

シクロヘキセンのマウスを用いた
吸入による 13 週間毒性試験報告書

試験番号：0364

TABLES

TABLES

TABLE 1 EXPERIMENTAL DESIGN AND MATERIALS AND METHODS IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

TABLE 2 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF MALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

TABLE 3 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF FEMALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

TABLE 4 FOOD CONSUMPTION CHANGES OF MALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

TABLE 5 FOOD CONSUMPTION CHANGES OF FEMALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

TABLE 1 EXPERIMENTAL DESIGN AND MATERIALS AND METHODS
IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

<Method of Administration>	Inhalation
<Number of Groups>	Male 6, Female 6
<Size of Groups>	10 males and 10 females of each group
<Animals>	Strain and Species Crj:BDF ₁ mouse
	Animal Source Charles River Japan, Inc.
	Duration Held Before Study 2 wk
	Age When Placed on Study 6 wk
	Age When Killed 19 wk
<Doses>	Male and Female 0, 20, 40, 75, 150 or 300ppm
<Duration of Dosing>	6 h/d, 5 d/wk for 13 wk
<Animal Maintenance>	Feed CRF-1 (Oriental Yeast Co., Ltd.) Sterilized by γ -ray Available <i>ad libitum</i>
	Water Filtrated and sterilized by ultraviolet ray Automatic watering system Available <i>ad libitum</i>
	Animal per Cage Single (stainless steel wire)
	Animal Room Environment Barrier system Temperature : 21±2°C Humidity : 55±15% Fluorescent light 12 h/d 15~17 room air changes /h
	Chamber Environment Barrier system Temperature : 20~24°C Humidity : 30~70% 12±1 air changes /h
<Type and Frequency of Observation>	Clinical Sign Observed 1 per day for mortality, Detailed clinical observation performed on once weekly before exposure.
	Body Weight Weighed 1 per wk
	Food Consumption Weighed 1 per wk

TABLE 1 EXPERIMENTAL DESIGN AND MATERIALS AND METHODS
(Continued) IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

<Urinalysis>

Urinalysis performed on all animals that survived to end of dosing period using fresh urine collection.

The following measurement parameters were examined;

pH, Protein, Glucose, Ketone body, Occult blood, Urobilinogen.

<Hematology>

Hematological examination performed on scheduled sacrificed animals.

The following measurement parameters were examined;

Red blood cell (RBC), Hemoglobin, Hematocrit,
Mean Corpuscular Volume (MCV),
Mean Corpuscular hemoglobin (MCH),
Mean Corpuscular hemoglobin concentrate (MCHC),
Platelet, White blood cell (WBC), Differential WBC.

<Biochemistry>

Biochemical examination performed on scheduled sacrificed animals.

The following measurement parameters were examined;

Total protein, Albumin, A/G ratio,
Total bilirubin, Glucose, Total cholesterol,
Triglyceride, Phospholipid,
Glutamic oxaloacetic transaminase (GOT),
Glutamic pyruvic transaminase (GPT),
Lactate dehydrogenase (LDH),
Alkaline phosphatase (ALP),
 γ -Glutamyl transpeptidase (γ -GTP),
Creatine phosphokinase (CPK),
Urea nitrogen, Sodium, Potassium, Chloride,
Calcium, Inorganic phosphorus.

<Necropsy>

Necropsy performed on all animals.

<Organ Weight>

Organ weight measurement performed on scheduled sacrificed animals.

The following organs were weighed;

thymus, adrenal, testis, ovary, heart, lung, kidney, spleen, liver, brain.

<Histopathologic Examination>

Histopathologic examination performed on all animals.

The following organs were examined;

skin, nasal cavity, nasopharynx, larynx, trachea, lung,
bone marrow, lymph node, thymus, spleen, heart, tongue,
salivary gland, esophagus, stomach, small intestine,
large intestine, liver, gall bladder, pancreas, kidney, urinary bladder,
pituitary, thyroid, parathyroid, adrenal, testis, epididymis, seminal vesicle,
prostate, ovary, uterus, vagina, mammary gland,
brain, spinal cord, peripheral nerve, eye, harderian gland, muscle, bone.

TABLE 2 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF MALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

Week-Day on Study	0ppm		20ppm		40ppm		75ppm		150ppm		300ppm						
	Av.Wt.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.			
	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>			
0-0	23.0 (10)	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10
1-7	24.9 (10)	10/10	24.4 (10)	98	10/10	24.1 (10)	97	10/10	24.5 (10)	98	10/10	24.9 (10)	100	10/10	24.2 (10)	97	10/10
2-7	26.0 (10)	10/10	25.7 (10)	99	10/10	25.0 (10)	96	10/10	24.8 (10)	95	10/10	25.4 (10)	98	10/10	25.3 (10)	97	10/10
3-7	26.7 (10)	10/10	26.2 (10)	98	10/10	25.7 (10)	96	10/10	25.4 (10)	95	10/10	26.0 (10)	97	10/10	25.8 (10)	97	10/10
4-7	27.4 (10)	10/10	26.5 (10)	97	10/10	25.9 (10)	95	10/10	25.6 (10)	93	10/10	26.5 (10)	97	10/10	26.4 (10)	96	10/10
5-7	28.2 (10)	10/10	27.2 (10)	96	10/10	26.7 (10)	95	10/10	26.4 (10)	94	10/10	27.0 (10)	96	10/10	26.7 (10)	95	10/10
6-7	29.2 (10)	10/10	28.1 (10)	96	10/10	27.1 (10)	93	10/10	27.2 (10)	93	10/10	27.6 (10)	95	10/10	27.1 (10)	93	10/10
7-7	29.6 (10)	10/10	28.0 (10)	95	10/10	27.6 (10)	93	10/10	27.8 (10)	94	10/10	28.1 (10)	95	10/10	27.6 (10)	93	10/10
8-7	30.4 (10)	10/10	29.1 (10)	96	10/10	27.9 (10)	92	10/10	28.5 (10)	94	10/10	28.6 (10)	94	10/10	28.1 (10)	92	10/10
9-7	31.0 (10)	10/10	29.4 (10)	95	10/10	28.5 (10)	92	10/10	28.9 (10)	93	10/10	29.3 (10)	95	10/10	28.2 (10)	91	10/10
10-7	31.6 (10)	10/10	30.1 (10)	95	10/10	29.4 (10)	93	10/10	30.0 (10)	95	10/10	29.9 (10)	95	10/10	29.2 (10)	92	10/10
11-7	32.3 (10)	10/10	30.7 (10)	95	10/10	29.8 (10)	92	10/10	30.7 (10)	95	10/10	30.4 (10)	94	10/10	29.3 (10)	91	10/10
12-7	33.0 (10)	10/10	31.8 (10)	96	10/10	30.3 (10)	92	10/10	31.1 (10)	94	10/10	31.4 (10)	95	10/10	30.1 (10)	91	10/10
13-7	33.6 (10)	10/10	31.9 (10)	95	10/10	30.6 (10)	91	10/10	30.8 (10)	92	10/10	31.1 (10)	93	10/10	30.1 (10)	90	10/10

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

TABLE 3 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF FEMALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

Week-Day on Study	0ppm		20ppm		40ppm		75ppm		150ppm		300ppm						
	Av.Wt.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.			
		<10>															
0-0	19.0 (10)	10/10	19.1 (10)	101	10/10	19.1 (10)	101	10/10	19.1 (10)	101	10/10	19.1 (10)	101	10/10	19.1 (10)	101	10/10
1-7	20.0 (10)	10/10	19.5 (10)	98	10/10	19.9 (10)	100	10/10	20.2 (10)	101	10/10	19.7 (10)	99	10/10	19.5 (10)	98	10/10
2-7	21.1 (10)	10/10	21.3 (10)	101	10/10	20.8 (10)	99	10/10	21.1 (10)	100	10/10	20.7 (10)	98	10/10	20.2 (10)	96	10/10
3-7	21.6 (10)	10/10	21.6 (10)	100	10/10	21.9 (10)	101	10/10	21.7 (10)	100	10/10	21.2 (10)	98	10/10	21.1 (10)	98	10/10
4-7	22.2 (10)	10/10	22.1 (10)	100	10/10	21.7 (10)	98	10/10	22.2 (10)	100	10/10	21.6 (10)	97	10/10	21.3 (10)	96	10/10
5-7	22.7 (10)	10/10	22.8 (10)	100	10/10	21.8 (10)	96	10/10	22.8 (10)	100	10/10	22.4 (10)	99	10/10	21.8 (10)	96	10/10
6-7	23.2 (10)	10/10	23.8 (10)	103	10/10	22.9 (10)	99	10/10	23.7 (10)	102	10/10	23.3 (10)	100	10/10	22.0 (10)	95	10/10
7-7	23.3 (10)	10/10	23.7 (10)	102	10/10	23.0 (10)	99	10/10	23.7 (10)	102	10/10	23.1 (10)	99	10/10	23.3 (10)	100	10/10
8-7	24.2 (10)	10/10	23.8 (10)	98	10/10	23.5 (10)	97	10/10	23.9 (10)	99	10/10	23.3 (10)	96	10/10	23.4 (10)	97	10/10
9-7	24.2 (10)	10/10	24.3 (10)	100	10/10	23.8 (10)	98	10/10	24.0 (10)	99	10/10	23.7 (10)	98	10/10	23.1 (10)	95	10/10
10-7	24.4 (10)	10/10	24.8 (10)	102	10/10	23.9 (10)	98	10/10	24.6 (10)	101	10/10	24.3 (10)	100	10/10	23.7 (10)	97	10/10
11-7	24.4 (10)	10/10	24.8 (10)	102	10/10	24.2 (10)	99	10/10	24.4 (10)	100	10/10	24.5 (10)	100	10/10	24.3 (10)	100	10/10
12-7	25.2 (10)	10/10	25.3 (10)	100	10/10	24.5 (10)	97	10/10	24.9 (10)	99	10/10	24.8 (10)	98	10/10	24.5 (10)	97	10/10
13-7	25.2 (10)	10/10	25.4 (10)	101	10/10	24.9 (10)	99	10/10	25.1 (10)	100	10/10	24.6 (10)	98	10/10	24.4 (10)	97	10/10

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

TABLE 4 FOOD CONSUMPTION CHANGES OF MALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

Week-Day on Study	0ppm		20ppm		40ppm		75ppm		150ppm		300ppm						
	Av.FC.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.			
	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>			
1-7	4.1 (10)	10/10	4.2 (10)	102	10/10	4.1 (10)	100	10/10	4.2 (10)	102	10/10	4.3 (10)	105	10/10	4.1 (10)	100	10/10
2-7	4.2 (10)	10/10	4.1 (10)	98	10/10	4.0 (10)	95	10/10	4.0 (10)	95	10/10	4.0 (10)	95	10/10	4.0 (10)	95	10/10
3-7	4.1 (10)	10/10	4.2 (10)	102	10/10	4.0 (10)	98	10/10	4.0 (10)	98	10/10	4.0 (10)	98	10/10	4.0 (10)	98	10/10
4-7	4.2 (10)	10/10	4.3 (10)	102	10/10	4.1 (10)	98	10/10	4.2 (10)	100	10/10	4.2 (10)	100	10/10	4.1 (10)	98	10/10
5-7	4.2 (10)	10/10	4.5 (10)	107	10/10	4.3 (10)	102	10/10	4.2 (10)	100	10/10	4.2 (10)	100	10/10	4.0 (10)	95	10/10
6-7	4.4 (10)	10/10	4.4 (10)	100	10/10	4.2 (10)	95	10/10	4.3 (10)	98	10/10	4.3 (10)	98	10/10	4.1 (10)	93	10/10
7-7	4.3 (10)	10/10	4.4 (10)	102	10/10	4.3 (10)	100	10/10	4.3 (10)	100	10/10	4.3 (10)	100	10/10	4.2 (10)	98	10/10
8-7	4.4 (10)	10/10	4.6 (10)	105	10/10	4.3 (10)	98	10/10	4.4 (10)	100	10/10	4.4 (10)	100	10/10	4.2 (10)	95	10/10
9-7	4.4 (10)	10/10	4.6 (10)	105	10/10	4.4 (10)	100	10/10	4.4 (10)	100	10/10	4.5 (10)	102	10/10	4.2 (10)	95	10/10
10-7	4.6 (10)	10/10	4.6 (10)	100	10/10	4.4 (10)	96	10/10	4.5 (10)	98	10/10	4.4 (10)	96	10/10	4.2 (10)	91	10/10
11-7	4.5 (10)	10/10	4.7 (10)	104	10/10	4.3 (10)	96	10/10	4.4 (10)	98	10/10	4.5 (10)	100	10/10	4.2 (10)	93	10/10
12-7	4.6 (10)	10/10	4.9 (10)	107	10/10	4.6 (10)	100	10/10	4.5 (10)	98	10/10	4.7 (10)	102	10/10	4.5 (10)	98	10/10
13-7	4.5 (10)	10/10	4.6 (10)	102	10/10	4.5 (10)	100	10/10	4.3 (10)	96	10/10	4.5 (10)	100	10/10	4.4 (10)	98	10/10

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

TABLE 5 FOOD CONSUMPTION CHANGES OF FEMALE MICE IN THE 13-WEEK INHALATION STUDY OF CYCLOHEXENE

Week-Day on Study	0ppm		20ppm		40ppm		75ppm		150ppm		300ppm						
	Av.FC.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.	Av.FC.	% of cont.	No.of Surviv.			
	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>	<10>			
1-7	3.4 (10)	10/10	3.6 (10)	106	10/10	3.5 (10)	103	10/10	3.5 (10)	103	10/10	3.5 (10)	103	10/10	3.5 (10)	103	10/10
2-7	3.5 (10)	10/10	3.7 (10)	106	10/10	3.4 (10)	97	10/10	3.6 (10)	103	10/10	3.6 (10)	103	10/10	3.3 (10)	94	10/10
3-7	3.6 (10)	10/10	3.8 (10)	106	10/10	3.5 (10)	97	10/10	3.7 (10)	103	10/10	3.6 (10)	100	10/10	3.5 (10)	97	10/10
4-7	3.8 (10)	10/10	4.1 (10)	108	10/10	3.7 (10)	97	10/10	4.0 (10)	105	10/10	3.9 (10)	103	10/10	3.6 (10)	95	10/10
5-7	3.8 (10)	10/10	4.2 (10)	111	10/10	3.8 (10)	100	10/10	4.1 (10)	108	10/10	4.0 (10)	105	10/10	3.8 (10)	100	10/10
6-7	4.0 (10)	10/10	4.3 (10)	108	10/10	3.9 (10)	98	10/10	4.1 (10)	103	10/10	4.1 (10)	103	10/10	3.9 (10)	98	10/10
7-7	4.0 (10)	10/10	4.3 (10)	108	10/10	4.0 (10)	100	10/10	4.3 (10)	108	10/10	4.2 (10)	105	10/10	4.0 (10)	100	10/10
8-7	4.1 (10)	10/10	4.3 (10)	105	10/10	4.2 (10)	102	10/10	4.2 (10)	102	10/10	4.2 (10)	102	10/10	4.0 (10)	98	10/10
9-7	4.2 (10)	10/10	4.3 (10)	102	10/10	4.2 (10)	100	10/10	4.3 (10)	102	10/10	4.2 (10)	100	10/10	4.0 (10)	95	10/10
10-7	4.1 (10)	10/10	4.3 (10)	105	10/10	4.1 (10)	100	10/10	4.4 (10)	107	10/10	4.3 (10)	105	10/10	4.1 (10)	100	10/10
11-7	4.1 (10)	10/10	4.3 (10)	105	10/10	4.1 (10)	100	10/10	4.3 (10)	105	10/10	4.2 (10)	102	10/10	3.9 (10)	95	10/10
12-7	4.2 (10)	10/10	4.6 (10)	110	10/10	4.4 (10)	105	10/10	4.5 (10)	107	10/10	4.4 (10)	105	10/10	4.4 (10)	105	10/10
13-7	4.1 (10)	10/10	4.3 (10)	105	10/10	4.3 (10)	105	10/10	4.3 (10)	105	10/10	4.4 (10)	107	10/10	4.1 (10)	100	10/10

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