FIG. 1

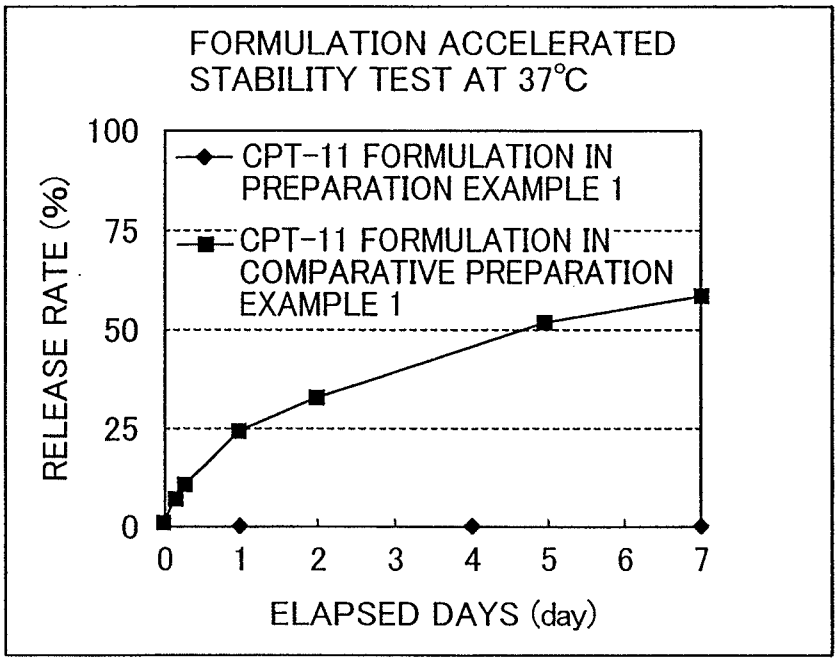

FORMULATION ACCELERATED STABILITY TEST AT 37°C
RELEASE RATE (%)
100
75
50
25
0
CPT-11 FORMULATION IN PREPARATION EXAMPLE 1
CPT-11 FORMULATION IN COMPARATIVE PREPARATION EXAMPLE 1
0
1
2
3
4
5
6
7
ELAPSED DAYS (day)

FIG. 2

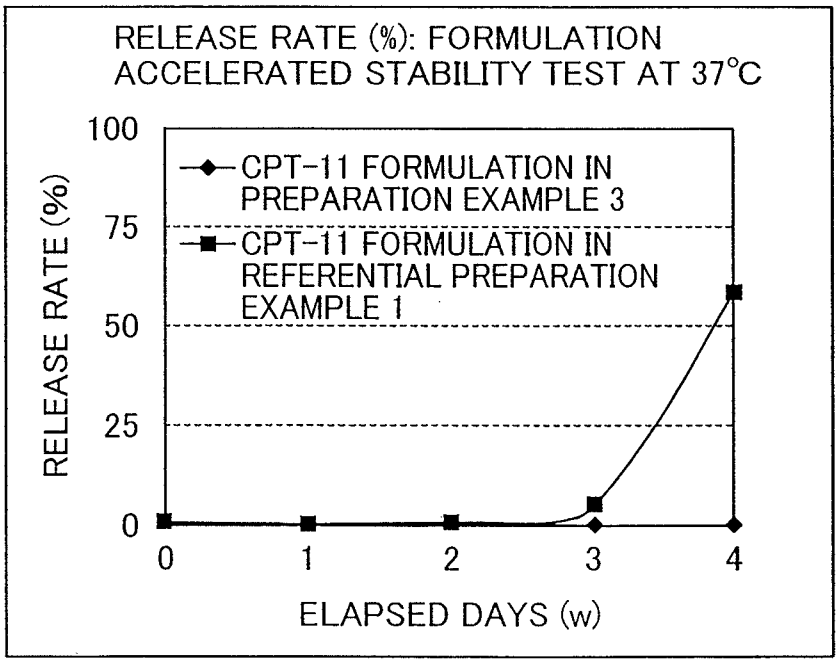

RELEASE RATE (%): FORMULATION ACCELERATED STABILITY TEST AT 37°C
RELEASE RATE (%)
100
75
50
25
0
CPT-11 FORMULATION IN PREPARATION EXAMPLE 3
CPT-11 FORMULATION IN REFERENTIAL PREPARATION EXAMPLE 1
0
1
2
3
4
ELAPSED DAYS (w)

FIG. 3

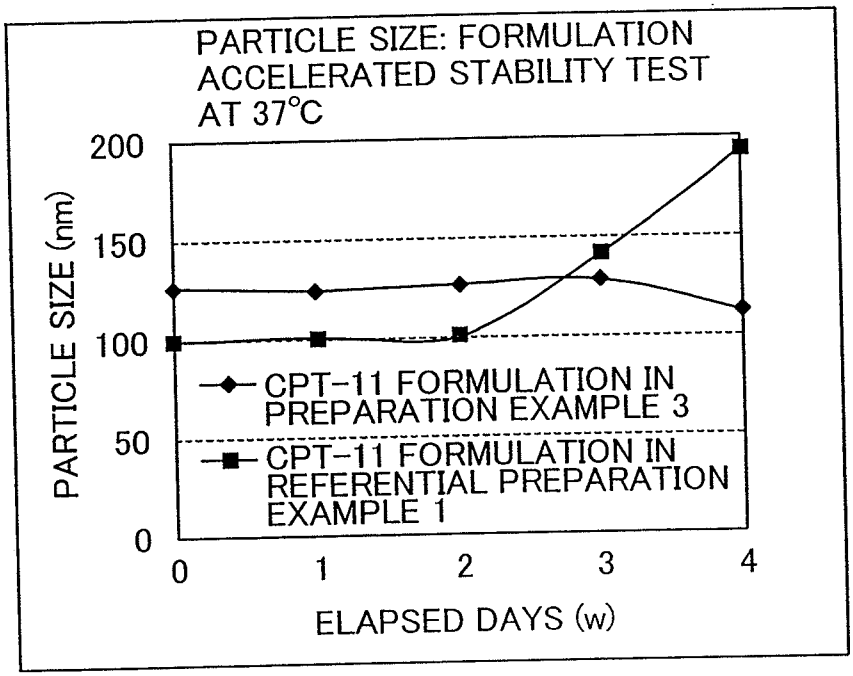

PARTICLE SIZE: FORMULATION ACCELERATED STABILITY TEST AT 37℃
PARTICLE SIZE (nm)
200
150
100
50
0
0
1
2
3
4
ELAPSED DAYS (w)
CPT-11 FORMULATION IN PREPARATION EXAMPLE 3
CPT-11 FORMULATION IN REFERENTIAL PREPARATION EXAMPLE 1

FIG. 4

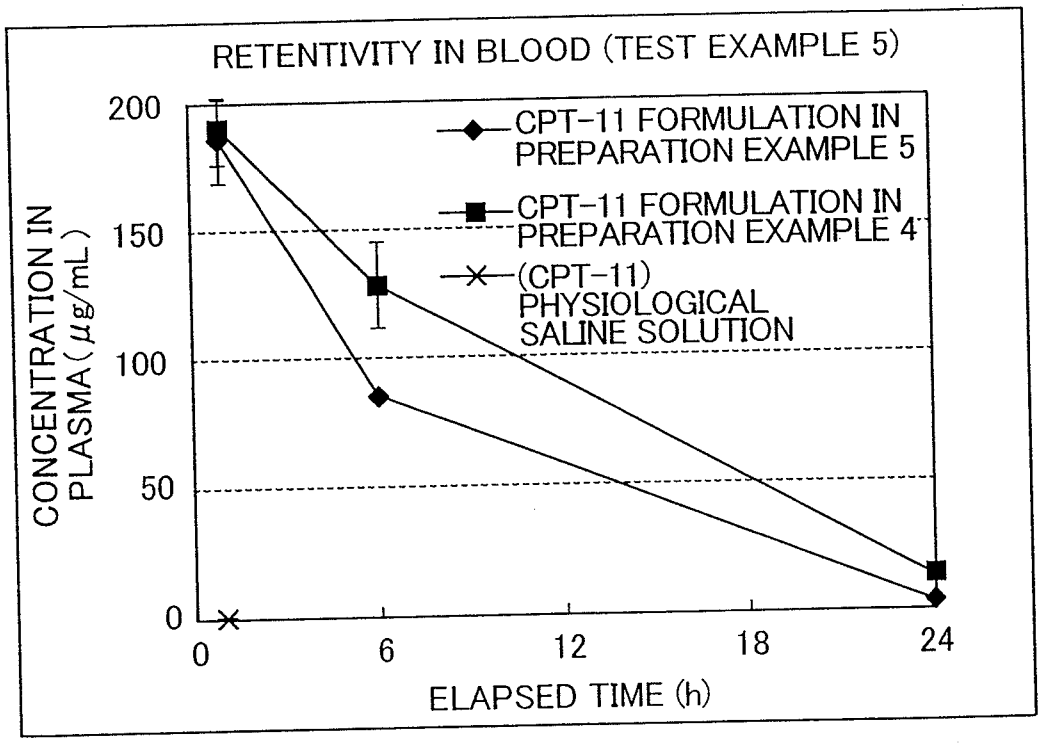

RETENTIVITY IN BLOOD (TEST EXAMPLE 5)
CONCENTRATION IN PLASMA(μg/mL)
200
150
100
50
0
0
6
12
18
24
ELAPSED TIME (h)
CPT-11 FORMULATION IN PREPARATION EXAMPLE 5
CPT-11 FORMULATION IN PREPARATION EXAMPLE 4
(CPT-11) PHYSIOLOGICAL SALINE SOLUTION

FIG. 5

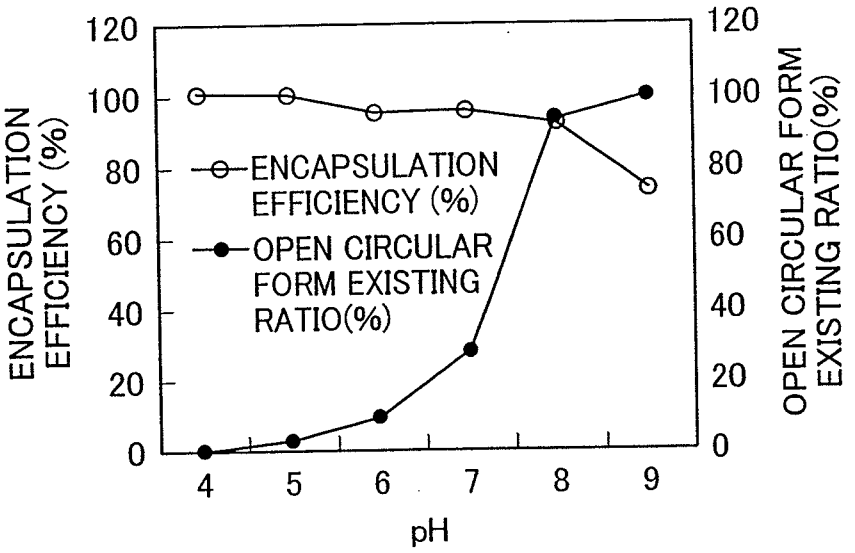

120
100
80
60
40
20
0
ENCAPSULATION EFFICIENCY (%)
ENCAPSULATION EFFICIENCY (%)
OPEN CIRCULAR FORM EXISTING RATIO(%)
4
5
6
7
8
9
pH
OPEN CIRCULAR FORM EXISTING RATIO(%)

FIG. 6

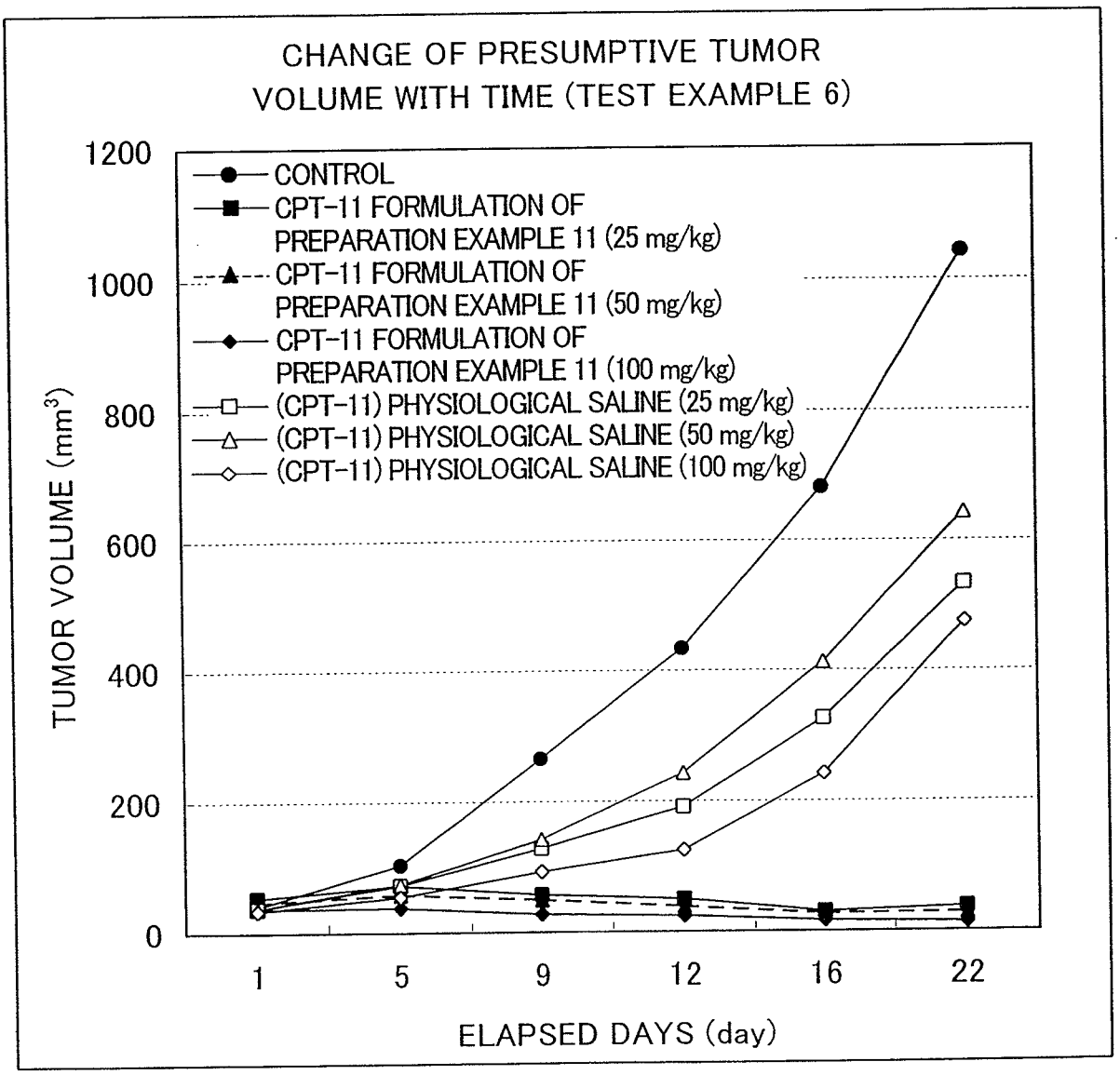

CHANGE OF PRESUMPTIVE TUMOR VOLUME WITH TIME (TEST EXAMPLE 6)
TUMOR VOLUME (mm3)
1200
1000
800
600
400
200
0
1
5
9
12
16
22
ELAPSED DAYS (day)
CONTROL
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)

FIG. 7

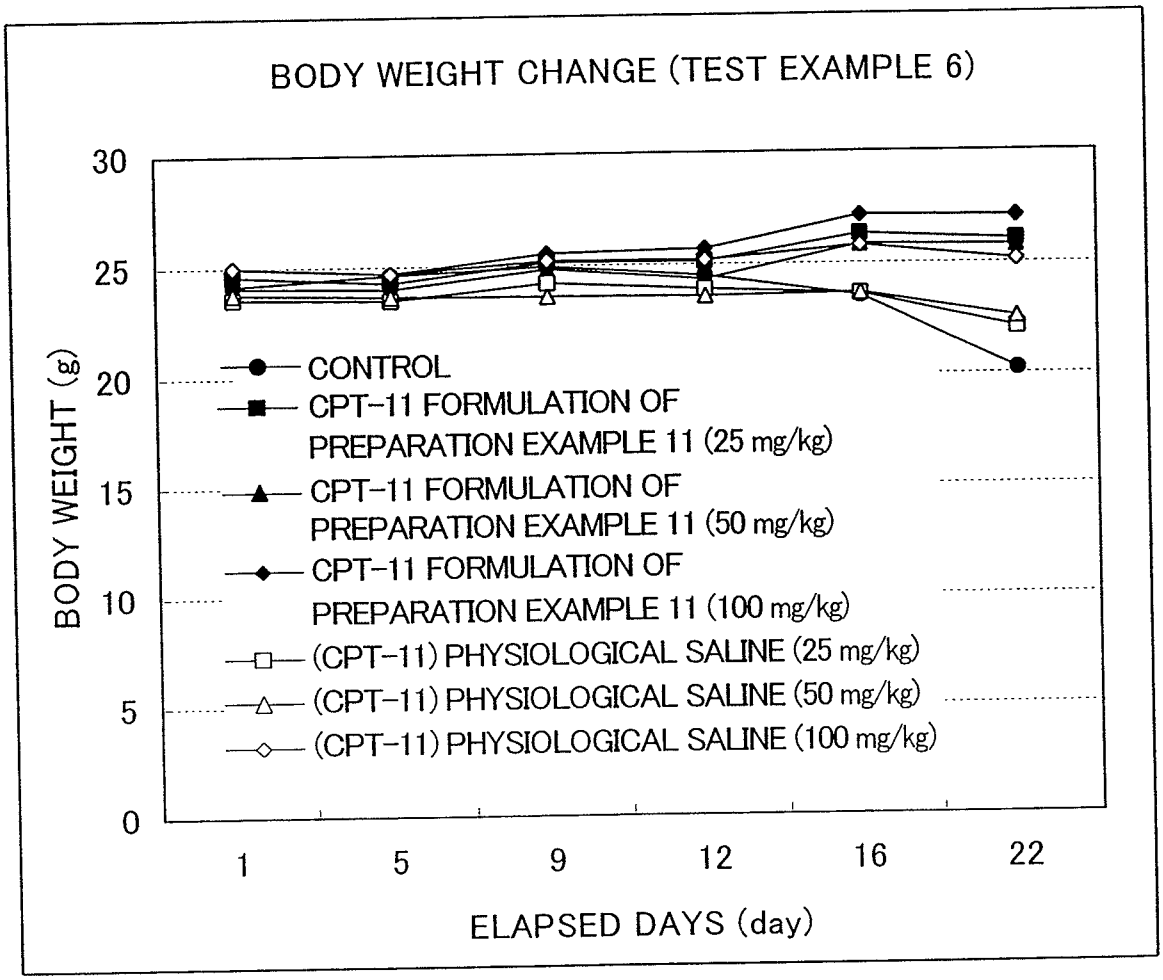

BODY WEIGHT CHANGE (TEST EXAMPLE 6)
BODY WEIGHT (g)
30
25
20
15
10
5
0
1
5
9
12
16
22
ELAPSED DAYS (day)
CONTROL
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)

FIG. 8

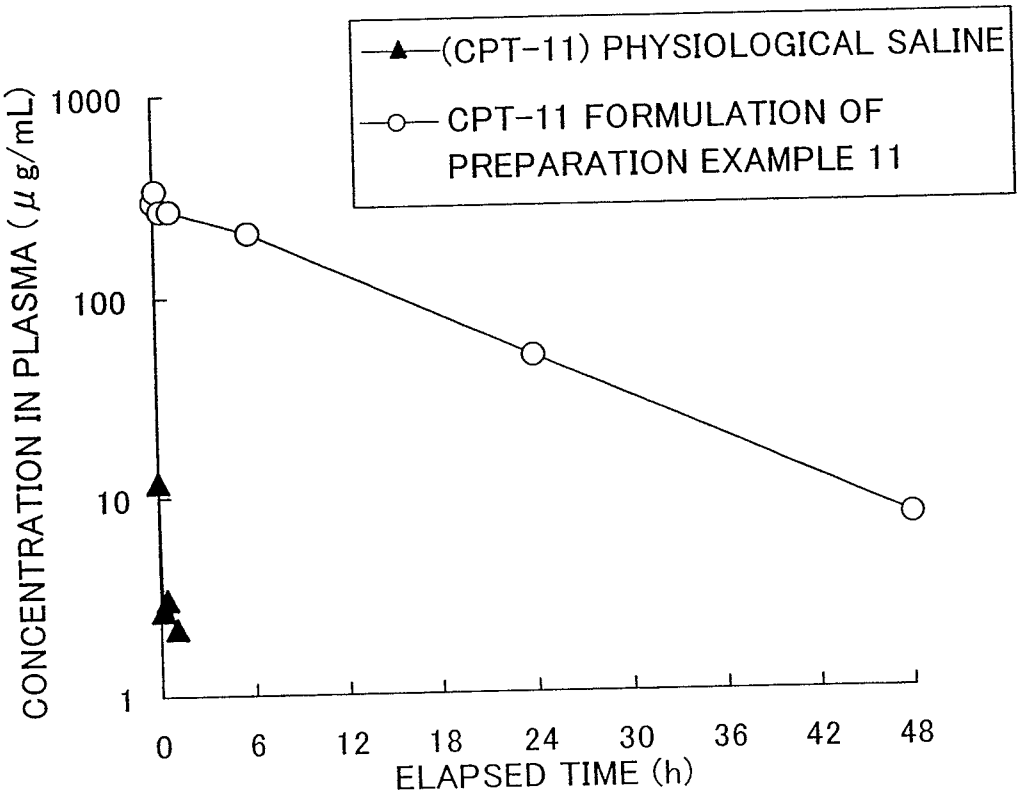

(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11
CONCENTRATION IN PLASMA (μg/mL)
1000
100
10
1
0
6
12
18
24
30
36
42
48
ELAPSED TIME (h)

FIG. 9

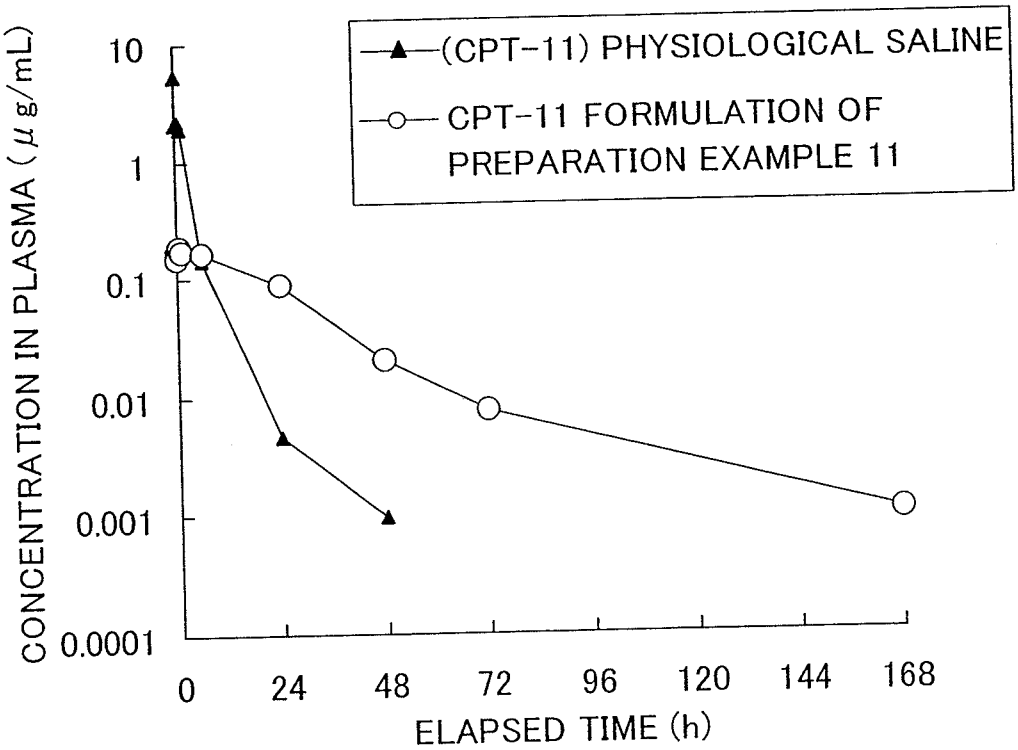

(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11
CONCENTRATION IN PLASMA (μg/mL)
10
1
0.1
0.01
0.001
0.0001
0
24
48
72
96
120
144
168
ELAPSED TIME (h)

FIG. 10

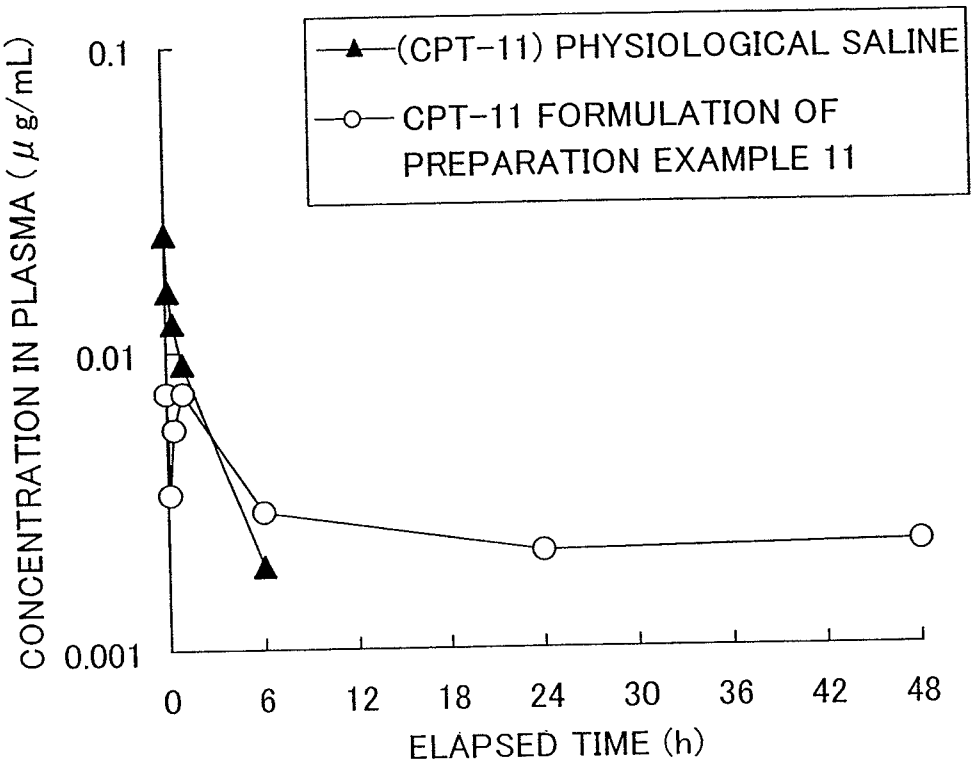

(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11
CONCENTRATION IN PLASMA (μg/mL)
0.1
0.01
0.001
0
6
12
18
24
30
36
42
48
ELAPSED TIME (h)

FIG. 11

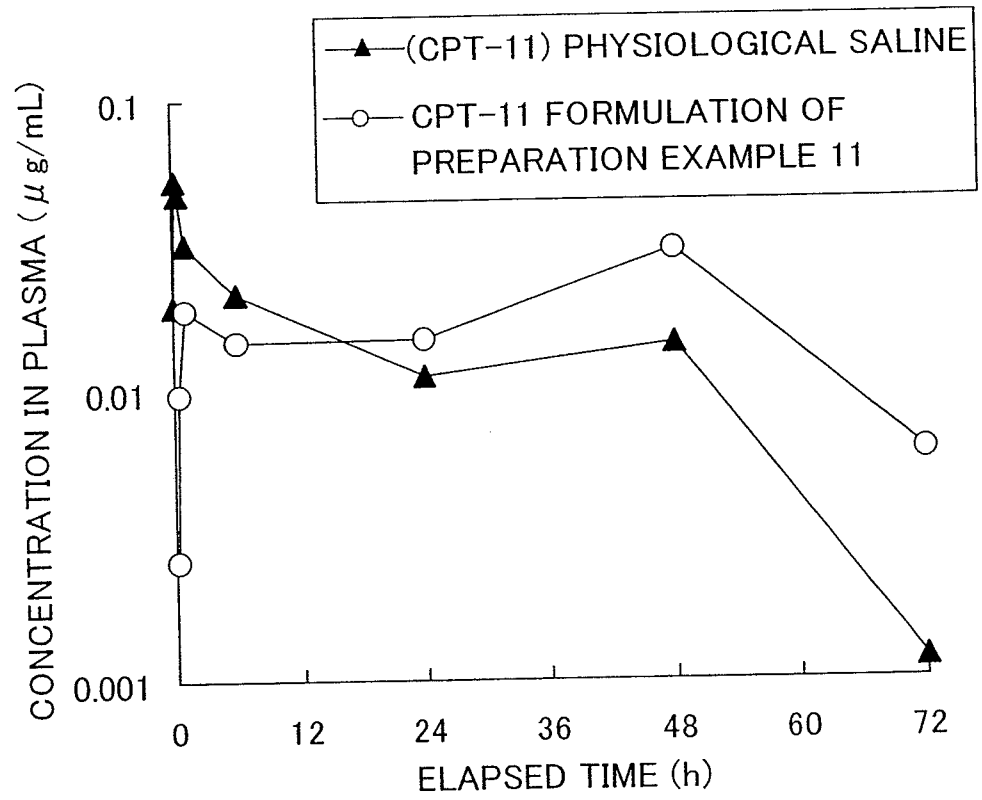

(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11
CONCENTRATION IN PLASMA (μg/mL)
0.1
0.01
0.001
0
12
24
36
48
60
72
ELAPSED TIME (h)

FIG. 12

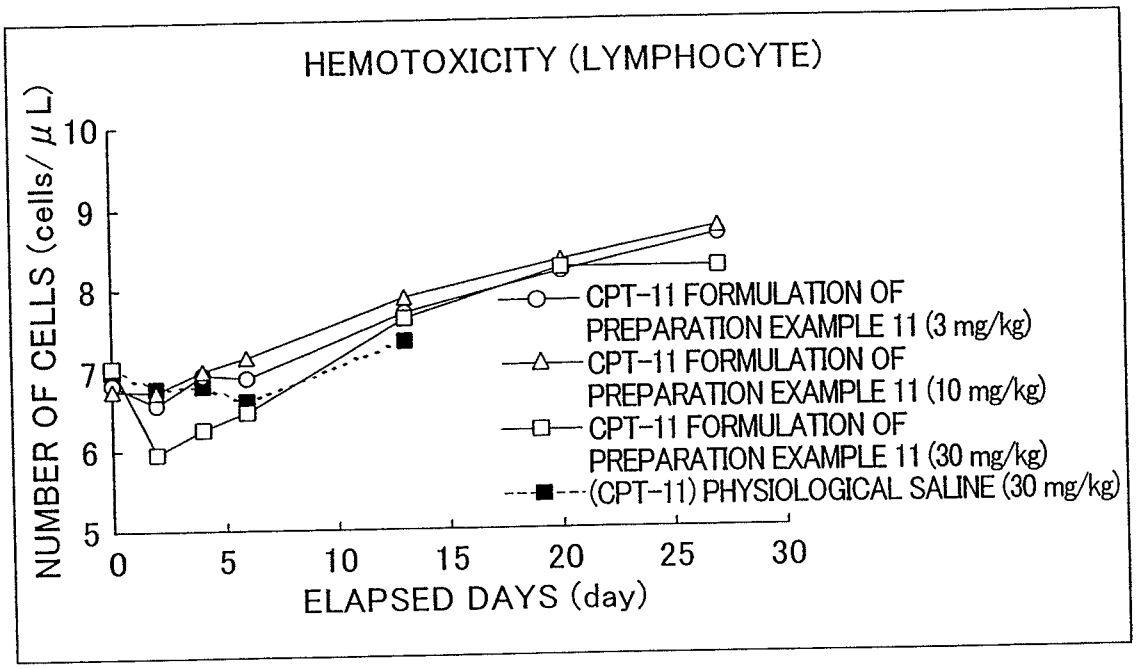

HEMOTOXICITY (LYMPHOCYTE)
NUMBER OF CELLS (cells/μL)
10
9
8
7
6
5
0
5
10
15
20
25
30
ELAPSED DAYS (day)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)

FIG. 13

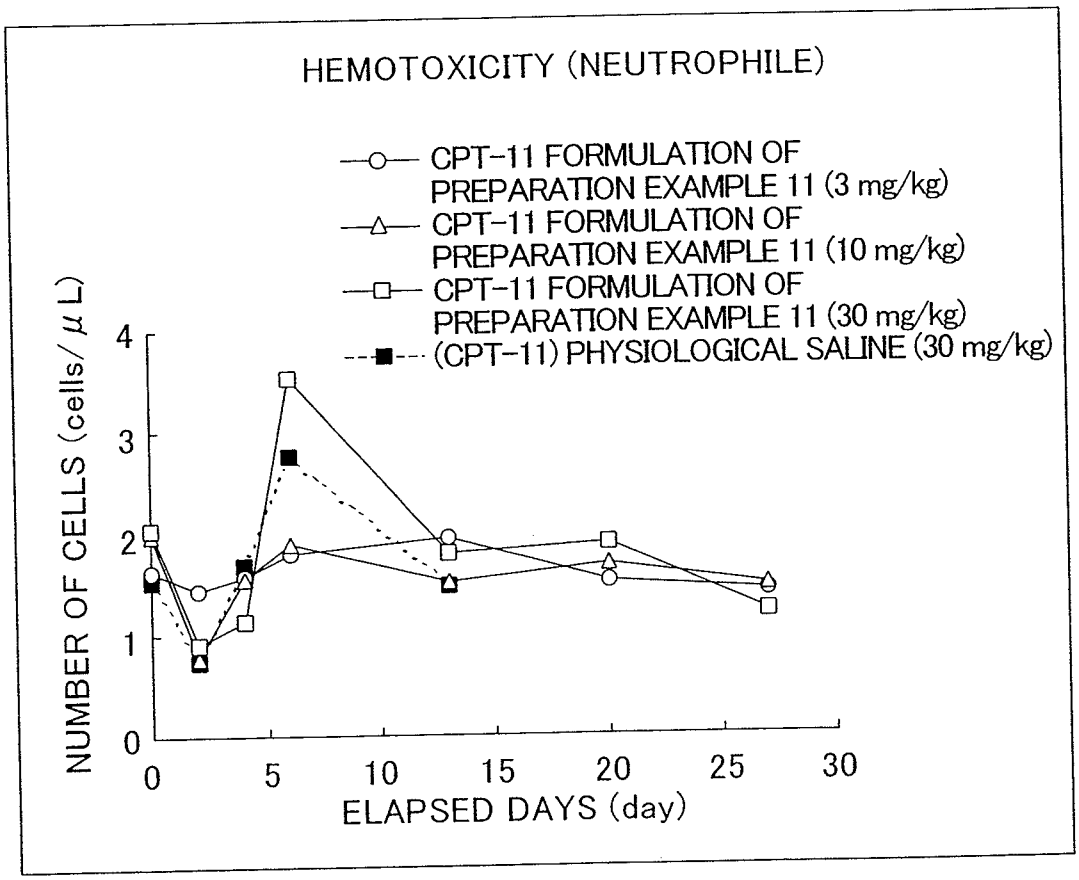

HEMOTOXICITY (NEUTROPHILE)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 11 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)
NUMBER OF CELLS (cells/μL)
4
3
2
1
0
0
5
10
15
20
25
30
ELAPSED DAYS (day)

FIG. 14

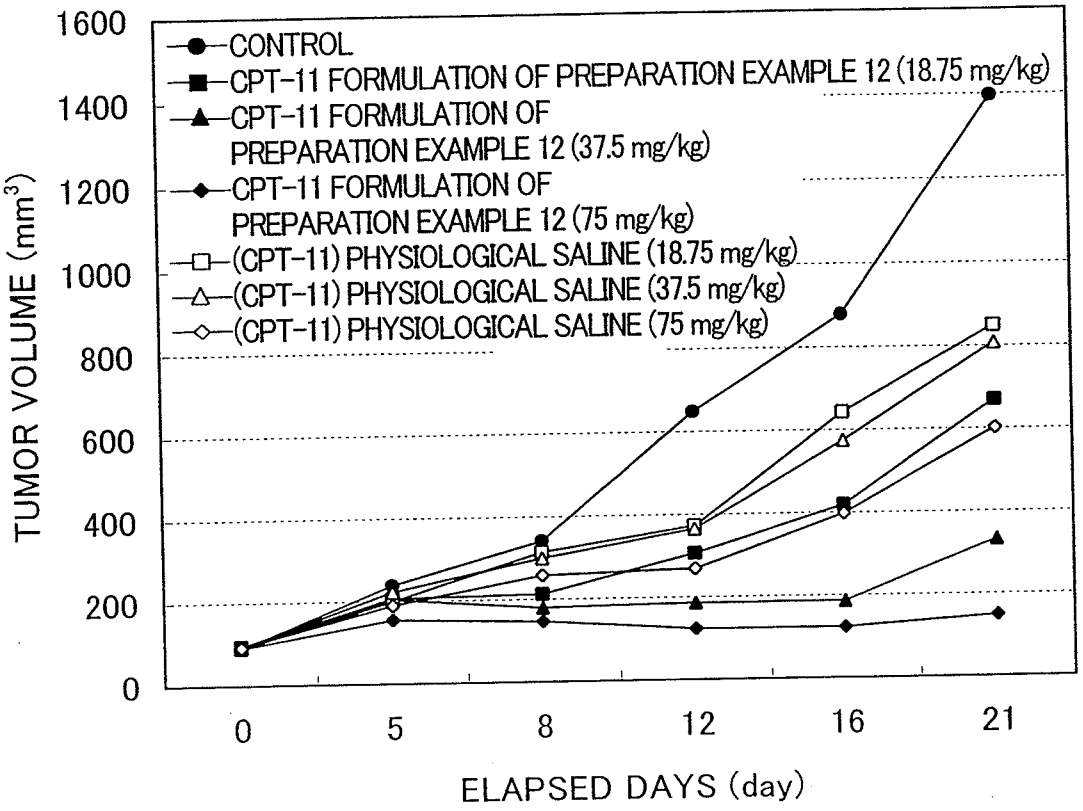

1600
1400
1200
1000
800
600
400
200
0
TUMOR VOLUME (mm3)
0
5
8
12
16
21
ELAPSED DAYS (day)
CONTROL
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (18.75 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (37.5 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (75 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (18.75 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (37.5 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (75 mg/kg)

FIG. 15

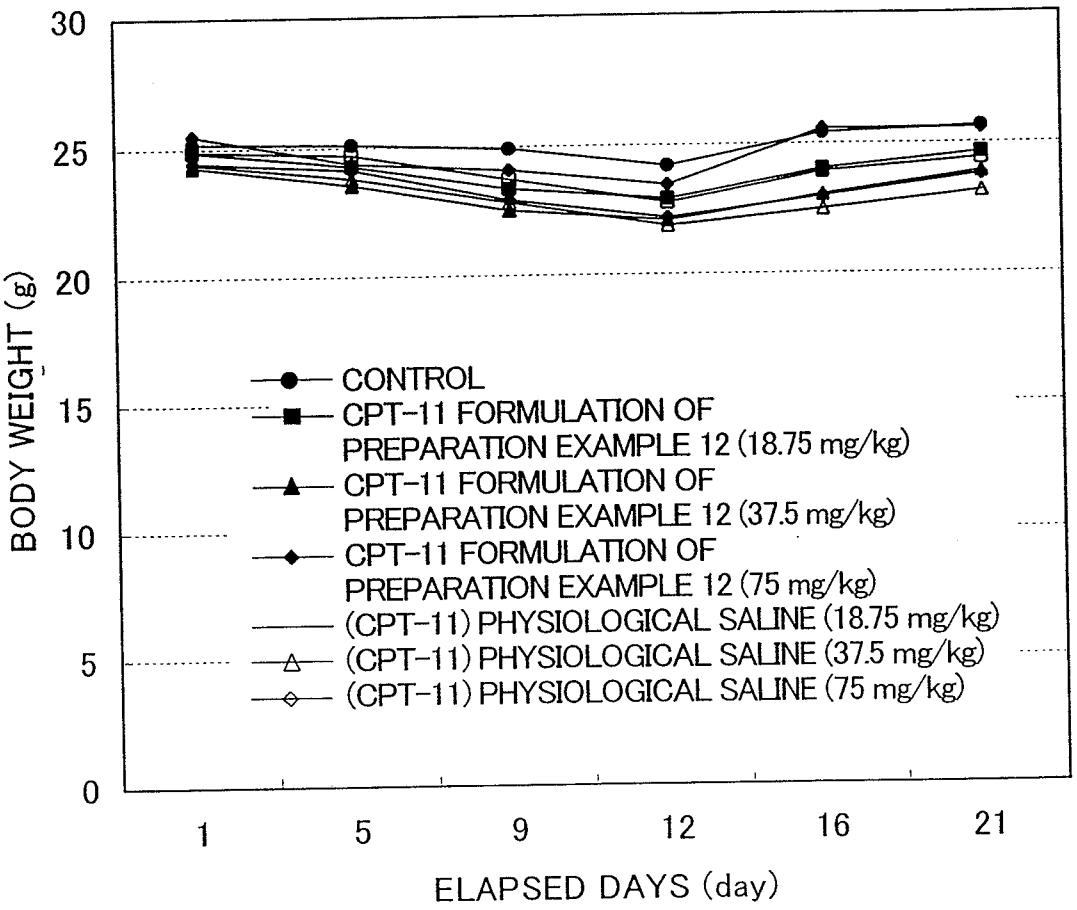

30
25
20
15
10
5
0
BODY WEIGHT (g)
1
5
9
12
16
21
ELAPSED DAYS (day)
CONTROL
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (18.75 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (37.5 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (75 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (18.75 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (37.5 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (75 mg/kg)

FIG.16

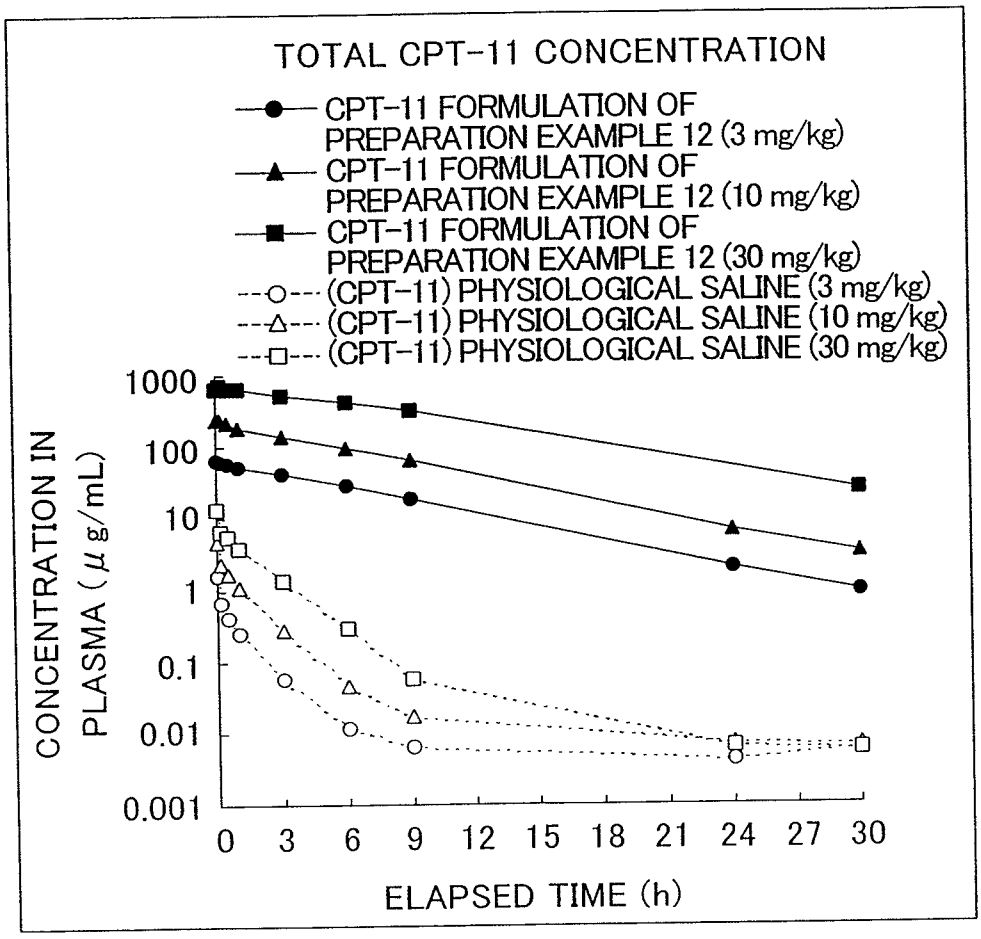

TOTAL CPT-11 CONCENTRATION
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (3 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (10 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)
CONCENTRATION IN PLASMA (μg/mL)
1000
100
10
1
0.1
0.01
0.001
0 3 6 9 12 15 18 21 24 27 30
ELAPSED TIME (h)

FIG. 17

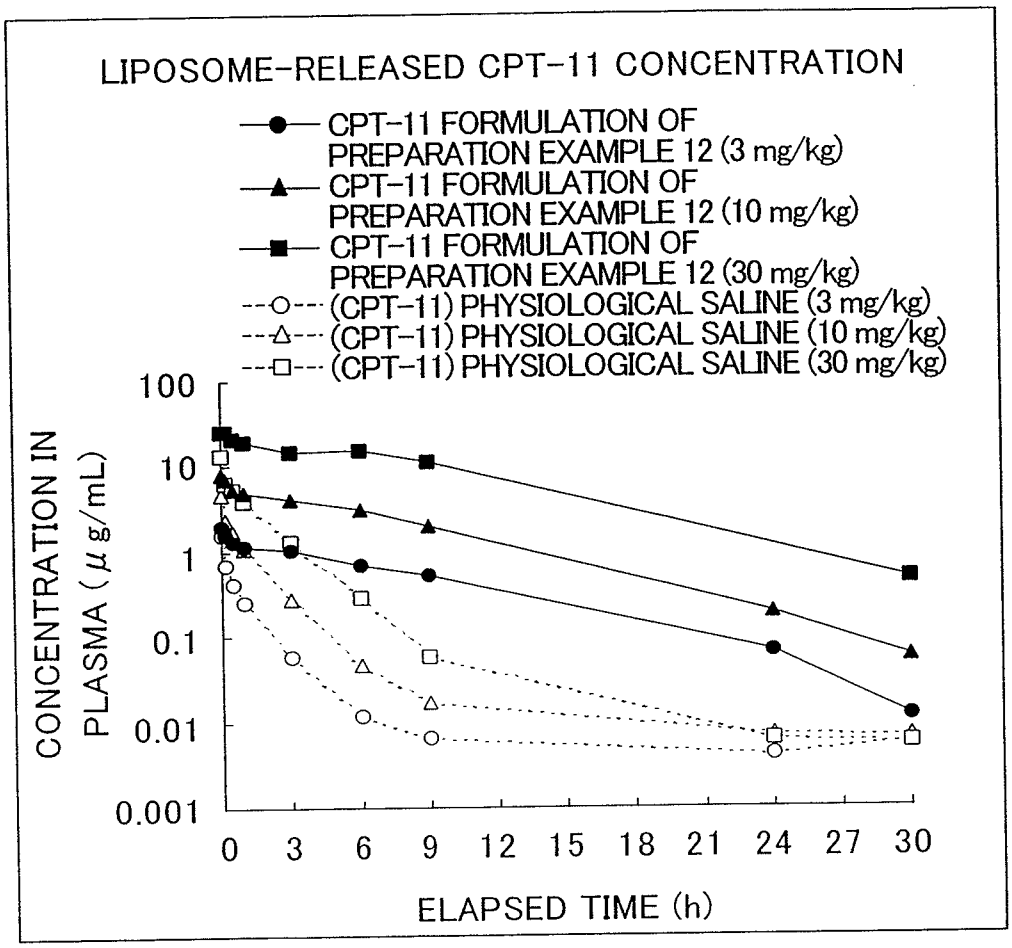

LIPOSOME-RELEASED CPT-11 CONCENTRATION
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (3 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (10 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)
CONCENTRATION IN PLASMA (μg/mL)
100
10
1
0.1
0.01
0.001
0
3
6
9
12
15
18
21
24
27
30
ELAPSED TIME (h)

FIG. 18

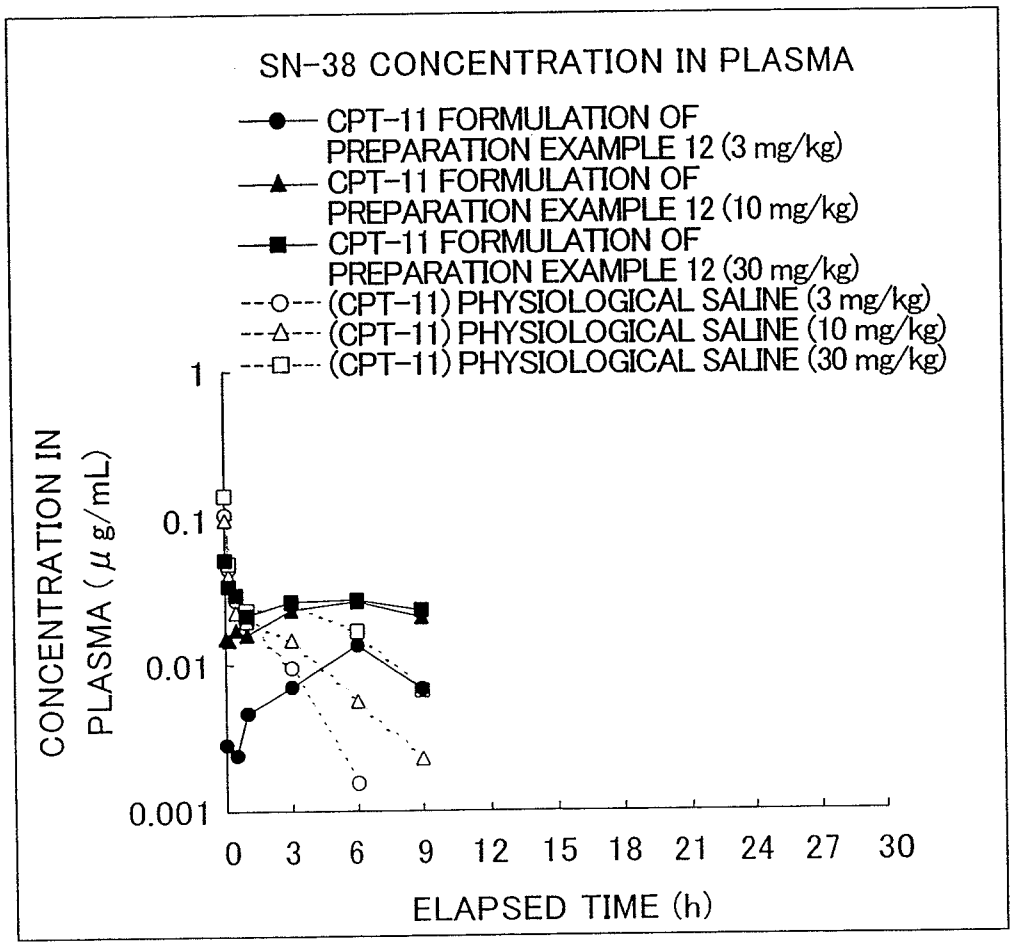

SN-38 CONCENTRATION IN PLASMA
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (3 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (10 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)
CONCENTRATION IN PLASMA (μg/mL)
1
0.1
0.01
0.001
0 3 6 9 12 15 18 21 24 27 30
ELAPSED TIME (h)

FIG. 19

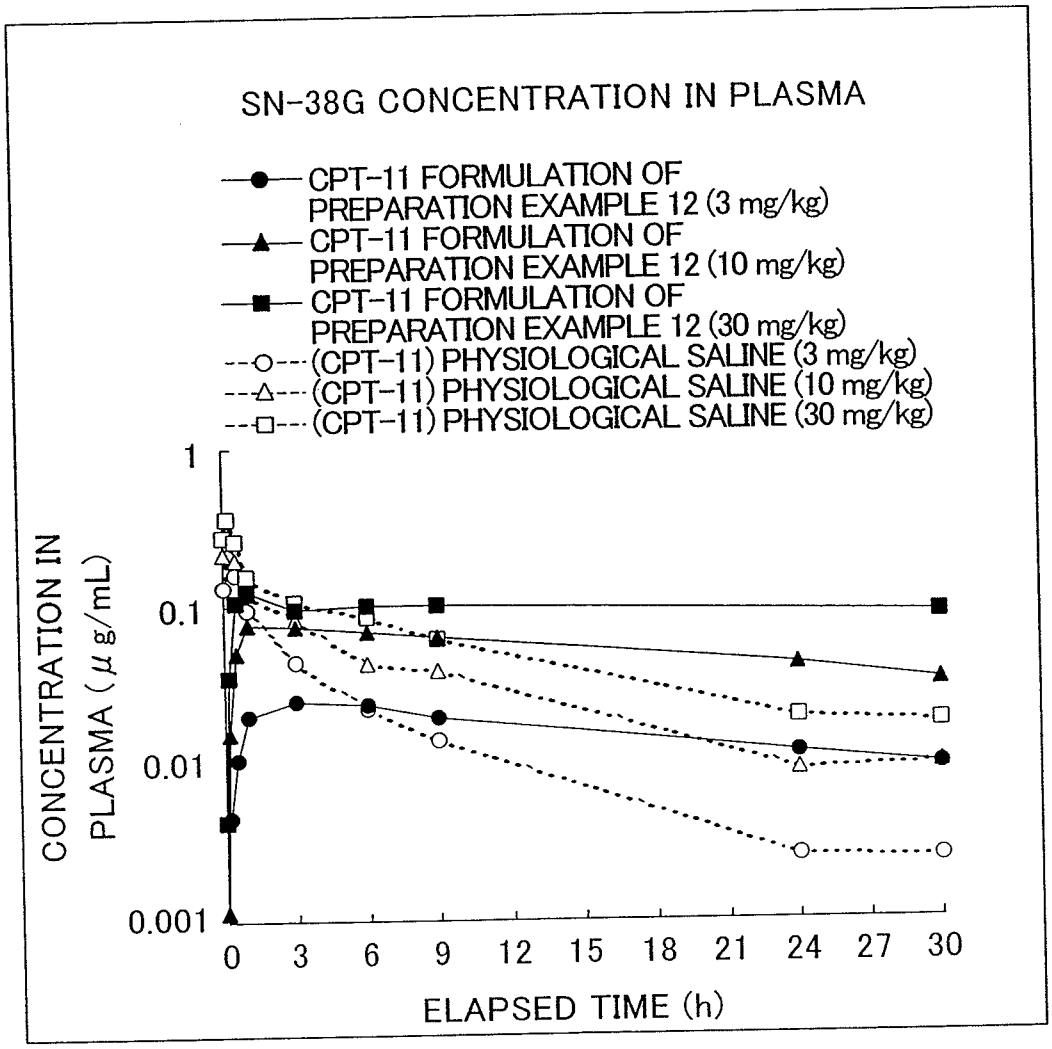

SN-38G CONCENTRATION IN PLASMA
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (3 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (10 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 12 (30 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (3 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (10 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (30 mg/kg)
CONCENTRATION IN PLASMA (μg/mL)
1
0.1
0.01
0.001
0
3
6
9
12
15
18
21
24
27
30
ELAPSED TIME (h)

FIG. 20

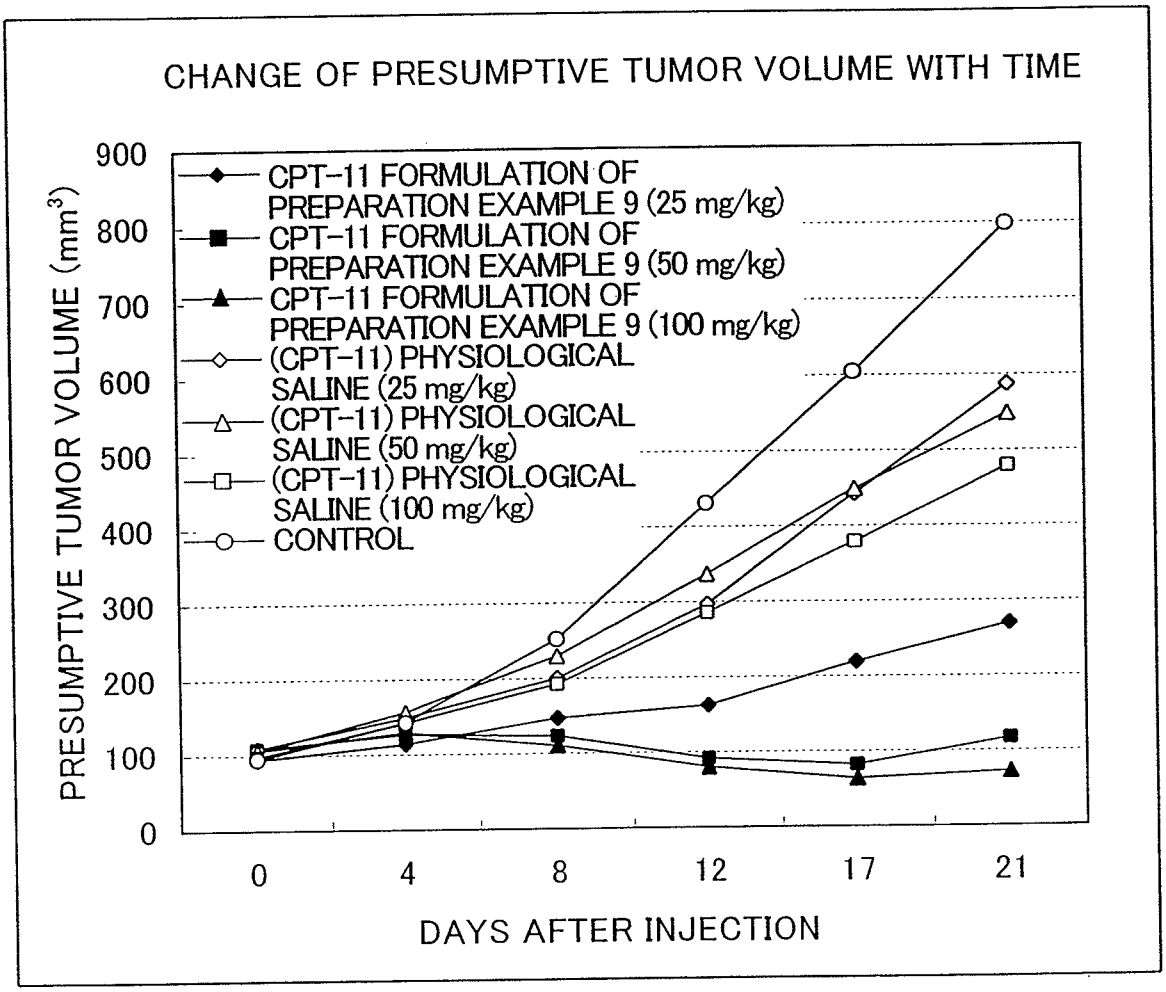

CHANGE OF PRESUMPTIVE TUMOR VOLUME WITH TIME
PRESUMPTIVE TUMOR VOLUME (mm3)
900
800
700
600
500
400
300
200
100
0
0
4
8
12
17
21
DAYS AFTER INJECTION
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)
CONTROL

FIG. 21

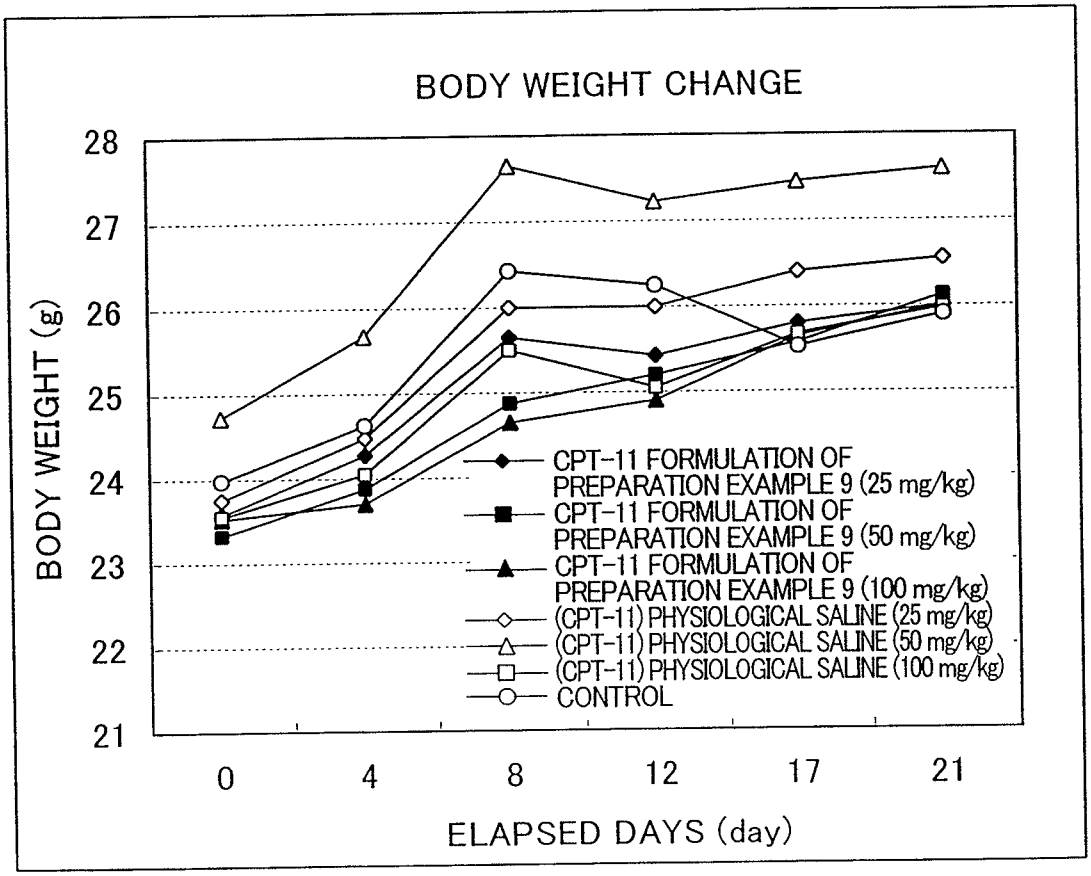

BODY WEIGHT CHANGE
BODY WEIGHT (g)
28
27
26
25
24
23
22
21
0
4
8
12
17
21
ELAPSED DAYS (day)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)
CONTROL

FIG. 22

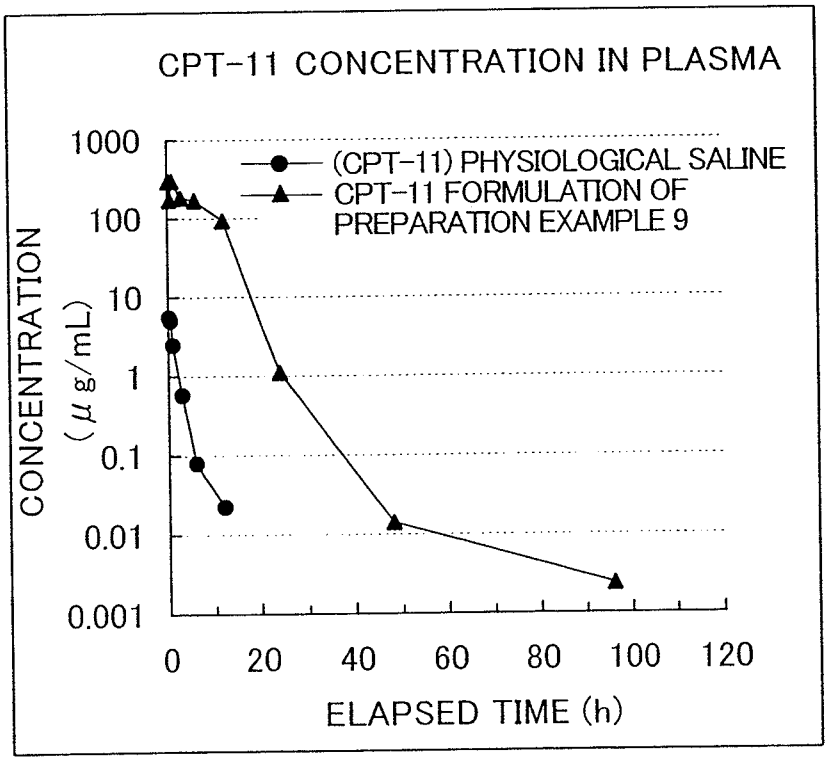

CPT-11 CONCENTRATION IN PLASMA
CONCENTRATION (μg/mL)
1000
100
10
1
0.1
0.01
0.001
0
20
40
60
80
100
120
ELAPSED TIME (h)
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9

FIG. 23

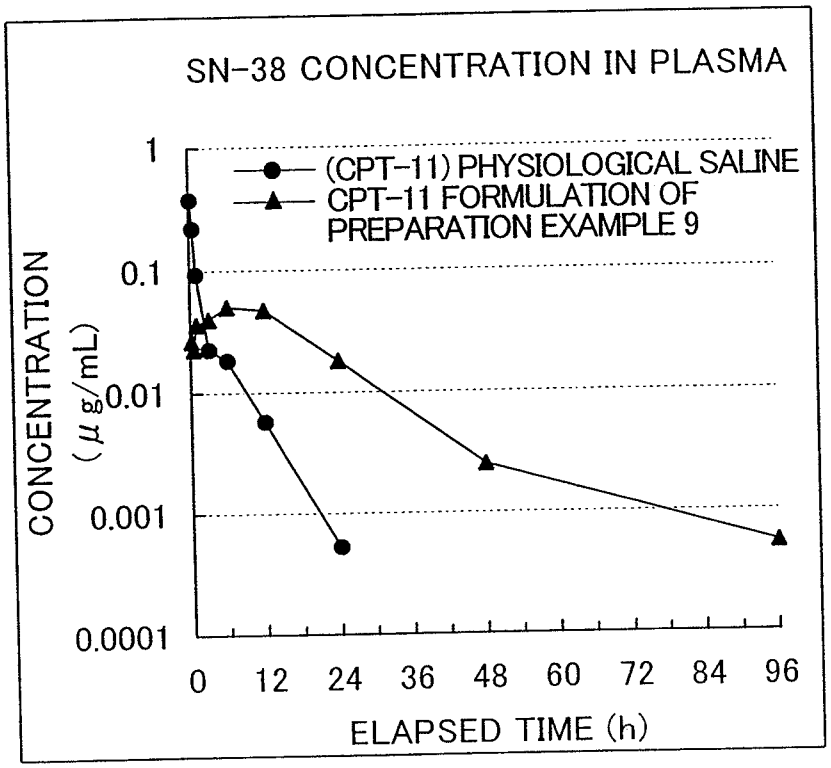

SN-38 CONCENTRATION IN PLASMA
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9
CONCENTRATION (μg/mL)
1
0.1
0.01
0.001
0.0001
0
12
24
36
48
60
72
84
96
ELAPSED TIME (h)

FIG. 24

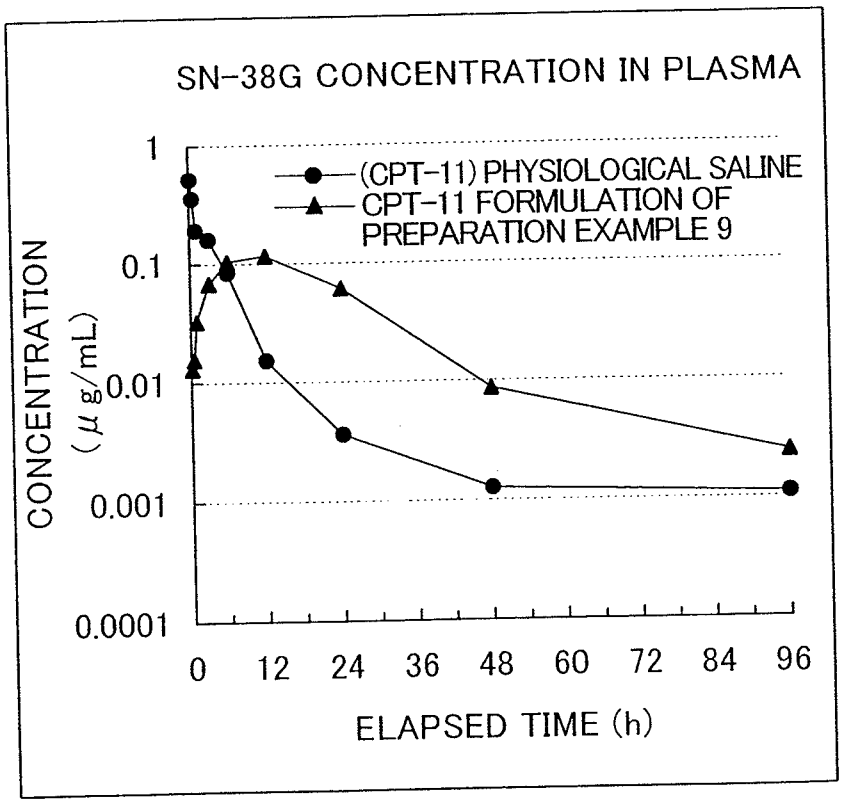

SN-38G CONCENTRATION IN PLASMA
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9
CONCENTRATION (μg/mL)
1
0.1
0.01
0.001
0.0001
0
12
24
36
48
60
72
84
96
ELAPSED TIME (h)

FIG. 25

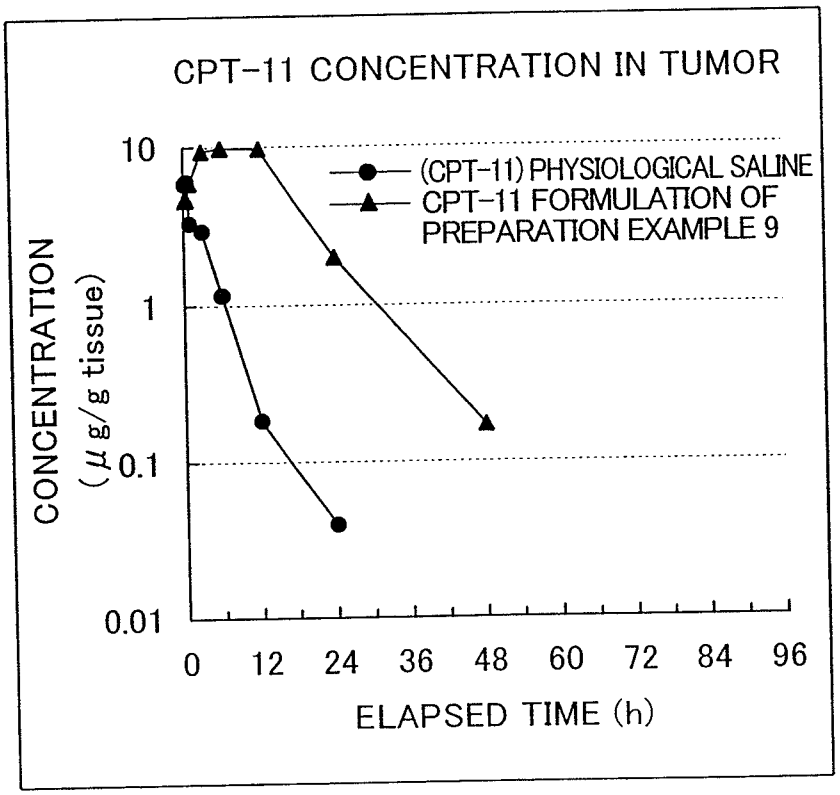

CPT-11 CONCENTRATION IN TUMOR
CONCENTRATION (μg/g tissue)
10
1
0.1
0.01
0
12
24
36
48
60
72
84
96
ELAPSED TIME (h)
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9

FIG. 26

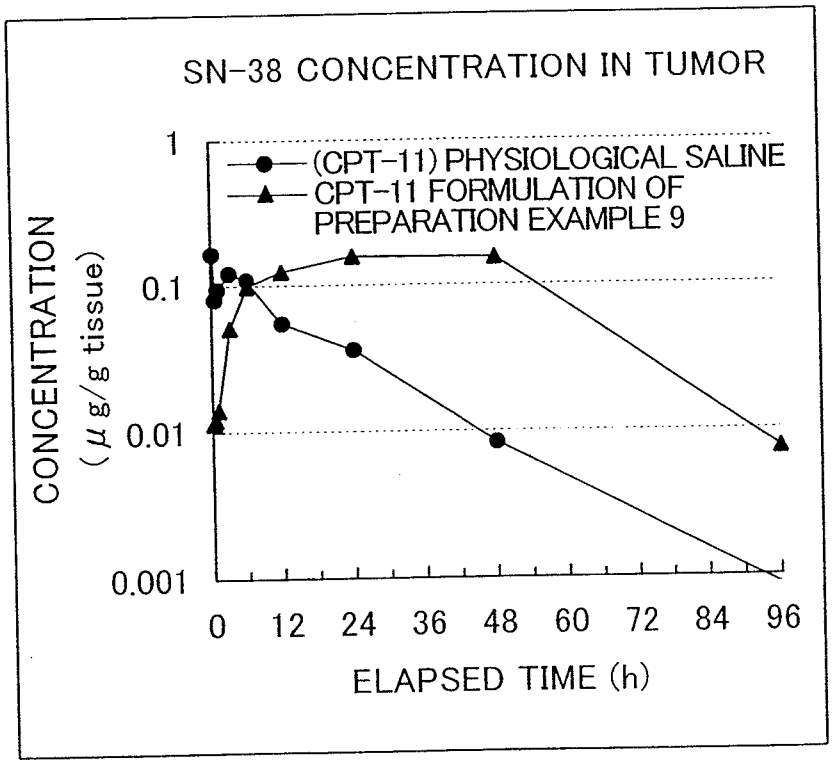

SN-38 CONCENTRATION IN TUMOR
CONCENTRATION (μg/g tissue)
1
0.1
0.01
0.001
0
12
24
36
48
60
72
84
96
ELAPSED TIME (h)
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF PREPARATION EXAMPLE 9

FIG.27

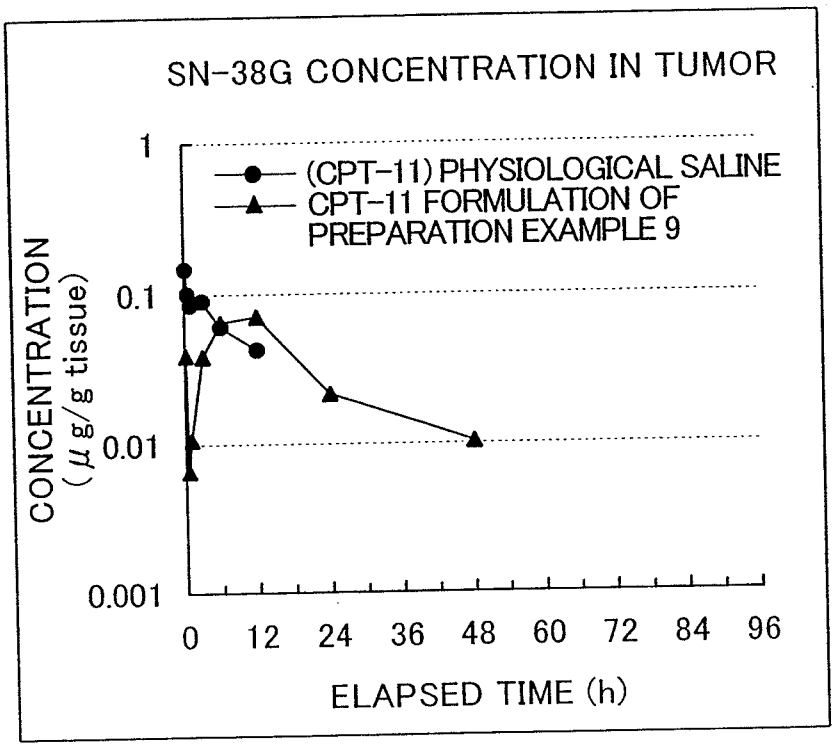

SN-38G CONCENTRATION IN TUMOR
1
0.1
0.01
0.001
CONCENTRATION
(μg/g tissue)
(CPT-11) PHYSIOLOGICAL SALINE
CPT-11 FORMULATION OF
PREPARATION EXAMPLE 9
0
12
24
36
48
60
72
84
96
ELAPSED TIME (h)

FIG. 28

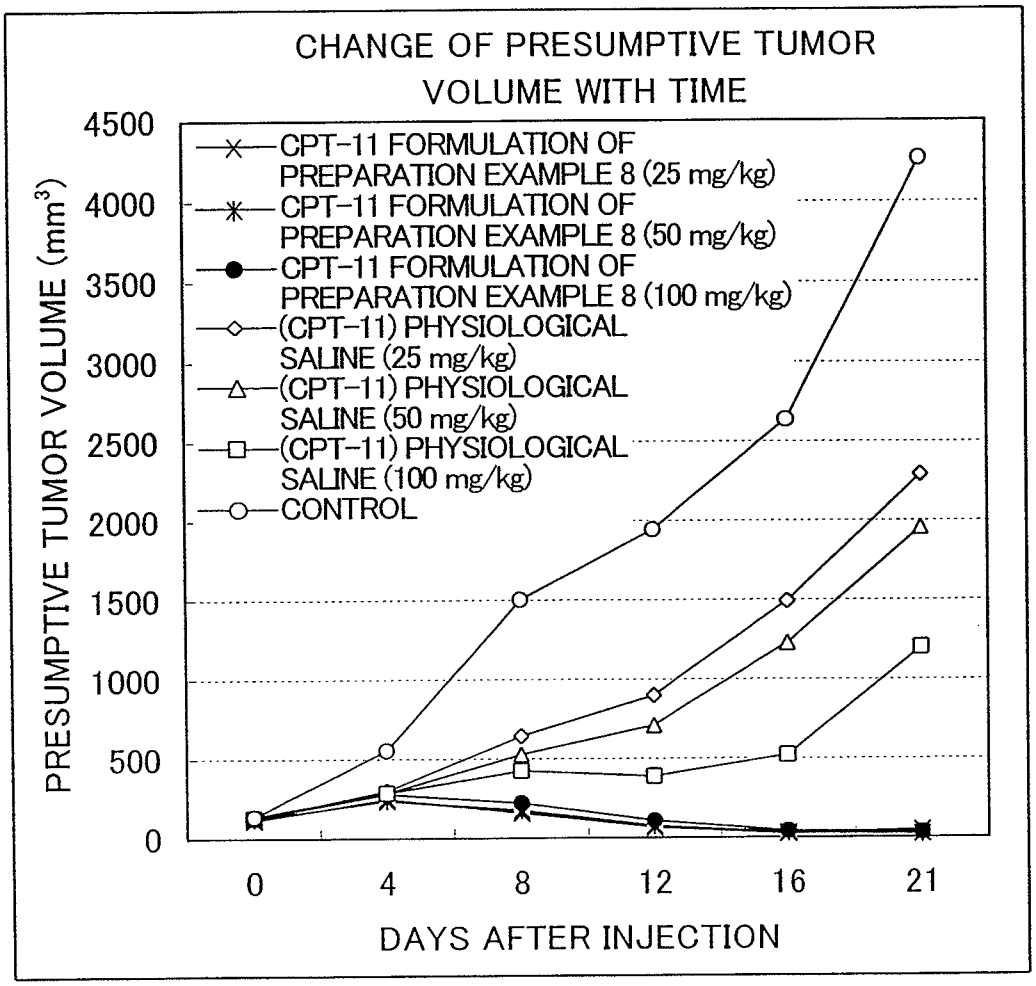
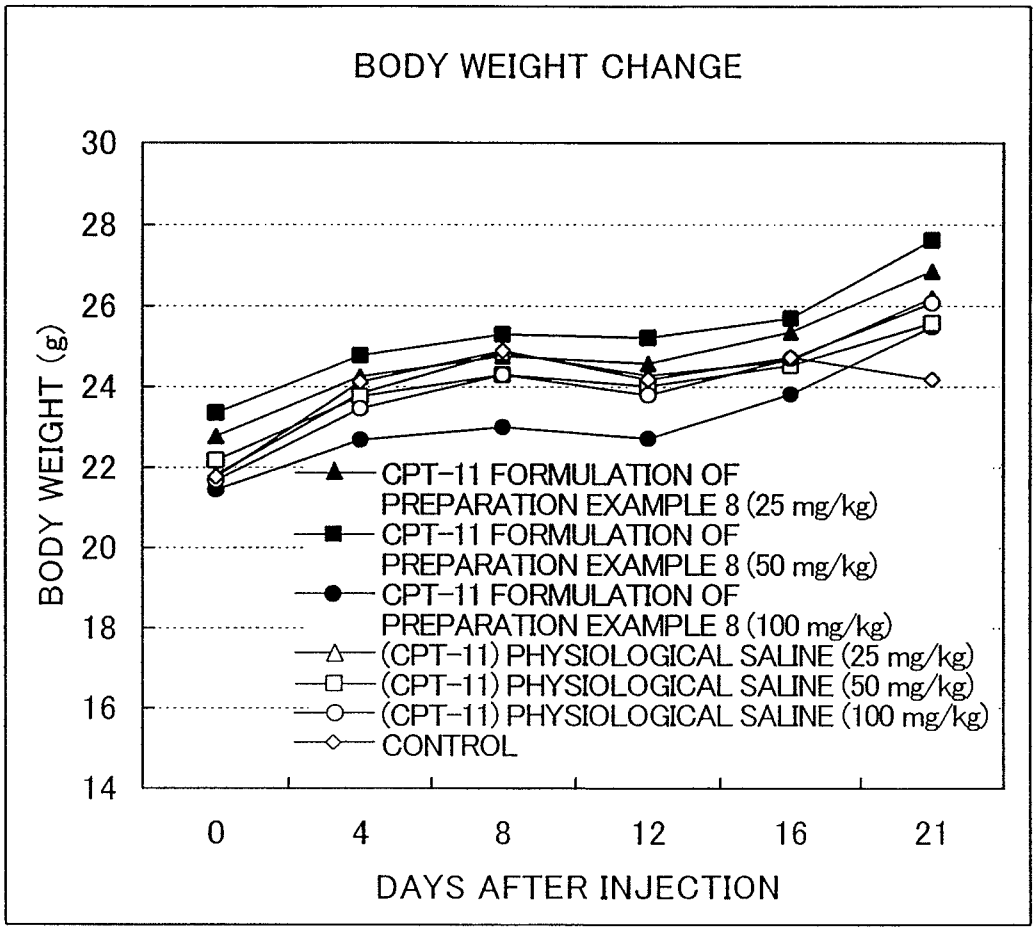

CHANGE OF PRESUMPTIVE TUMOR VOLUME WITH TIME
PRESUMPTIVE TUMOR VOLUME (mm3)
4500
4000
3500
3000
2500
2000
1500
1000
500
0
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)
CONTROL
0
4
8
12
16
21
DAYS AFTER INJECTION

FIG. 29


BODY WEIGHT CHANGE
BODY WEIGHT (g)
30
28
26
24
22
20
18
16
14
0
4
8
12
16
21
DAYS AFTER INJECTION
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (25 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (50 mg/kg)
CPT-11 FORMULATION OF PREPARATION EXAMPLE 8 (100 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (25 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (50 mg/kg)
(CPT-11) PHYSIOLOGICAL SALINE (100 mg/kg)
CONTROL