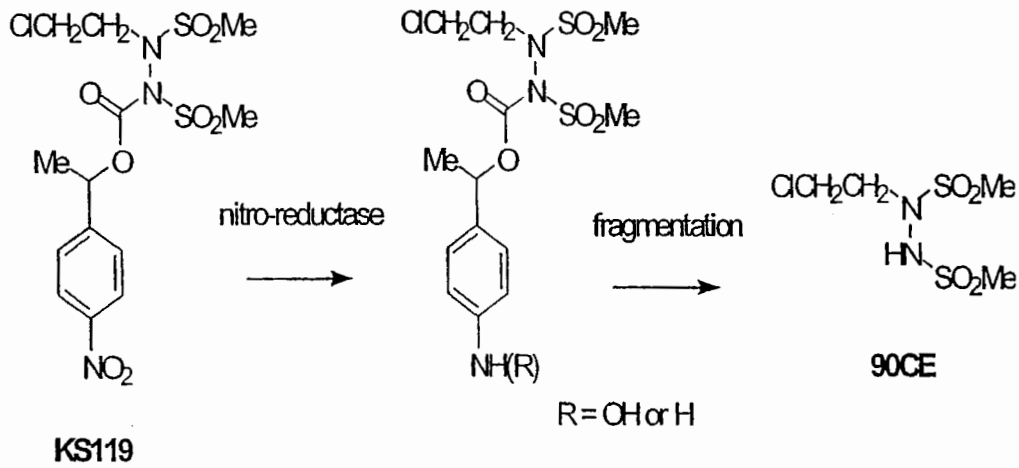
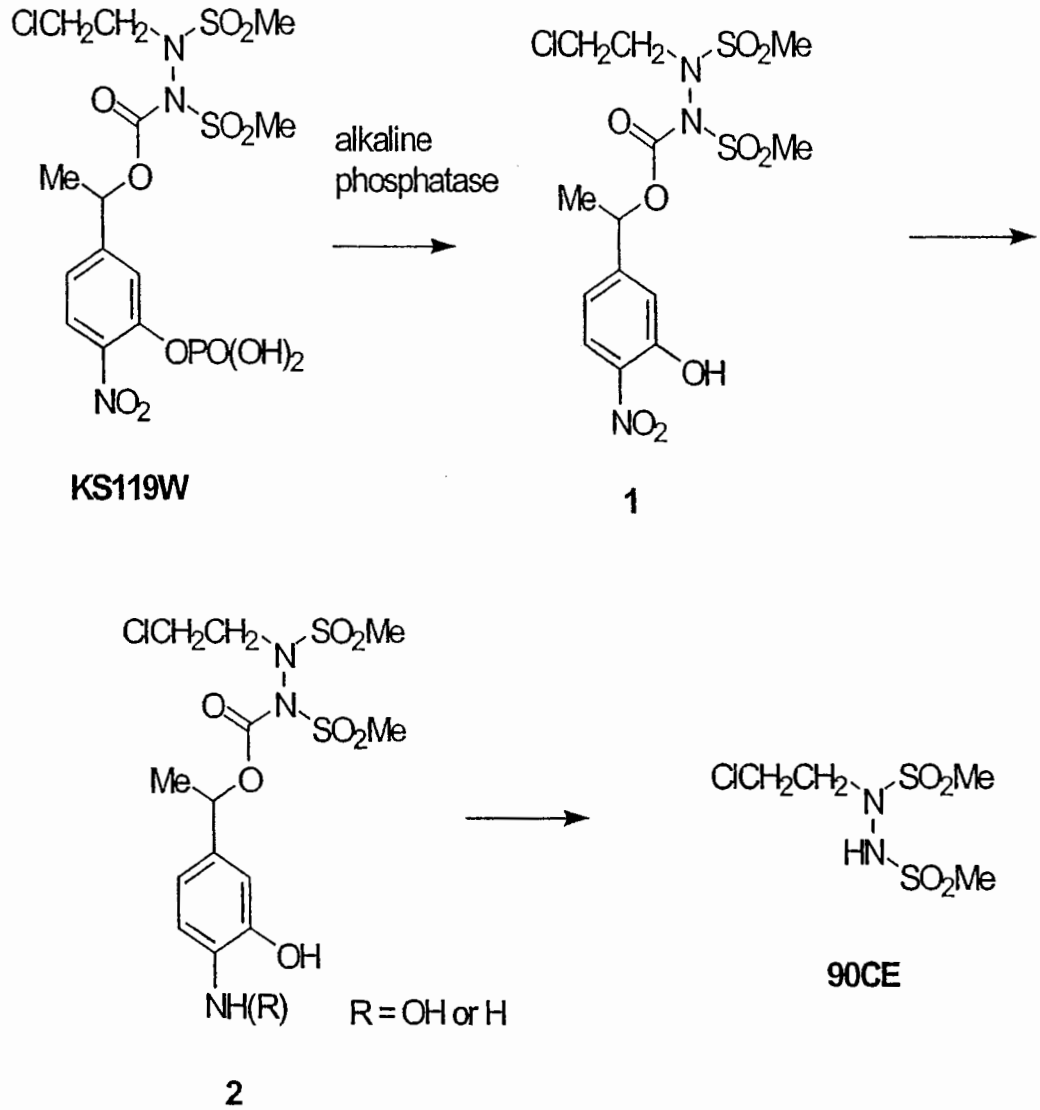
Figure 1.



KS119

90CE

Figure 2.



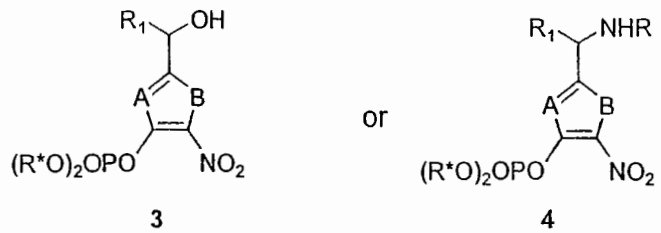
KS119W

1

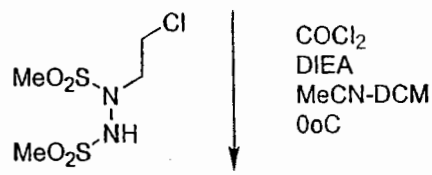
2

90CE

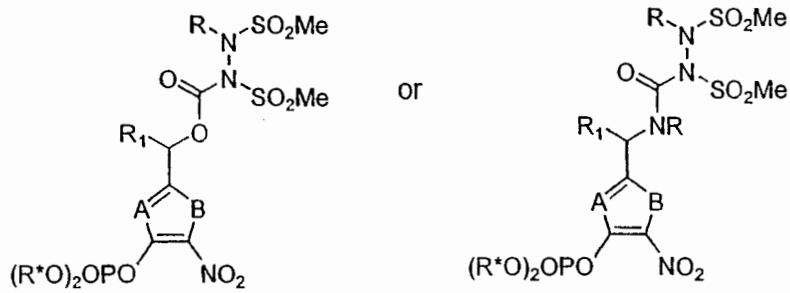
Figure 3



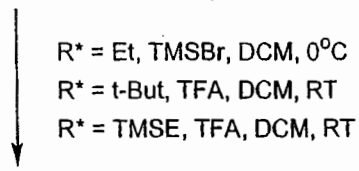
3 4

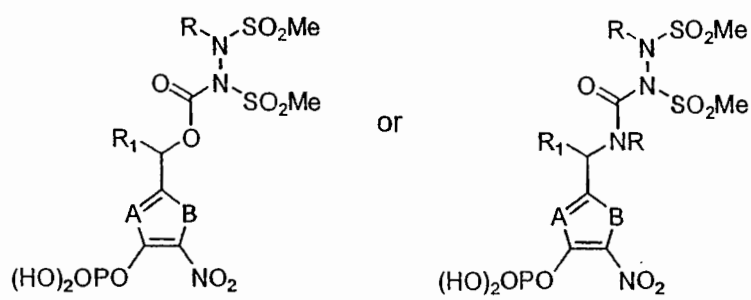


(90CE)



5 6





7 8

Figure 4

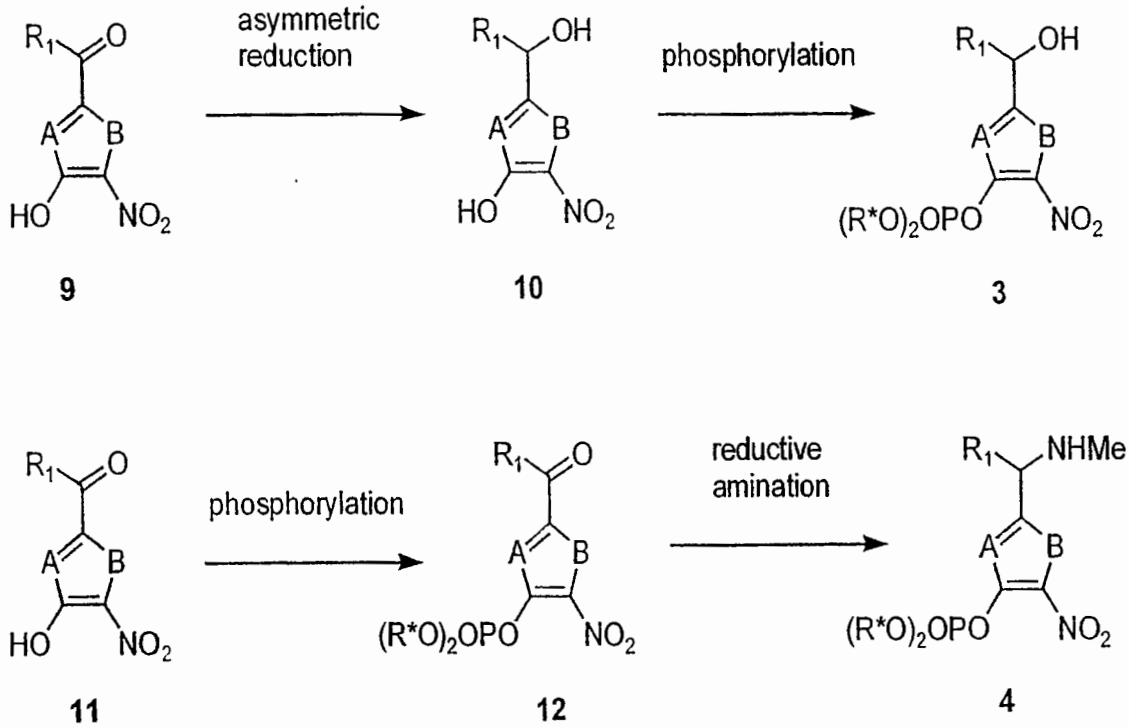

asymmetric reduction
phosphorylation
phosphorylation
reductive amination
9
10
3
11
12
4

Figure 5

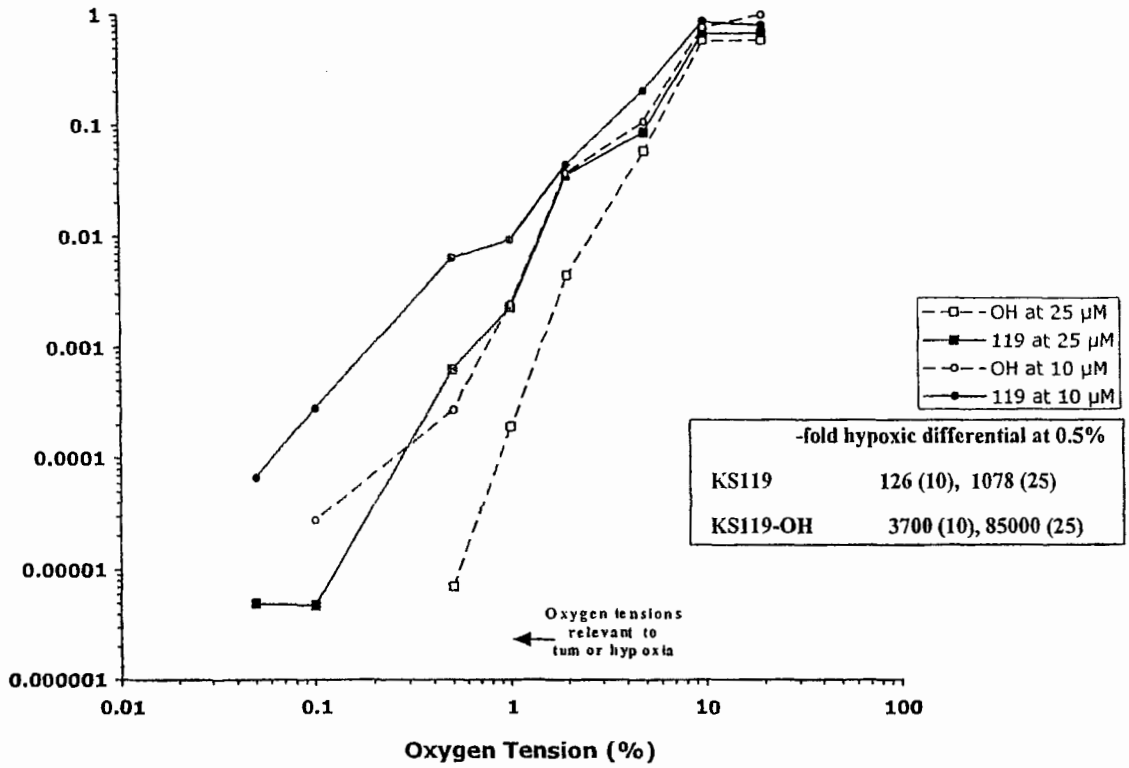

1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0.01
0.1
1
10
100
Oxygen Tension (%)
Oxygen tensions relevant to tum or hyp oxia
OH at 25 µM
119 at 25 µM
OH at 10 µM
119 at 10 µM
-fold hypoxic differential at 0.5%
KS119 126 (10), 1078 (25)
KS119-OH 3700 (10), 85000 (25)

Figure 6

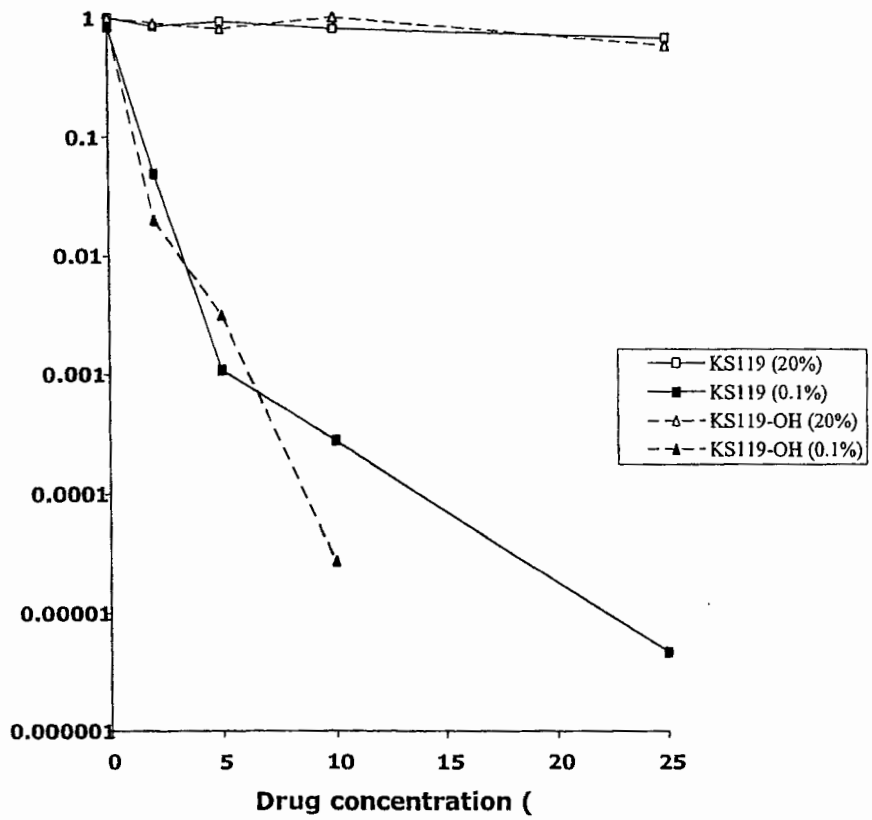

1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
5
10
15
20
25
Drug concentration (
KS119 (20%)
KS119 (0.1%)
KS119-OH (20%)
KS119-OH (0.1%)

Figure 7

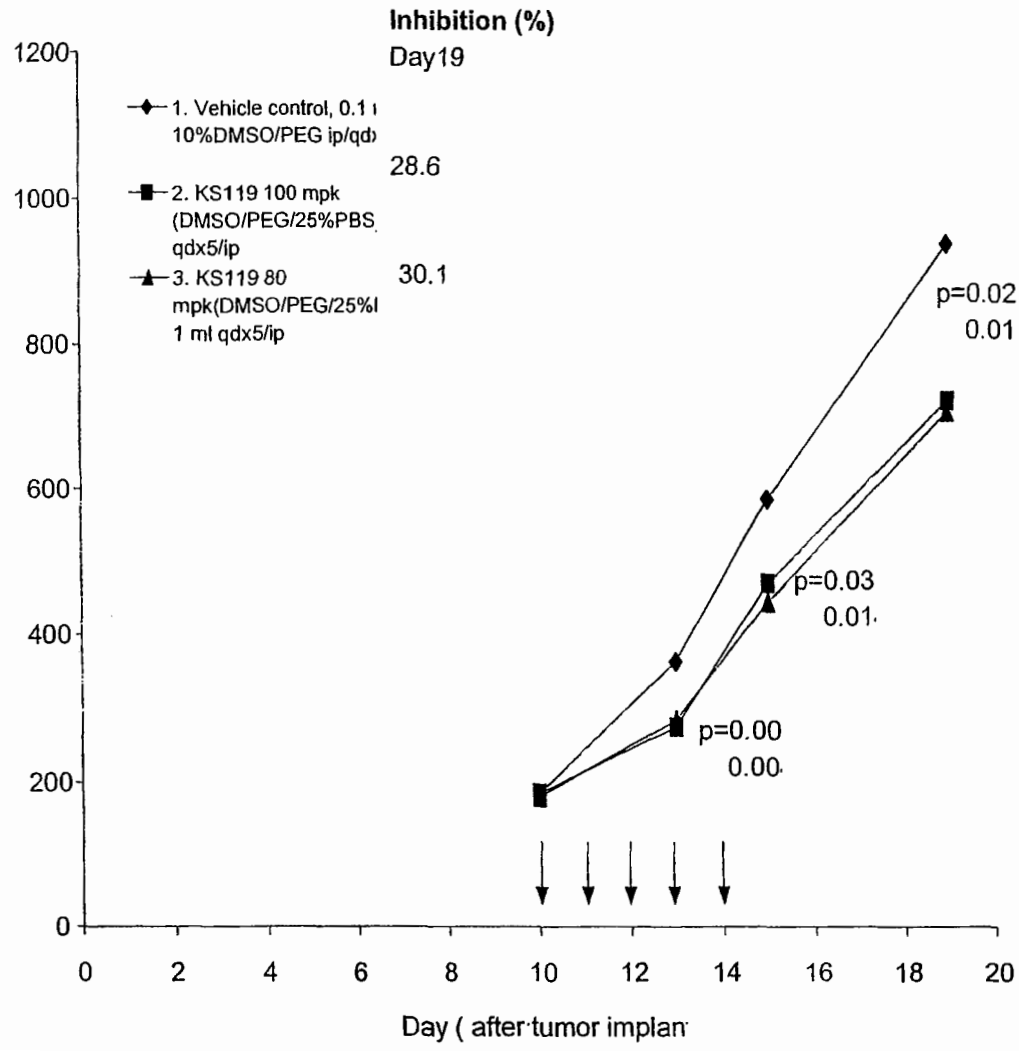

Inhibition (%)
Day19
1. Vehicle control, 0.1 r
10%DMSO/PEG ip/qd
28.6
2. KS119 100 mpk
(DMSO/PEG/25%PBS
qdx5/ip
3. KS119 80
mpk(DMSO/PEG/25%I
1 ml qdx5/ip
30.1
p=0.02
0.01
p=0.03
0.01
p=0.00
0.00
1200
1000
800
600
400
200
0
0 2 4 6 8 10 12 14 16 18 20
Day (after tumor implan

Figure 8

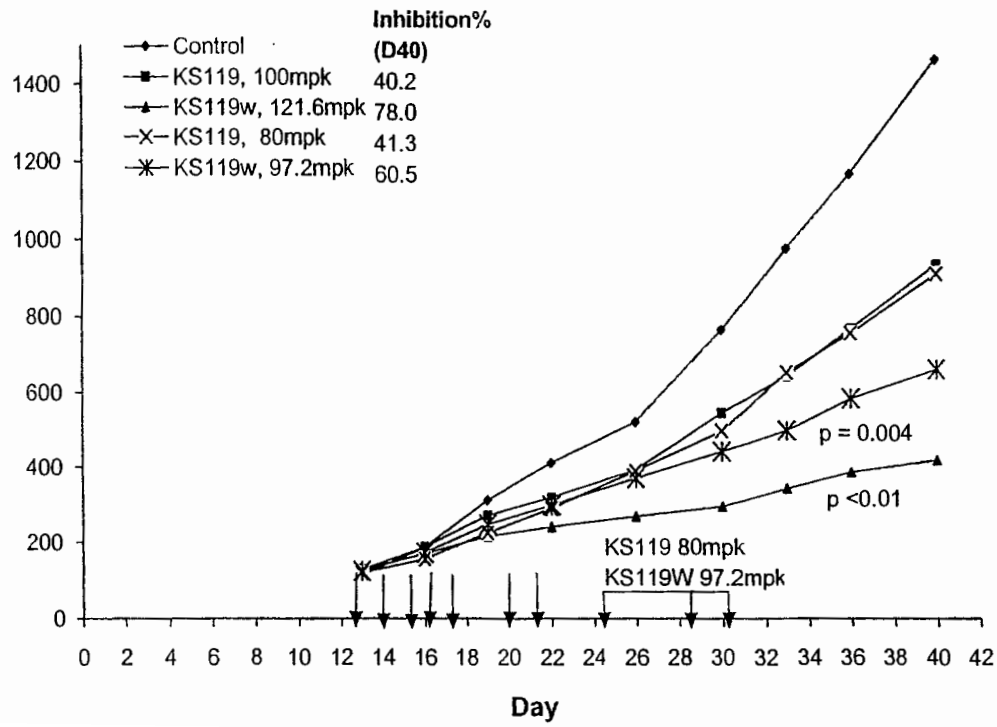
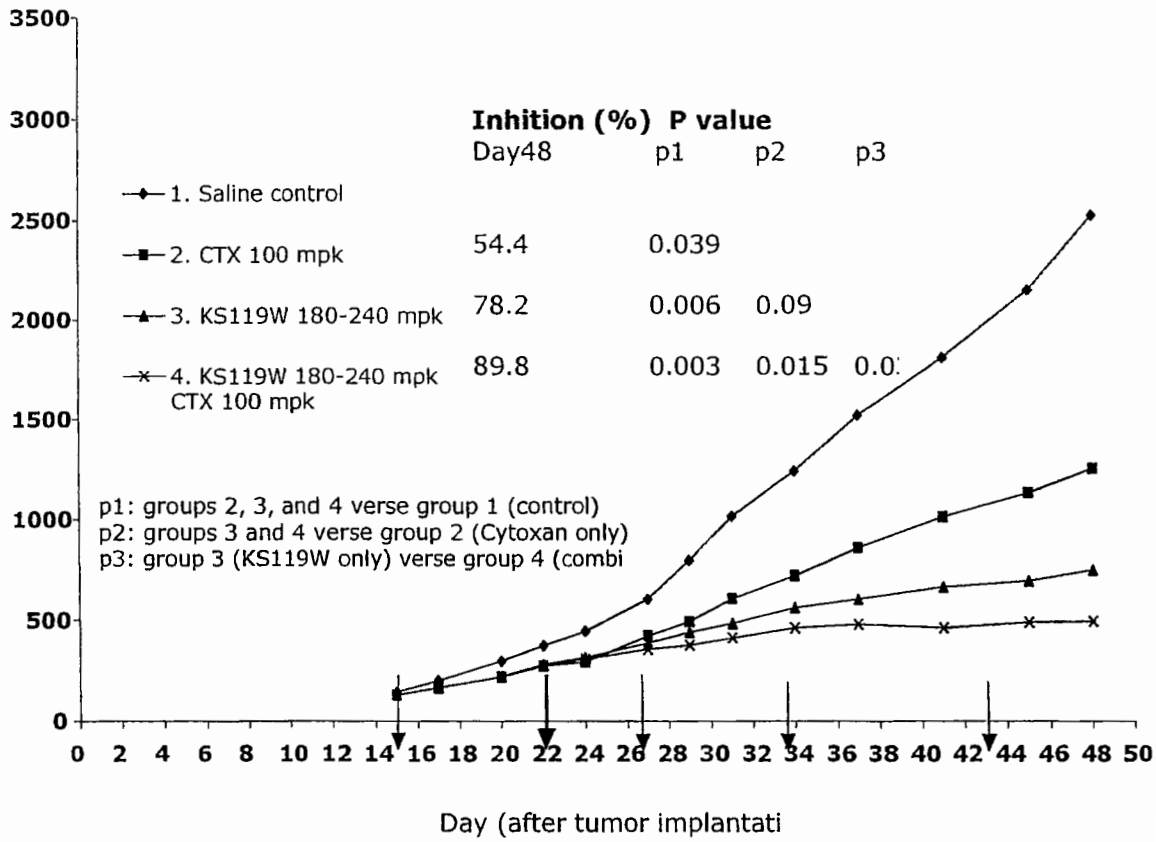

Inhibition%
(D40)
Control
KS119, 100mpk 40.2
KS119w, 121.6mpk 78.0
KS119, 80mpk 41.3
KS119w, 97.2mpk 60.5
1400
1200
1000
800
600
400
200
0
0 2 4 6 8 10 12 14 16 18 20 22 24 26 28 30 32 34 36 38 40 42
Day
p = 0.004
p <0.01
KS119 80mpk
KS119W 97.2mpk

Figure 9


3500
3000
2500
2000
1500
1000
500
0
0 2 4 6 8 10 12 14 16 18 20 22 24 26 28 30 32 34 36 38 40 42 44 46 48 50
Day (after tumor implantati
Inhition (%) P value
Day48 p1 p2 p3
1. Saline control
2. CTX 100 mpk 54.4 0.039
3. KS119W 180-240 mpk 78.2 0.006 0.09
4. KS119W 180-240 mpk CTX 100 mpk 89.8 0.003 0.015 0.0
p1: groups 2, 3, and 4 verse group 1 (control)
p2: groups 3 and 4 verse group 2 (Cytoxan only)
p3: group 3 (KS119W only) verse group 4 (combi