0.1	0.07±	0.03	178±	33	62±	17	27±	9
40 ppm	25	5.3±	0.2	3.0±	0.2	1.3±	0.1	0.07±	0.03	197±	25*	64±	15	26±	8

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

REPORT TYPE : A1

PAGE : 5

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	131±	24	90±	36	30±	19	251±	97	363±	71	0.5±	0.3	230±	387
10 ppm	24	132±	22	89±	32	25±	5	240±	62	376±	64	0.5±	0.4	202±	274
20 ppm	25	132±	33	90±	40	25±	8	220±	54	368±	74	0.4±	0.3	224±	236
40 ppm	25	137±	27	84±	31	27±	12	220±	58	347±	58	0.7±	0.6	143±	148

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

Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 MEASURE. TIME : 1
 SEX : FEMALE REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (27W)

Group Name	NO. of Animals	UREA NITROGEN mg/dl		SODIUM mEq/l		POTASSIUM mEq/l		CHLORIDE mEq/l		CALCIUM mg/dl		INORGANIC PHOSPHORUS mg/dl	
Control	24	24.1±	3.1	151±	2	3.8±	0.3	120±	3	8.9±	0.2	6.1±	1.4
10 ppm	24	25.1±	3.2	151±	2	3.9±	0.4	120±	2	8.9±	0.3	6.0±	1.3
20 ppm	25	24.0±	3.0	151±	2	3.9±	0.4	120±	3	8.9±	0.3	6.3±	1.3
40 ppm	25	24.6±	3.0	151±	2	3.8±	0.3	119±	3	8.9±	0.3	6.0±	1.2

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

Test of Dunnett

TABLE I 1

GROSS FINDINGS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

GROSS FINDINGS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name NO. of Animals	Control			
			25 (%)	10 ppm 25 (%)	20 ppm 25 (%)	40 ppm 25 (%)
thymus	atrophic		0 (0)	0 (0)	1 (4)	1 (4)
spleen	black zone		1 (4)	3 (12)	1 (4)	2 (8)
stomach	glandular stomach:red zone		0 (0)	0 (0)	0 (0)	1 (4)
	glandular stomach:thick		0 (0)	1 (4)	2 (8)	1 (4)
liver	nodule		0 (0)	4 (16)	0 (0)	0 (0)
	deformed		0 (0)	1 (4)	0 (0)	0 (0)
kidney	white zone		0 (0)	0 (0)	0 (0)	1 (4)
	cyst		0 (0)	0 (0)	1 (4)	1 (4)
Harder gl	nodule		1 (4)	0 (0)	0 (0)	0 (0)
peritoneum	red zone		1 (4)	0 (0)	0 (0)	0 (0)

TABLE I 2

GROSS FINDINGS : FEMALE

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
ALL ANIMALS (0- 27W)

Organ	Findings	Group Name NO. of Animals	Control			
			25 (%)	10 ppm 25 (%)	20 ppm 25 (%)	40 ppm 25 (%)
skin/app	erosion		0 (0)	1 (4)	0 (0)	1 (4)
	crust		0 (0)	0 (0)	0 (0)	1 (4)
subcutis	red zone		0 (0)	0 (0)	1 (4)	0 (0)
thymus	atrophic		1 (4)	0 (0)	0 (0)	1 (4)
spleen	enlarged		1 (4)	1 (4)	0 (0)	1 (4)
	black zone		1 (4)	2 (8)	2 (8)	1 (4)
stomach	glandular stomach:red zone		1 (4)	0 (0)	1 (4)	0 (0)
	glandular stomach:thick		2 (8)	0 (0)	1 (4)	4 (16)
liver	enlarged		0 (0)	1 (4)	0 (0)	0 (0)
	nodule		1 (4)	0 (0)	0 (0)	2 (8)
ovary	cyst		0 (0)	0 (0)	0 (0)	1 (4)

TABLE J1

ORGAN WEIGHT, ABSOLUTE : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE
 UNIT: g

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

Group Name	NO. of Animals	Body Weight	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	25	33.4 ± 3.3	0.041 ± 0.008	0.012 ± 0.003	0.219 ± 0.017	0.174 ± 0.017	0.169 ± 0.018
10 ppm	25	32.8 ± 3.0	0.041 ± 0.007	0.011 ± 0.002	0.217 ± 0.015	0.172 ± 0.016	0.169 ± 0.016
20 ppm	25	32.0 ± 3.6	0.037 ± 0.010	0.012 ± 0.004	0.213 ± 0.015	0.172 ± 0.021	0.186 ± 0.019**
40 ppm	24	30.3 ± 3.3**	0.036 ± 0.008	0.011 ± 0.002	0.212 ± 0.015	0.169 ± 0.019	0.197 ± 0.023**

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

TABLE J2

ORGAN WEIGHT, ABSOLUTE : FEMALE

TABLE K1

ORGAN WEIGHT, RELATIVE : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE
 UNIT: %

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

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	TESTES	HEART	LUNGS
Control	25	33.4 ± 3.3	0.122 ± 0.018	0.036 ± 0.011	0.663 ± 0.088	0.524 ± 0.058	0.509 ± 0.069
10 ppm	25	32.8 ± 3.0	0.126 ± 0.017	0.032 ± 0.006	0.666 ± 0.075	0.528 ± 0.069	0.516 ± 0.061
20 ppm	25	32.0 ± 3.6	0.116 ± 0.024	0.037 ± 0.013	0.677 ± 0.117	0.543 ± 0.085	0.589 ± 0.089**
40 ppm	24	30.3 ± 3.3**	0.118 ± 0.018	0.037 ± 0.007	0.708 ± 0.092	0.561 ± 0.061	0.654 ± 0.064**

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE
 UNIT : %

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

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	25	1.451 ± 0.224	0.241 ± 0.046	3.905 ± 0.443	1.440 ± 0.161
10 ppm	25	1.426 ± 0.212	0.238 ± 0.036	4.031 ± 0.423	1.446 ± 0.161
20 ppm	25	1.545 ± 0.257	0.248 ± 0.043	3.812 ± 0.425	1.502 ± 0.226
40 ppm	24	1.556 ± 0.173	0.255 ± 0.041	3.799 ± 0.470	1.574 ± 0.152*

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

TABLE K2

ORGAN WEIGHT, RELATIVE : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE
 UNIT : %

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

Group Name	NO. of Animals	Body Weight (g)	THYMUS	ADRENALS	OVARIES	HEART	LUNGS
Control	24	24.4 ± 2.1	0.177 ± 0.022	0.065 ± 0.013	0.132 ± 0.035	0.611 ± 0.081	0.667 ± 0.072
10 ppm	24	24.0 ± 1.9	0.173 ± 0.022	0.061 ± 0.007	0.122 ± 0.024	0.608 ± 0.064	0.674 ± 0.048
20 ppm	25	24.4 ± 1.9	0.169 ± 0.035	0.062 ± 0.013	0.127 ± 0.028	0.607 ± 0.075	0.742 ± 0.063**
40 ppm	25	24.0 ± 2.0	0.170 ± 0.028	0.062 ± 0.010	0.128 ± 0.031	0.595 ± 0.078	0.813 ± 0.099**

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE
 UNIT : %

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

Group Name	NO. of Animals	KIDNEYS	SPLEEN	LIVER	BRAIN
Control	24	1.591 ± 0.177	0.388 ± 0.061	4.232 ± 0.511	2.005 ± 0.146
10 ppm	24	1.626 ± 0.149	0.387 ± 0.070	4.226 ± 0.549	2.034 ± 0.150
20 ppm	25	1.627 ± 0.175	0.404 ± 0.053	4.185 ± 0.518	1.982 ± 0.146
40 ppm	25	1.569 ± 0.183	0.432 ± 0.265	4.220 ± 0.592	1.999 ± 0.148

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

TABLE L1

HISTOPATHOLOGICAL FINDINGS :

NEOPLASTIC LESIONS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of animals on Study	Control 25	10 ppm 25	20 ppm 25	40 ppm 25
{Respiratory system}						
lung	bronchiolar-alveolar adenoma		<25> 0 (0%)	<25> 0 (0%)	<25> 0 (0%)	<25> 2 (8%)
{Digestive system}						
stomach	squamous cell papilloma		<25> 0 (0%)	<25> 0 (0%)	<25> 0 (0%)	<25> 1 (4%)

< 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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS : NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of animals on Study	Control 25	10 ppm 25	20 ppm 25	40 ppm 25
{Integumentary system/appandage}						
subcutis	hemangioma		<25> 0 (0%)	<25> 0 (0%)	<25> 1 (4%)	<25> 0 (0%)
{Respiratory system}						
lung	bronchiolar-alveolar adenoma		<25> 1 (4%)	<25> 0 (0%)	<25> 0 (0%)	<25> 1 (4%)
{Hematopoietic system}						
spleen	hemangioma		<25> 0 (0%)	<25> 0 (0%)	<25> 1 (4%)	<25> 0 (0%)
{Digestive system}						
liver	histiocytic sarcoma		<25> 0 (0%)	<25> 1 (4%)	<25> 0 (0%)	<25> 0 (0%)

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

TABLE M

NEOPLASTIC LESIONS-INCIDENCE
AND STATISTICAL ANALYSIS : MALE

STUDY No. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 SEX : MALE

NEOPLASTIC LESIONS-INCIDENCE AND STATISTICAL ANALYSIS

Group Name	Control	10 ppm	20 ppm	40 ppm
SITE : lung				
TUMOR : bronchiolar-alveolar adenoma				
Tumor rate				
Overall rates(a)	0/25(0.0)	0/25(0.0)	0/25(0.0)	2/25(8.0)
Adjusted rates(b)	0.00	0.00	0.00	8.33
Terminal rates(c)	0/25(0.0)	0/25(0.0)	0/25(0.0)	2/24(8.3)
Statistical analysis				
Peto test				
Standard method(d)	P = -----			
Prevalence method(d)	P = 0.0165* ?			
Combined analysis(d)	P = -----			
Cochran-Armitage test(e)	P = 0.0298*			
Fisher Exact test(e)		P = N. C.	P = N. C.	P = 0.2449

(HPT360A)

BAIS6

- (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 outcomes cannot be estimated or this P-value is beyond the estimated P-value.
 ----- : There is no data which should be statistical analysis.
 Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$
 N. C. : Statistical value cannot be calculated and was not significant.

TABLE N1

**NUMBER OF ANIMALS WITH TUMORS AND
NUMBER OF TUMORS-TIME RELATED : MALE**

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

Time-related Weeks	Items	Group Name	Control	10 ppm	20 ppm	40 ppm
1 - 26	NO. OF EXAMINED ANIMALS		0	0	0	1
	NO. OF ANIMALS WITH TUMORS		0	0	0	0
	NO. OF ANIMALS WITH SINGLE TUMORS		0	0	0	0
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	0	0
	NO. OF BENIGN TUMORS		0	0	0	0
	NO. OF MALIGNANT TUMORS		0	0	0	0
	NO. OF TOTAL TUMORS		0	0	0	0
27 - 27	NO. OF EXAMINED ANIMALS		25	25	25	24
	NO. OF ANIMALS WITH TUMORS		0	0	0	3
	NO. OF ANIMALS WITH SINGLE TUMORS		0	0	0	3
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	0	0
	NO. OF BENIGN TUMORS		0	0	0	3
	NO. OF MALIGNANT TUMORS		0	0	0	0
	NO. OF TOTAL TUMORS		0	0	0	3
1 - 27	NO. OF EXAMINED ANIMALS		25	25	25	25
	NO. OF ANIMALS WITH TUMORS		0	0	0	3
	NO. OF ANIMALS WITH SINGLE TUMORS		0	0	0	3
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	0	0
	NO. OF BENIGN TUMORS		0	0	0	3
	NO. OF MALIGNANT TUMORS		0	0	0	0
	NO. OF TOTAL TUMORS		0	0	0	3

TABLE N2

**NUMBER OF ANIMALS WITH TUMORS AND
NUMBER OF TUMORS-TIME RELATED : FEMALE**

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

NUMBER OF ANIMALS WITH TUMORS AND NUMBER OF TUMORS - TIME RELATED

Time-related Weeks	Items	Group Name	Control	10 ppm	20 ppm	40 ppm
1 - 26	NO. OF EXAMINED ANIMALS		1	1	0	0
	NO. OF ANIMALS WITH TUMORS		0	1	0	0
	NO. OF ANIMALS WITH SINGLE TUMORS		0	1	0	0
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	0	0
	NO. OF BENIGN TUMORS		0	0	0	0
	NO. OF MALIGNANT TUMORS		0	1	0	0
	NO. OF TOTAL TUMORS		0	1	0	0
27 - 27	NO. OF EXAMINED ANIMALS		24	24	25	25
	NO. OF ANIMALS WITH TUMORS		1	0	1	1
	NO. OF ANIMALS WITH SINGLE TUMORS		1	0	0	1
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	1	0
	NO. OF BENIGN TUMORS		1	0	2	1
	NO. OF MALIGNANT TUMORS		0	0	0	0
	NO. OF TOTAL TUMORS		1	0	2	1
1 - 27	NO. OF EXAMINED ANIMALS		25	25	25	25
	NO. OF ANIMALS WITH TUMORS		1	1	1	1
	NO. OF ANIMALS WITH SINGLE TUMORS		1	1	0	1
	NO. OF ANIMALS WITH MULTIPLE TUMORS		0	0	1	0
	NO. OF BENIGN TUMORS		1	0	2	1
	NO. OF MALIGNANT TUMORS		0	1	0	0
	NO. OF TOTAL TUMORS		1	1	2	1

TABLE O

HISTOPATHOLOGICAL FINDINGS :

METASTASIS OF TUMOR : FEMALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

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

Organ	Findings	Group Name No. of Animals on Study	Control 25	10 ppm 25	20 ppm 25	40 ppm 25
[Hematopoietic system]						
spleen	metastasis:liver tumor		<25> 0	<25> 1	<25> 0	<25> 0
[Urinary system]						
kidney	metastasis:liver tumor		<25> 0	<25> 1	<25> 0	<25> 0
[Body cavities]						
mediastinum	metastasis:liver tumor		<25> 0	<25> 1	<25> 0	<25> 0

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

TABLE P1

HISTOPATHOLOGICAL FINDINGS :
NON-NEOPLASTIC LESIONS : MALE

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Control				10 ppm				20 ppm				40 ppm			
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
Grade		1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Respiratory system]																	
nasal cavit																	
	eosinophilic change:olfactory epithelium	<25>				<25>				<25>				<25>			
		0	0	0	0	0	0	0	0	0	0	0	0	17	4	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(68)	(16)	(0)	(0)
	eosinophilic change:respiratory epithelium	5	0	0	0	5	0	0	0	20	0	0	0	2	23	0	0
		(20)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(80)	(0)	(0)	(0)	(8)	(92)	(0)	(0)
	respiratory metaplasia:gland	15	0	0	0	20	0	0	0	20	0	0	0	21	0	0	0
		(60)	(0)	(0)	(0)	(80)	(0)	(0)	(0)	(80)	(0)	(0)	(0)	(84)	(0)	(0)	(0)
	inflammation:transitional epithelium	0	0	0	0	0	0	0	0	1	0	0	0	8	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(32)	(0)	(0)	(0)
	atrophy:olfactory epithelium	0	0	0	0	0	0	0	0	3	0	0	0	17	6	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(68)	(24)	(0)	(0)
	exudate:olfactory region	0	0	0	0	0	0	0	0	3	0	0	0	21	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(84)	(0)	(0)	(0)
nasopharynx																	
	eosinophilic change:respiratory epithelium	<25>				<25>				<25>				<25>			
		3	0	0	0	1	0	0	0	5	0	0	0	11	13	0	0
		(12)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(44)	(52)	(0)	(0)
lung																	
	hemorrhage	<25>				<25>				<25>				<25>			
		0	0	0	0	1	0	0	0	1	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(4)	(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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control 25				10 ppm 25				20 ppm 25				40 ppm 25									
			1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)						
[Respiratory system]																								
lung	edema		<25>				<25>				<25>				<25>									
			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)	1 (4)						
	inflammatory cell infiltration:focal		0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	9 (36)	1 (4)	0 (0)	0 (0)	**	19 (76)	0 (0)	0 (0)	0 (0)	0 (0)	**	
[Hematopoietic system]																								
thymus	ectopic tissue		<25>				<25>				<25>				<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)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	atrophy		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)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	cyst		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)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
spleen	atrophy		<25>				<25>				<25>				<25>									
			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)	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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control				10 ppm				20 ppm				40 ppm			
			25				25				25				25			
			1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Hematopoietic system]																		
spleen	deposit of melanin		<25>				<25>				<25>				<25>			
			1	0	0	0	3	0	0	0	1	0	0	0	2	0	0	0
			(4)	(0)	(0)	(0)	(12)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)
[Digestive system]																		
salivary gl	atrophy		<25>				<25>				<25>				<25>			
			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)
	hypertrophy		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)
stomach	hyperplasia:forestomach		<25>				<25>				<25>				<25>			
			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)
	erosion:glandular stomach		0	0	0	0	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)
liver	angiectasis		<25>				<25>				<25>				<25>			
			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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Control				10 ppm				20 ppm				40 ppm				
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				
Grade		1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	
{Digestive system}																		
liver	fatty change	11 (44)	0 (0)	0 (0)	0 (0)	10 (40)	0 (0)	0 (0)	0 (0)	8 (32)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0) **
	inflammatory cell nest	3 (12)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	3 (12)	0 (0)	0 (0)	0 (0)	4 (16)	0 (0)	0 (0)	0 (0)	0 (0)
	acidophilic cell focus	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)	0 (0)
pancreas	degeneration	1 (4)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
	<25>																	
{Urinary system}																		
kidney	malformation	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)	0 (0)
	hydronephrosis	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)	0 (0)

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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : MALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control				10 ppm				20 ppm				40 ppm			
			25				25				25				25			
			1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Endocrine system]																		
adrenal	spindle-cell hyperplasia		10 (40)	0 (0)	0 (0)	0 (0)	9 (36)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	7 (28)	0 (0)	0 (0)	0 (0)
			<25>				<25>				<25>				<25>			
[Reproductive system]																		
semin ves	hyperplasia		0 (0)	0 (0)	0 (0)	0 (0)	1 (4)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
			<25>				<25>				<25>				<25>			
[Musculoskeletal system]																		
bone	osteofibrosis		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)
			<25>				<25>				<25>				<25>			
[Body cavities]																		
peritoneum	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)	0 (0)	0 (0)	0 (0)
			<25>				<25>				<25>				<25>			

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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control 25				10 ppm 25				20 ppm 25				40 ppm 25			
			1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)
[Integumentary system/appandage]																		
skin/app	ulcer		<25>				<25>				<25>				<25>			
			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)	
	erosion		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)	
[Respiratory system]																		
nasal cavit	eosinophilic change:olfactory epithelium		<25>				<25>				<25>				<25>			
			2 (8)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)	0 (0)	11 (44)	0 (0)	0 (0)	0 (0)	0 (0)	17 (68)	8 (32)
	eosinophilic change:respiratory epithelium		6 (24)	0 (0)	0 (0)	0 (0)	5 (20)	1 (4)	0 (0)	0 (0)	0 (0)	21 (84)	0 (0)	0 (0)	0 (0)	0 (0)	2 (8)	23 (92)
	respiratory metaplasia:gland		25 (100)	0 (0)	0 (0)	0 (0)	22 (88)	0 (0)	0 (0)	0 (0)	0 (0)	24 (96)	0 (0)	0 (0)	0 (0)	0 (0)	25 (100)	0 (0)
	inflammation:transitional 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)	0 (0)	7 (28)	2 (8)
	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)	0 (0)	0 (0)	17 (68)	2 (8)

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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Control				10 ppm				20 ppm				40 ppm			
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
Grade		1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Respiratory system]																	
nasal cavit	exudate:olfactory region	<25>				<25>				<25>				<25>			
		0	0	0	0	0	0	0	0	1	0	0	0	19	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(76)	(0)	(0)	(0)
nasopharynx	eosinophilic change:respiratory epithelium	<25>				<25>				<25>				<25>			
		1	0	0	0	1	2	0	0	1	0	0	0	11	5	0	0
		(4)	(0)	(0)	(0)	(4)	(8)	(0)	(0)	(4)	(0)	(0)	(0)	(44)	(20)	(0)	(0)
lung	inflammation	<25>				<25>				<25>				<25>			
		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)	(0)	(0)
	inflammatory cell infiltration:focal	0	0	0	0	0	0	0	0	5	0	0	0	10	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(20)	(0)	(0)	(0)	(40)	(0)	(0)	(0)
	accumulation:macrophage	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)
[Hematopoietic system]																	
bone marrow	fatty marrow	<25>				<25>				<25>				<25>			
		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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Control				10 ppm				20 ppm				40 ppm			
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
Grade		1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Hematopoietic system]																	
bone marrow	granulopoiesis: increased	<25>				<25>				<25>				<25>			
		0	0	0	0	0	0	0	0	0	1	0	0	1	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(4)	(0)	(0)	(0)
thymus	atrophy	<25>				<25>				<25>				<25>			
		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)
	cyst	0	0	0	0	0	0	0	0	1	0	0	0	1	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
spleen	deposit of melanin	<25>				<25>				<25>				<25>			
		1	0	0	0	2	0	0	0	2	0	0	0	1	0	0	0
		(4)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(4)	(0)	(0)	(0)
	extramedullary hematopoiesis	2	0	1	0	0	0	0	0	2	0	0	0	1	0	1	0
		(8)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(8)	(0)	(0)	(0)	(4)	(0)	(4)	(0)
[Circulatory system]																	
heart	mineralization	<25>				<25>				<25>				<25>			
		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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Control				10 ppm				20 ppm				40 ppm			
		No. of Animals on Study				No. of Animals on Study				No. of Animals on Study				No. of Animals on Study			
Grade		1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
		(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Digestive system]																	
salivary gl	atrophy	<25>				<25>				<25>				<25>			
		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)
	vacuolic change	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)
stomach	hyperplasia:forestomach	<25>				<25>				<25>				<25>			
		0	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
		(0)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
	erosion:glandular stomach	1	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
		(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)	(4)	(0)	(0)	(0)	(0)	(0)	(0)	(0)
liver	necrosis:focal	<25>				<25>				<25>				<25>			
		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)
	inflammatory cell nest	5	0	0	0	10	0	0	0	7	0	0	0	7	0	0	0
		(20)	(0)	(0)	(0)	(40)	(0)	(0)	(0)	(28)	(0)	(0)	(0)	(28)	(0)	(0)	(0)
pancreas	vacuolic change	<25>				<25>				<25>				<25>			
		2	0	0	0	1	0	0	0	1	0	0	0	0	0	0	0
		(8)	(0)	(0)	(0)	(4)	(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 : Number of animals with lesion
 (c) c : b / a * 100
 Significant difference : * : P ≤ 0.05 ** : P ≤ 0.01 Test of Chi Square

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control				10 ppm				20 ppm				40 ppm			
			25				25				25				25			
			1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+	1+	2+	3+	4+
			(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)	(%)
[Digestive system]																		
pancreas	degeneration		5 (20)	0 (0)	0 (0)	0 (0)	7 (28)	0 (0)	0 (0)	0 (0)	4 (16)	0 (0)	0 (0)	0 (0)	7 (28)	0 (0)	0 (0)	0 (0)
			<25>				<25>				<25>				<25>			
[Urinary system]																		
kidney	hyaline droplet		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)
			<25>				<25>				<25>				<25>			
	hydronephrosis		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)
[Endocrine system]																		
adrenal	spindle-cell hyperplasia		25 (100)	0 (0)	0 (0)	0 (0)	24 (96)	0 (0)	0 (0)	0 (0)	24 (96)	0 (0)	0 (0)	0 (0)	24 (96)	0 (0)	0 (0)	0 (0)
			<25>				<25>				<25>				<25>			
[Reproductive system]																		
ovary	cyst		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)
			<25>				<25>				<25>				<25>			

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

STUDY NO. : 0904
 ANIMAL : B6.129S2-Trp53tm1Tyj/J
 REPORT TYPE : A1
 SEX : FEMALE

HISTOPATHOLOGICAL FINDINGS :NON-NEOPLASTIC LESIONS (SUMMARY)
 ALL ANIMALS (0- 27W)

Organ	Findings	Group Name No. of Animals on Study Grade	Control 25				10 ppm 25				20 ppm 25				40 ppm 25				
			1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	1+ (%)	2+ (%)	3+ (%)	4+ (%)	
[Reproductive system]																			
uterus	atrophy		1 (4)	0 (0)	0 (0)	0 (0)	5 (20)	0 (0)	0 (0)	0 (0)	0 (0)	4 (16)	0 (0)	0 (0)	0 (0)	2 (8)	0 (0)	0 (0)	0 (0)
[Musculoskeletal system]																			
bone	osteofibrosis		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)	0 (0)

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

TABLE Q1

CAUSE OF DEATH : MALE

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
SEX : MALE

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

Group Name	Control	10 ppm	20 ppm	40 ppm
Number of Dead and Moribund Animal	0	0	0	1
no microscop confirm	0	0	0	1

(B10120)

BAIS6

TABLE Q2

CAUSE OF DEATH : FEMALE

STUDY NO. : 0904
ANIMAL : B6.129S2-Trp53tm1Tyj/J
SEX : FEMALE

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

PAGE : 2

Group Name	Control	10 ppm	20 ppm	40 ppm
Number of Dead and Moribund Animal	1	1	0	0
no microscop confirm tumor d:liver	1 0	0 1	0 0	0 0

(BI0120)

BAIS6