FIG. 1

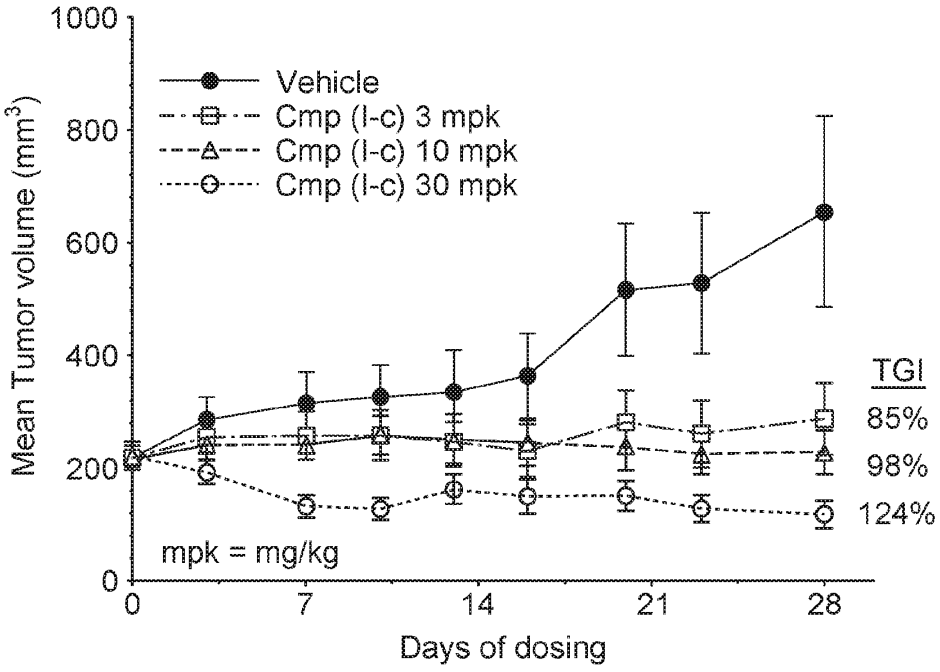

1000
800
600
400
200
0
Mean Tumor volume (mm3)
Vehicle
Cmp (I-c) 3 mpk
Cmp (I-c) 10 mpk
Cmp (I-c) 30 mpk
mpk = mg/kg
0
7
14
21
28
Days of dosing
TGI
85%
98%
124%

FIG. 2

WESTERN BLOT PD (18 hours post last dose)	% ER REDUCTION
3 mpk	95
10 mpk	97
30 mpk	94

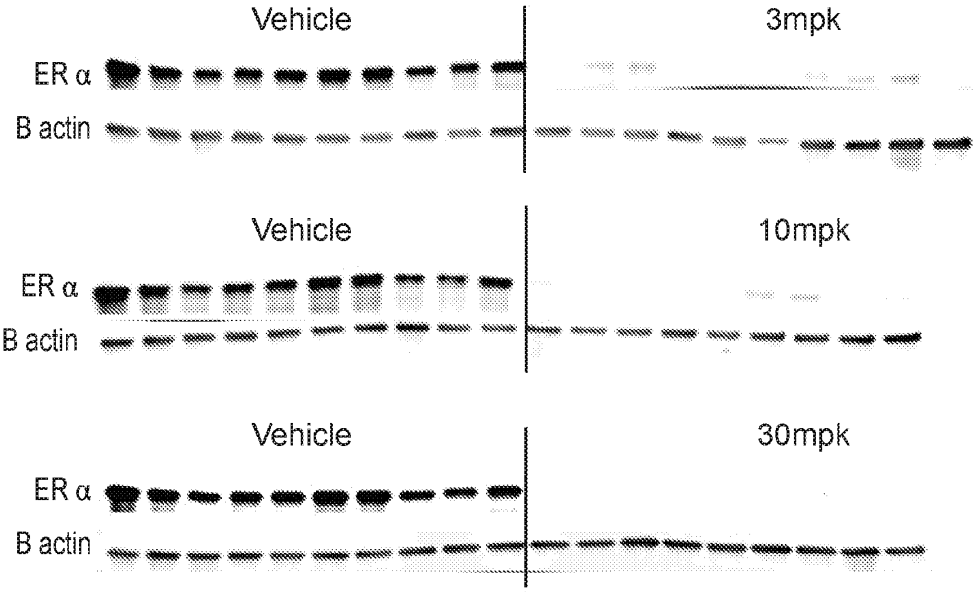


FIG. 3

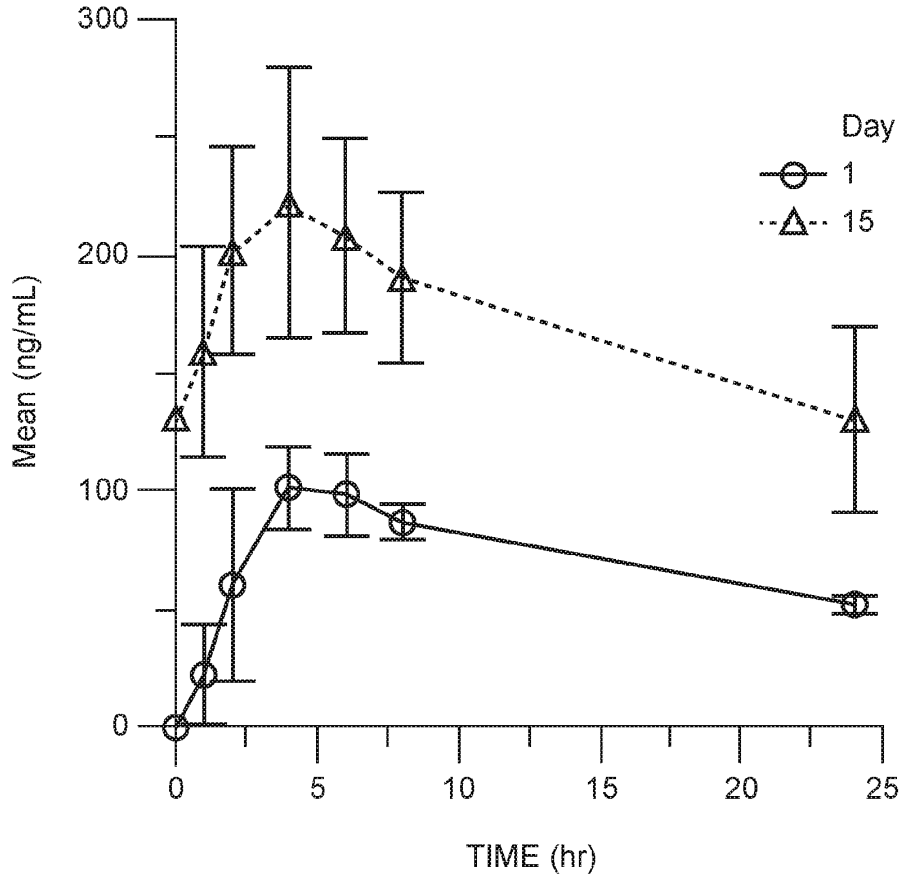


FIG. 4

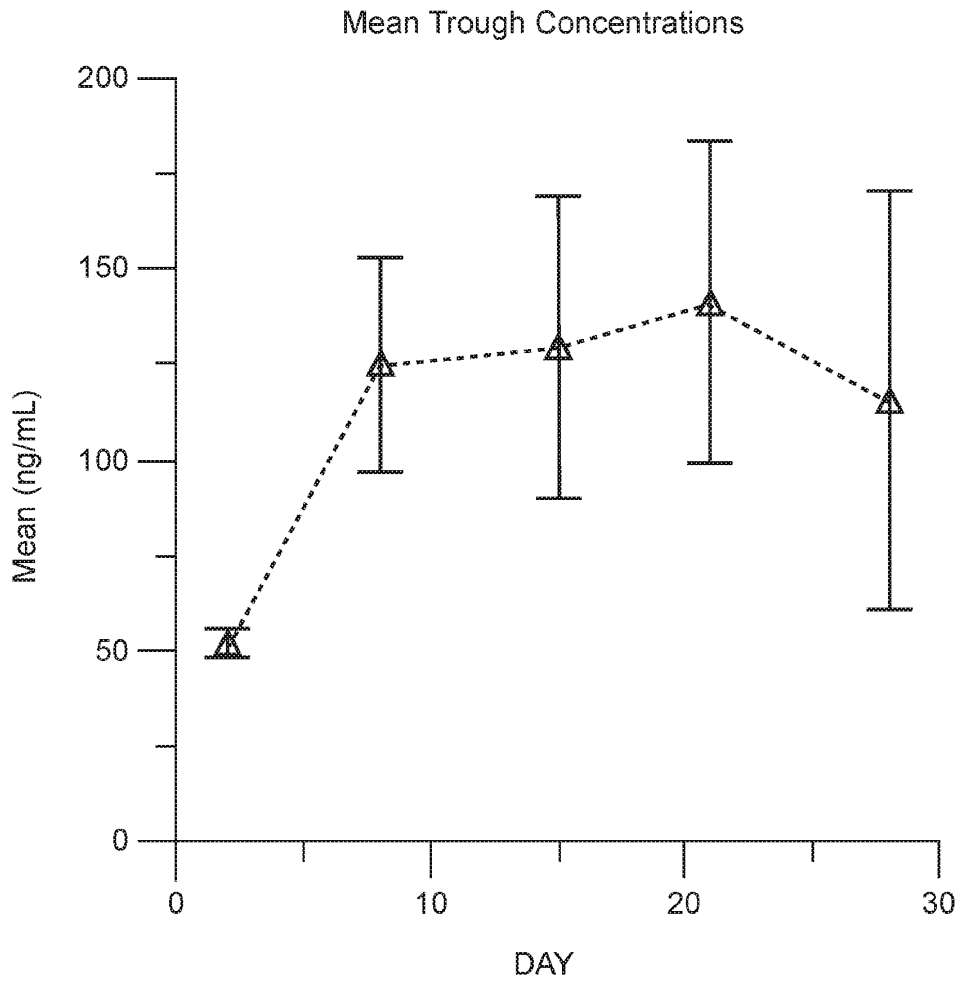

Mean Trough Concentrations
Mean (ng/mL)
200
150
100
50
0
0
10
20
30
DAY

FIG. 5

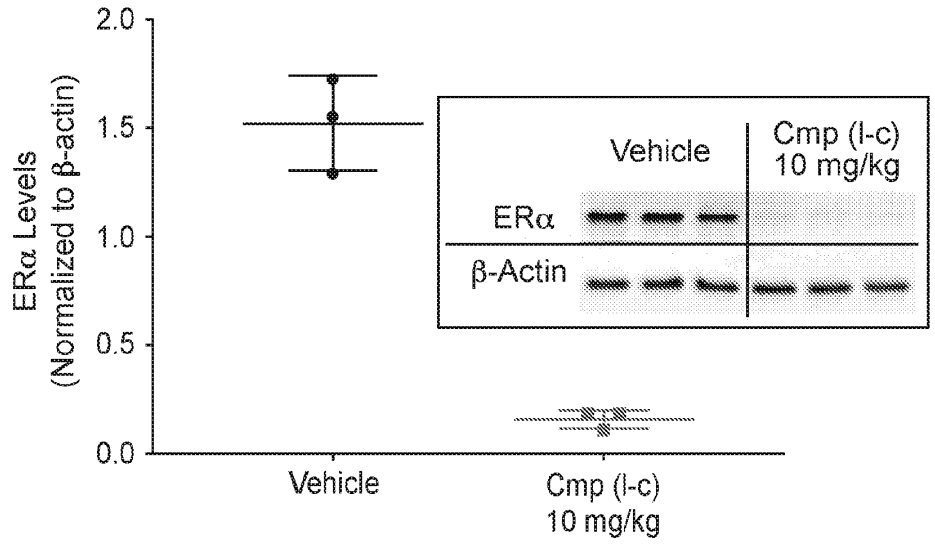

2.0
1.5
1.0
0.5
0.0
ERα Levels
(Normalized to β-actin)
Vehicle
Cmp (I-c)
10 mg/kg
Vehicle
Cmp (I-c)
10 mg/kg
ERα
β-Actin

FIG. 6

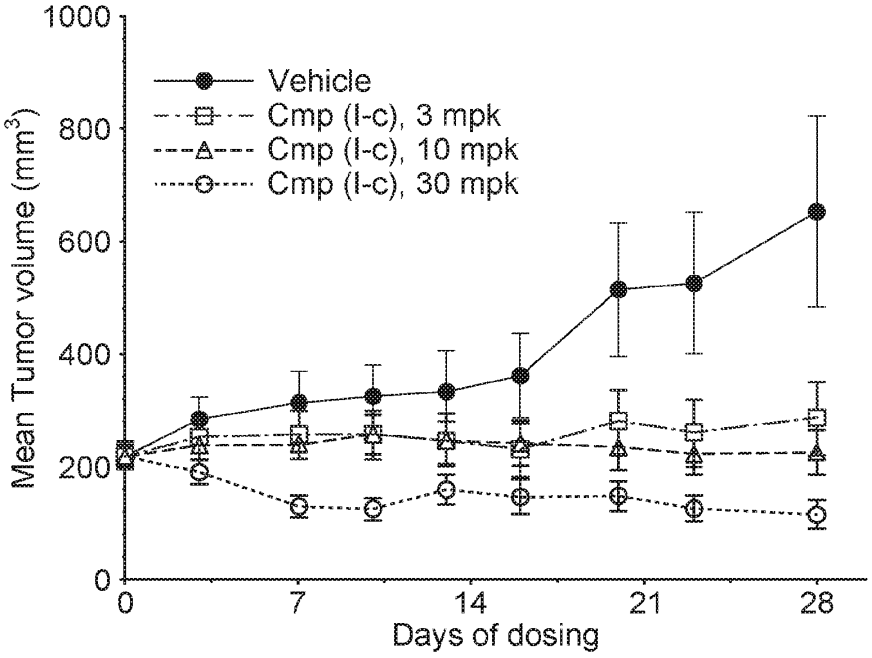

1000
800
600
400
200
0
Mean Tumor volume (mm3)
Vehicle
Cmp (I-c), 3 mpk
Cmp (I-c), 10 mpk
Cmp (I-c), 30 mpk
0
7
14
21
28
Days of dosing

FIG. 7

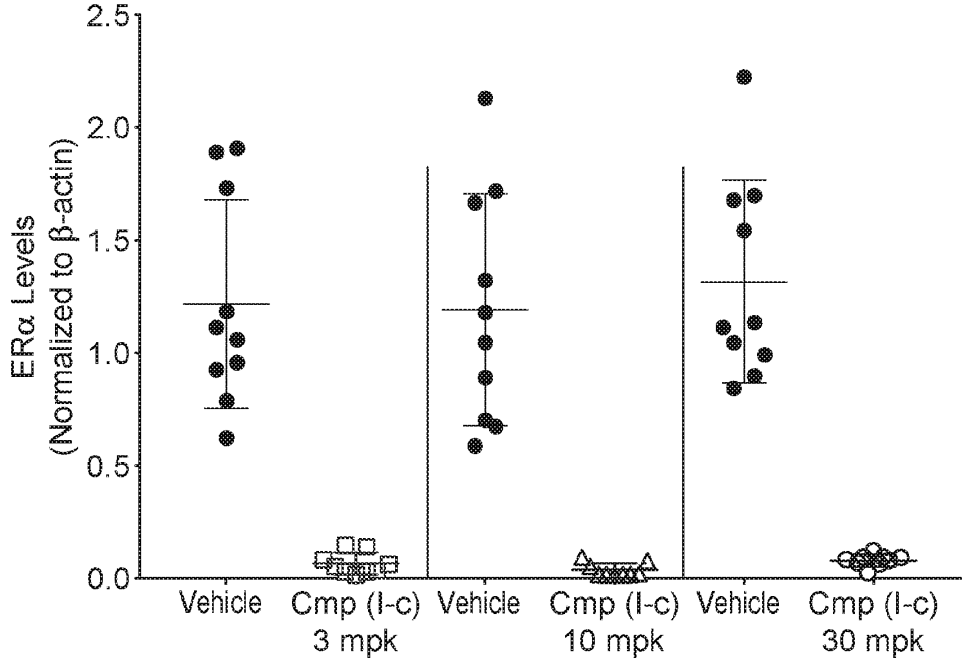

2.5
2.0
1.5
1.0
0.5
0.0
ERα Levels
(Normalized to β-actin)
Vehicle
Cmp (I-c)
3 mpk
Vehicle
Cmp (I-c)
10 mpk
Vehicle
Cmp (I-c)
30 mpk

FIG. 8

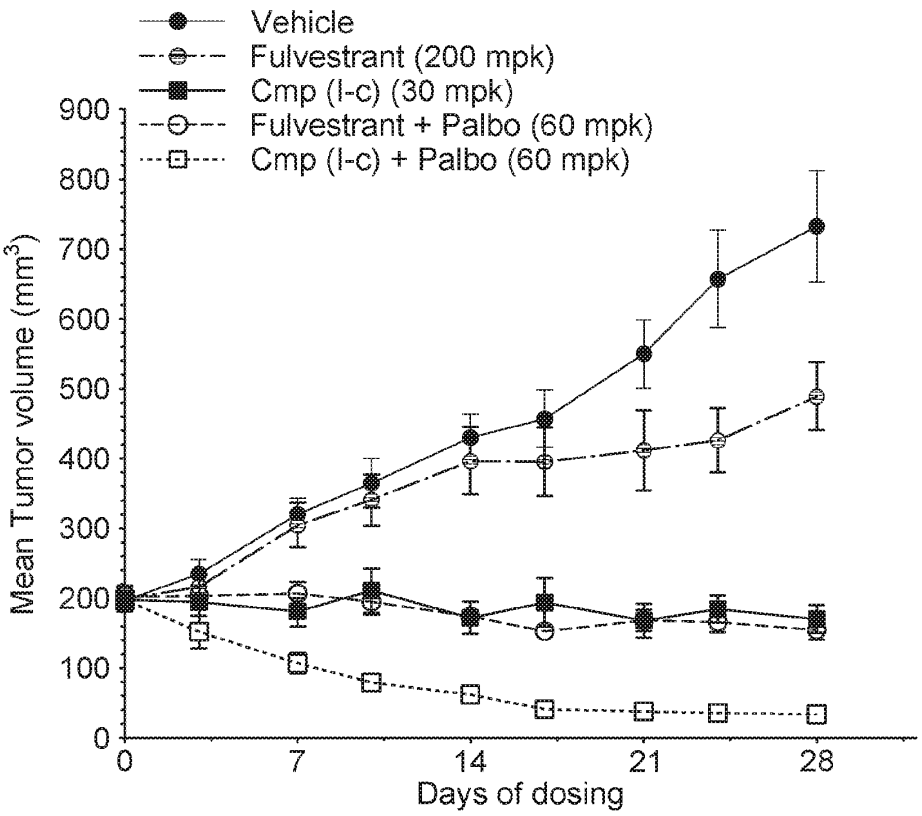

Vehicle
Fulvestrant (200 mpk)
Cmp (I-c) (30 mpk)
Fulvestrant + Palbo (60 mpk)
Cmp (I-c) + Palbo (60 mpk)
Mean Tumor volume (mm3)
900
800
700
600
500
400
300
200
100
0
0
7
14
21
28
Days of dosing

FIG. 9

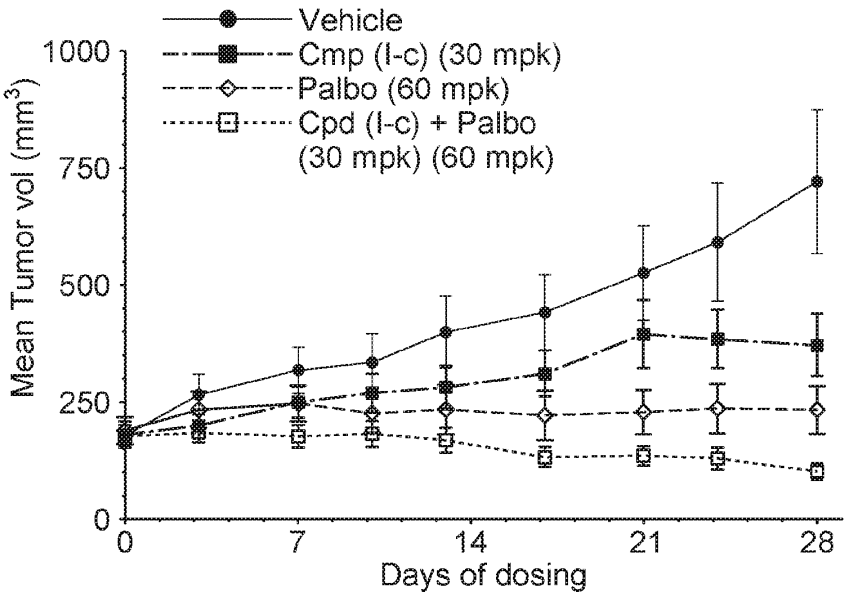

Vehicle
Cmp (I-c) (30 mpk)
Palbo (60 mpk)
Cpd (I-c) + Palbo
(30 mpk) (60 mpk)
Mean Tumor vol (mm3)
1000
750
500
250
0
0
7
14
21
28
Days of dosing

FIG. 10

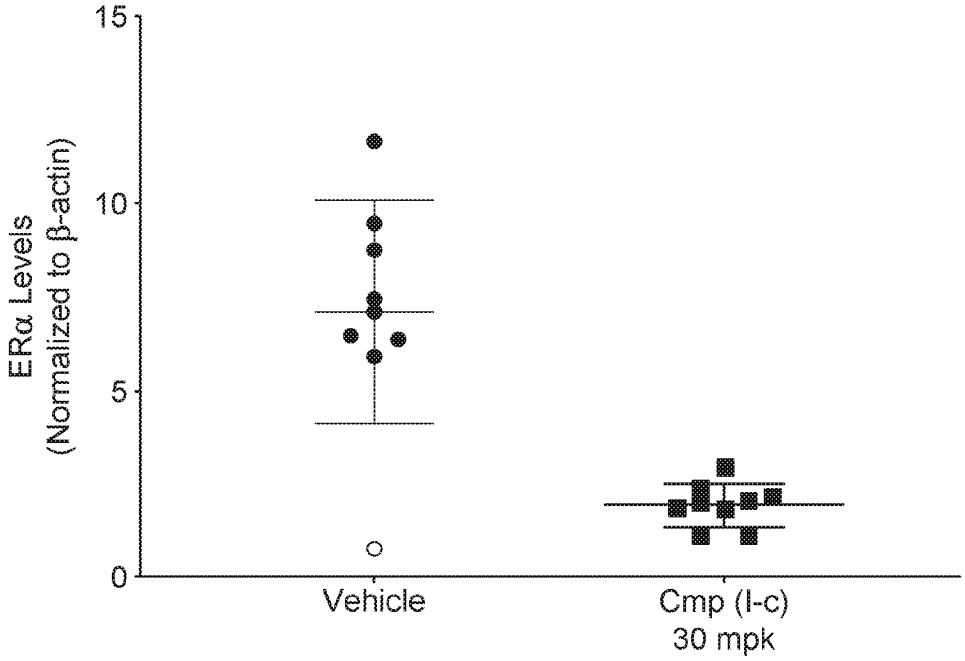

ERα Levels
(Normalized to β-actin)
15
10
5
0
Vehicle
Cmp (I-c)
30 mpk

FIG. 11

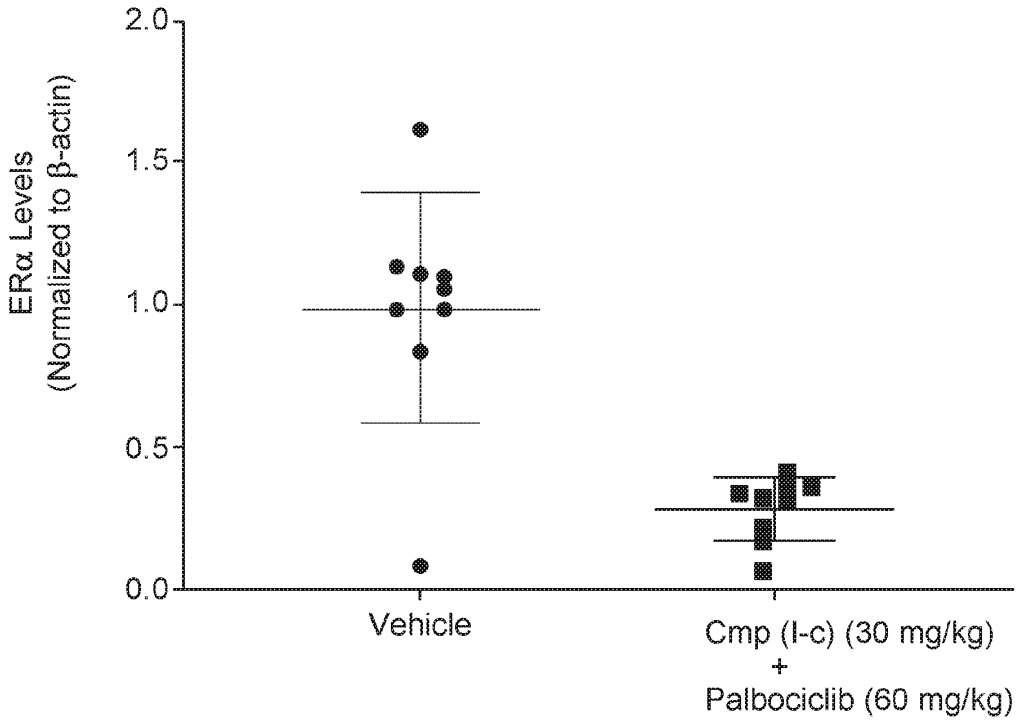

ERα Levels
(Normalized to β-actin)
2.0
1.5
1.0
0.5
0.0
Vehicle
Cmp (I-c) (30 mg/kg)
+
Palbociclib (60 mg/kg)

FIG. 12

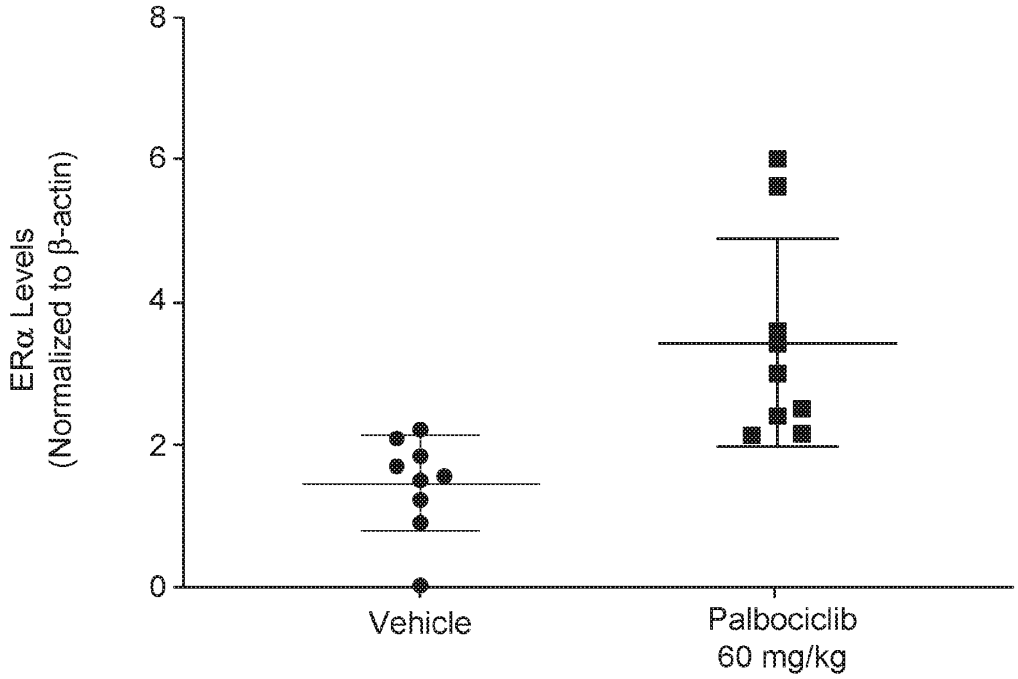

ERα Levels
(Normalized to β-actin)
8
6
4
2
0
Vehicle
Palbociclib
60 mg/kg

FIG. 13

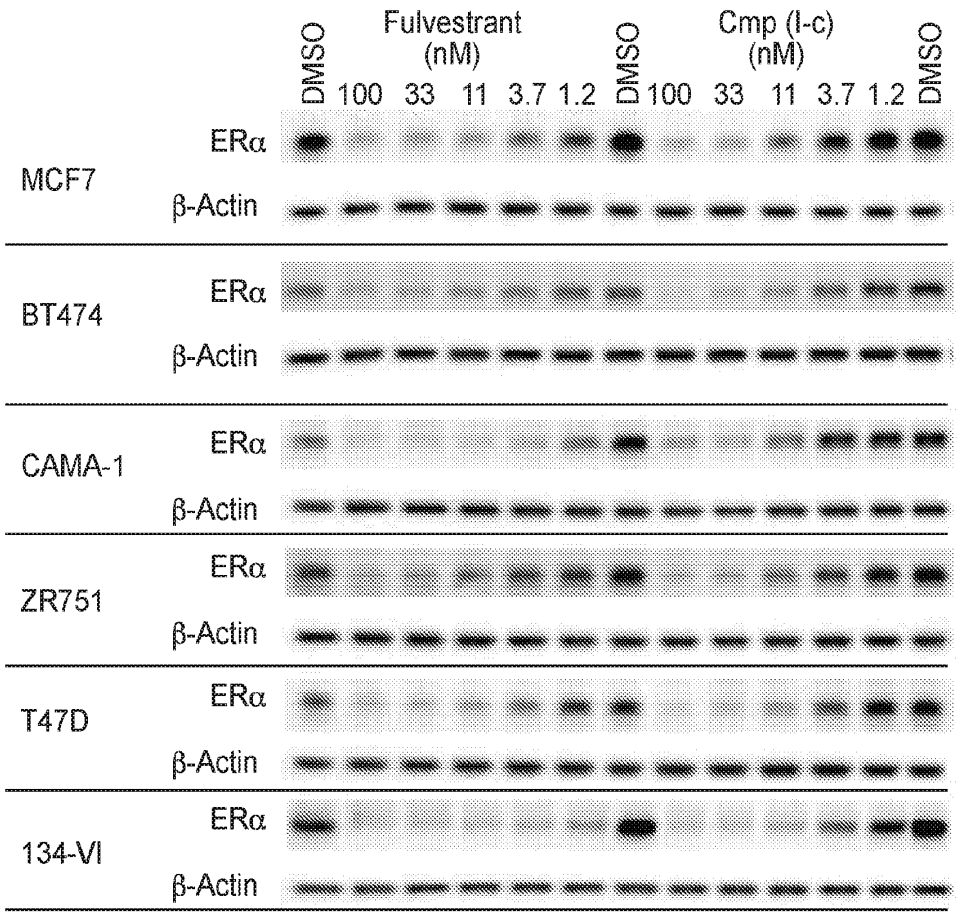

DMSO
Fulvestrant (nM)
100
33
11
3.7
1.2
DMSO
Cmp (I-c) (nM)
100
33
11
3.7
1.2
DMSO
MCF7
ERα
β-Actin
BT474
ERα
β-Actin
CAMA-1
ERα
β-Actin
ZR751
ERα
β-Actin
T47D
ERα
β-Actin
134-VI
ERα
β-Actin

FIG. 14

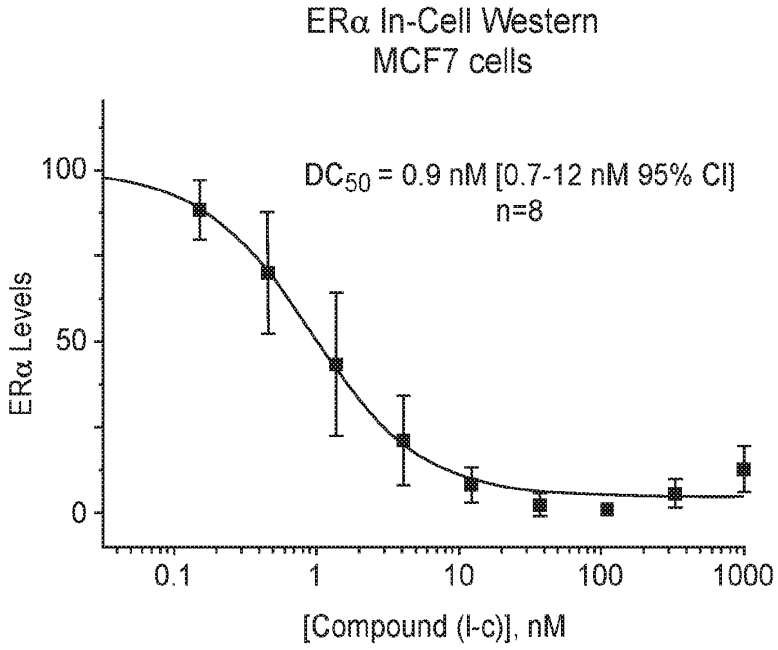

ERα In-Cell Western
MCF7 cells
DC50 = 0.9 nM [0.7-12 nM 95% CI]
n=8
ERα Levels
100
50
0
0.1
1
10
100
1000
[Compound (I-c)], nM

FIG. 15

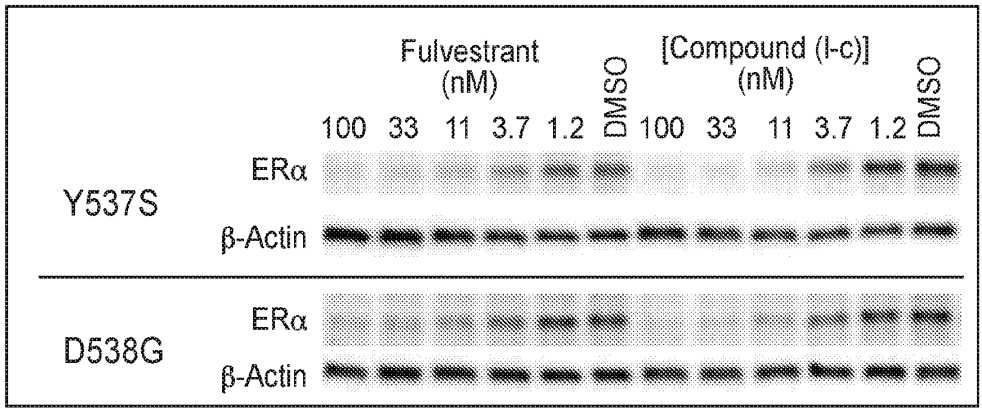

Fulvestrant (nM)
DMSO
[Compound (I-c)] (nM)
DMSO
100 33 11 3.7 1.2
100 33 11 3.7 1.2
Y537S
ERα
β-Actin
D538G
ERα
β-Actin

FIG. 16

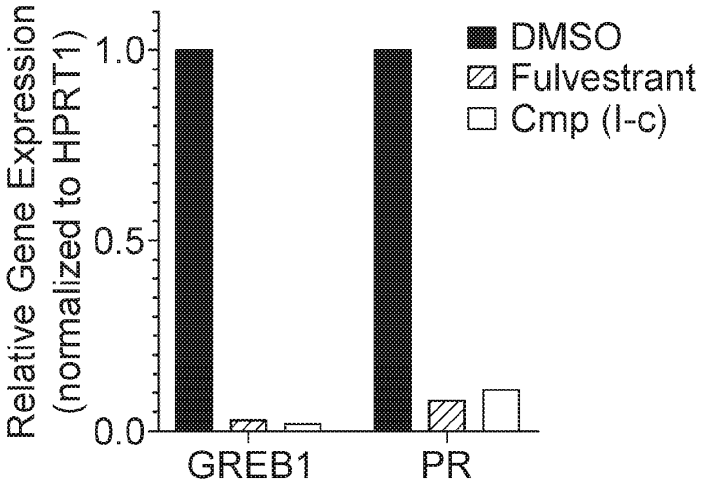

Relative Gene Expression
(normalized to HPRT1)
1.0
0.5
0.0
DMSO
Fulvestrant
Cmp (I-c)
GREB1
PR

FIG. 17

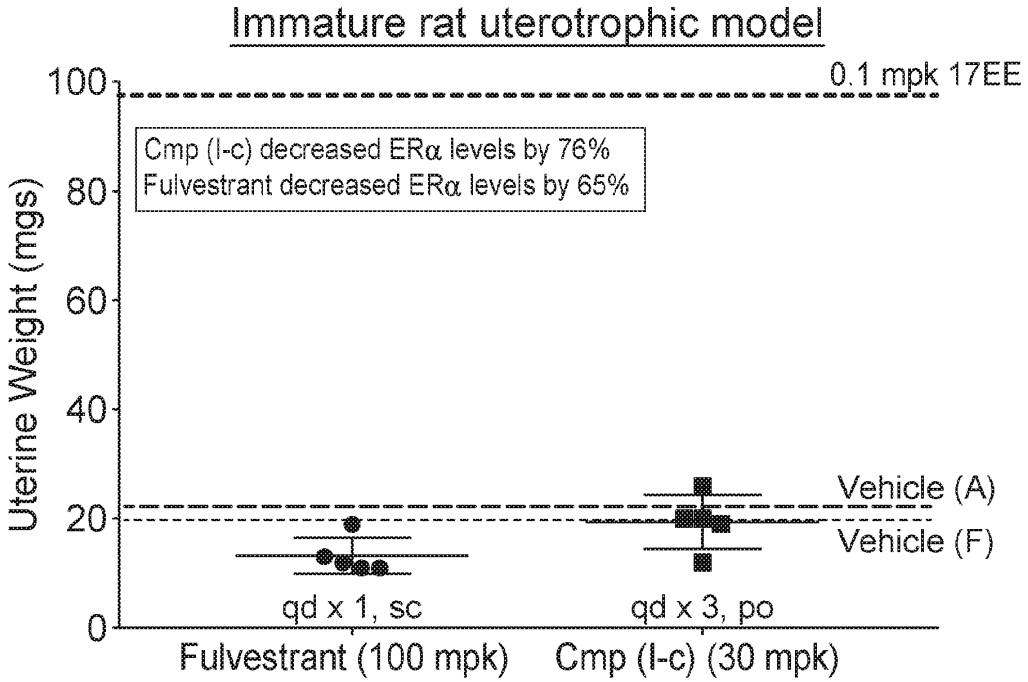

Immature rat uterotrophic model
0.1 mpk 17EE
Cmp (I-c) decreased ERα levels by 76%
Fulvestrant decreased ERα levels by 65%
Uterine Weight (mgs)
100
80
60
40
20
0
Vehicle (A)
Vehicle (F)
qd x 1, sc
qd x 3, po
Fulvestrant (100 mpk)
Cmp (I-c) (30 mpk)

FIG. 18

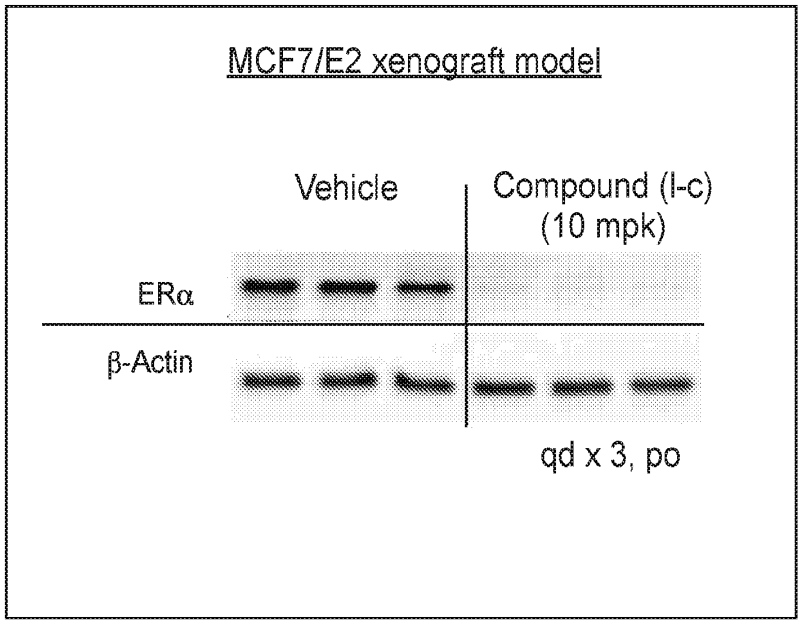

MCF7/E2 xenograft model
Vehicle
Compound (I-c)
(10 mpk)
ERα
β-Actin
qd x 3, po

FIG. 19

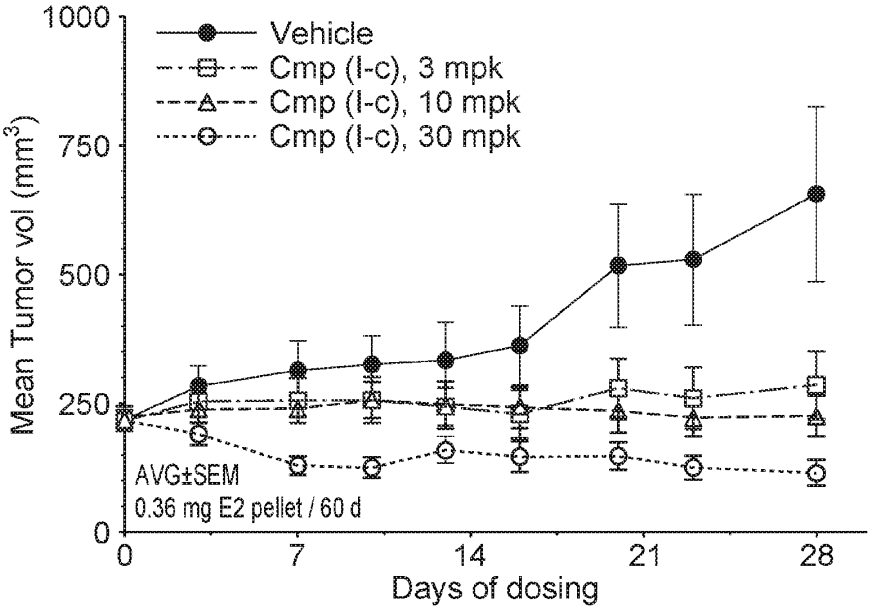

1000
750
500
250
0
Mean Tumor vol (mm3)
Vehicle
Cmp (I-c), 3 mpk
Cmp (I-c), 10 mpk
Cmp (I-c), 30 mpk
AVG±SEM
0.36 mg E2 pellet / 60 d
0
7
14
21
28
Days of dosing

FIG. 20

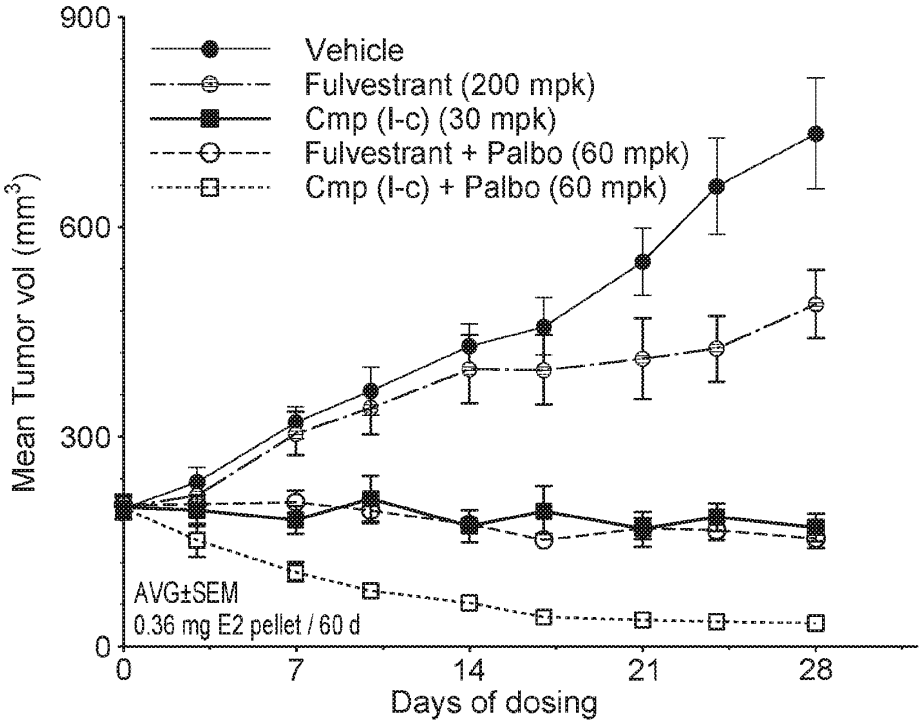

900
600
300
0
Mean Tumor vol (mm3)
Vehicle
Fulvestrant (200 mpk)
Cmp (I-c) (30 mpk)
Fulvestrant + Palbo (60 mpk)
Cmp (I-c) + Palbo (60 mpk)
AVG±SEM
0.36 mg E2 pellet / 60 d
0
7
14
21
28
Days of dosing

FIG. 21

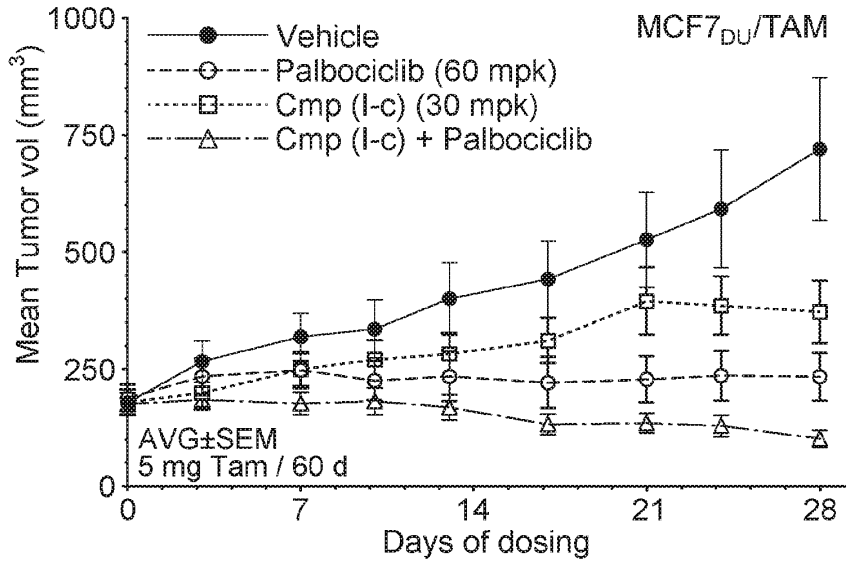
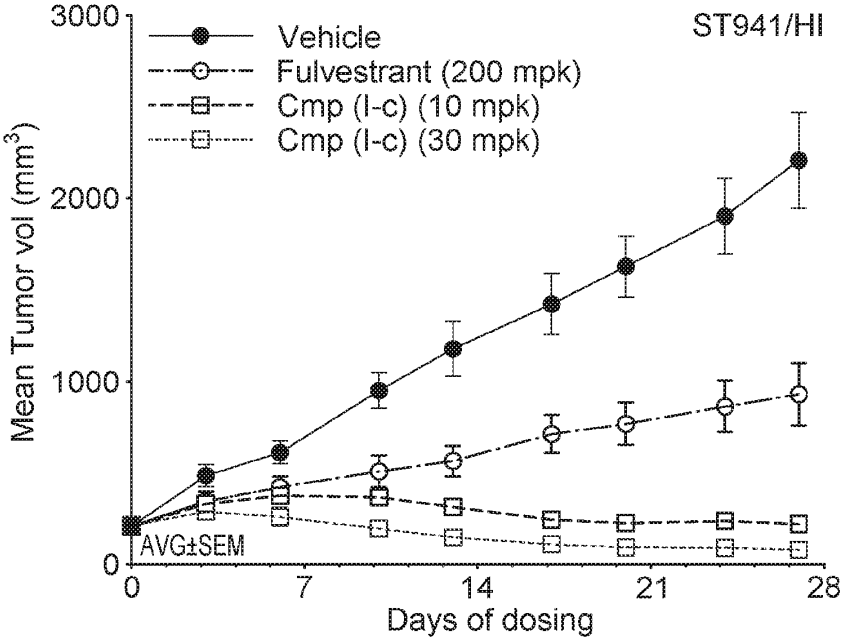

Vehicle
Palbociclib (60 mpk)
Cmp (I-c) (30 mpk)
Cmp (I-c) + Palbociclib
MCF7DU/TAM
Mean Tumor vol (mm3)
1000
750
500
250
0
AVG±SEM
5 mg Tam / 60 d
0
7
14
21
28
Days of dosing

FIG. 22


Vehicle
Fulvestrant (200 mpk)
Cmp (I-c) (10 mpk)
Cmp (I-c) (30 mpk)
ST941/HI
Mean Tumor vol (mm3)
3000
2000
1000
0
AVG±SEM
0
7
14
21
28
Days of dosing