アクリル酸=2 - ヒドロキシエチルのラットを用いた経口投与によるがん原性試験(混水試験)報告書

試験番号:0347

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TABLE 1 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

·											
Group	Co	ontrol		320 ppm			800 ppm		:	2000 ppm	
<del>-</del>		:50>		<50>			<49>			<50>	
Week	Survival		Survival	BW	0.4	Survival	BW		Survival	BW	0.4
on Study	No.	g 	No.	g 	%	No.	g 	%	No.	g	% 
0	50	123 ( 50 )	50	123 ( 50 )	100	49	123 ( 49 )	100	50	123 ( 50 )	100
1	50	156 ( 50 )	50	153 ( 50 )	98	49	151 (49)	97 **	50	141 ( 50 )	90 **
2	50	185 ( 50 )	50	182 (50)	98	49	179 ( 49 )	97 **	50	167 ( 50 )	90 **
3	50	207 ( 50 )	50	204 (50)	99	49	201 (49)	97 **	50	189 ( 50 )	91 **
4	50	224 ( 50 )	50	222 (50)	99	49	218 ( 49 )	97 **	50	205 ( 50 )	92 **
5	50	239 (50)	50	236 (50)	99	49	232 ( 49 )	97 **	50	220 ( 50 )	92 **
6	50	251 (50)	50	248 (50)	99	49	242 ( 49 )	96 **	50	230 ( 50 )	92 **
7	50	262 ( 50 )	50	258 (50)	98	49	253 ( 49 )	97 **	50	239 ( 50 )	91 **
8	50	270 (50)	50	267 (50)	99	49	261 (49)	97 **	50	246 ( 50 )	91 **
9	50	279 ( 50 )	50	274 (50)	98	49	268 ( 49 )	96 **	50	253 ( 50 )	91 **
10	50	285 (50)	50	280 (50)	98	49	275 ( 49 )	96 **	50	259 ( 50 )	91 **
11	50	291 (50)	50	285 ( 50 )	98	49	280 (49)	96 **	50	264 ( 50 )	91 **
12	50	296 (50)	50	291 (50)	98	49	285 (49)	96 **	50	269 ( 50 )	91 **
13	50	302 ( 50 )	50	295 (50)	98 *	49	290 ( 49 )	96 **	50	273 ( 50 )	90 **
14	50	307 (50)	50	299 ( 50 )	97 *	49	294 ( 49 )	96 **	50	277 ( 50 )	90 **
18	50	322 ( 50 )	50	314 ( 50 )	98 **	49	308 ( 49 )	96 **	50	292 ( 50 )	91 **
22	50	335 ( 50 )	50	326 (50)	97 **	49	321 (49)	96 **	50	305 ( 50 )	91 **
26	50	348 ( 50 )	50	336 (50)	97 **	49	332 ( 49 )	95 **	50	317 ( 50 )	91 **
30	50	355 ( 50 )	50 -	343 ( 50 )	97 **	49	340 ( 49 )	96 **	50	324 ( 50 )	91 **
34	50	364 ( 50 )	50	350 ( 50 )	96 **	49	345 ( 49 )	95 **	50	331 ( 50 )	91 **
38	50	374 ( 50 )	50	357 (50)	95 **	49	353 (49)	94 **	50	338 ( 50 )	90 **
42	50	382 ( 50 )	50	363 ( 50 )	95 **	49	359 (49)	94 **	49	343 ( 49 )	90 **
46	50	388 (50)	50	369 ( 50 )	95 **	49	365 (49)	94 **	49	348 ( 49 )	90 **
50	50	392 ( 50 )	50	373 (50)	95 **	49	369 (49)	94 **	49	351 (49)	90 **
54	50	398 ( 50 )	50	379 ( 50 )	95 **	49	374 ( 49 )	94 **	49	355 ( 49 )	89 **
58	50	403 ( 50 )	50	383 ( 50 )	95 **	49	377 (49)	94 **	49	357 ( 49 )	89 **
62	50	406 (50)	50	387 (50)	95 **	48	381 (48)	94 **	49	356 ( 49 )	88 **
66	50	409 ( 50 )	50	392 ( 50 )	96 **	48	385 (48)	94 **	47	357 ( 47 )	87 **
70	49	406 ( 49 )	49	393 (49)	97 *	47	385 (47)	95 **	46	355 (46)	87 **
74	49	416 ( 49 )	48	400 (48)	96 **	47	387 (47)	93 **	46	355 ( 46 )	85 **
78	49	419 (49)	47	402 (47)	96 *	46	388 ( 46 )	93 **	45	356 ( 45 )	85 **
82	48	428 (48)	47	407 (47)	95 **	46	388 (46)	91 **	45	357 ( 45 )	83 **
86	48	425 (48)	47	405 (47)	95 **	44	385 (44)	91 **	44	348 ( 44 )	82 **
90	47	423 (47)	47	404 (47)	96 **	42	382 ( 42 )	90 **	43	341 (43)	81 **
94	45	419 ( 45 )	47	399 (47)	95 **	41	376 (41)	90 **	41	332 (41)	79 **
98	41	419 (41)	47	393 (47)	94 **	41	367 (41)	88 **	39	325 ( 39 )	78 **
102	40	412 ( 40 )	44	388 (44)	94 **	39	357 ( 39 )	87 **	36	318 ( 36 )	77 **
104	40	408 ( 40 )	44	383 (44)	94 **	37	353 ( 37 )	87 **	35	316 ( 35 )	77 **

<sup>&</sup>lt; > : No.of effective animals, ( ): No.of measured animals %:% of control group
Significant Difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$ , Test of Dunnett

TABLE 2 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Co	Control		320 ppm	320 ppm			800 ppm			
•	<	:50>		<50>			<50>			<50>	
Week	Survival	BW	Survival	BW		Survival	BW		Survival	BW	
on Study	No.	g	No.	g	<b>%</b>	No.	g	%	No.	g	%
0	50	96 (50)	50	96 ( 50 )	100	50	96 (50)	100	50	96 (50)	100
1	50	115 (50)	50	113 (50)	98	50	112 (50)	97 **	50	108 (50)	94 **
2	50	126 (50)	50	124 (50)	98	50	123 (50)	98 *	50	119 (50)	94 *
3	50	134 (50)	50	132 (50)	99	50	132 (50)	99	50	127 (50)	95 *
4	50	140 (50)	50	138 (50)	99	50	138 (50)	99	50	132 (50)	94 *
5	50	147 (50)	50	146 (50)	99	50	144 (50)	98	50	137 (50)	93 *
6	50	151 (50)	50	149 (50)	99	50	149 (50)	99	50	141 (50)	93 *
7	50	154 (50)	50	153 (50)	99	50	152 (50)	99	50	143 (50)	93 *
8	50	157 (50)	50	156 (50)	99	50	155 (50)	99	50	146 (50)	93 *
9	50	160 (50)	50	160 (50)	100	50	158 (50)	99	50	$149 (50^{\circ})$	93 *
10	50	163 (50)	50	163 (50)	100	50	161 (50)	99	50	150 (50)	92 *
11	50	166 (50)	50	165 (50)	99	50	164 (50)	99	50	154 (50)	93 *
12	50	168 (50)	50	168 (50)	100	50	167 (50)	99	50	155 (50)	92 *
13	50	170 (50)	50	171 (50)	101	50	169 (50)	99	50	158 (50)	93 *
14	50	171 (50)	50	173 (50)	101	50	171 (50)	100	50	159 (50)	93 *
18	50	179 (50)	50	180 (50)	101	50	177 (50)	99	50	165 (50)	92 *
22	50	185 (50)	50	187 (50)	101	50	184 (50)	99	50	171 (50)	92 *
26	50	189 (50)	50	191 (50)	101	50	188 (50)	99	50	175 (50)	93 *
30	50	194 (50)	50	197 (50)	102	50	193 (50)	99	50	179 (50)	92 *
34	50	198 (50)	50	200 (50)	101	50	195 (50)	98	50	181 (50)	91 *
38	50	200 (50)	50	204 (50)	102	50	197 (50)	99	50	183 (50)	92 *
42	50	205 (50)	50	208 (50)	101	50	201 (50)	98	50	186 (50)	91 *
46	50	210 (50)	50	212 (50)	101	50	205 (50)	98	50	188 (50)	90 *
50	49	213 (49)	50	215 (50)	101	49	208 (49)	98	50	191 (50)	90 *
54	49	217 (49)	50	219 (50)	101	49	212 (49)	98	50	193 (50)	89 *
58	49	220 (49)	50	223 (50)	101	49	215 (49)	98	50	194 (50)	88 *
62	49	224 (49)	49	228 (49)	102	48	219 (48)	98	50	195 (50)	87 *
66	49	230 (49)	49	233 (49)	101	48	223 (48)	97	50	198 (50)	86 *
70	48	234 (48)	49	234 (49)	100	48	225 (48)	96	50	199 (50)	85 *
74	47	241 (47)	48	241 (48)	100	48	229 (48)	95	50	199 (50)	83 *
78	47	247 (47)	48	247 (48)	100	48	233 (48)	94 *	50	202 (50)	82 *
82	46	253 (46)	48	254 (48)	100	47	240 (47)	95 *	49	206 (49)	81 *
86	44	257 (44)	48	255 (48)	99	47	242 (47)	94 *	47	210 (47)	82 *
90	42	262 (42)	47	257 (47)	98	47	242 (47)	92 **	46	211 (46)	81 *
94	42	262 (42)	46	257 (46)	98	46	243 (46)	93 **	46	211 (46)	81 *
98	40	265 (40)	45	257 (45)	97	44	242 (44)	91 **	46	212 (46)	80 *
102	37	268 (37)	40	264 (40)	99	42	242 (42)	90 **	46	212 (46)	79 *
104	37	266 (37)	40	261 (40)	98	42	240 (42)	90 **	42	208 (42)	78 *

<sup>&</sup>lt; >: No.of effective animals, ( ): No.of measured animals %:% of control group
Significant Difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$ , Test of Dunnett

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TABLE 3 WATER CONSUMPTION CHANGES OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Co	ontrol		320 ppm				800 ppm				2000 ppm		
Week	<u> </u>	<50>		<50>				<49>			<50>			
on Study	Survival		Survival	WC			Survival	WC			Survival	WC		
	No.	g 	No.	g 	%		No.	g	%		No.	g 	%	
1	50	18.0 ( 50 )	50	15.6 ( 50 )	87	**	49	14.2 ( 49 )	79	**	50	11.7 ( 50 )	65	**
2	50	18.8 ( 50 )	50	16.8 ( 50 )	89	**	49	15.1 (48)	80	**	50	12.1 ( 50 )	64	**
3	50	20.3 ( 50 )	50	16.9 ( 50 )	83	**	49	15.4 ( 49 )	76	**	50	13.3 ( 50 )	66	**
4	50	19.9 ( 50 )	50	17.3 ( 50 )	87	**	49	15.5 ( 48 )	78	**	50	13.4 ( 50 )	67	**
5	50	18.9 ( 50 )	50	16.4 ( 50 )	87	**	49	14.9 ( 49 )	79	**	50	12.8 ( 50 )	68	**
6	50	18.6 ( 50 )	50	16.1 ( 50 )	87	**	49	14.6 ( 49 )	78	**	50	12.2 ( 50 )	66	**
7	50	18.5 ( 50 )	50	16.9 ( 50 )	91	**	49	14.5 ( 49 )	78	**	50	12.2 ( 50 )	66	**
8	50	18.2 ( 50 )	50	16.4 ( 50 )	90	**	49	13.9 ( 49 )	76	**	50	11.6 ( 50 )	64	**
9	50	17.9 ( 50 )	50	15.8 ( 50 )	88	**	49	13.9 ( 49 )	78	**	50	11.9 ( 50 )	66	**
10	50	17.8 ( 50 )	50	16.0 ( 50 )	90	**	49	14.6 ( 49 )	82	**	50	12.1 ( 50 )	68	**
11	50	17.1 ( 50 )	50	15.2 ( 50 )	89	**	49	13.5 ( 49 )	79	**	50	11.6 ( 50 )	68	**
12	50	16.9 ( 50 )	50	15.0 ( 50 )	89	**	49	13.6 ( 49 )	80	**	50	11.5 ( 50 )	68	**
13	50	17.7 ( 50 )	50	16.0 ( 50 )	90	**	49	14.2 ( 49 )	80	**	50	11.8 ( 50 )	67	**
14	50	17.5 ( 50 )	50	15.3 ( 50 )	87	**	49	13.9 ( 49 )	79	**	50	11.7 ( 50 )	67	**
18	50	16.5 ( 50 )	50	14.8 ( 50 )	90	**	49	13.8 ( 49 )	84	**	50	11.7 ( 50 )	71	**
22	50	16.2 ( 50 )	50	14.9 ( 50 )	92	**	49	13.1 ( 49 )	81	**	50	11.4 ( 50 )	70	**
26	50	16.7 ( 50 )	50	14.6 ( 50 )	87	**	49	13.3 ( 49 )	80	**	50	11.6 ( 49 )	69	**
30	50	16.5 ( 50 )	50	15.0 ( 50 )	91	**	49	13.5 ( 49 )	82	**	50	11.6 ( 50 )	70	**
34	50	16.5 ( 50 )	50	14.4 ( 50 )	87	**	49	13.3 ( 49 )	81	**	50	11.7 ( 50 )	71	**
38	50	16.4 ( 50 )	50	14.4 ( 50 )	88	**	49	13.6 ( 49 )	83	**	50	12.3 ( 50 )	75	**
42	50	16.3 ( 50 )	50	14.4 ( 50 )	88	**	49	13.7 ( 49 )	84	**	49	12.0 ( 49 )	74	**
46	50	16.4 ( 50 )	50	14.7 ( 50 )	90	**	49	13.7 ( 49 )	84	**	49	12.6 ( 49 )	77	**
50	50	16.5 ( 50 )	50	15.4 ( 50 )	93	**	49	14.1 ( 49 )	85	**	49	12.6 ( 49 )	76	**
<b>5</b> 4	50	17.5 ( 50 )	50	15.8 ( 50 )	90	**	49	14.9 ( 49 )	85	**	49	13.1 ( 49 )	75	**
58	50	16.8 ( 50 )	50	15.3 ( 50 )	91	**	49	14.5 ( 49 )	86	**	49	12.8 ( 49 )	76	**
62	50	17.5 ( 50 )	50	15.9 ( 50 )	91	**	48	15.4 ( 48 )	88	**	49	13.5 ( 49 )	77	**
66	50	17.0 ( 50 )	50	15.7 ( 50 )	92	**	48	15.0 (48)	88	**	47	13.3 ( 47 )	78	**
70	49	18.6 ( 49 )	49	16.6 ( 49 )	89	**	47	15.7 (47)	84	**	46	13.7 ( 46 )	74	**
74	49	18.3 ( 49 )	48	16.4 (48)	90	**	47	15.5 ( 47 )	85	**	46	13.7 ( 46 )	75	**
78	49	18.3 ( 49 )	47	16.4 ( 47 )	90	**	46	15.3 ( 45 )	84	**	45	13.7 ( 45 )	75	**
82	48	17.8 (48)	47	16.4 ( 47 )	92	**	46	15.3 ( 46 )	86	**	45	13.6 ( 45 )	76	**
86	48	18.0 (48)	47	16.2 ( 47 )	90	**	44	15.1 ( 43 )	84	**	44	13.7 ( 43 )	76	**
90	47	18.4 ( 47 )	47	16.7 ( 47 )	91	**	42	16.4 ( 42 )	89	**	43	14.1 ( 43 )	77	**
94	45	18.9 ( 45 )	47	17.0 ( 47 )	90	**	41	16.5 ( 41 )	87	**	41	14.1 (41)		
98	41	19.7 (41)	47	17.2 ( 47 )	87	**	41	17.4 ( 41 )	88	**	39	15.3 ( 39 )	78	**
102	40	20.4 ( 40 )	44	17.7 ( 44 )	87	**	39	17.6 ( 38 )	86	**	36	15.8 ( 36 )	77	**
104	40	20.4 ( 40 )	44	17.5 ( 44 )	86	**	37	17.7 ( 37 )	87	**	35	15.8 ( 35 )	77	**

< > : No.of effective animals, ( ) : No.of measured animals % : % of control group
Significant Difference, \*\*:  $p \le 0.01$ , Test of Dunnett

TABLE 4 WATER CONSUMPTION CHANGES OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Co	ntrol		320 ppm				800 ppm				2000 ppm		
Week		<50>		<50>				<50>				<50>		
on Study	Survival No.	WC g	Survival No.	WC g	%		Survival No.	WC g	%		Survival No.	WC g	%	
7	=		50	<del></del>	90	**			70	**	E0.	10.0 ( 50 )	C E	**
$rac{1}{2}$	50 50	15.5 ( 50 )	50 50	13.3 ( 50 ) 14.0 ( 50 )		**	50 50	12.2 ( 50 ) 11.8 ( 50 )		**	50 50	9.6 (50)		
3	50 50	16.5 ( 49 ) 16.9 ( 47 )	50 50	14.8 (50)		**	50 50	12.1 (49)		**	50	9.9 ( 50 )		
4	50	17.7 (49)	50	15.8 (48)			50	12.2 ( 50 )		**	50	9.8 (50)		
5	50	16.3 (48)	50	14.9 (50)			50	11.9 (50)		**	50	10.0 (50)		
6	50	16.8 (48)	50	15.0 (49)			50	12.3 (49)		**	50	10.3 (50)		
7	50	15.9 (46)	50	14.3 (50)			50	11.8 ( 50 )		**	50	8.8 ( 50 )		
8	50	15.9 (46)	50	14.3 (48)			50	11.5 ( 50 )		**	50	8.6 ( 50 )		
9	50	16.5 (48)	50	14.8 (46)			50	11.4 (47)		**	50	9.2 ( 50 )		
10	50	16.7 (48)	50	14.7 (47)		*	50	11.4 (48)		**	50	8.5 ( 50 )		
11	50	15.2 (48)	50	14.1 (48)			50	11.7 (49)		**	50	8.9 ( 50 )		
12	50	16.5 (48)	50	14.5 (48)		*	50	12.1 (48)	73	**	50	9.2 ( 50 )	56	**
13	50	17.1 ( 49 )	50	15.1 ( 46 )			50	11.6 ( 49 )	68	**	50	8.8 ( 50 )	51	**
14	50	17.4 ( 50 )	50	15.1 ( 46 )	87	*	50	11.4 ( 48 )	66	**	50	8.8 ( 50 )	51	**
18	50	17.5 (46)	50	15.3 ( 47 )	87		50	11.8 ( 48 )	67	**	50	8.8 ( 50 )	50	**
22	50	16.6 (47)	50	14.4 ( 44 )	87	*	50	12.3 ( 49 )	74	**	50	9.0 ( 50 )	54	**
26	50	16.1 (49)	50	14.8 ( 49 )	92		50	12.0 ( 50 )	75	**	50	8.9 ( 50 )	55	**
30	50	17.1 ( 49 )	50	15.3 ( 47 )	89		50	12.9 ( 50 )	75	**	50	9.0 ( 50 )	53	**
34	50	16.6 ( 50 )	50	14.7 ( 49 )	89		50	11.2 ( 49 )	67	**	50	8.8 ( 50 )	53	**
38	50	16.1 (49)	50	15.2 ( 50 )	94		50	11.3 ( 49 )	70	**	50	9.0 (48)	56	**
42	50	16.8 ( 50 )	50	16.0 ( 50 )	95		50	12.4 ( 50 )	74	**	50	9.0 ( 50 )	54	**
46	50	15.5 (49)	50	15.3 ( 50 )	99		50	12.0 ( 50 )	77	**	50	9.3 ( 50 )	60	**
50	49	15.3 ( 49 )	50	13.9 ( 50 )	91		49	11.3 ( 49 )	74	**	50	9.2 ( 50 )	60	**
54	49	16.6 ( 49 )	50	14.8 ( 50 )	89		49	12.1 ( 49 )	73	**	50	9.6 ( 50 )	58	**
58	49	14.4 ( 49 )	50	13.6 ( 50 )	94		49	10.6 (49)	74	**	50	9.4 ( 50 )		
62	49	16.5 ( 49 )	49	14.8 ( 49 )	90		48	11.8 (48)		**	50	10.2 ( 50 )		**
66	49	15.8 (47)	49	15.1 ( 49 )			48	12.3 (48)	78	**	50	10.4 ( 50 )		**
70	48	16.0 (48)	49	14.0 ( 49 )	88	*	48	12.0 (48)		**	50	10.8 ( 50 )		**
74	47	15.7 (47)	48	14.1 ( 48 )	90		48	11.7 ( 48 )	75	**	50	10.8 ( 50 )		
78	47	15.1 (47)	48	13.2 ( 47 )			48	11.8 ( 48 )			50	11.1 ( 50 )		
82	46	15.7 ( 46 )	48	13.0 ( 48 )			47	11.5 ( 47 )			49	11.0 ( 49 )		
86	44	15.4 ( 43 )	48	13.8 ( 48 )			47	11.5 ( 47 )			47	11.3 (47)		
90	42	15.9 ( 40 )	47	14.2 ( 47 )			47	12.6 (47)			46	12.0 (46)		
94	42	16.5 ( 42 )	46	14.7 ( 46 )			46	12.0 (46)			46	12.8 (46)		
98	40	17.4 ( 40 )	45	15.3 (45)			44	12.4 (44)			46	13.5 (46)		
102	37	18.1 ( 36 )	40	16.5 ( 40 )			42	13.0 (42)			46	13.7 (46)		
104	37	17.4 ( 37 )	40	14.5 ( 40 )	83	**	42	12.7 ( 42 )	73	**	42	13.2 ( 42 )	76	**

<sup>&</sup>lt; > : No.of effective animals, ( ) : No.of measured animals % : % of control group
Significant Difference, \*: p  $\leq$  0.05, \*\* : p  $\leq$  0.01, Test of Dunnett

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TABLE 5 FOOD CONSUMPTION CHANGES OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	C	ontrol		320 ppm			800 ppm			2000 ppm	
-		<50>		<50>			<49>			<50>	
Week	Surviva		Survival	FC		Survival	FC		Survival	FC	
on Study	No.	g	No.	g	%	No.	g	%	No.	g	%
1	50	14.3 ( 50 )	50	13.6 ( 50 )	100 *	49	13.7 ( 49 )	96 *	50	11.9 ( 50 )	83 **
2	50	15.4 ( 50 )	50	15.1 ( 50 )	98	49	14.6 ( 49 )	95 **	50	13.4 ( 50 )	87 **
3	50	15.9 ( 49 )	50	15.5 (48)	97 *	49	15.2 ( 49 )	96 **	50	14.2 ( 50 )	89 **
4	50	15.8 ( 50 )	50	15.5 ( 50 )	98	49	15.2 ( 49 )	96 **	50	14.3 ( 50 )	91 **
5	50	15.6 ( 50 )	50	15.2 ( 50 )	97 *	49	14.9 ( 49 )	96 **	50	14.3 ( 50 )	92 **
6	50	15.1 ( 50 )	50	14.7 ( 50 )	97 *	49	14.4 ( 49 )	95 **	50	13.8 ( 50 )	91 **
7	50	15.5 ( 50 )	50	15.0 ( 50 )	97 *	49	14.7 ( 49 )	95 **	50	14.0 ( 50 )	90 **
8	50	14.9 ( 50 )	50	14.4 ( 50 )	97 **	49	14.1 ( 49 )	95 **	50	13.5 ( 50 )	91 **
9	50	14.9 ( 50 )	50	14.4 ( 50 )	97 *	49	14.1 ( 49 )	95 **	50	13.6 ( 50 )	91 **
10	50	14.7 ( 49 )	50	14.2 ( 50 )	97 *	49	13.9 ( 49 )	95 **	50	13.1 ( 50 )	89 **
11	50	14.5 ( 50 )	50	14.1 ( 50 )	97 *	49	13.7 ( 49 )	94 **	50	13.3 ( 50 )	92 **
12	50	14.2 ( 50 )	50	13.8 ( 50 )	97 *	49	13.5 ( 49 )	95 **	50	13.3 ( 50 )	94 **
13	50	14.4 ( 50 )	50	13.8 ( 50 )	96 **	49	13.8 ( 49 )	96 **	50	13.3 ( 50 )	92 **
14	50	14.0 ( 50 )	50	13.4 ( 50 )	96 **	49	13.5 ( 49 )	96 **	50	12.9 ( 49 )	92 **
18	50	14.6 ( 50 )	50	14.2 ( 50 )	97 *	49	13.9 ( 49 )	95 **	50	13.5 ( 50 )	92 **
22	50	14.6 ( 50 )	50	14.0 ( 50 )	96 **	49	14.0 ( 49 )	96 **	50	13.6 ( 50 )	93 **
26	50	15.4 ( 50 )	50	14.7 ( 50 )	95 **	49	14.5 ( 49 )	94 **	50	14.2 ( 50 )	92 **
30	50	14.9 ( 50 )	50	14.4 ( 50 )	97 **	49	14.4 ( 49 )	97 *	50	14.1 ( 50 )	95 **
34	50	15.4 ( 50 )	50	14.7 ( 50 )	95 **	49	14.7 ( 49 )	95 **	50	14.7 ( 50 )	95 **
38	50	15.6 ( 50 )	50	14.6 ( 50 )	94 **	49	14.8 ( 49 )	95 **	50	14.9 ( 50 )	96 **
42	50	15.6 ( 50 )	50	14.7 ( 50 )	94 **	49	15.2 ( 49 )	97	49	14.9 ( 49 )	96 *
46	50	15.4 ( 50 )	50	14.8 ( 50 )	96 **	49	15.0 ( 49 )	97	49	15.1 ( 49 )	98
50	50	15.5 ( 50 )	50	14.9 ( 50 )	96	49	15.1 ( 49 )	97	49	15.2 ( 49 )	98
54	50	15.9 ( 50 )	50	15.1 ( 50 )	95 **	49	15.3 ( 49 )	96	49	15.2 ( 49 )	96 **
58	50	15.8 ( 50 )	50	15.1 ( 50 )	96 **	49	15.5 ( 49 )	98	49	15.0 ( 49 )	95 **
62	50	15.8 ( 50 )	50	15.2 ( 50 )	96 **	48	15.4 ( 48 )	97	49	14.8 ( 49 )	94 **
66	50	15.5 ( 50 )	50	15.2 ( 50 )	98	48	15.4 ( 48 )	99	47	14.9 ( 47 )	96 **
70	49	16.2 ( 49 )	49	15.5 ( 49 )	96 **	47	15.8 ( 47 )	98	46	15.3 ( 46 )	94 **
74	49	16.2 ( 49 )	48	15.7 (48)	97	47	15.6 (47)	96 *	46	15.2 ( 46 )	94 **
78	49	15.9 ( 49 )	47	15.2 ( 47 )	96 **	46	15.5 ( 46 )	97	45	15.0 ( 45 )	94 **
82	48	15.9 ( 48 )	47	15.4 (47)	97	46	15.5 ( 46 )	97	45	15.1 ( 45 )	95 **
86	48	15.8 ( 48 )	47	15.4 ( 47 )	97	44	15.6 ( 44 )	99	44	15.0 ( 44 )	95 **
90	47	15.7 ( 47 )	47	15.4 ( 47 )	98	42	15.2 ( 42 )	97	43	14.8 ( 43 )	94 **
94	45	15.9 ( 45 )	47	15.7 (47)	99	41	15.7 (41)	99	41	15.0 (41)	94 **
98	41	16.2 (41)	47	15.4 ( 47 )	95	41	15.1 ( 41 )	93 *	39	14.5 ( 39 )	90 **
102	40	16.4 ( 40 )	44	15.6 ( 44 )	95	39	15.2 ( 39 )	93 *	36	14.9 ( 36 )	91 **
104	40	15.7 ( 40 )	44	15.2 ( 44 )	97	37	15.1 ( 37 )	96	35	14.7 (35)	94

<sup>&</sup>lt; >: No.of effective animals, ( ): No.of measured animals %:% of control group
Significant Difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$ , Test of Dunnett

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TABLE 6 FOOD CONSUMPTION CHANGES OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Co	ontrol		320 ppm			800 ppm			2000 ppm	
-		<50>		<50>			<50>			<50>	
Week	Survival	FC	Survival	FC		Survival	FC		Survival	FC	
on Study	No.	g	No.	g 	%	No.	g	%	No.	g	%
1	50	10.6 ( 50 )	50	10.4 ( 50 )	98	50	10.1 ( 50 )	95 **	50	9.2 ( 50 )	87 **
2	50	10.5 ( 50 )	50	10.6 ( 50 )	101	50	10.1 ( 50 )	96 **	50	9.7 (50)	92 **
3	50	10.8 ( 50 )	50	10.6 ( 50 )	98	50	10.5 ( 50 )	97	50	10.0 ( 50 )	93 **
4	50	10.9 ( 50 )	50	10.7 ( 50 )	98	50	10.4 ( 50 )	95 **	50	10.1 (50)	93 **
5	50	10.7 ( 50 )	50	10.7 ( 50 )	100	50	10.3 ( 50 )	96 *	50	9.9 ( 50 )	93 **
6	50	10.3 ( 50 )	50	10.3 ( 50 )	100	50	10.0 ( 50 )	97	50	9.4 ( 50 )	91 **
7	50	10.2 ( 50 )	50	10.2 ( 50 )	100	50	10.0 ( 50 )	98	50	9.3 (50)	91 **
8	50	9.9 ( 50 )	50	9.8 ( 50 )	99	50	9.6 ( 50 )	97	50	9.1 (50)	92 **
9	50	10.1 ( 50 )	50	10.0 ( 50 )	99	50	9.5 ( 50 )	94 **	50	9.0 ( 50 )	89 **
10	50	9.8 ( 50 )	50	9.8 ( 50 )	100	50	9.5 ( 50 )	97	50	8.7 ( 50 )	89 **
11	50	9.8 ( 50 )	50	9.8 ( 50 )	100	50	9.5 ( 50 )	97	50	8.7 ( 50 )	89 **
12	50	9.8 ( 50 )	50	9.8 ( 50 )		50	9.6 ( 50 )	98	50	8.8 ( 50 )	90 **
13	50	9.7 ( 50 )	50	10.0 ( 50 )		50	9.6 ( 50 )	99	50	8.8 ( 50 )	91 **
14	50	9.7 ( 50 )	50	9.9 ( 50 )		50	9.6 ( 50 )	99	50	9.0 (50)	93 **
18	50	10.3 (49)	50	10.2 ( 50 )	99	50	9.9 ( 50 )	96 *	50	9.1 ( 50 )	88 **
22	50	10.1 ( 50 )	50	10.3 ( 50 )		50	10.0 ( 50 )	99	50	9.3 ( 50 )	92 **
26	50	10.2 ( 50 )	50	10.3 ( 50 )		50	10.1 ( 50 )	99	50	9.5 ( 50 )	93 **
30	50	10.6 ( 50 )	50	10.8 ( 50 )		50	10.4 ( 50 )	98	50	9.6 ( 50 )	91 **
34	50	10.5 (49)	50	10.7 (50)		50	10.2 ( 50 )	97	50	9.6 ( 50 )	91 **
38	50	10.6 ( 50 )	50	10.9 ( 50 )		50	10.2 ( 50 )	96	50	9.8 (48)	92 **
42	50	10.7 (49)	50	10.9 ( 50 )		50	10.5 ( 50 )	98	50	10.0 ( 50 )	93 **
46	50	10.8 ( 50 )	50	11.0 (50)		50	10.8 ( 50 )		50	10.0 ( 50 )	93 **
50	49	10.7 (49)	50	10.9 (50)		49	10.4 ( 49 )	97	50	9.9 ( 50 )	93 **
54	49	11.2 (49)	50	11.4 ( 50 )		49	11.0 (49)	98	50	10.3 ( 50 )	92 **
58	49	10.9 (49)	50	11.0 (50)		49	10.6 (49)	97	50	9.9 ( 50 )	91 **
62	49	11.3 (49)	49	11.4 (49)		48	11.0 (48)	97	50	10.3 ( 50 )	91 **
66	49	11.5 (49)	49	11.8 (49)		48	11.0 (48)	96	50	10.4 ( 50 )	90 **
70	48	11.4 ( 48 )	49	11.4 (49)		48	11.1 (48)	97	50	10.4 ( 50 )	91 **
74	47	11.7 (47)	48	11.9 (48)		48	11.1 (48)	95 *	50	10.3 ( 50 )	
78	47	11.9 (47)	48	11.7 (48)		48	11.1 (48)	93 **	50	10.1 ( 50 )	85 **
82	46	11.6 (46)	48	11.9 (48)		47	11.4 (47)	98	49	10.6 (49)	91 **
86	44	11.7 (44)	48	11.8 (48)		47	11.2 (47)	96 **	47	10.8 (47)	92 **
90	42	11.7 (44)	47	11.7 (47)		47	11.1 (47)	94 **	46	10.4 (46)	88 **
94	42	11.8 (42)	46	11.9 (46)		46	11.4 (46)	97	46	10.7 (46)	91 **
98	40	12.0 (40)	45	11.6 (45)	97	44	11.4 (40)	92 **	46	10.9 (46)	91 **
102	40 37	12.0 (40)	40	12.0 (40)	99	42	11.3 (42)	93 *	46	10.6 (46)	88 **
102	37	11.6 (37)	40	11.5 (40)	99	42	11.0 (42)		42	10.6 (42)	

<sup>&</sup>lt; > : No.of effective animals, ( ) : No.of measured animals % : % of control group
Significant Difference, \*:  $p \le 0.05$ , \*\* :  $p \le 0.01$ , Test of Dunnett

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TABLE 7 INCIDENCE AND TIME OF MASS OCCURRENCE IN CLINICAL OBSERVATION OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Time of mass occurrence (wee	k) 0-13	14-26	27-39	40-52	53-65	66-78	79-91	92-104	0-104
External mass									
Control	0/50	0/50	0/50	0/50	2/50	1/50	6/49	16/47	17/50 (6/10)
320 ppm	0/50	0/50	0/50	0/50	1/50	1/50	6/47	14/47	14/50 (1/6)
800 ppm	0/49	0/49	0/49	1/49	2/49	2/48	6/47	10/42	13/49 (5/12)
2000 ppm	0/50	0/50	0/50	2/50	2/49	1/47	3/45	7/42	9/50 (2/15)
Internal mass									
Control	0/50	0/50	0/50	0/50	0/50	0/50	0/49	2/47	2/50 (1/10)
320 ppm	0/50	0/50	0/50	0/50	0/50	0/50	0/47	1/47	1/50 (0/ 6)
800 ppm	0/49	0/49	0/49	0/49	0/49	0/48	0/47	0/42	0/49 (0/12)
2000 ppm	0/50	0/50	0/50	0/50	0/49	0/47	0/45	0/42	0/50 (0/15)

No. of animals with mass / No. of survival animals at first week on each period.

(No. of dead and moribund animals with mass / No. of dead and moribund animals)

TABLE 8 INCIDENCE AND TIME OF MASS OCCURRENCE IN CLINICAL OBSERVATION OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Time of mass occurrence (week)	0-13	14-26	27-39	40-52	53-65	66-78	79-91	92-104	0-104
External mass						- "			
Control	0/50	0/50	0/50	1/50	1/49	2/49	3/46	4/42	6/50 (3/13)
$320~\mathrm{ppm}$	0/50	0/50	0/50	0/50	0/50	1/49	2/48	3/47	3/50 (1/10)
$800~\mathrm{ppm}$	0/50	0/50	0/50	0/50	0/49	2/48	6/48	9/46	10/50 (1/8)
$2000~\mathrm{ppm}$	0/50	0/50	0/50	0/50	0/50	1/50	4/50	7/46	7/50 (1/8)
Internal mass									
Control	0/50	0/50	0/50	0/50	0/49	1/49	1/46	1/42	3/50 (3/13)
$320~\mathrm{ppm}$	0/50	0/50	0/50	0/50	0/50	0/49	0/48	1/47	1/50 (1/10)
800 ppm	0/50	0/50	0/50	0/50	0/49	0/48	0/48	0/46	0/50 (0/ 8)
$2000~{ m ppm}$	0/50	0/50	0/50	0/50	0/50	0/50	0/50	0/46	0/50 (0/ 8)

No. of animals with mass / No. of survival animals at first week on each period.

(No. of dead and moribund animals with mass / No. of dead and moribund animals)

TABLE 9 HEMATOLOGY OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group	Control	320 ppm	800 ppm	2000 ppm
No. of animals examined	40	44	37	34
Red blood cell $(10^6/\mu\mathrm{L})$	$8.00 \pm 1.60$	$8.50 \pm 1.26$	$8.36 \pm 1.38$	$8.47 \pm 1.60$
Hemoglobin (g/dL)	$13.5~\pm~2.8$	$14.1 \pm 1.8$	$13.7 \pm 2.1$	$13.4 ~\pm~ 2.4$
Hematocrit (%)	$41.2~\pm~7.2$	$42.9~\pm~4.7$	$42.0~\pm~5.3$	$41.8 \pm 6.6$
MCV (fL)	$52.2~\pm~6.5$	$51.0 \pm 5.7$ *	50.9 ± 5.2 **	* 49.9 ± 3.9 **
MCH (pg)	$17.0 \pm 1.6$	$16.8 \pm 1.6$	$16.5 \pm 1.1$ *	$15.9 \pm 1.0  **$
MCHC (g/dL)	$32.6 \pm 1.7$	$32.9 \pm 1.1$	$32.5 \pm 1.6$	32.0 ± 1.2 **
Platelet $(10^3/\mu L)$	$891 ~\pm~ 277$	$861 \pm 228$	$932 \pm 148$	$856 \pm 163$
WBC $(10^3/\mu\mathrm{L})$	$8.19 \pm 11.57$	$7.02 ~\pm~ 2.63$	$6.96~\pm~2.68$	$11.15 \pm 23.07$

Significant difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$ , Test of Dunnett

TABLE 10 HEMATOLOGY OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group	Control	320 ppm	800 ppm	2000 ppm			
No. of animals examined	37	37	41	42			
Red blood cell $(10^6/\mu$ L)	$8.11 \pm 0.74$	$8.00 \pm 0.85$	$7.14~\pm~1.92$	** 7.09 ±	1.05 **		
Hemoglobin (g/dL)	$14.8~\pm~1.3$	$14.6~\pm~1.3$	$13.1 \pm 3.3$	** 13.1 ±	1.5 **		
Hematocrit (%)	$43.6~\pm~2.9$	$43.1~\pm~3.3$	$39.2~\pm~8.5$	** 39.5 ±	4.0 **		
MCV (fL)	$53.9~\pm~2.4$	$54.2~\pm~3.2$	$57.3 \pm 10.6$	56.4 ±	4.6 **		
MCH (pg)	$18.3~\pm~0.5$	$18.3 \pm 0.8$	$18.7~\pm~2.5$	$18.7 \pm$	1.2		
MCHC (g/dL)	$34.0~\pm~0.9$	$33.9~\pm~0.7$	$33.0~\pm~2.3$	** 33.2 ±	0.8 **		
Platelet $(10^3/\mu$ L)	$618~\pm~103$	$645~\pm~93$	$641~\pm~169$	* 718 ±	77 **		
WBC $(10^3/\mu L)$	$13.51 \pm 59.34$	$4.04 \pm 5.69$	$5.03 \pm 11.32$	$4.79 \pm$	13.31		

Data represent means  $\pm$  S.D.

TABLE 11 BIOCHEMISTRY OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Control	320 ppm	800 ppm	2000 ppm
No. of animals examined	40	44	37	34
Total protein (g/dL)	$6.5~\pm~0.5$	$6.7 \pm 0.4$ **	$6.6~\pm~0.4$	$6.5 \pm 0.3$
Albumin (g/dL)	$3.3~\pm~0.4$	$3.4~\pm~0.3$	$3.3~\pm~0.3$	$3.4~\pm~0.2$
A/G ratio	$1.1~\pm~0.1$	$1.1~\pm~0.2$	$1.0~\pm~0.1$	$1.1 ~\pm~ 0.1$
T-Bilirubin (mg/dL)	$0.22~\pm~0.44$	$0.66 \pm 3.18$ *	$0.22~\pm~0.15~**$	$0.33 \pm 0.35 **$
Glucose (mg/dL)	$153~\pm~18$	$155~\pm~19$	$142~\pm~24~$	$146~\pm~17$
T-Cholesterol (mg/dL)	$164~\pm~42$	218 ± 50 **	$252 \pm 72  **$	285 ± 43 **
Triglyceride (mg/dL)	$73 \pm 51$	108 ± 83 *	115 ± 82 **	$195 \pm 148$ **
Phospholipid (mg/dL)	$232 \pm 71$	$301 \pm 70  **$	341 ± 87 **	409 ± 67 **
GOT (IU/L)	$95 \pm 62$	$106~\pm~87$	$104 \pm 24  *$	171 ± 69 **
GPT (IU/L)	$41 \pm 16$	$52 \pm 50$	$47 \pm 15$	66 ± 19 **
LDH (IU/L)	$208~\pm~58$	$196~\pm~39$	$177 \pm 42  **$	193 ± 134 **
ALP (IU/L)	$231~\pm~115$	$283 \pm 95$ *	335 ± 116 **	484 ± 171 **
γ -GTP (IU/L)	$12 \pm 8$	24 ± 11 **	46 ± 29 **	111 ± 33 **
CPK (IU/L)	$106~\pm~73$	$94~\pm~14$	$94~\pm~21$	$107~\pm~71$
Urea nitrogen (mg/L)	$19.2 \pm 6.9$	$18.7~\pm~3.1$	$21.4~\pm~4.0~^{**}$	23.0 ± 4.0 **
Creatinine (mg/dL)	$0.5~\pm~0.1$	$0.5~\pm~0.1$	$0.6~\pm~0.1$	$0.5 \pm 0.1$
Sodium (mEq/L)	$142~\pm~2$	$141  \pm  2$	141 ± 1 **	140 ± 1 **
Potassium (mEq/L)	$3.7~\pm~0.4$	$3.8~\pm~0.5$	$3.8~\pm~0.4$	$4.0 \pm 0.4$ *
Chloride (mEq/L)	$107 \pm 2$	$105 \pm 2 \qquad **$	105 ± 2 **	105 ± 2 *
Calcium (mg/dL)	$10.2~\pm~0.4$	$10.2~\pm~0.9$	$10.3 \pm 0.3$ *	$10.4~\pm~0.3~$
Inorganic phosphorus (mg/dL)	$4.3~\pm~0.7$	$4.3~\pm~1.3$	$4.4~\pm~0.5$	$4.2~\pm~0.5$

TABLE 12 BIOCHEMISTRY OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group	Control	320 ppm	800 ppm	2000 ppm
No. of animals examined	37	37	41	42
Total protein (g/dL)	$6.9~\pm~0.5$	$6.8~\pm~0.4$	$6.7~\pm~0.5$	$6.3 \pm 0.4$ **
Albumin (g/dL)	$3.9~\pm~0.3$	$3.9~\pm~0.2$	$3.9~\pm~0.3$	$3.8 \pm 0.2$ **
A/G ratio	$1.3~\pm~0.1$	$1.3~\pm~0.1$	$1.4~\pm~0.1$	$1.5~\pm~0.2~~^{**}$
T-Bilirubin (mg/dL)	$0.17~\pm~0.11$	$0.16~\pm~0.2$	$0.47 ~\pm~ 1.42$	$0.17~\pm~0.11$
Glucose (mg/dL)	$145~\pm~13$	$150~\pm~14$	$140~\pm~22$	$144 ~\pm~ 16$
T-Cholesterol (mg/dL)	$130 \pm 26$	$140~\pm~29$	$154 \pm 36$ **	$155 \pm 23$ **
Triglyceride (mg/dL)	$64~\pm~54$	$62~\pm~44$	$92 \pm 98$	$75 \pm 116$
Phospholipid (mg/dL)	$231 ~\pm~ 48$	$235~\pm~46$	$263 \pm 63$ *	261 ± 43 *
GOT (IU/L)	$159 \pm 111$	$110~\pm~39$	$172~\pm~199$	$120 \pm 79$ *
GPT (IU/L)	$63 \pm 35$	$45 \pm 18$ *	$52 \pm 41$ *	$42 \pm 17$ **
LDH (IU/L)	$330~\pm~225$	$263~\pm~78$	$379~\pm~378$	$261~\pm~104$
ALP (IU/L)	$125~\pm~75$	$115~\pm~34$	$144~\pm~101$	$154 \pm 58$ **
γ -GTP (IU/L)	$5 \pm 4$	$5 \pm 2$	8 ± 6 *	12 ± 8 **
CPK (IU/L)	$150~\pm~290$	$96~\pm~21$	$158~\pm~320$	$110 \pm 47$
Urea nitrogen (mg/L)	$17.3 \pm 1.7$	$17.4 \pm 5.8$	$17.5~\pm~3.1$	$20.3 \pm 3.2  **$
Creatinine (mg/dL)	$0.5~\pm~0.1$	$0.5~\pm~0.1$	$0.5~\pm~0.1$	$0.5~\pm~0.1$
Sodium (mEq/L)	$140 \pm 1$	$140  \pm  2$	$140 \pm 2$	$140 \pm 2$
Potassium (mEq/L)	$3.8~\pm~0.5$	$3.7~\pm~0.4$	$3.9~\pm~0.4$	$4.0 \pm 0.5$ *
Chloride (mEq/L)	$105~\pm~2$	$105~\pm~2$	$105 \pm 3$	$106 \pm 2$
Calcium (mg/dL)	$10.2~\pm~0.3$	$10.1~\pm~0.4$	$10.2~\pm~0.4$	$10.1~\pm~0.3$
Inorganic phosphorus (mg/dL)	$4.0~\pm~0.8$	$3.9~\pm~0.8$	$4.2~\pm~0.6$	$4.4 \pm 0.5$ *

TABLE 13 URINALYSIS OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group		Control	320 ppm	800 ppm	2000 ppm	
No. of animals examined		40	44	38	35	
pН	6.0	1	0 **	2	2	
•	6.5	3	0	$\overline{3}$	ō	
	7.0	12	5	7	6	
	7.5	20	22	14	16	
	8.0	3	16	12	11	
	8.5	1	1	0	0	
	(Grade)					
Protein	` <u>-</u> ′	0	0	0 **	0	
	±	0	0	0	0	
	+	0	0	0	0	
	2+	2	<b>2</b>	0	0	
	3+	27	21	12	18	
	4+	11	21	26	17	
Occult blood	-	39	42	37	27 *	
	±.	1	0	0	<b>2</b>	
	+	0	0	0	0	
	2+	0	1	1	3	
	3+	0	1	0	3	

Significant difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$  Chi square test

TABLE 14 URINALYSIS OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group		$\operatorname{Control}$	320 ppm	800 ppm	2000 ppm
No. of animals exar	nined	37	40	42	45
pН	6.0	0	0	0	1
	6.5	3	2	5	12
	7.0	11	11	12	8
	7.5	12	13	13	10
	8.0	9	13	10	13
	8.5	2	1	2	1
	(Grade)				
Protein	<u> </u>	0	0 **	0 **	0 **
	±	2	0	0	0
	+	10	1	0	1
	2+	16	16	9	6
	3+	6	11	21	27
	4+	3	12	12	11
Occult blood	-	35	37	30 *	9 **
	±	1	1	4	2
	+	0	0	0	0
	2+	1	1	0	0
	3+	0	1	8	34

Significant difference, \*:  $p \le 0.05$ , \*\*:  $p \le 0.01$  Chi square test

TABLE 15 ORGAN WEIGHTS OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group		Control	320 ppm	800 ppm	2000 ppm
No. of animexami		<40>	<44>	<37>	<35>
Adrenal	(g) (%)	$0.070 \pm 0.015$ $0.018 \pm 0.003$	$0.081 \pm 0.066$ $0.023 \pm 0.019$	$0.083 \pm 0.121$ $0.027 \pm 0.048$	$0.090 \pm 0.170 * 0.030 \pm 0.054 **$
Testis	(g) (%)	$2.665 \pm 1.057$ $0.706 \pm 0.295$	$2.599 \pm 1.004$ $0.726 \pm 0.275$	$2.597 \pm 0.877$ $0.782 \pm 0.231$	$2.787 \pm 0.748$ $0.937 \pm 0.213$ **
Heart	(g) (%)	$1.178 \pm 0.118$ $0.309 \pm 0.035$	$1.127 \pm 0.105$ $0.316 \pm 0.032$	$1.103 \pm 0.091$ ** $0.338 \pm 0.051$ **	$1.023 \pm 0.089$ ** $0.350 \pm 0.049$ **
Lung	(g) (%)	$1.442 \pm 0.373$ $0.380 \pm 0.125$	$1.370 \pm 0.155$ $0.385 \pm 0.057$	$1.347 \pm 0.168$ $0.415 \pm 0.094$ **	$1.377 \pm 0.400$ ** $0.486 \pm 0.261$ **
Kidney	(g) (%)	$2.515 \pm 0.197$ $0.660 \pm 0.071$	$2.666 \pm 0.276$ ** $0.750 \pm 0.117$ **	$2.764 \pm 0.210$ ** $0.846 \pm 0.108$ **	$2.791 \pm 0.217$ ** $0.956 \pm 0.150$ **
Spleen	(g) (%)	$1.570 \pm 3.764$ $0.442 \pm 1.186$	$1.566 \pm 3.674$ $0.456 \pm 1.151$	$1.035 \pm 0.762$ $0.314 \pm 0.232$ *	$1.380 \pm 2.353$ $0.469 \pm 0.805$ **
Liver	(g) (%)	$10.574 \pm 2.260$ $2.770 \pm 0.669$	$11.038 \pm 1.625$ $3.091 \pm 0.485$ **	11.425 ± 1.535 ** 3.473 ± 0.403 **	$11.545 \pm 2.291$ ** $3.922 \pm 0.757$ **
Brain	(g) (%)	$2.017 \pm 0.063$ $0.530 \pm 0.048$	$2.020 \pm 0.050$ $0.568 \pm 0.051$ *	$2.023 \pm 0.053$ $0.619 \pm 0.065$ **	$2.019 \pm 0.054$ $0.693 \pm 0.105$ **

TABLE 16 ORGAN WEIGHTS OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

Group		Control	320 ppm	800 ppm	2000 ppm
No. of anir		<37>	<40>	<42>	<42>
Adrenal	(g) (%)	$0.067 \pm 0.011$ $0.027 \pm 0.004$	$0.067 \pm 0.009 \\ 0.027 \pm 0.004$	$0.065 \pm 0.010$ $0.029 \pm 0.005$	$0.058 \pm 0.013$ ** $0.030 \pm 0.010$
Ovaries	(g) (%)	$0.129 \pm 0.019$ $0.052 \pm 0.009$	$0.135 \pm 0.033$ $0.056 \pm 0.016$	$0.142 \pm 0.102$ $0.063 \pm 0.040$	$0.122 \pm 0.027$ $0.063 \pm 0.015$ **
Heart	(g) (%)	$0.842 \pm 0.100$ $0.340 \pm 0.040$	$0.837 \pm 0.077$ $0.344 \pm 0.027$	$0.818 \pm 0.092$ $0.370 \pm 0.074$ *	$0.745 \pm 0.050$ ** $0.385 \pm 0.035$ **
Lung	(g) (%)	$1.066 \pm 0.186$ $0.431 \pm 0.079$	$1.046 \pm 0.191$ $0.434 \pm 0.109$	$1.030 \pm 0.157$ $0.466 \pm 0.104$	$0.949 \pm 0.115$ ** $0.492 \pm 0.077$ *
Kidney	(g) (%)	$1.666 \pm 0.103$ $0.672 \pm 0.046$	$1.897 \pm 0.300$ ** $0.784 \pm 0.157$ **	$1.902 \pm 0.230$ ** $0.854 \pm 0.116$ **	$1.786 \pm 0.128$ ** $0.924 \pm 0.086$ **
Spleen	(g) (%)	$0.854 \pm 1.247$ $0.346 \pm 0.488$	$0.743 \pm 0.869$ $0.308 \pm 0.374$	$1.313 \pm 2.631$ $0.619 \pm 1.269$	$0.599 \pm 0.406$ $0.315 \pm 0.230$ *
Liver	(g) (%)	$6.493 \pm 1.190$ $2.606 \pm 0.388$	$6.592 \pm 1.002$ $2.697 \pm 0.283$	$6.702 \pm 1.160$ $2.996 \pm 0.503$ **	$6.347 \pm 0.981$ $3.277 \pm 0.512$ **
Brain	(g) (%)	$1.861 \pm 0.047$ $0.754 \pm 0.078$	$1.848 \pm 0.042$ $0.766 \pm 0.097$	$1.835 \pm 0.046$ * $0.830 \pm 0.118$ **	$1.835 \pm 0.039$ * $0.953 \pm 0.104$ **

TABLE 17 NEOPLASTIC LESIONS OF MALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group	$\operatorname{Control}$	$320~\mathrm{ppm}$	$800~\mathrm{ppm}$	$2000~\mathrm{ppm}$	
No. of animals examined	<50>	<50>	<49>	<50>	
Skin					
Keratoacanthoma	3 ( 6%) a)	2 (4%)	1 ( 2%)	2 ( 4%)	
Squamous cell papilloma	1 (2%)	2(4%)	1 (2%)	1 ( 2%)	
Subcutis					
Fibroma	7 (14%)	2 (4%)	4 ( 8%)	3 (6%)	
Lung					
Bronchiolar-alveolar adenoma	1 ( 2%)	3 (6%)	2 ( 4%)	1 (2%)	
Spleen					
Mononuclear cell leukemia	6 (12%)	4 (8%)	6 (12%)	5 (10%)	
Stomach					
Squamous cell papilloma	1 (2%)	1 ( 2%)	0 ( 0%)	0 ( 0%)	
Squamous cell carcinoma	0 ( 0%)	0 ( 0%)	0 ( 0%)	1 ( 2%)	
Liver					
Hepatocellular adenoma	1 (2%)	4 (8%)	4 (8%)	10 (20%) **	<b>វវ</b> ប្ប
Pituitary					
Adenoma	21~(42%)	22 (44%)	14 (29%)	13 (26%)	Û
Thyroid					
Č-cell adenoma	7 (14%)	10 (20%)	6 (12%)	11 (22%)	
Follicular adenoma	0 ( 0%)	0 ( 0%)	1 (2%)	$2\ (\ 4\%)$	
Follicular adenocarcinoma	0 ( 0%)	1 (2%)	0 ( 0%)	1 ( 2%)	
Follicular adenoma / adenocarcinoma	0 ( 0%)	1 ( 2%)	1 (2%)	3 (6%)	Ť
Testis					
Interstitial cell tumor	28 (56%)	31 (62%)	35 (71%)	35 (70%)	Ť

a) : No. of animals with bearing tumor (incidence; %)

<sup>\*\* :</sup> Statistically differenct from control group at p≤0.01 by Fisher exact test

**<sup>1</sup>** and **11**: The trend of treated groups statistically different from control group at  $p ext{ ≤ } 0.05$  and  $p ext{ ≤ } 0.01$  by Peto test, respectively.

 $<sup>\</sup>mathbb{Q}$  and  $\mathbb{Q}$   $\mathbb{Q}$ : The trend of treated groups statistically different from control group at  $p \le 0.05$  and  $p \le 0.01$  by Cochran-Armitage test, respectively.

TABLE 18 NEOPLASTIC LESIONS OF FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

Group No. of animals examined	Control <50>	320 ppm <50>	800 ppm <50>	2000 ppm <50>	
Spleen					
Mononuclear cell leukemia	7 (14%) <sup>a)</sup>	6 (12%)	6 (12%)	8 (16%)	
Stomach					
Squamous cell papilloma	0 ( 0%)	1 (2%)	0 ( 0%)	0 ( 0%)	
Liver					
Hepatocellular adenoma	0 ( 0%)	1(2%)	0 ( 0%)	3 (6%)	<b>f</b> û
Pituitary					
Adenoma	19 (38%)	15 (30%)	16 (32%)	15 (30%)	
Thyroid					
Č-cell adenoma	6 (12%)	4 (8%)	5 (10%)	8 (16%)	
Follicular adenoma	1 ( 2%)	2 (4%)	0 (0%)	0 ( 0%)	
Uterus					
Endometrial stromal polyp	7 (14%)	8 (16%)	11 (22%)	7 (14%)	
Endometrial stromal sarcoma	4 ( 8%)	1 (2%)	0 ( 0%)	0 ( 0%)	$\hat{\mathbf{U}}$
Mammary gland					
Fibroadenoma	3 (6%)	3 (6%)	6 (12%)	7 (14%)	

a) : No. of animals with bearing tumor (incidence; %)

î ↓: The trend of treated groups statistically different from control group at p≤0.05 by Cochran-Armitage test

TABLE 19 NON-NEOPLASTIC LESIONS OF MALE AND FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE (SELECTED)

				ale			Fer	nale	
Group No. of animals examined		Control <50>	320 ppm <50>	800 ppm <49>	2000 ppm <50>	Control <50>	320 ppm <50>	800 ppm <50>	2000 ppm <50>
Liver	Grade								
Basophilic cell focus	+ 2+ 3+	9 0 0	8 0 0	15 7 0	16 7 0	3 1 0	3 1 0	4 0 1	6 0 0
Kidney									
Chronic nephropathy	+ 2+ 3+ 4+	$10 \\ 25 \\ 13 \\ 0$	$egin{array}{c} 3 \\ 13 \\ 28 \\ 2 \end{array}$	4 15 26 3	3 7 36 2	15 2 2 0	18 6 7 0	16 13 6 0	31 6 0 0
Papillary necrosis	+ 2+ 3+	1 0 0	12 0 0	14 1 0	20 4 0	0 0 0	7 0 0	23 0 0	7 19 2
Mineralization : papilla	+ 2+	3 0	2 0	11 0	19 0	3 0	3	6 1	22 1
Urothelial hyperplasia : pelvis	+ 2+	16 0	18 0	25 0	.25 1	9	9	9	27 0
Stomach (Forestomach)									
Squamous cell hyperplasia	+	3	0	0	5	1	3	3	4
Basal cell hyperplasia	+	0	0	0	0	0	0	0	3

Grade +: Slight 2+: Moderate 3+: Marked 4+: Severe

TABLE 20 CAUSE OF DEATH OF MALE AND FEMALE RATS IN THE 2-YEAR DRINKING WATER STUDY OF 2-HYDROXYETHYL ACRYLATE

		M	ale		Female			
Group	Control	320 ppm	800 ppm	2000 ppm	Control	320 ppm	800 ppm	2000 ppm
No. of dead/moribund animals	<10>	<6>	<12>	<15>	<13>	<10>	<8>	<8>
Chronic nephropathy	0	0	0	3	0	0	0	0
Urinary retention	0	0	1	0	0	0	0	0
Thrombosis	0	0	0	1	0	0	0	0
Deglutition disorder	0	0	0	0	0	0	2	0
Tumor death:								
leukemia	<b>2</b>	2	5	3	1	3	2	4
skin/appendage	1	0	0	1	0	0	0	0
subcutis	2	0	0	0	1	0	0	0
tongue	0	0	0	1	0	0	0	0
salivary gland	0	0	2	0	0	0	0.	0
pancreas	0	0	0	0	0	0	1	0
pituitary	1	<b>2</b>	1	4	4	5	1	2
adrenal	1	0	1	0	0	0	0	0
uterus	-	-	-	-	5	0	0	0
mammary gland	0	0	0	0	0	0	0	1
clitoral gland	-	-	-	-	1	0	0	0
brain	0	0	0	0	0	0	1	0
spinal cord	0	0	0	0	0	1	0	0
Zymbal gland	0	0	0	1	0	0	0	0
bone	1	0	1	0	0	0	1	0
vertebrae	0	0	1	0	0	0	0	0
peritoneum	1	0	0	0	0	0	0	0
No microscopical confirmation	1	2	0	1	1	1	0	1

TABLE 21 HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER: F344/DuCrj MALE RATS

Organs Tu	umors	No. of animals examined	No. of animals with bearing tumor	Incidence (%)	Min Max. (%)
	eratoacanthoma Juamous cell papilloma	<1248>	39 14	3.1 1.1	0 - 8 0 - 4
Subcutis Fi	broma	<1249>	90	7.2	2 - 14
Lung Br	ronchiolar-alveolar adenoma	<1249>	37	3.0	0 - 8
Spleen Me	ononuclear cell leukemia	<1249>	152	12.2	4 - 22
	quamous cell papillpma quamous cell carcinoma	<1248>	2 0	0.2	0 - 2
Liver He	epatocellular adenoma	<1249>	20	1.6	0 - 6
Pituitary Ac	denoma	<1244>	439	35.3	18 - 66
Fo	-cell adenoma ollicular adenoma ollicular adenocarcinoma	<1243>	155 12 27	12.5 1.0 2.2	4 - 26 0 - 4 0 - 8
Testis In	nterstitial cell tumor	<1249>	1099	88.0	74 - 98

25 carcinogenicity studies examined in Japan Bioassay Research Center were used. Study No. : 0043, 0059, 0061, 0063, 0065, 0067, 0095, 0104, 0115, 0130, 0141, 0158, 0162, 0189, 0205, 0210, 0224, 0242, 0267, 0269 0284, 0288, 0294, 0296, 0318

TABLE 22 HISTORICAL CONTROL DATA OF SELECTED NEOPLASTIC LESIONS IN JAPAN BIOASSAY RESEARCH CENTER : F344/DuCrj FEMALE RATS

Organs Tumors	No. of animals examined	No. of animals with bearing tumor	Incidence (%)	Min Max. (%)
Spleen	<1197>			
Mononuclear cell leukemia		160	13.4	2 - 26
Stomach	<1197>			
Squamous cell papilloma		2	0.2	0 - 2
Liver	<1197>			
Hepatocellular adenoma		16	1.3	0 - 6
Pituitary	<1195>			
Adenoma		493	41.3	16 - 71
Thyroid	<1191>			
C-cell adenoma		115	9.7	0 - 16
Follicular adenoma		12	1.0	0 - 4
Uterus	<1197>			
Endometrial stromal polyp		172	14.4	2 - 28
Endometrial stromal sarcoma		7	0.6	0 - 2
Mammary gland	<1197>			
Fibroadenoma		130	10.9	0 - 20

24 carcinogenicity studies examined in Japan Bioassay Research Center were used. Study No.: 0043, 0059, 0061, 0063, 0065, 0067, 0095, 0104, 0115, 0130, 0141, 0158, 0162, 0189, 0205, 0210, 0224, 0242, 0267, 0269 0284, 0296, 0303, 0318