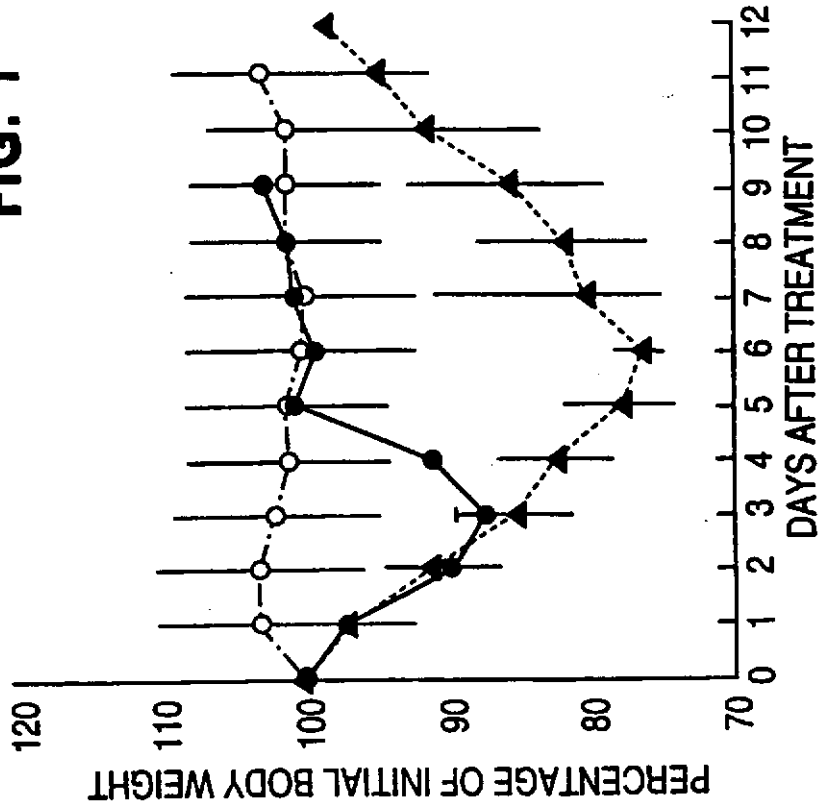
FIG. 1



N = 4	NO TREATMENT (DAY 3)	CPT-11 300 mg/kg (DAY 3)	CPT-11 400 mg/kg (DAY 6)
NO TREATMENT (DAY 3)	——	P < 0.01	P < 0.01
CPT-11 300 mg/kg (DAY 3)	P < 0.01	——	P < 0.05
CPT-11 400 mg/kg (DAY 6)	P < 0.01	P < 0.05	——

FIG. 2

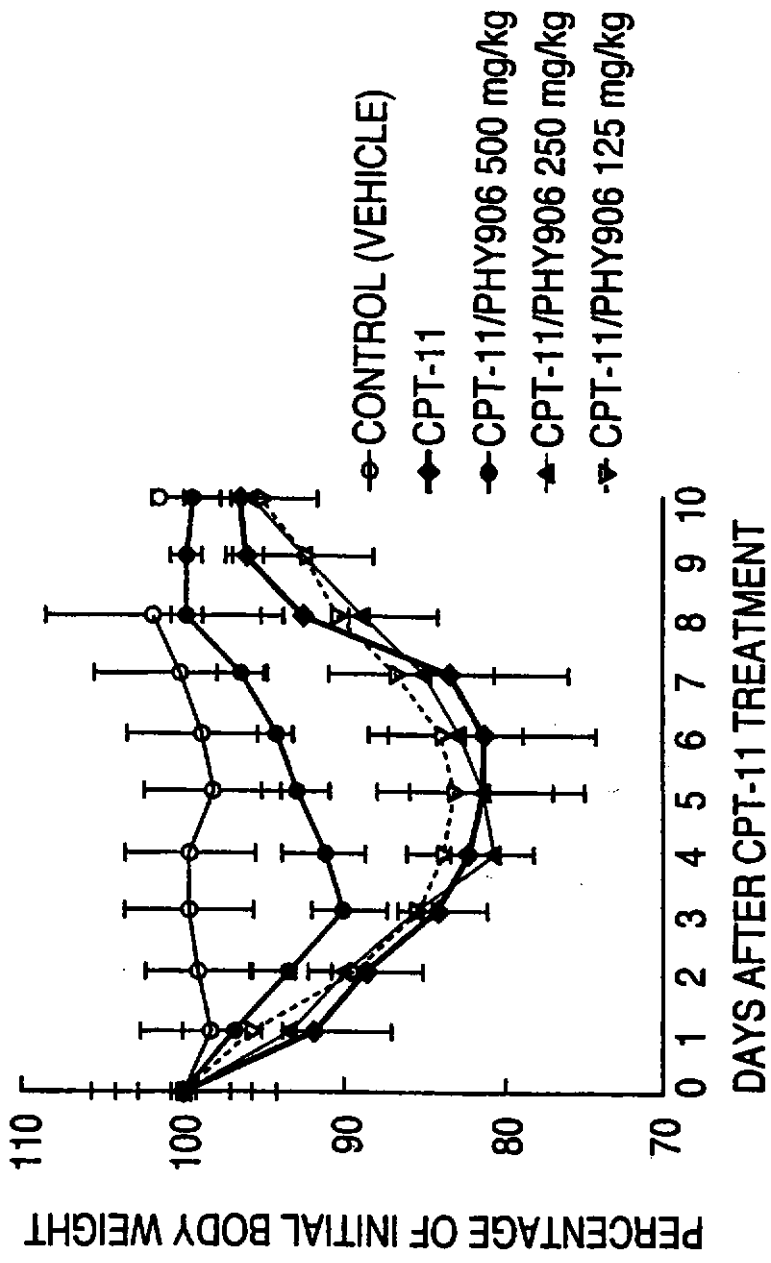
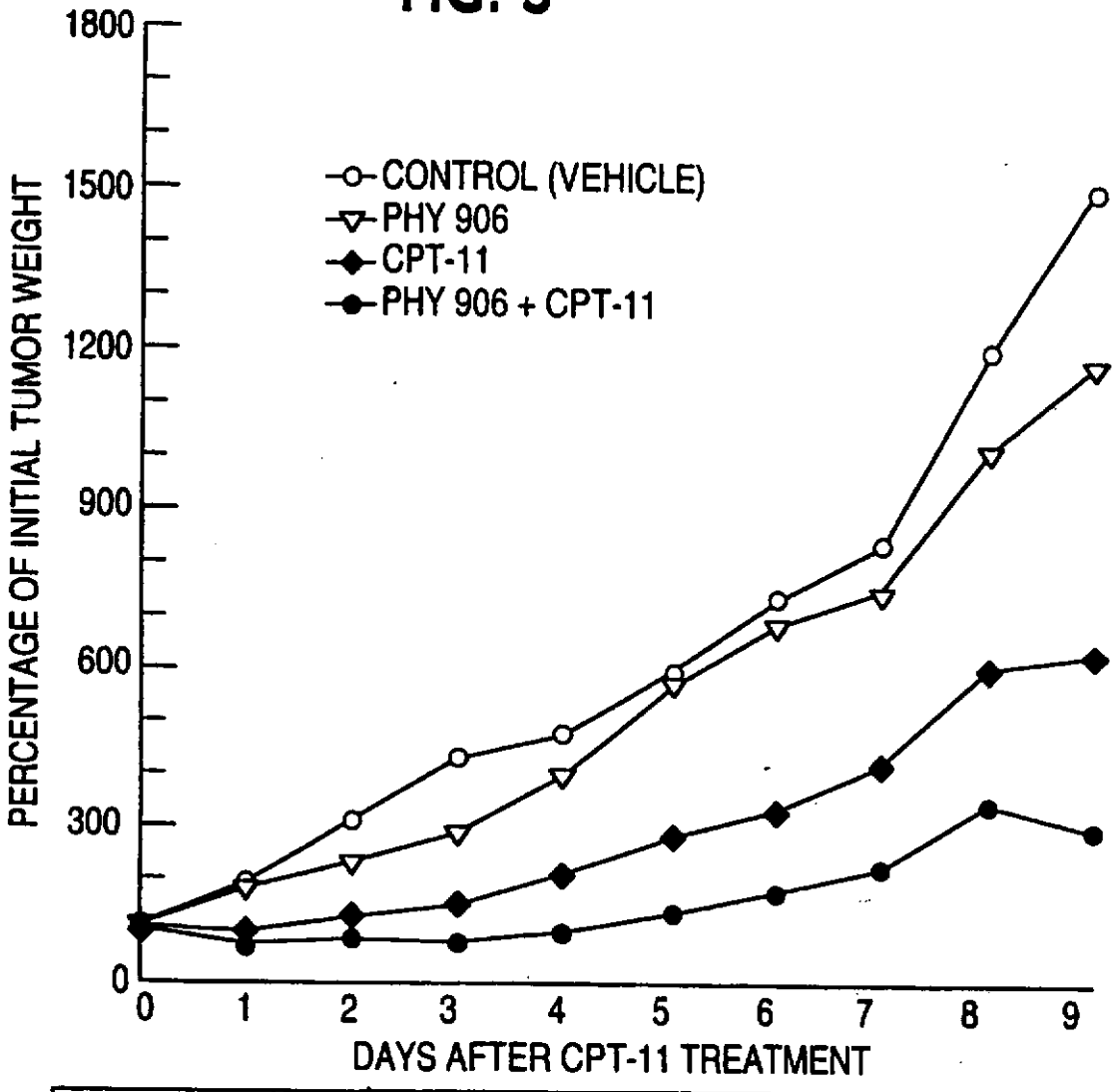

CONTROL (VEHICLE)
CPT-11
CPT-11/PHY906 500 mg/kg
CPT-11/PHY906 250 mg/kg
CPT-11/PHY906 125 mg/kg
110
100
90
80
70
0 1 2 3 4 5 6 7 8 9 10
DAYS AFTER CPT-11 TREATMENT
PERCENTAGE OF INITIAL BODY WEIGHT

FIG. 3



t-TEST P-VALUE AT DAY 6	CONTROL (VEHICLE)	PHY 906	CPT-11	CPT-11 WITH PHY 906
PHY 906	0.5950	——	0.0027	0.0001
CPT-11	0.0004	0.0027	——	0.0007
CPT-11 WITH PHY 906	0.0001	0.0001	0.0007	——

FIG. 4

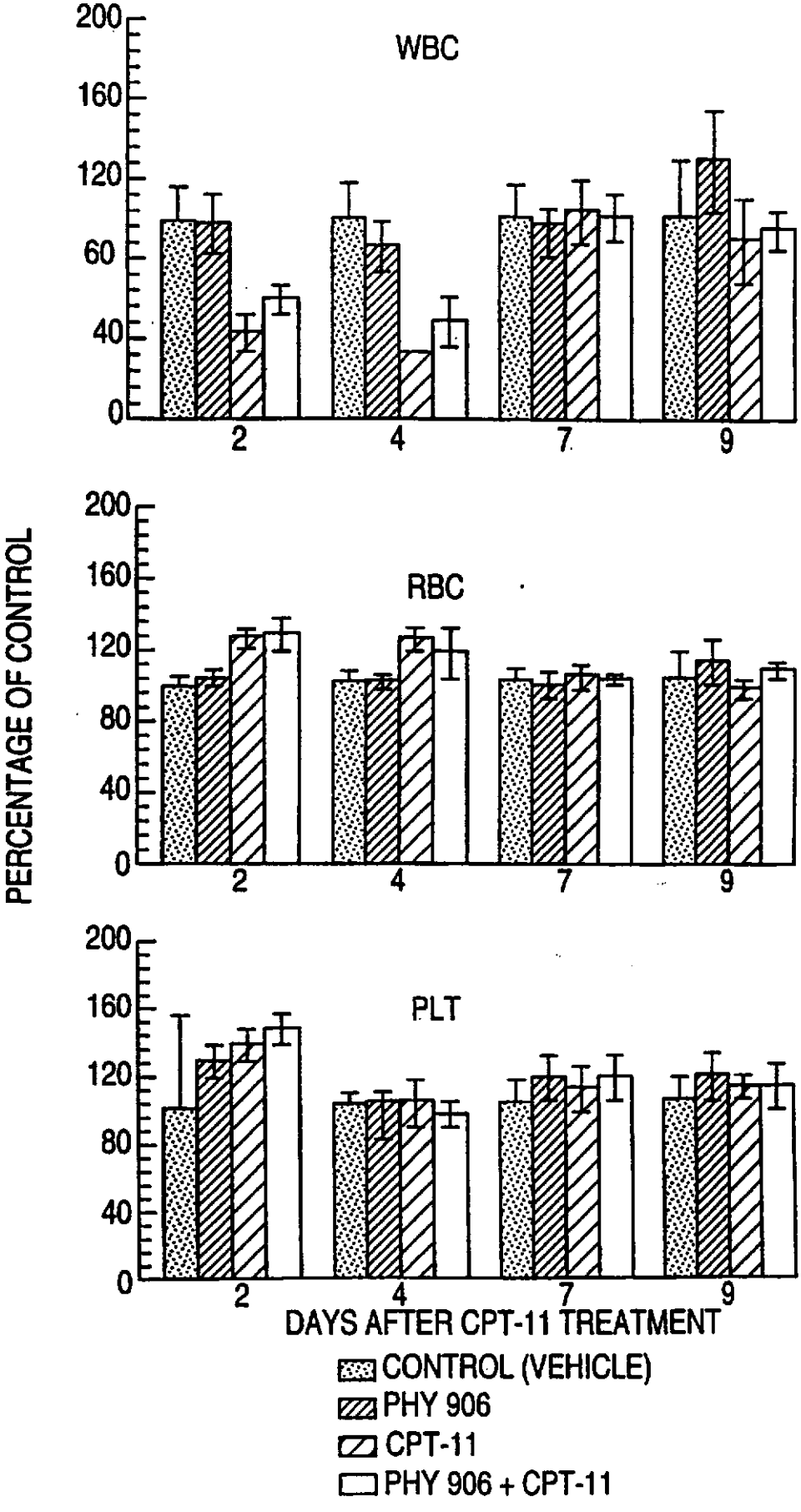

WBC
RBC
PLT
200
160
120
80
60
40
0
2
4
7
9
PERCENTAGE OF CONTROL
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
CPT-11
PHY 906 + CPT-11

FIG. 5

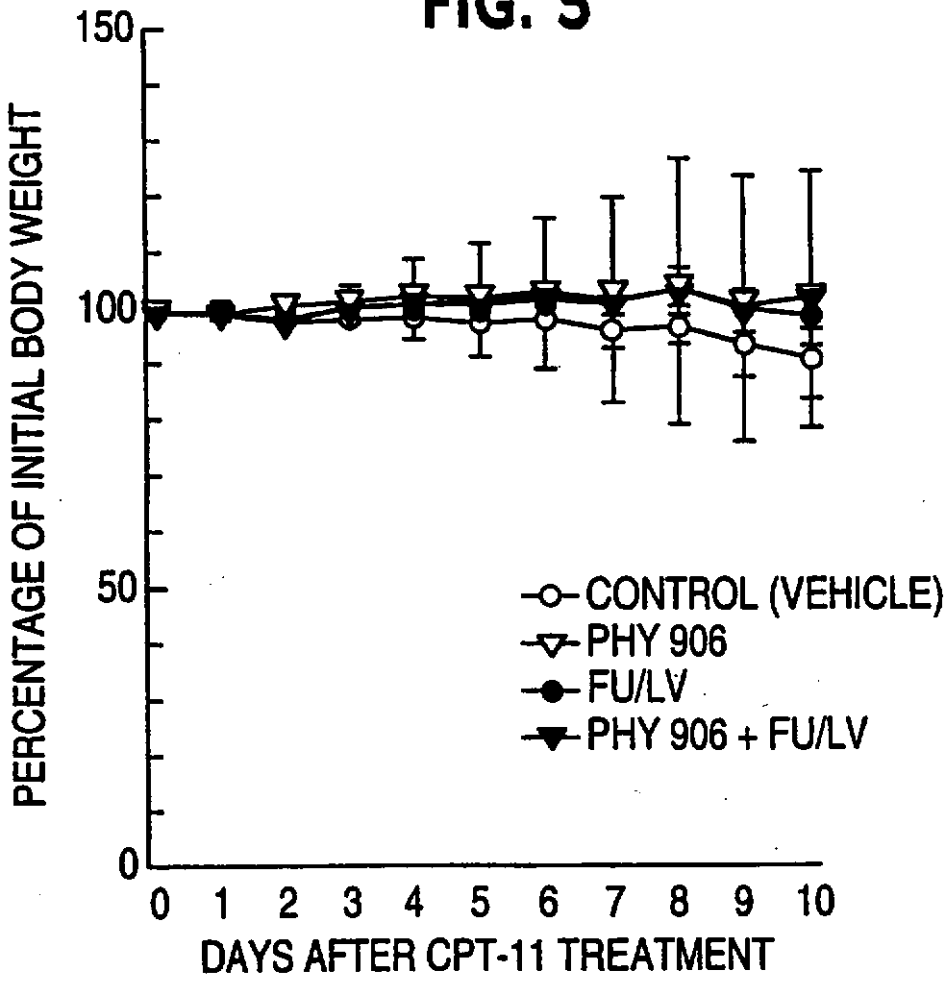

150
100
50
0
PERCENTAGE OF INITIAL BODY WEIGHT
0 1 2 3 4 5 6 7 8 9 10
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
FU/LV
PHY 906 + FU/LV

FIG. 6

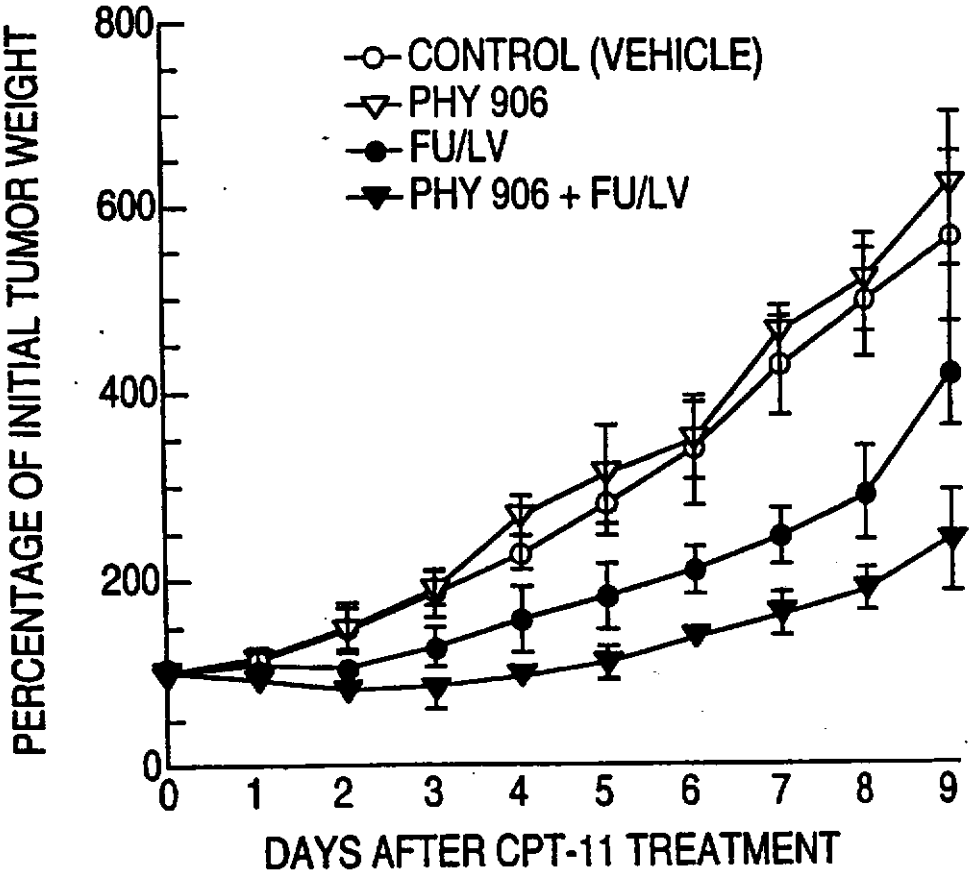

PERCENTAGE OF INITIAL TUMOR WEIGHT
800
600
400
200
0
CONTROL (VEHICLE)
PHY 906
FU/LV
PHY 906 + FU/LV
0 1 2 3 4 5 6 7 8 9
DAYS AFTER CPT-11 TREATMENT

FIG. 7

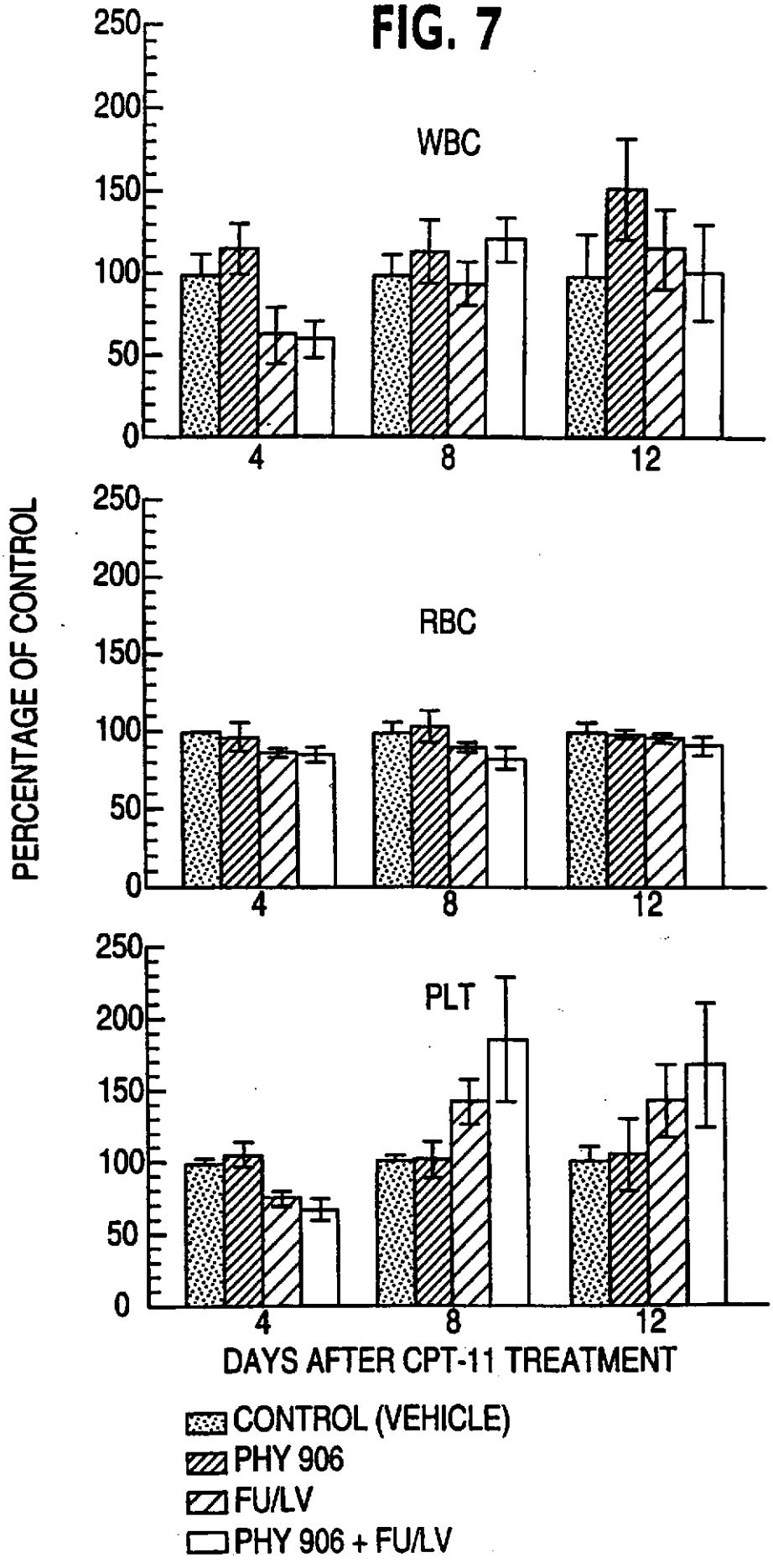

WBC
RBC
PLT
PERCENTAGE OF CONTROL
250
200
150
100
50
0
4
8
12
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
FU/LV
PHY 906 + FU/LV

FIG. 8

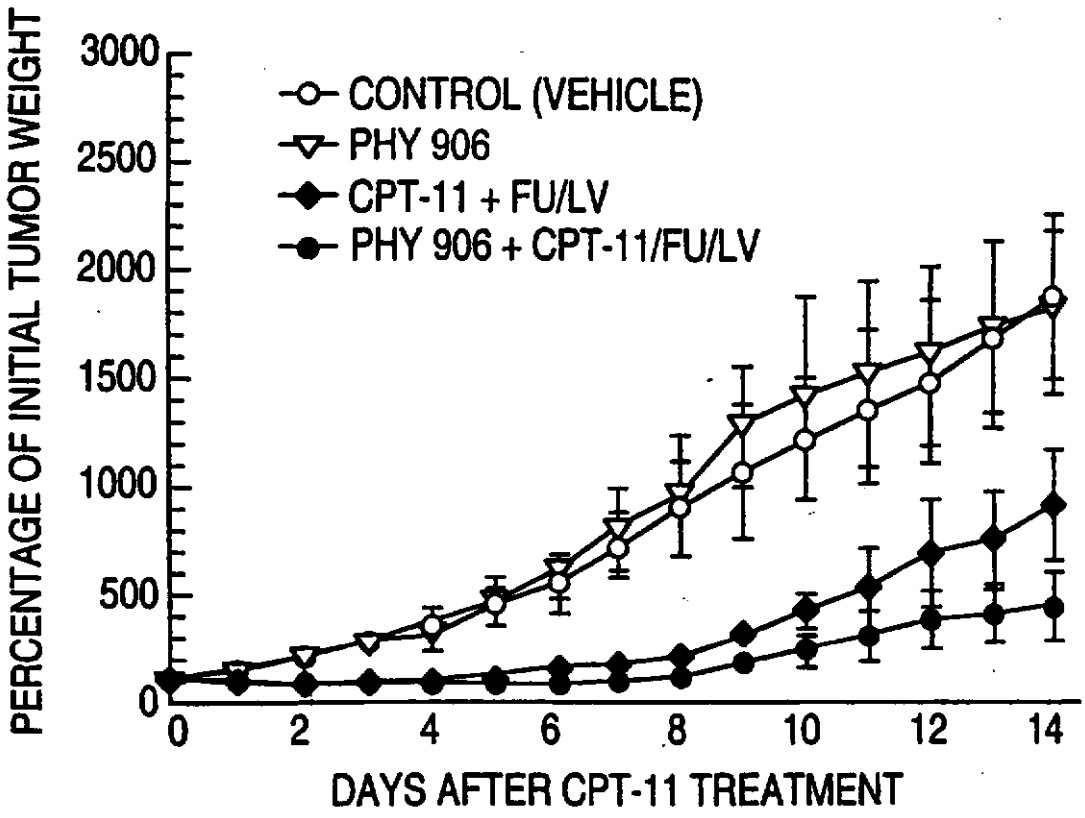

PERCENTAGE OF INITIAL TUMOR WEIGHT
3000
2500
2000
1500
1000
500
0
0
2
4
6
8
10
12
14
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
CPT-11 + FU/LV
PHY 906 + CPT-11/FU/LV

FIG. 9

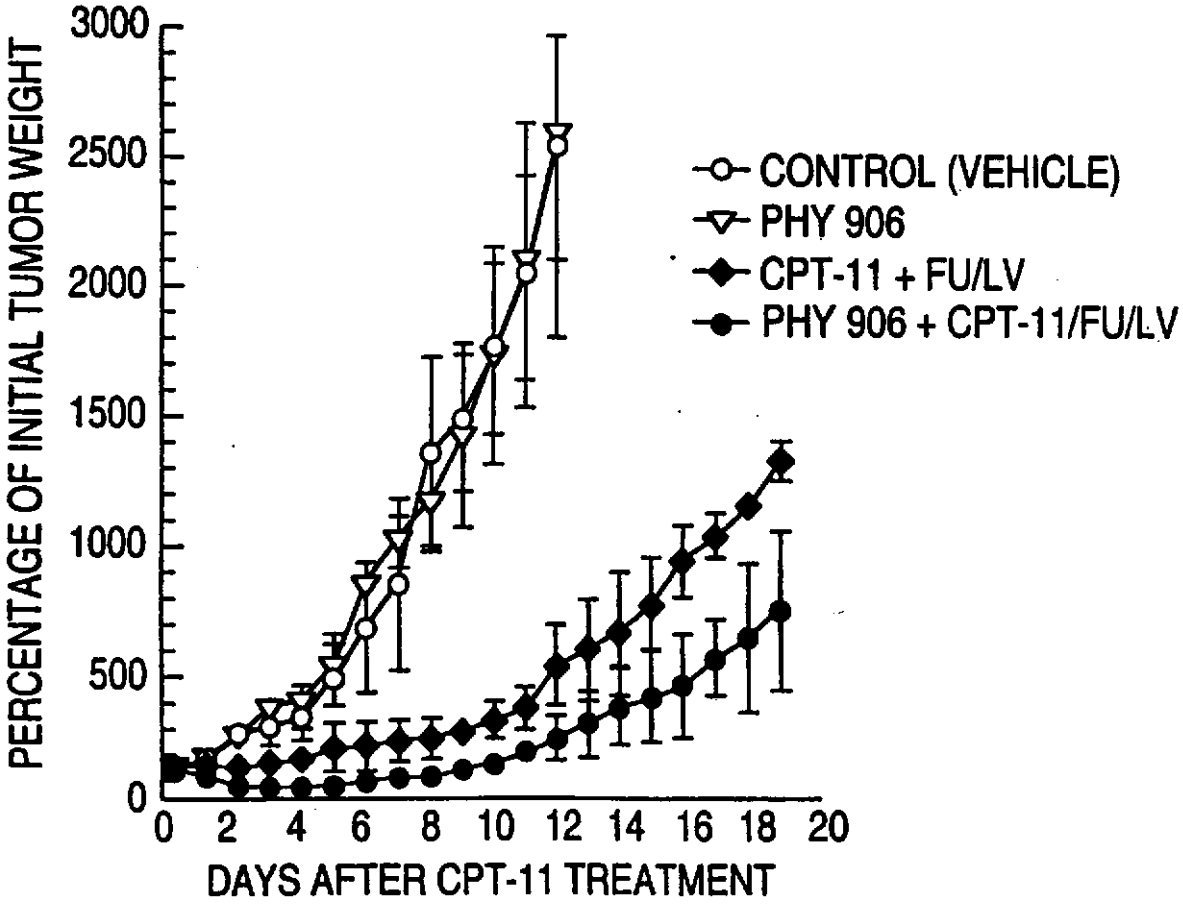

PERCENTAGE OF INITIAL TUMOR WEIGHT
3000
2500
2000
1500
1000
500
0
0 2 4 6 8 10 12 14 16 18 20
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
CPT-11 + FU/LV
PHY 906 + CPT-11/FU/LV

FIG. 10

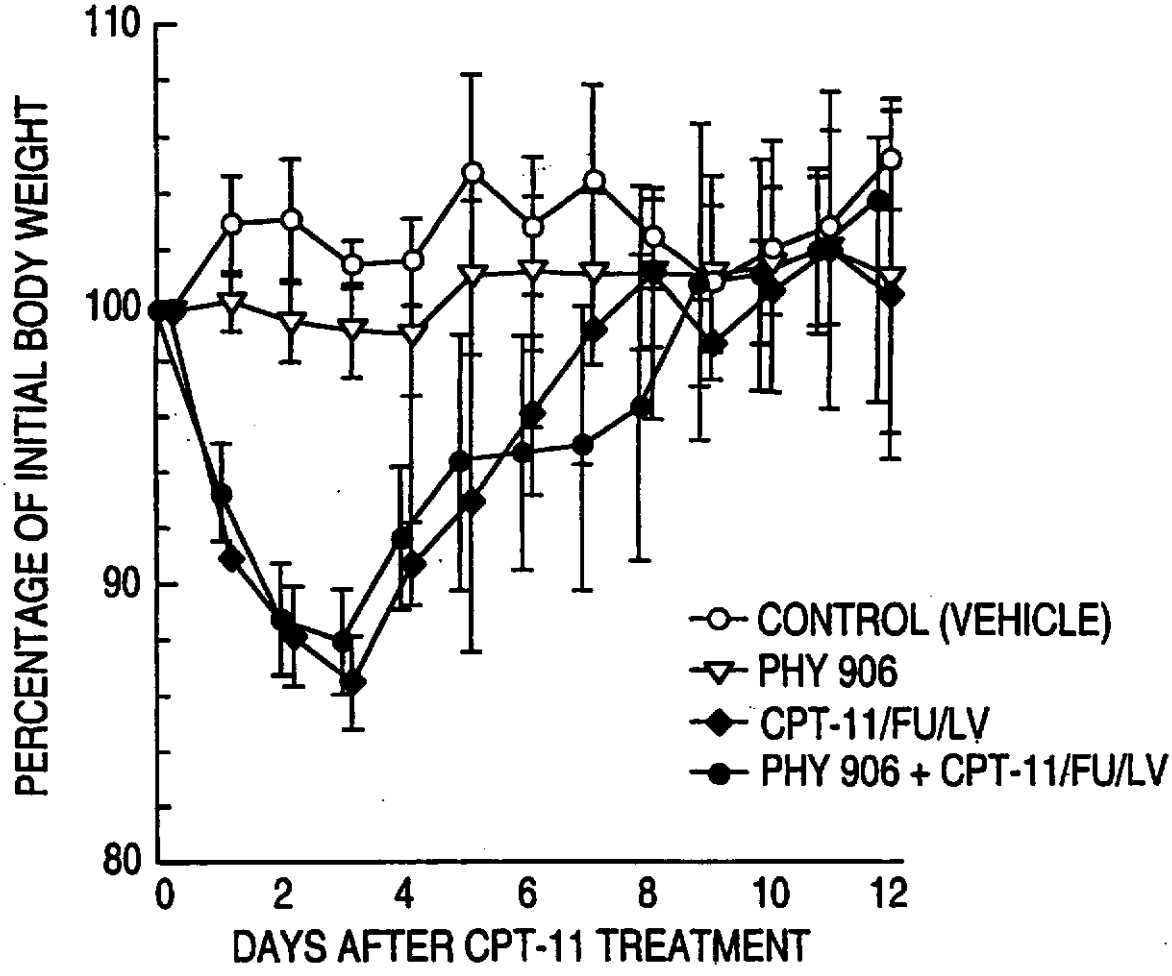
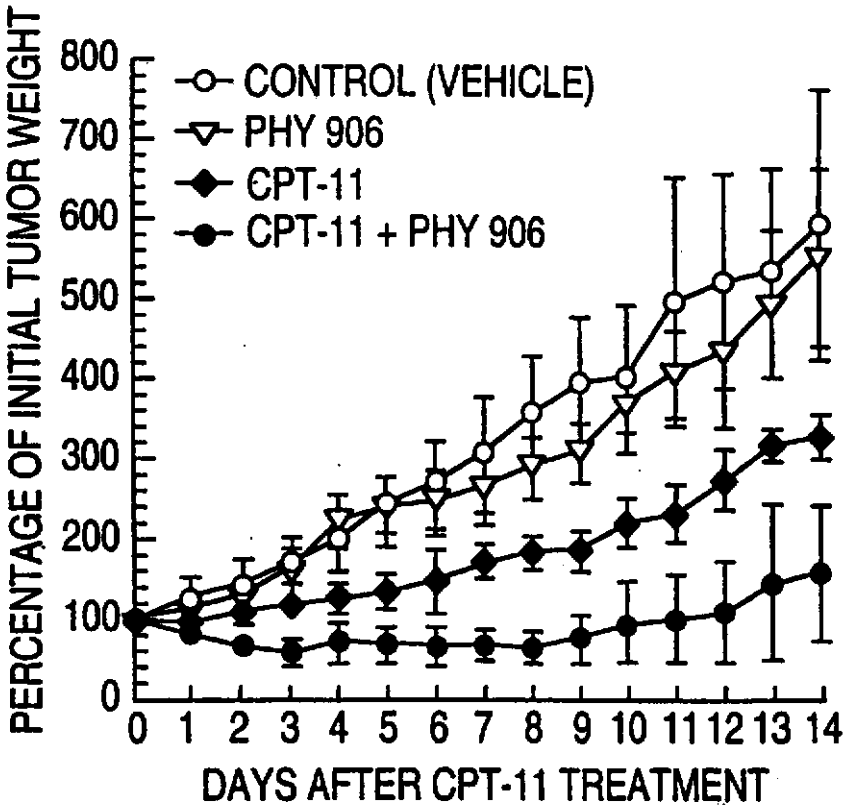

110
100
90
80
PERCENTAGE OF INITIAL BODY WEIGHT
0
2
4
6
8
10
12
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
CPT-11/FU/LV
PHY 906 + CPT-11/FU/LV

FIG. 11



t-tst (*PVALUE) AT DAY 14	CONTROL (VEHICLE)	CPT-11	PHY 906
CPT011	0.02	—	0.02
PHY 906	0.6	0.02	—
CPT-11 + PHY 906	0.002	0.01	0.003

FIG. 12

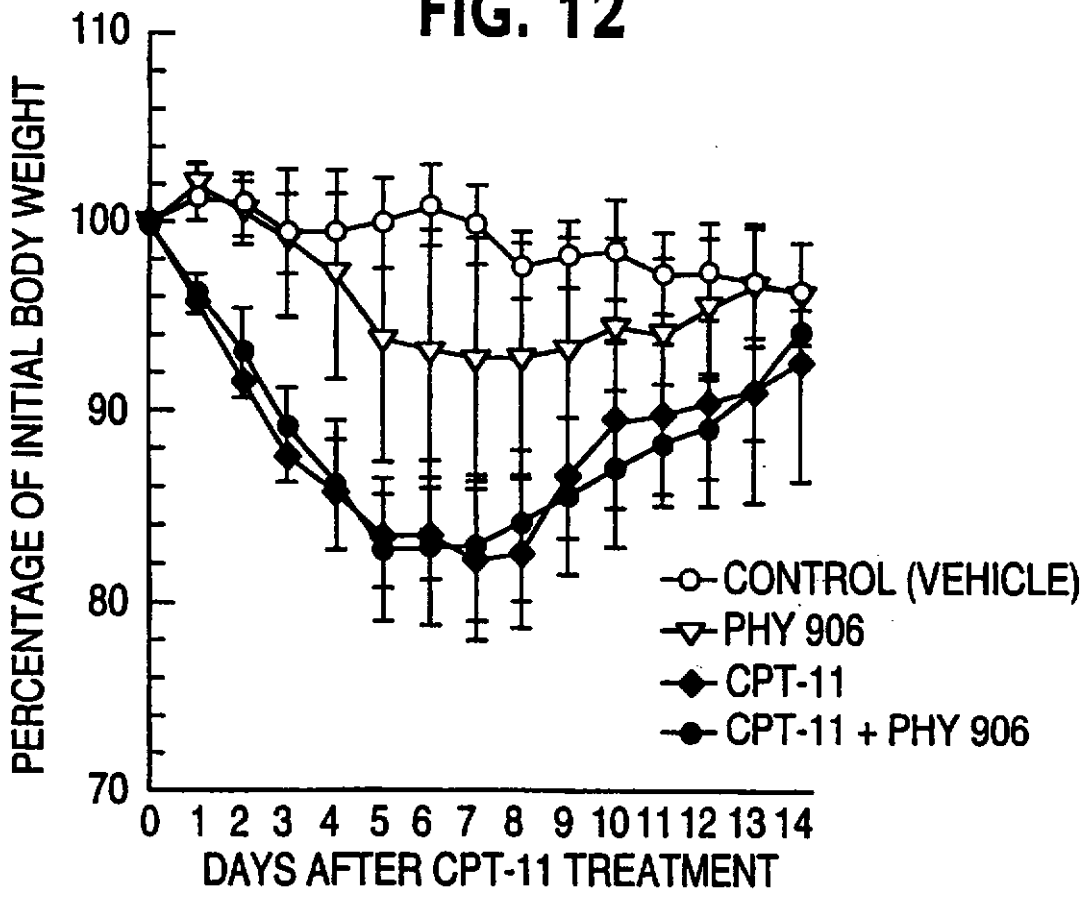

PERCENTAGE OF INITIAL BODY WEIGHT
110
100
90
80
70
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14
DAYS AFTER CPT-11 TREATMENT
CONTROL (VEHICLE)
PHY 906
CPT-11
CPT-11 + PHY 906

FIG. 13

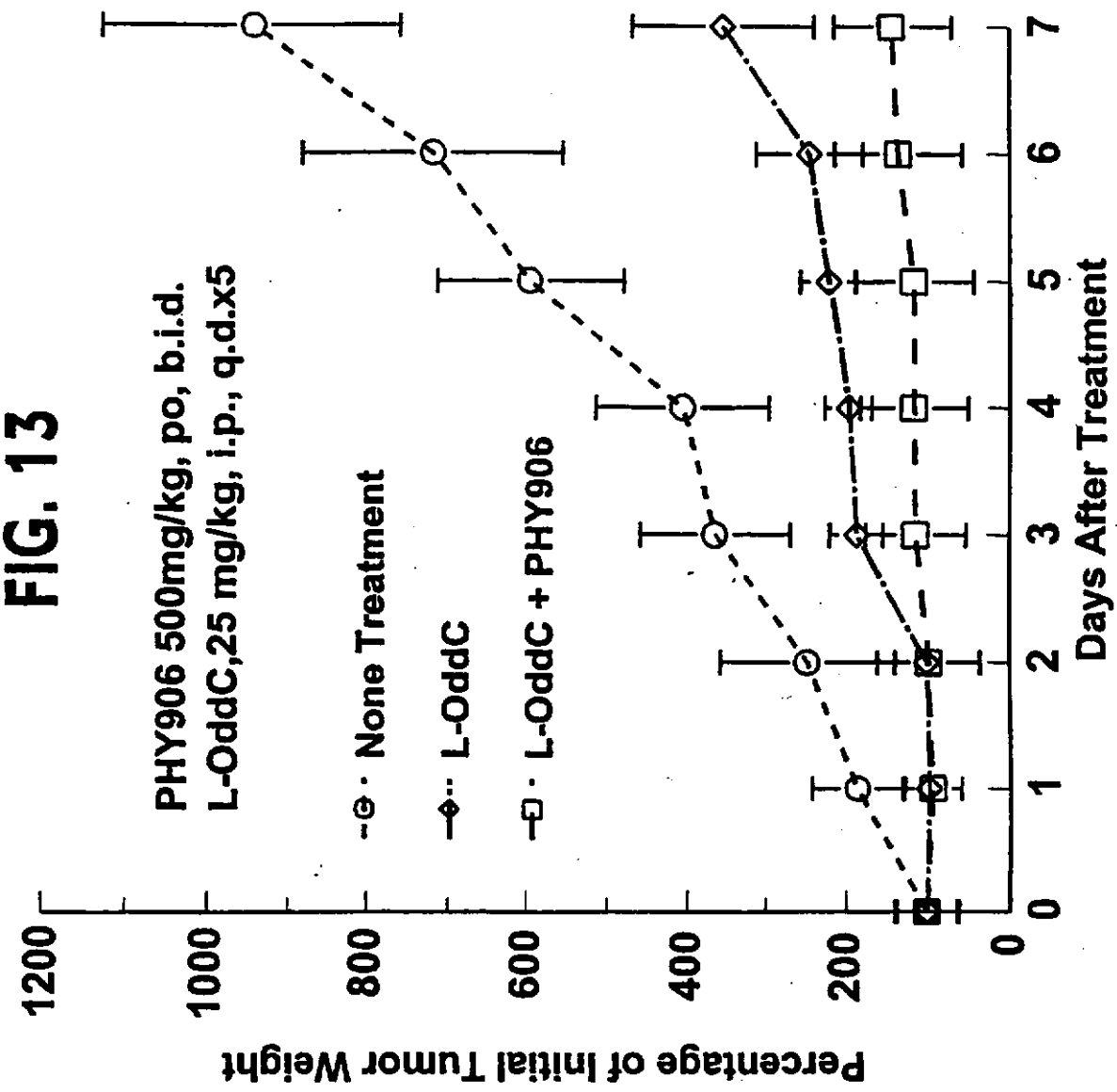

PHY906 500mg/kg, po, b.i.d.
L-OddC,25 mg/kg, i.p., q.d.x5
None Treatment
L-OddC
L-OddC + PHY906
Percentage of Initial Tumor Weight
Days After Treatment
0
200
400
600
800
1000
1200
0
1
2
3
4
5
6
7

FIG. 14

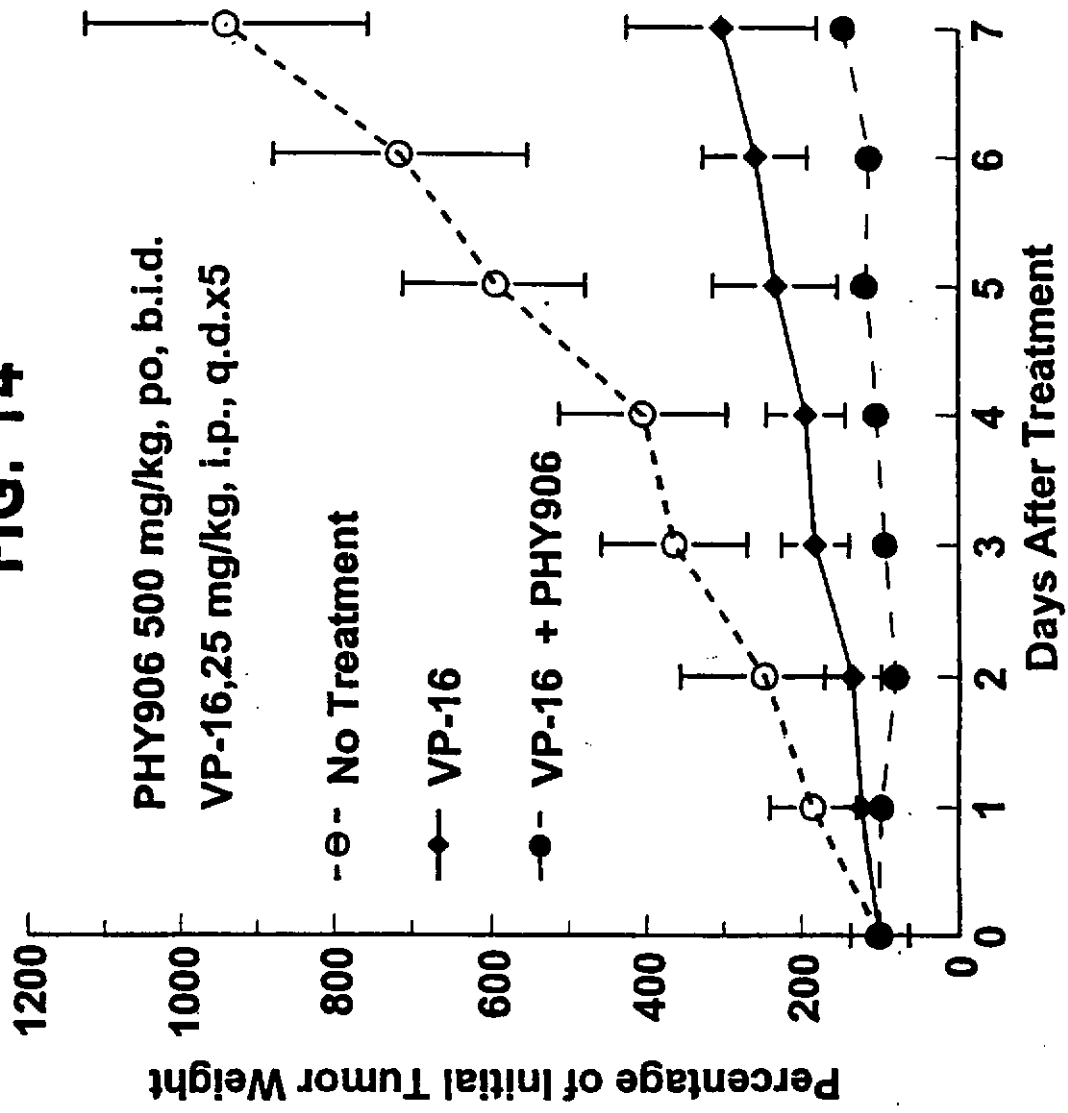

PHY906 500 mg/kg, po, b.i.d.
VP-16,25 mg/kg, i.p., q.d.x5
-e- No Treatment
VP-16
VP-16 + PHY906
Percentage of Initial Tumor Weight
Days After Treatment
1200
1000
800
600
400
200
0
0
1
2
3
4
5
6
7

FIG. 15

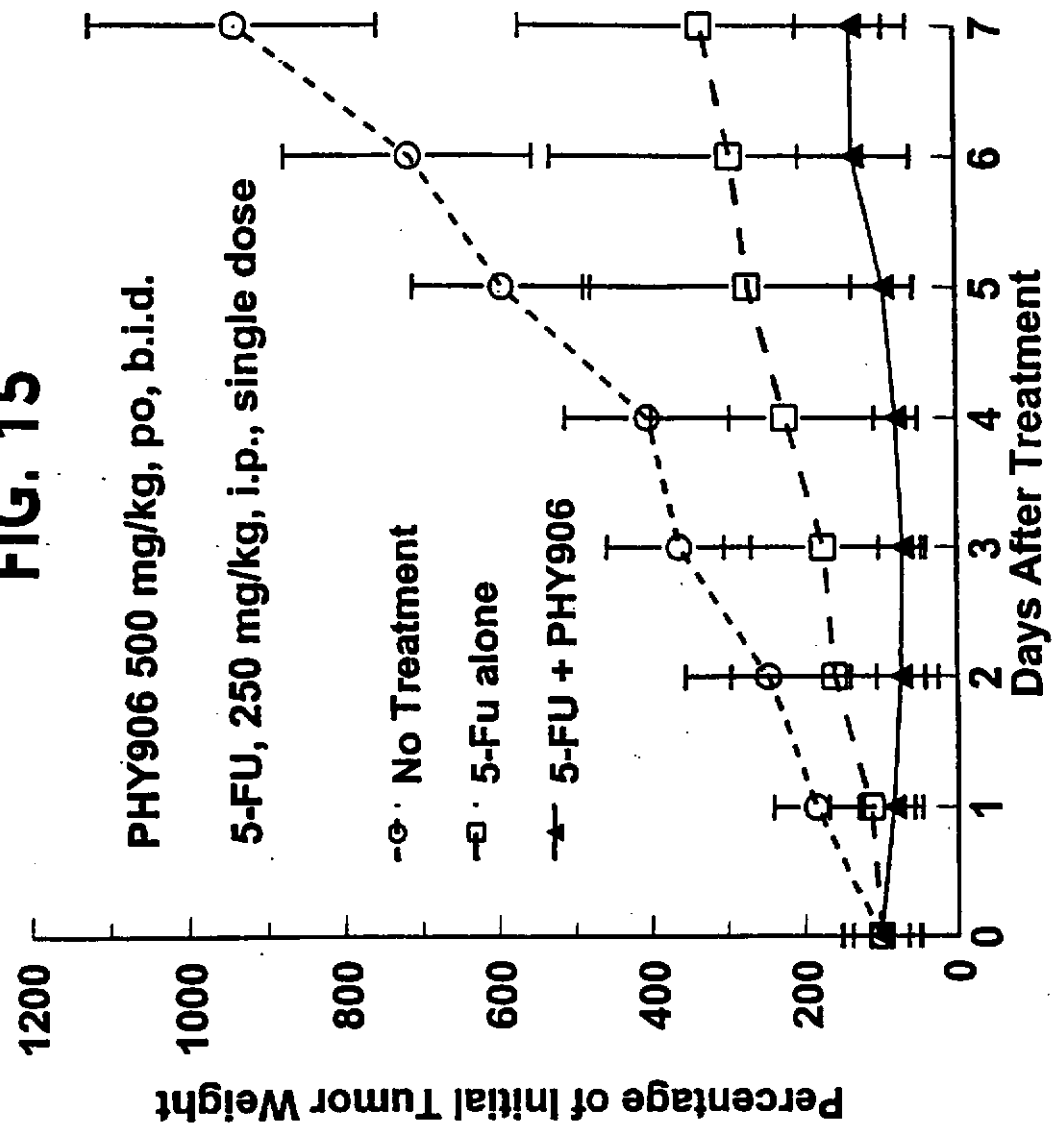

PHY906 500 mg/kg, po, b.i.d.
5-FU, 250 mg/kg, i.p., single dose
No Treatment
5-Fu alone
5-FU + PHY906
Percentage of Initial Tumor Weight
Days After Treatment
0
200
400
600
800
1000
1200
0
1
2
3
4
5
6
7

FIG. 16

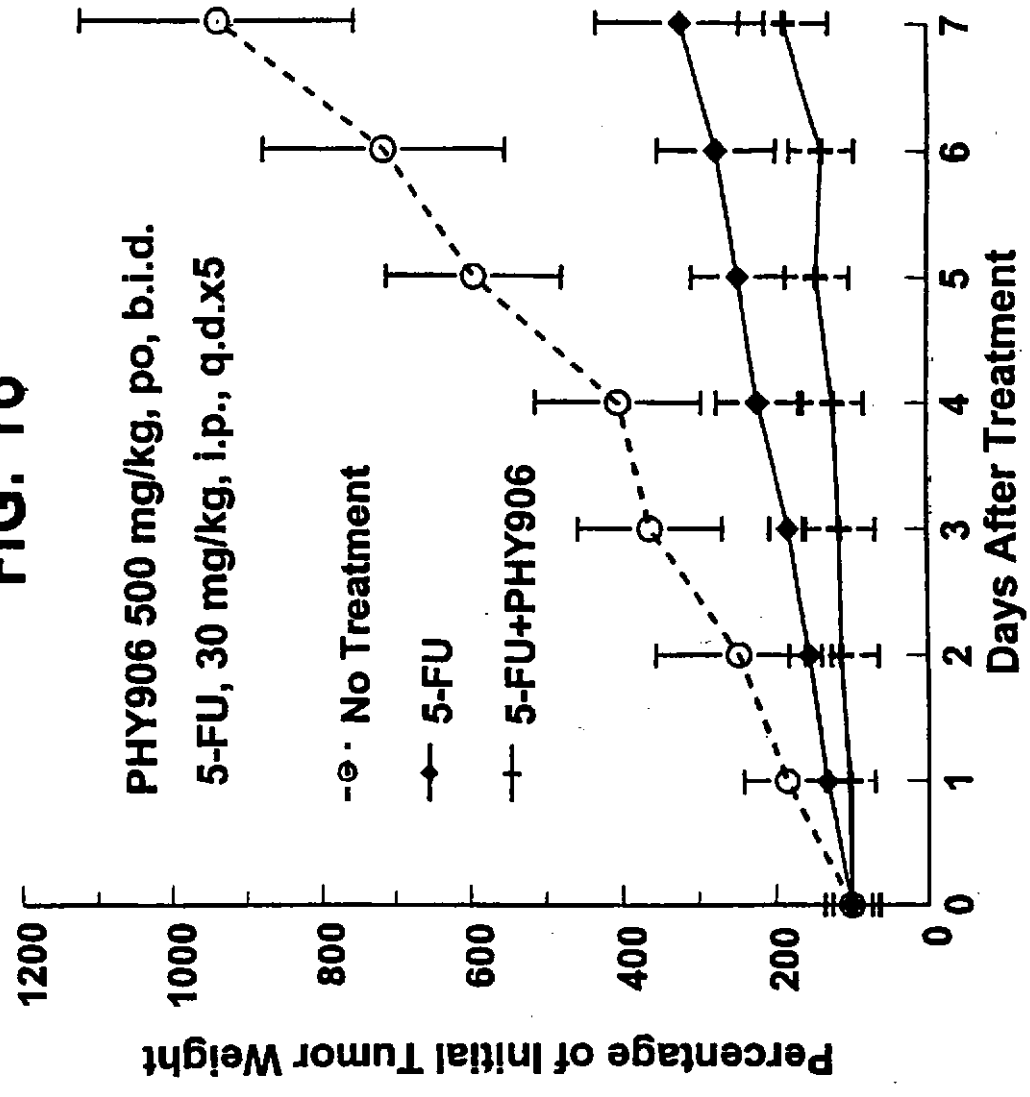

PHY906 500 mg/kg, po, b.i.d.
5-FU, 30 mg/kg, i.p., q.d.x5
No Treatment
5-FU
5-FU+PHY906
Percentage of Initial Tumor Weight
1200
1000
800
600
400
200
0
0
1
2
3
4
5
6
7
Days After Treatment

FIG. 17

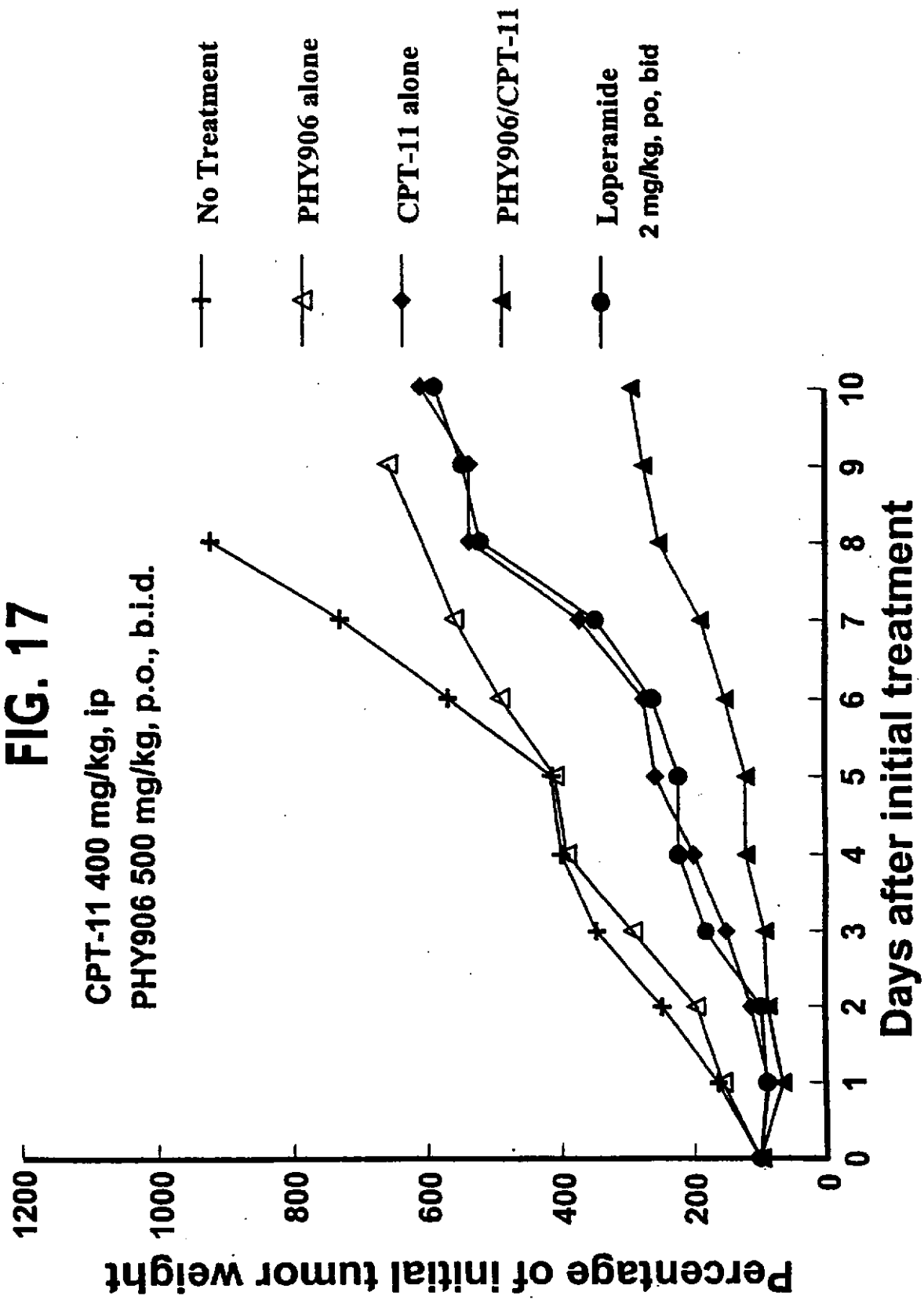

CPT-11 400 mg/kg, ip
PHY906 500 mg/kg, p.o., b.i.d.
No Treatment
PHY906 alone
CPT-11 alone
PHY906/CPT-11
Loperamide
2 mg/kg, po, bid
Percentage of initial tumor weight
1200
1000
800
600
400
200
0
0
1
2
3
4
5
6
7
8
9
10
Days after initial treatment

FIG. 18

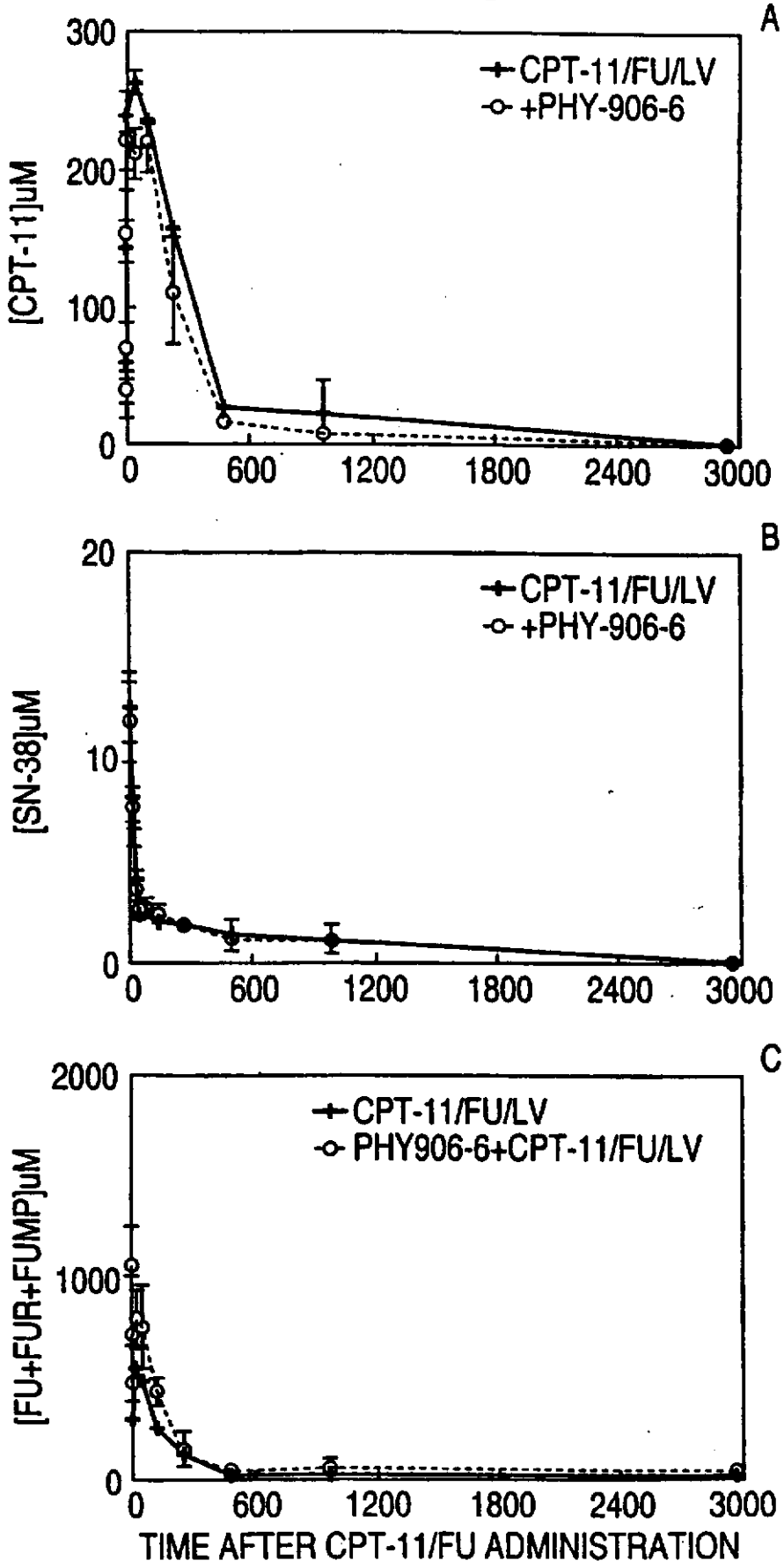

A
300
200
100
0
[CPT-11]uM
CPT-11/FU/LV
+PHY-906-6
0
600
1200
1800
2400
3000
B
20
10
0
[SN-38]uM
CPT-11/FU/LV
+PHY-906-6
0
600
1200
1800
2400
3000
C
2000
1000
0
[FU+FUR+FUMP]uM
CPT-11/FU/LV
PHY906-6+CPT-11/FU/LV
0
600
1200
1800
2400
3000
TIME AFTER CPT-11/FU ADMINISTRATION

FIG. 19

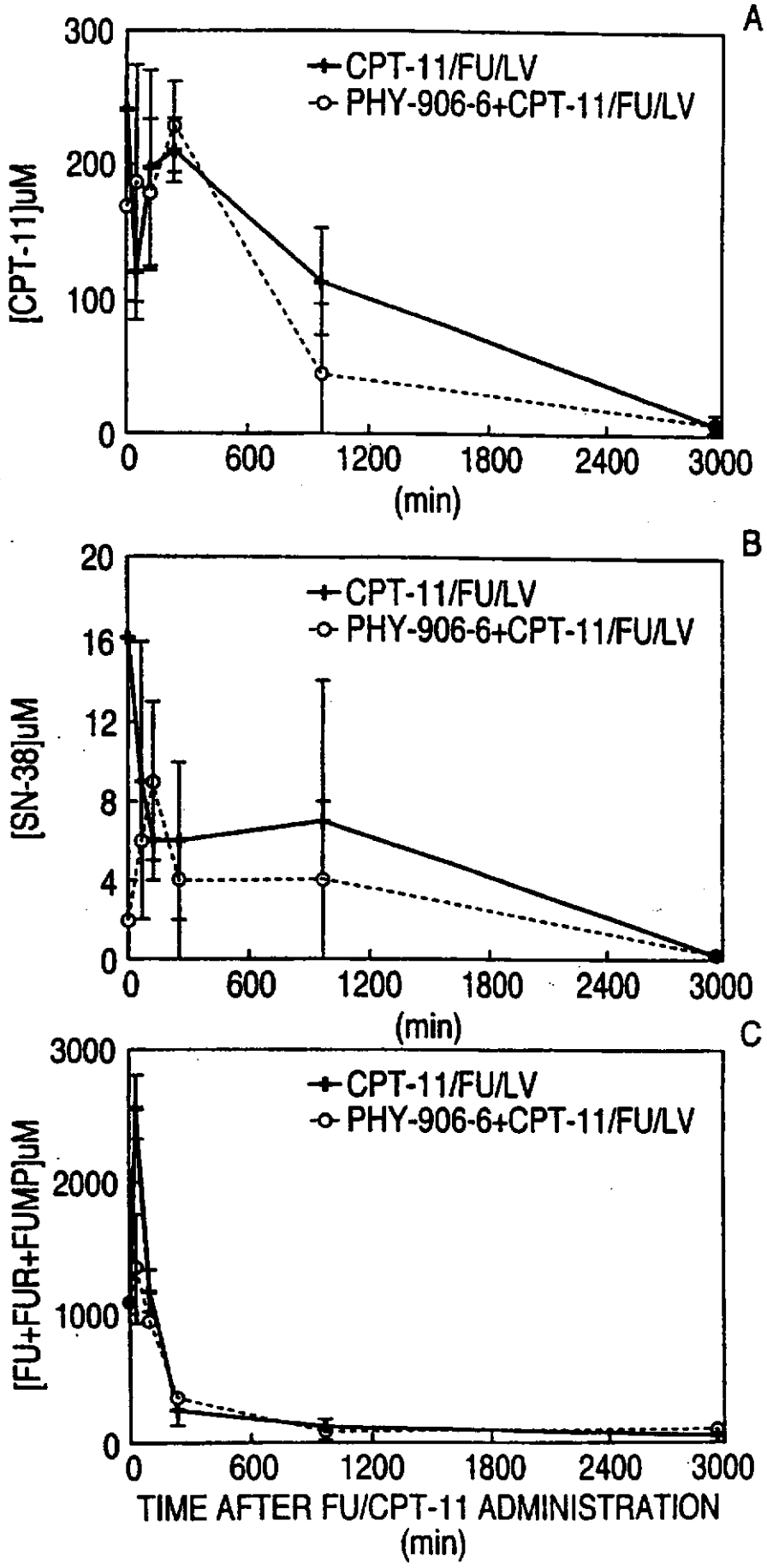
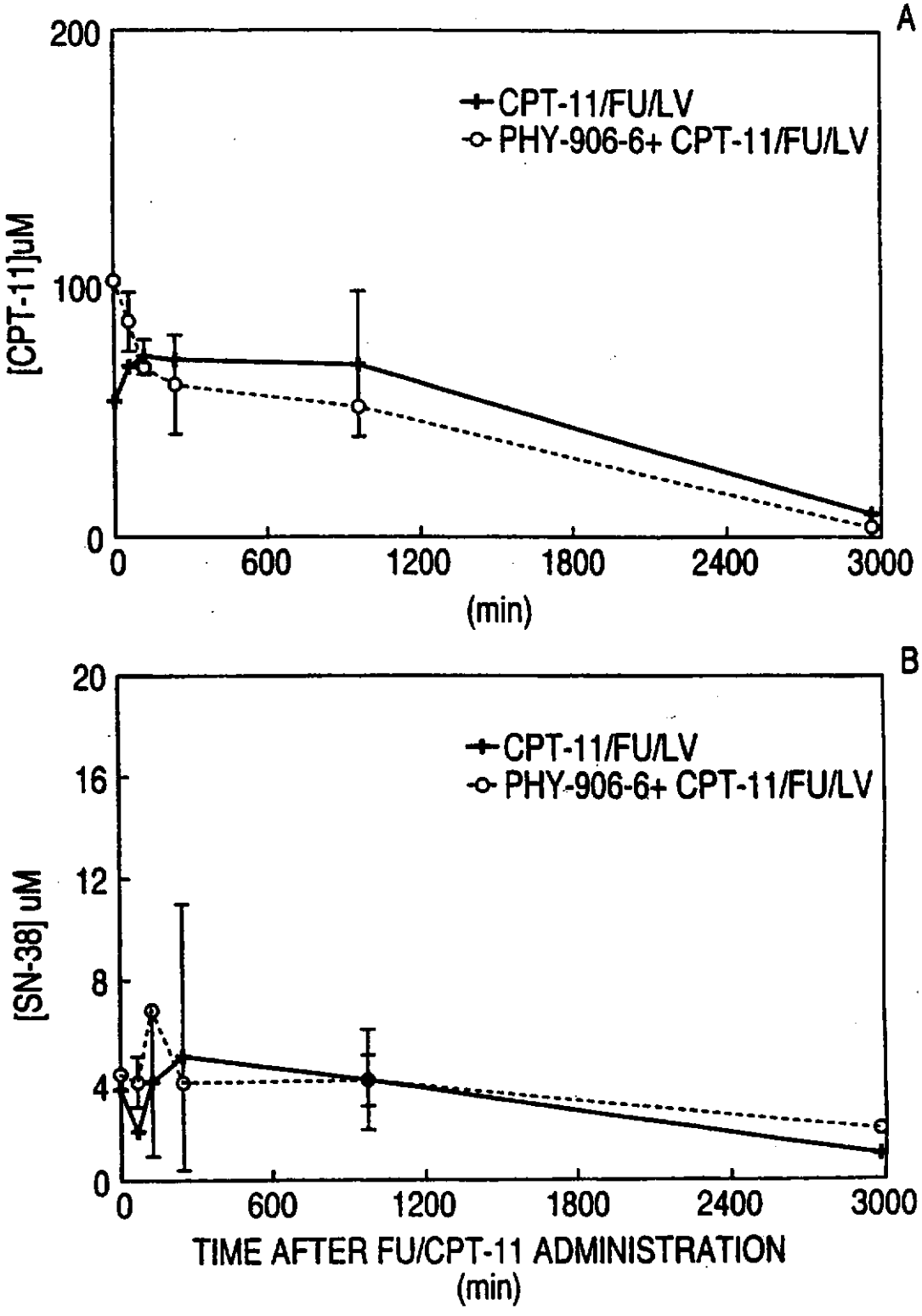

A
300
200
100
0
[CPT-11]μM
CPT-11/FU/LV
PHY-906-6+CPT-11/FU/LV
0
600
1200
1800
2400
3000
(min)
B
20
16
12
8
4
0
[SN-38]μM
CPT-11/FU/LV
PHY-906-6+CPT-11/FU/LV
0
600
1200
1800
2400
3000
(min)
C
3000
2000
1000
0
[FU+FUR+FUMP]μM
CPT-11/FU/LV
PHY-906-6+CPT-11/FU/LV
0
600
1200
1800
2400
3000
TIME AFTER FU/CPT-11 ADMINISTRATION
(min)

FIG. 20


A
200
100
0
[CPT-11]μM
CPT-11/FU/LV
PHY-906-6+ CPT-11/FU/LV
0
600
1200
1800
2400
3000
(min)
B
20
16
12
8
4
0
[SN-38] uM
CPT-11/FU/LV
PHY-906-6+ CPT-11/FU/LV
0
600
1200
1800
2400
3000
TIME AFTER FU/CPT-11 ADMINISTRATION
(min)