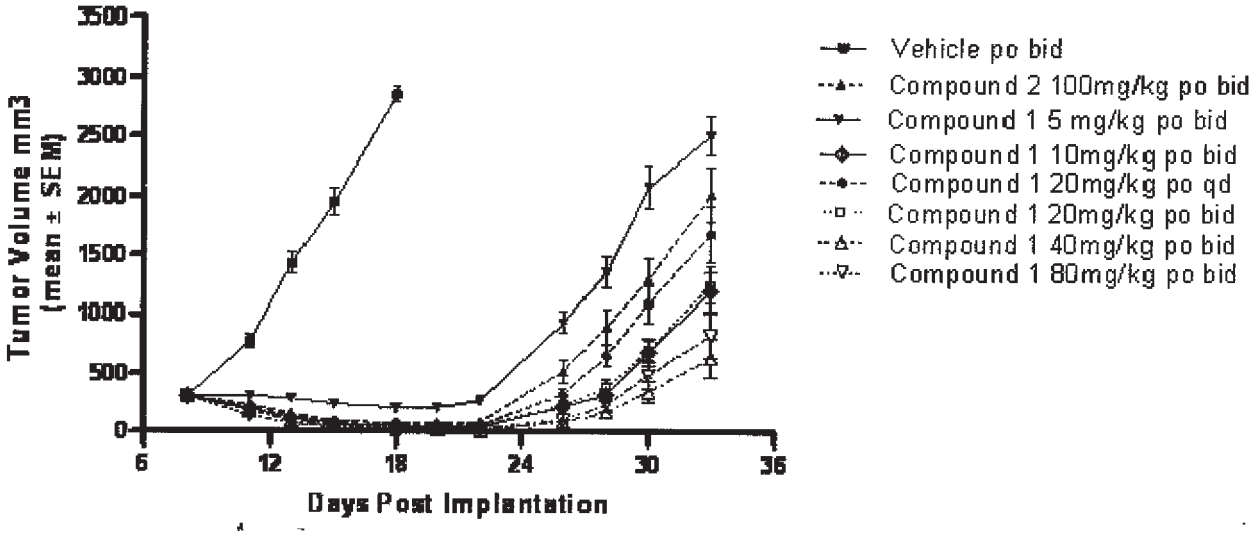

Tumor Volume mm3
(mean ± SEM)
3500
3000
2500
2000
1500
1000
500
0
6
12
18
24
30
36
Days Post Implantation
Vehicle po bid
Compound 2 100mg/kg po bid
Compound 1 5 mg/kg po bid
Compound 1 10mg/kg po bid
Compound 1 20mg/kg po qd
Compound 1 20mg/kg po bid
Compound 1 40mg/kg po bid
Compound 1 80mg/kg po bid

Figure 1

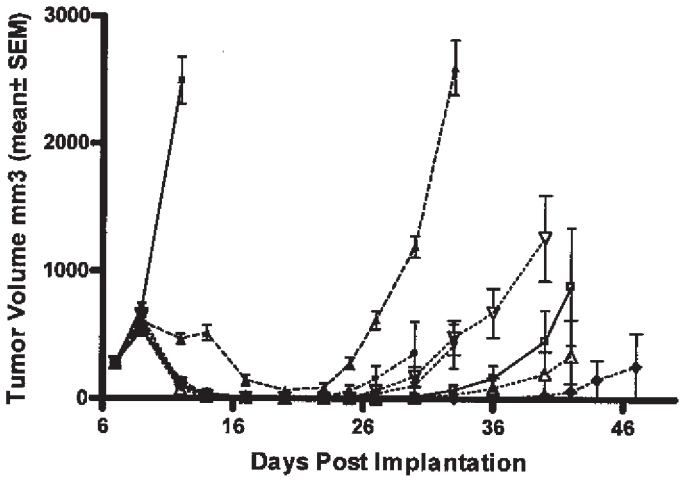

Tumor Volume mm3 (mean± SEM)
3000
2000
1000
0
6
16
26
36
46
Days Post Implantation

Figure 2

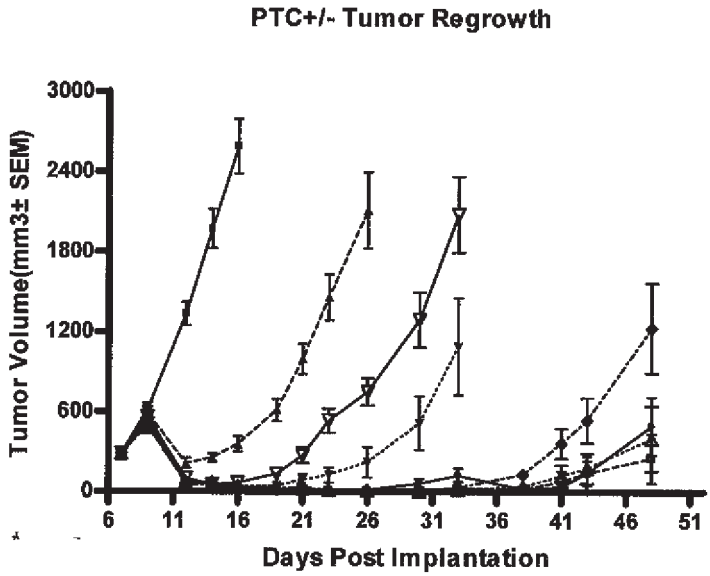

PTC+/- Tumor Regrowth
Tumor Volume(mm3± SEM)
3000
2400
1800
1200
600
0
6
11
16
21
26
31
36
41
46
51
Days Post Implantation

Figure 3

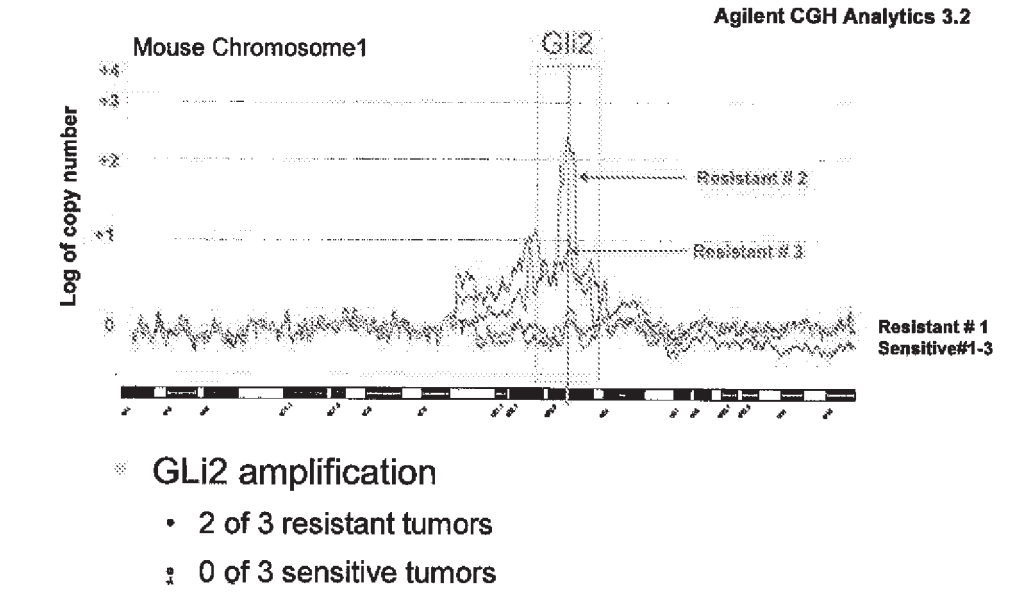

Agilent CGH Analytics 3.2
Mouse Chromosome1
Gli2
Log of copy number
+4
+3
+2
+1
0
Resistant # 2
Resistant # 3
Resistant # 1
Sensitive#1-3
GLi2 amplification
2 of 3 resistant tumors
0 of 3 sensitive tumors

Figure 4

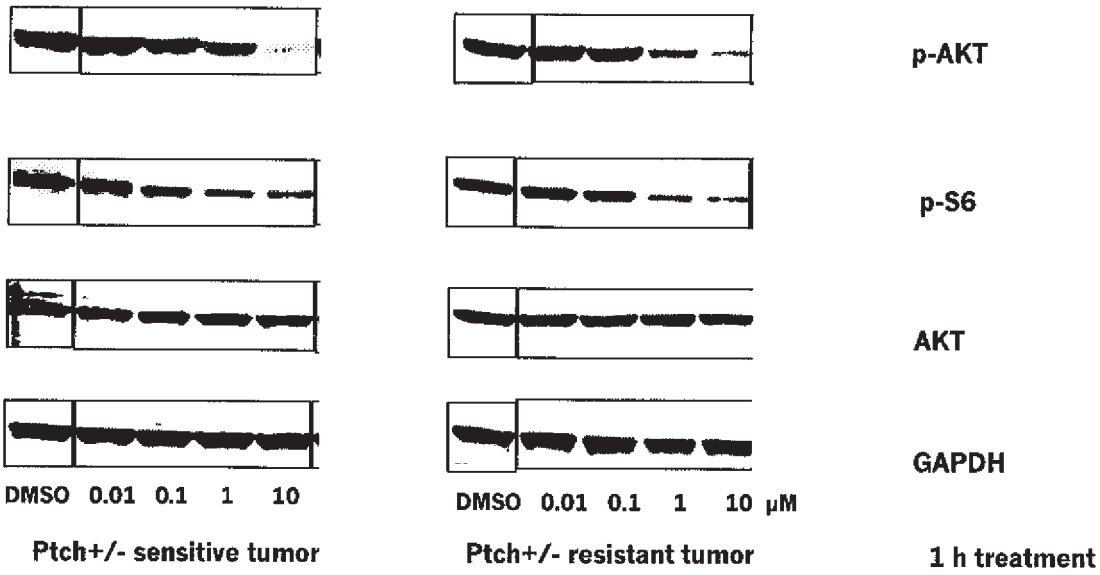

p-AKT
p-S6
AKT
GAPDH
DMSO 0.01 0.1 1 10
DMSO 0.01 0.1 1 10 µM
Ptch+/- sensitive tumor
Ptch+/- resistant tumor
1 h treatment

Figure 5

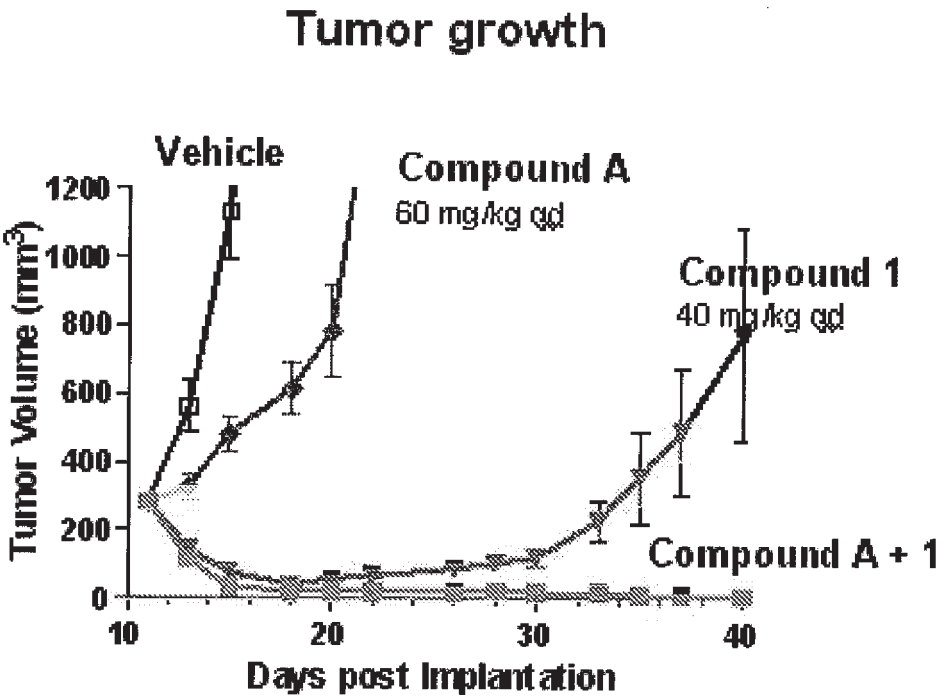


Figure 6A

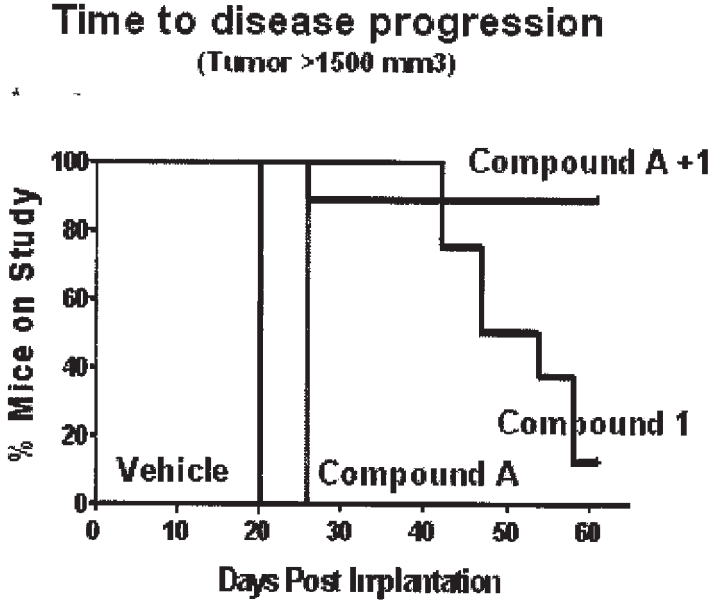


Figure 6B

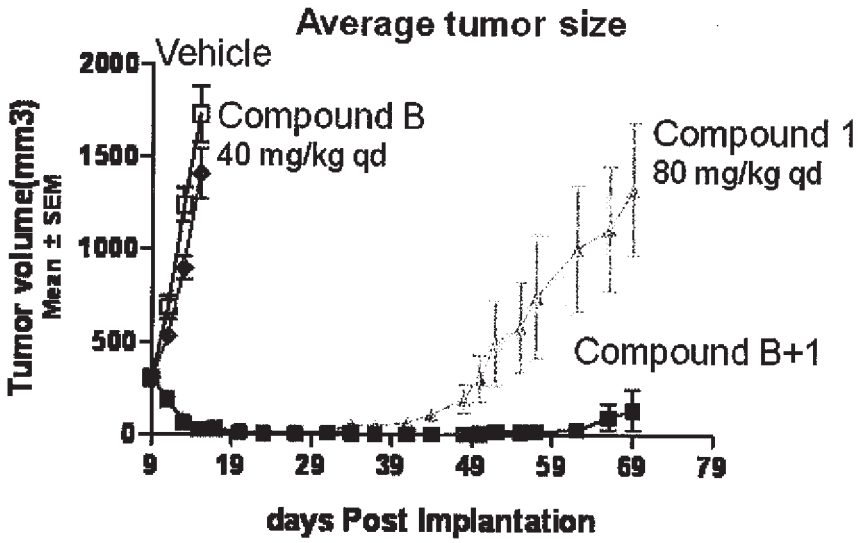


Figure 7A

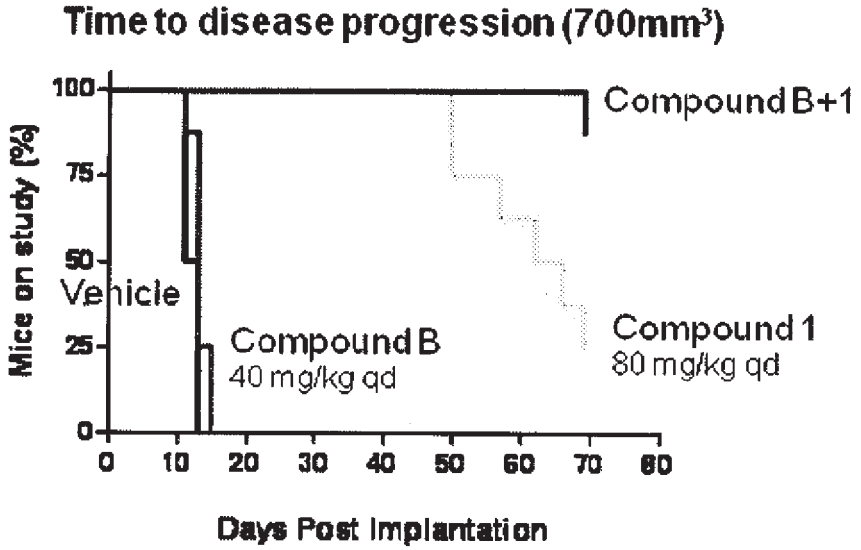


Figure 7B