Figure 1

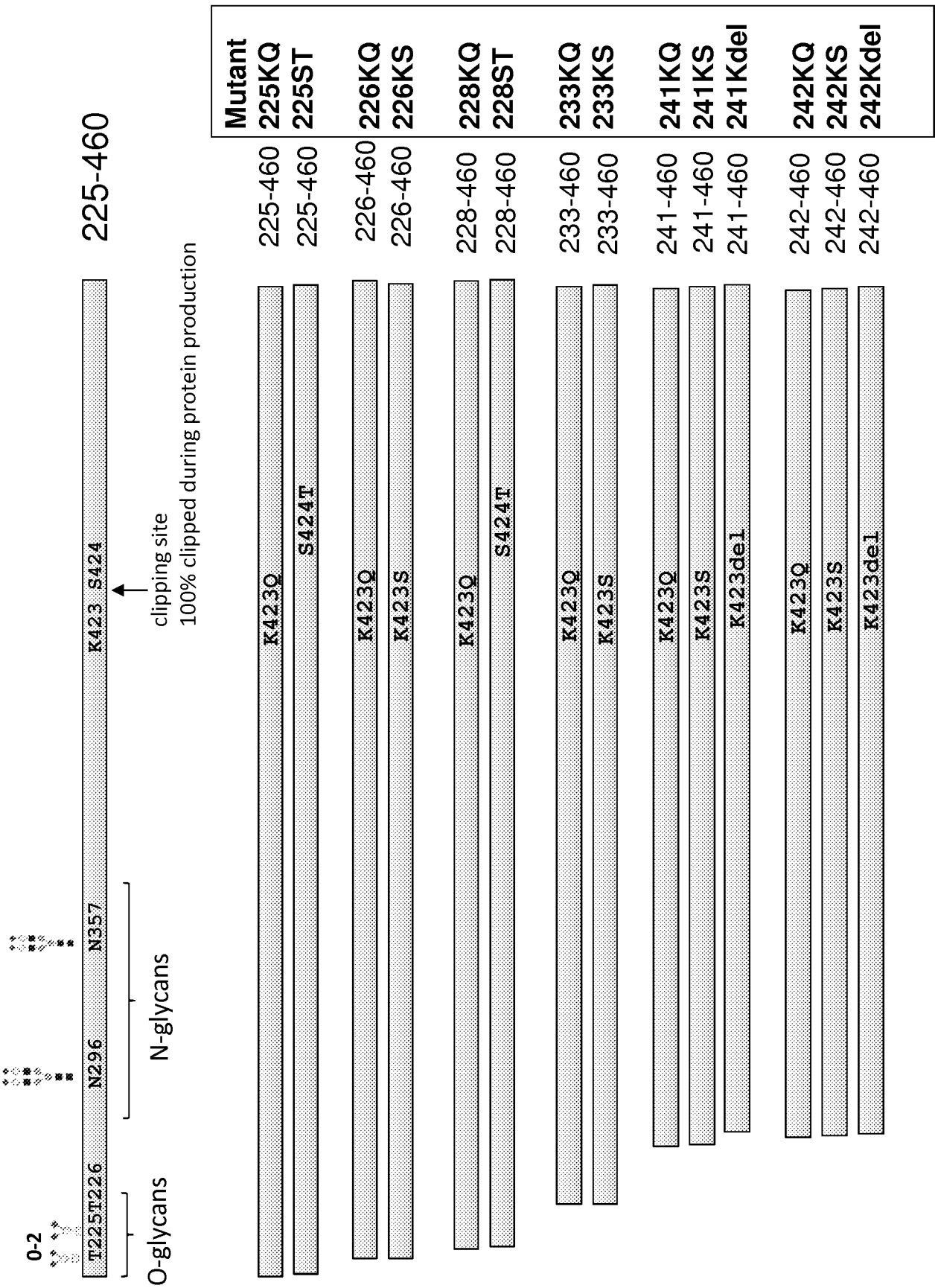
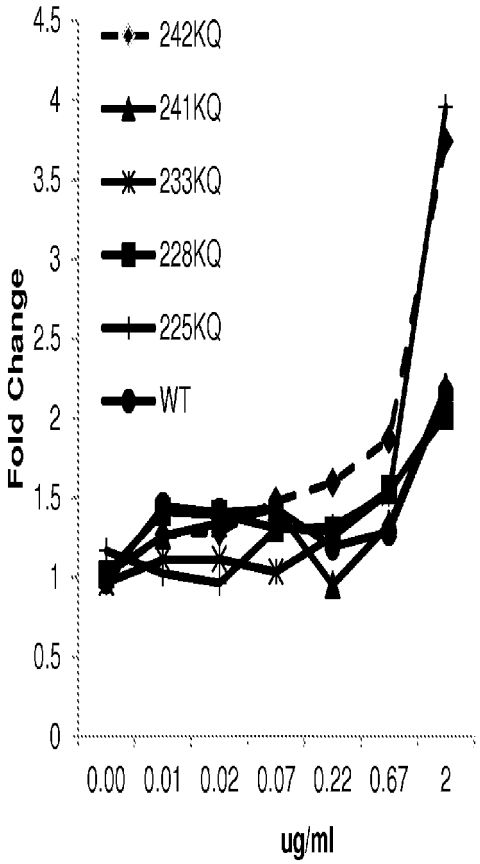
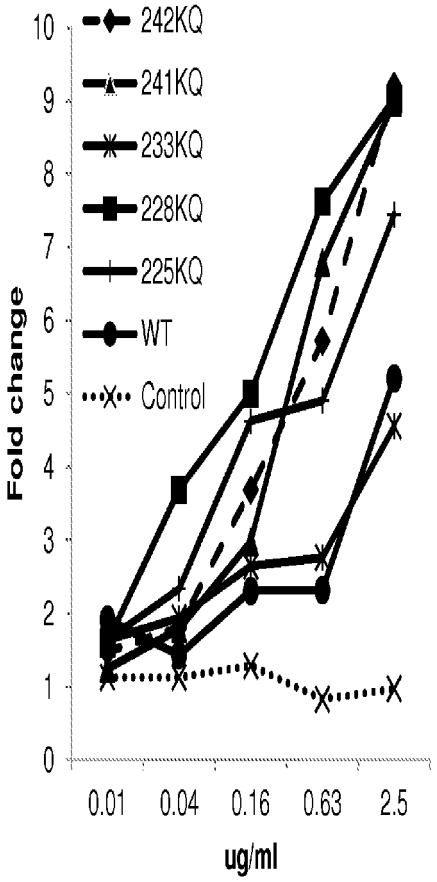

225-460
0-2
T225T226
N296
N357
K423 S424
clipping site
100% clipped during protein production
O-glycans
N-glycans
Mutant
225KQ
225ST
226KQ
226KS
228KQ
228ST
233KQ
233KS
241KQ
241KS
241Kdel
242KQ
242KS
242Kdel
225-460
225-460
226-460
226-460
228-460
228-460
233-460
233-460
241-460
241-460
241-460
242-460
242-460
242-460
K423Q
S424T
K423Q
K423S
K423Q
S424T
K423Q
K423S
K423Q
K423S
K423del
K423Q
K423S
K423del

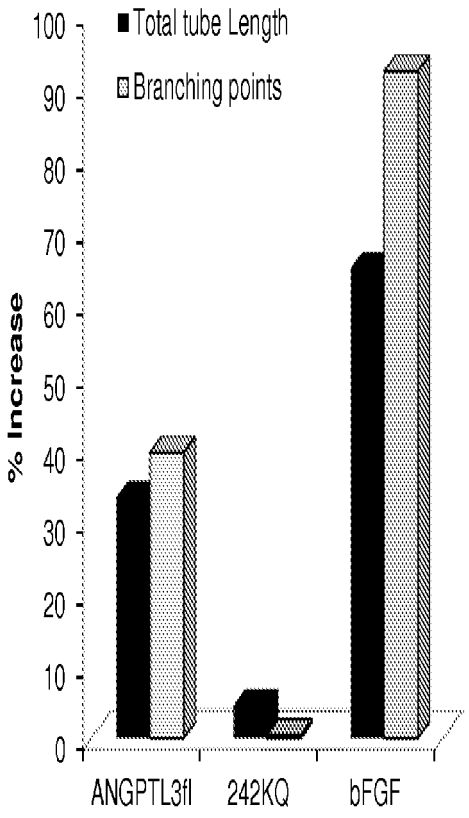
FIG 2


4.5
4
3.5
3
2.5
2
1.5
1
0.5
0
Fold Change
242KQ
241KQ
233KQ
228KQ
225KQ
WT
0.00 0.01 0.02 0.07 0.22 0.67 2
ug/ml

2A

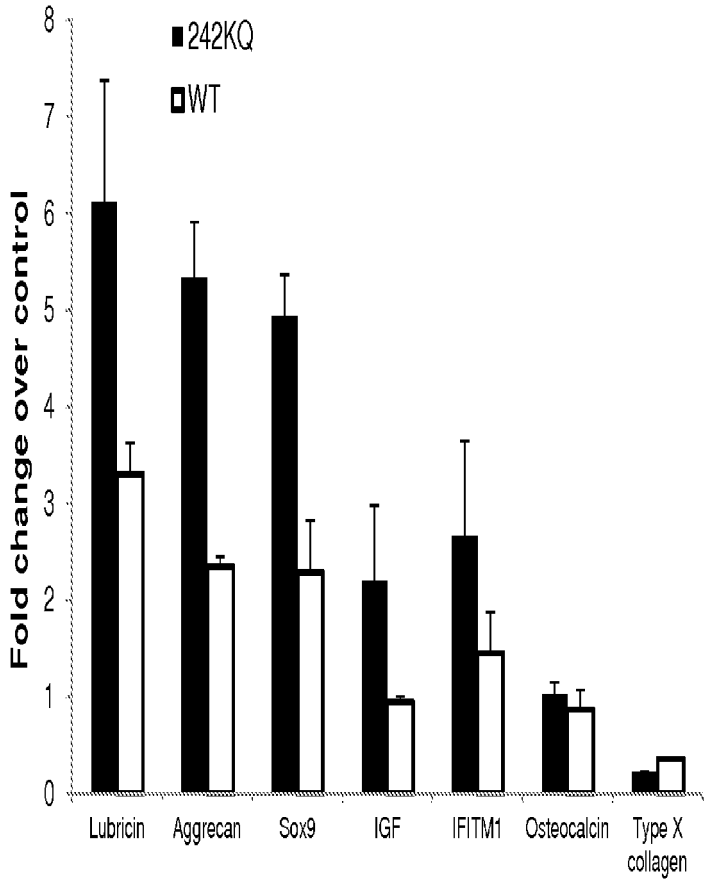

10
9
8
7
6
5
4
3
2
1
0
Fold change
242KQ
241KQ
233KQ
228KQ
225KQ
WT
Control
0.01 0.04 0.16 0.63 2.5
ug/ml

2B

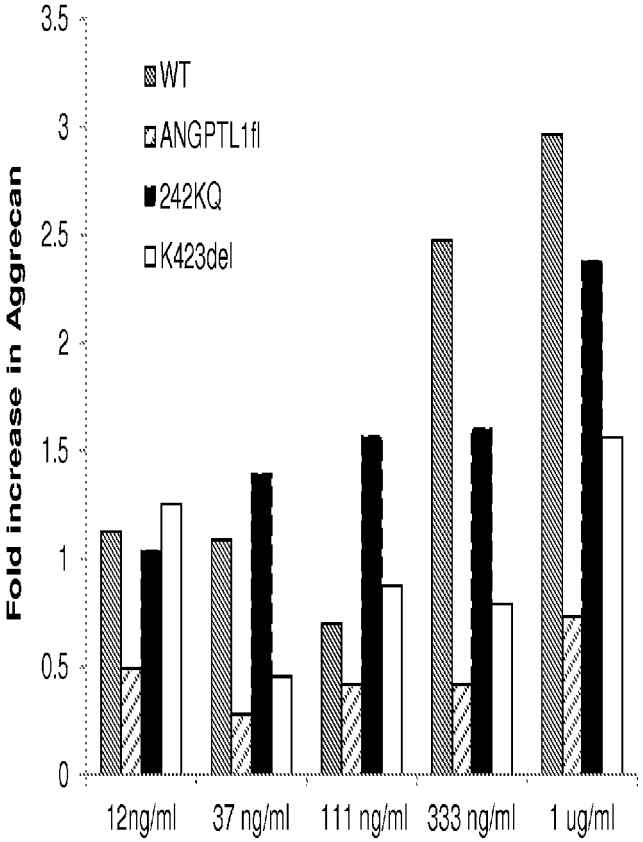

100
90
80
70
60
50
40
30
20
10
0
% Increase
Total tube Length
Branching points
ANGPTL3fl
242KQ
bFGF

2C

Figure 3

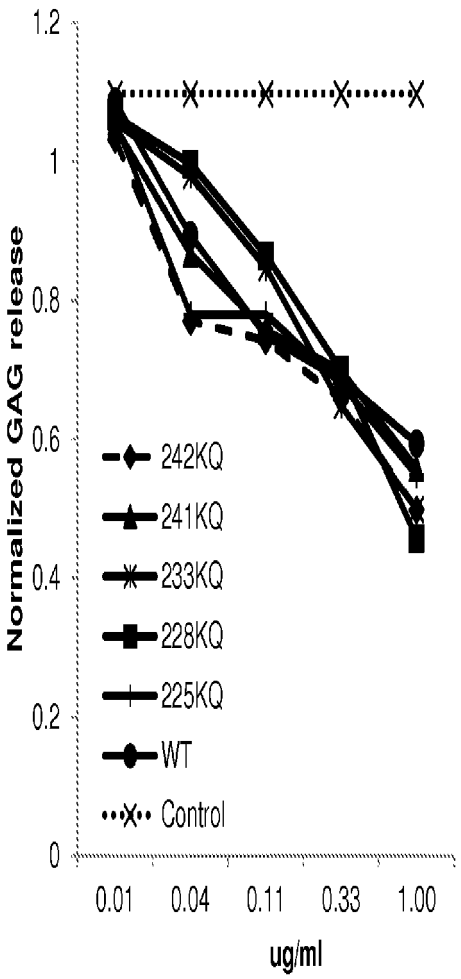


3A

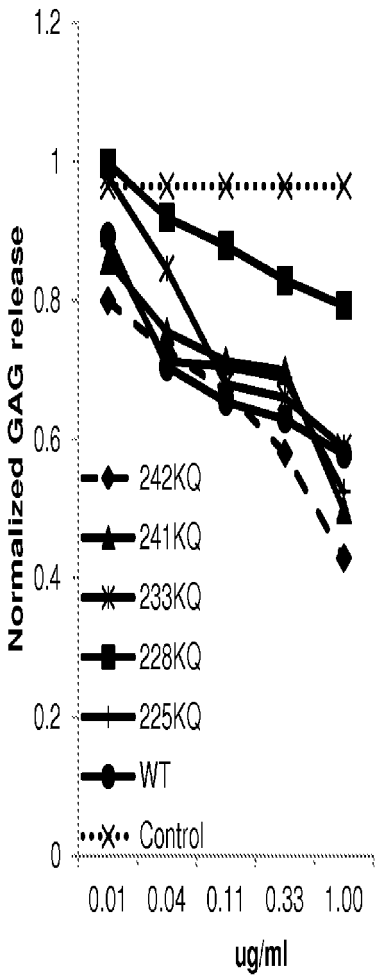


3B

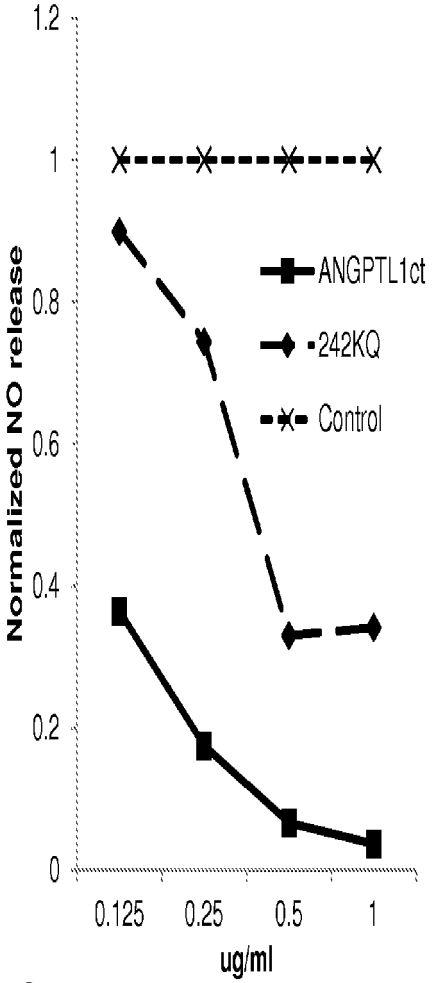
Figure 4



4A

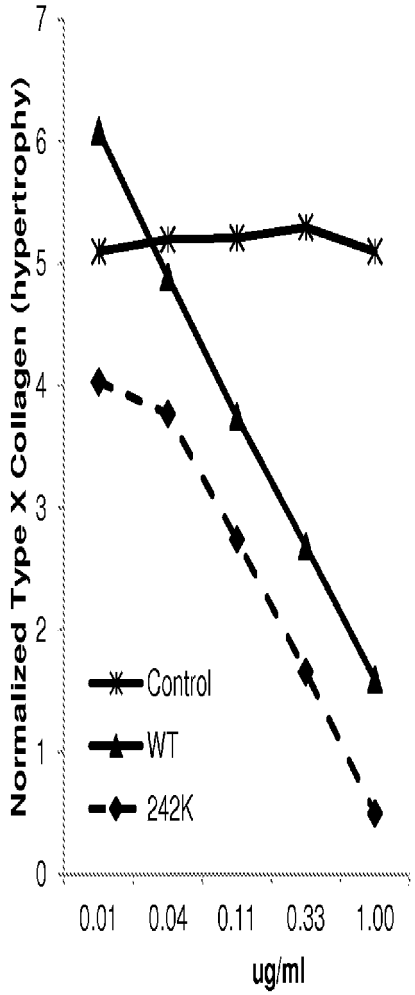


4B

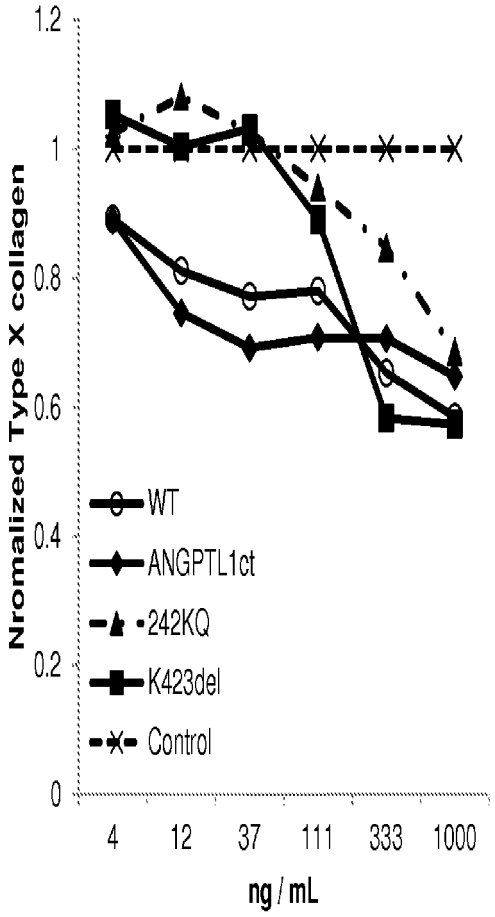


4C

Figure 5

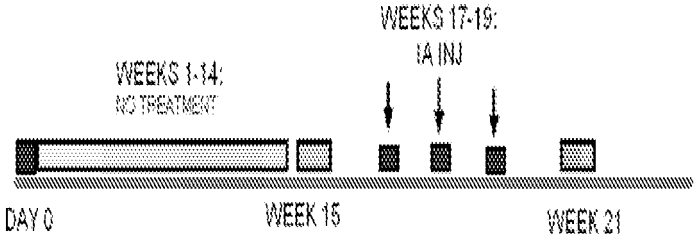


5A

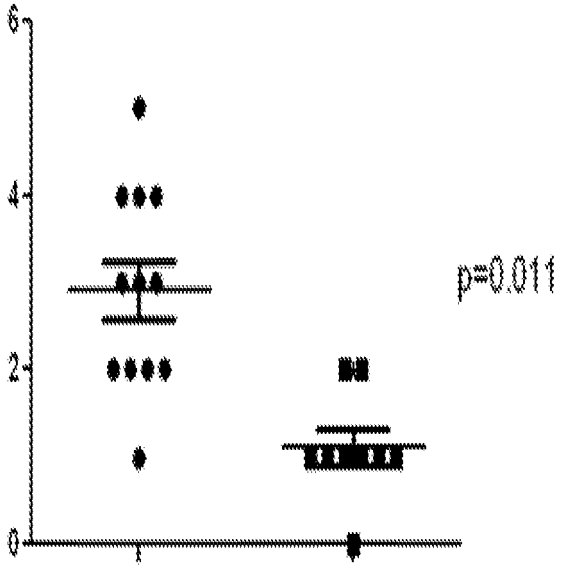


5B

Figure 6



6A



6B

Figure 7

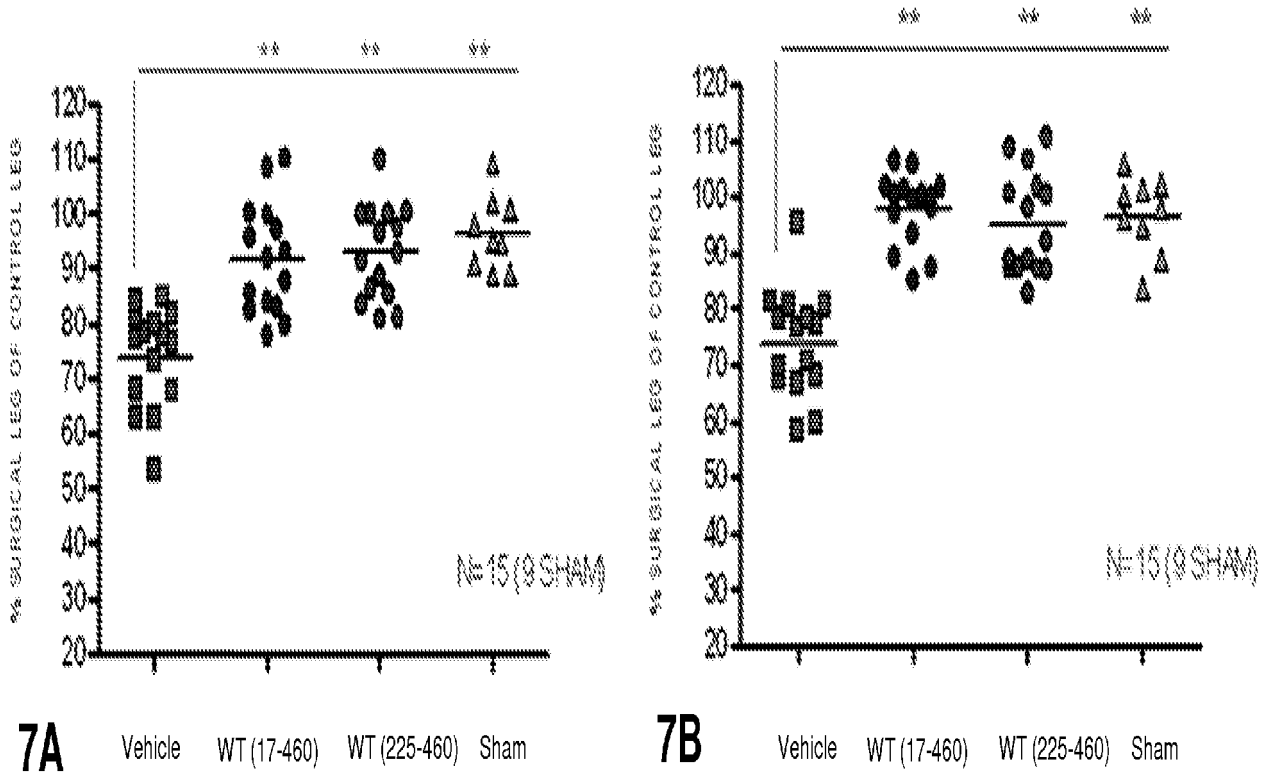

% SURGICAL LEG OF CONTROL LEG
120
110
100
90
80
70
60
50
40
30
20
**
**
**
N=15 (9 SHAM)
7A
Vehicle
WT (17-460)
WT (225-460)
Sham
% SURGICAL LEG OF CONTROL LEG
120
110
100
90
80
70
60
50
40
30
20
**
**
**
N=15 (9 SHAM)
7B
Vehicle
WT (17-460)
WT (225-460)
Sham

Figure 8

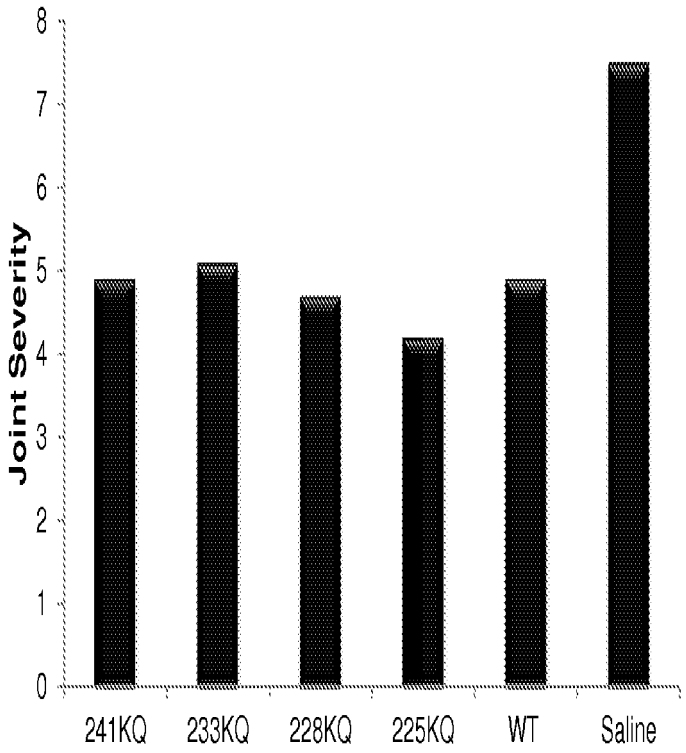

Joint Severity
0
1
2
3
4
5
6
7
8
241KQ
233KQ
228KQ
225KQ
WT
Saline

Figure 9

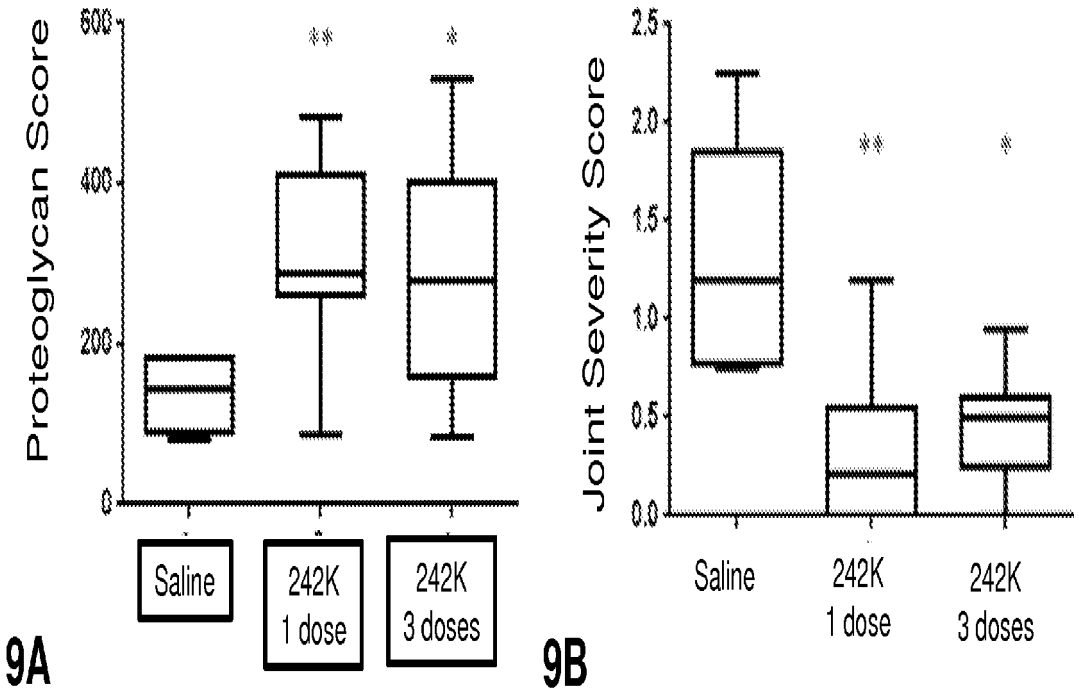
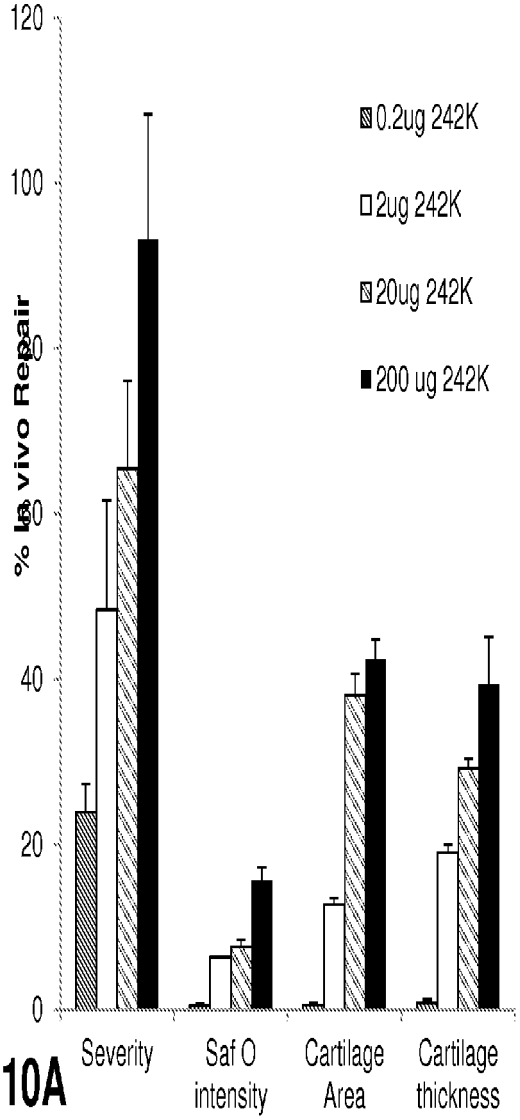
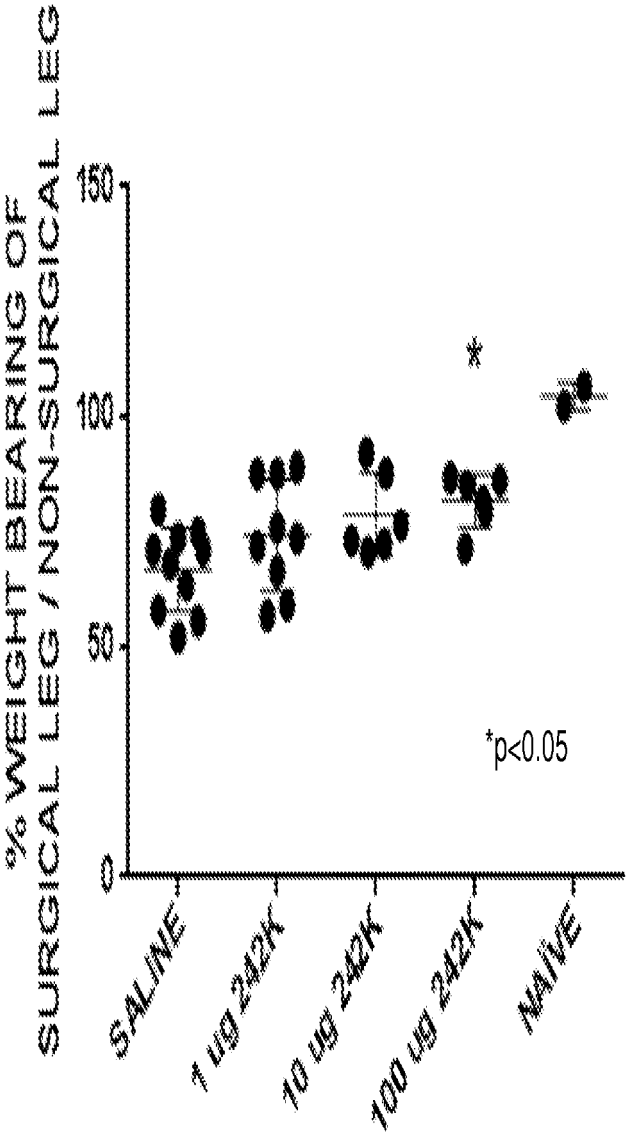

Proteoglycan Score
600
400
200
0
**
*
Saline
242K
1 dose
242K
3 doses
9A
Joint Severity Score
2.5
2.0
1.5
1.0
0.5
0.0
**
*
Saline
242K
1 dose
242K
3 doses
9B

Figure 10

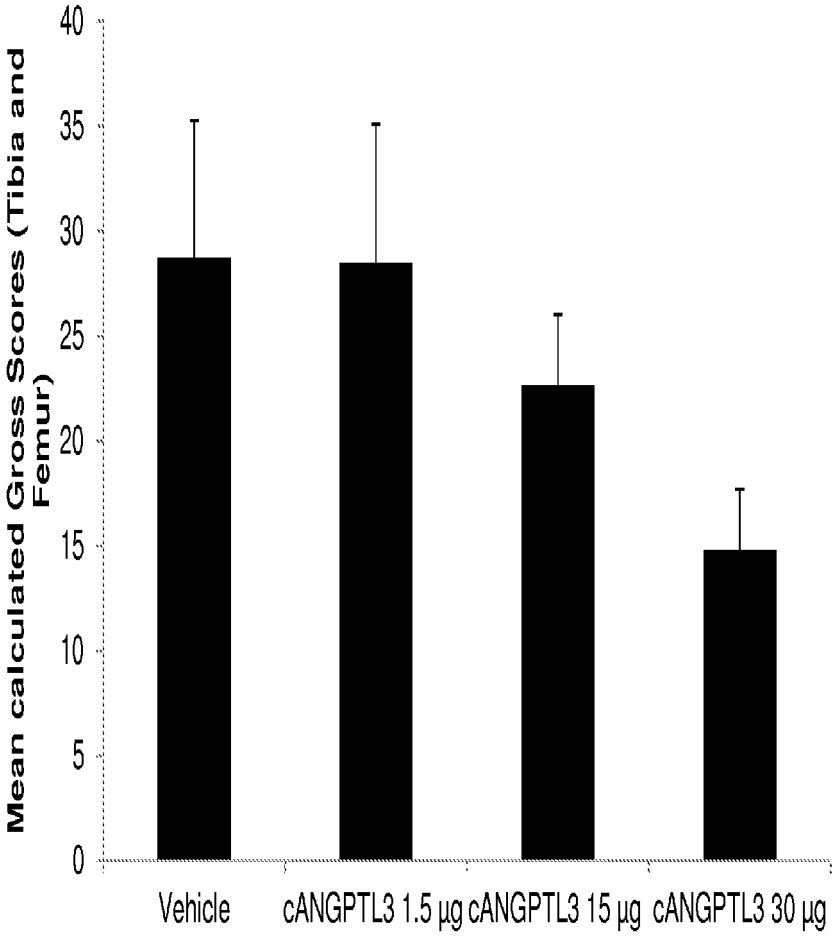


10A



10B

Figure 11


Mean calculated Gross Scores (Tibia and Femur)
40
35
30
25
20
15
10
5
0
Vehicle
cANGPTL3 1.5 µg
cANGPTL3 15 µg
cANGPTL3 30 µg