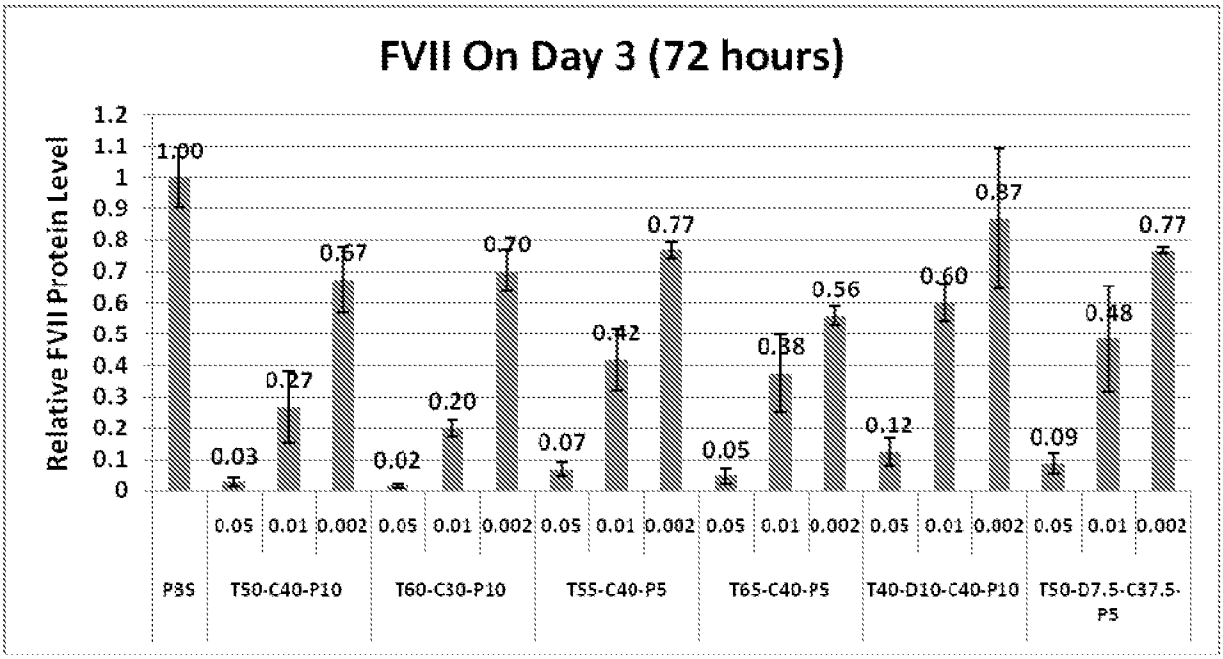

FVII On Day 3 (72 hours)
Relative FVII Protein Level
1.2
1.1
1
0.9
0.8
0.7
0.6
0.5
0.4
0.3
0.2
0.1
0
1.00
0.03
0.27
0.67
0.02
0.20
0.70
0.07
0.42
0.77
0.05
0.38
0.56
0.12
0.60
0.87
0.09
0.48
0.77
0.05
0.01
0.002
PBS
T50-C40-P10
T60-C30-P10
T55-C40-P5
T65-C40-P5
T40-D10-C40-P10
T50-D7.5-C37.5-P5

FIG. 1

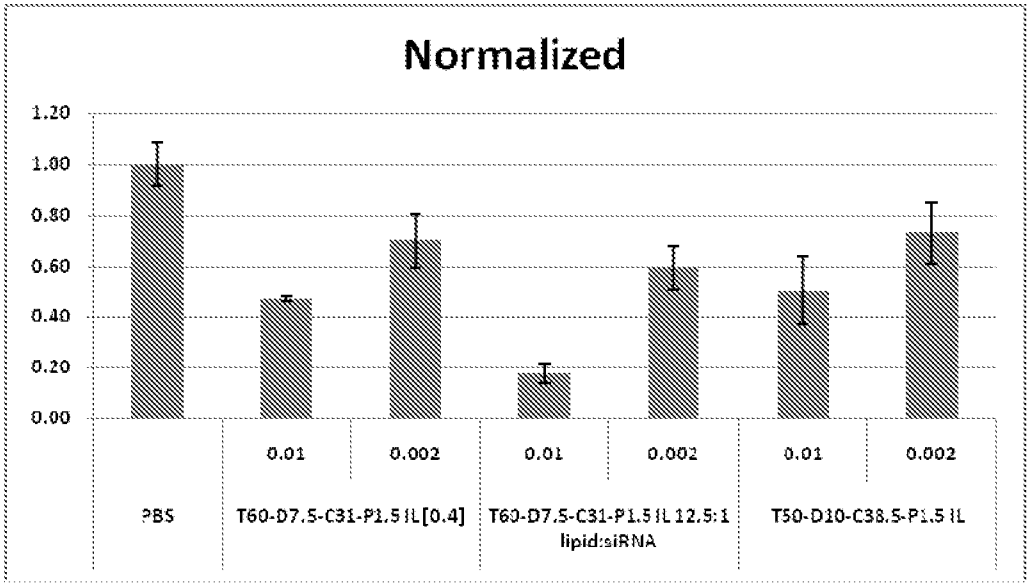

Normalized
1.20
1.00
0.80
0.60
0.40
0.20
0.00
0.01
0.002
0.01
0.002
0.01
0.002
PBS
T60-D7.5-C31-P1.5 iL[0.4]
T60-D7.5-C31-P1.5 iL 12.5:1 lipid:siRNA
T50-D10-C38.5-P1.5 iL

FIG. 2

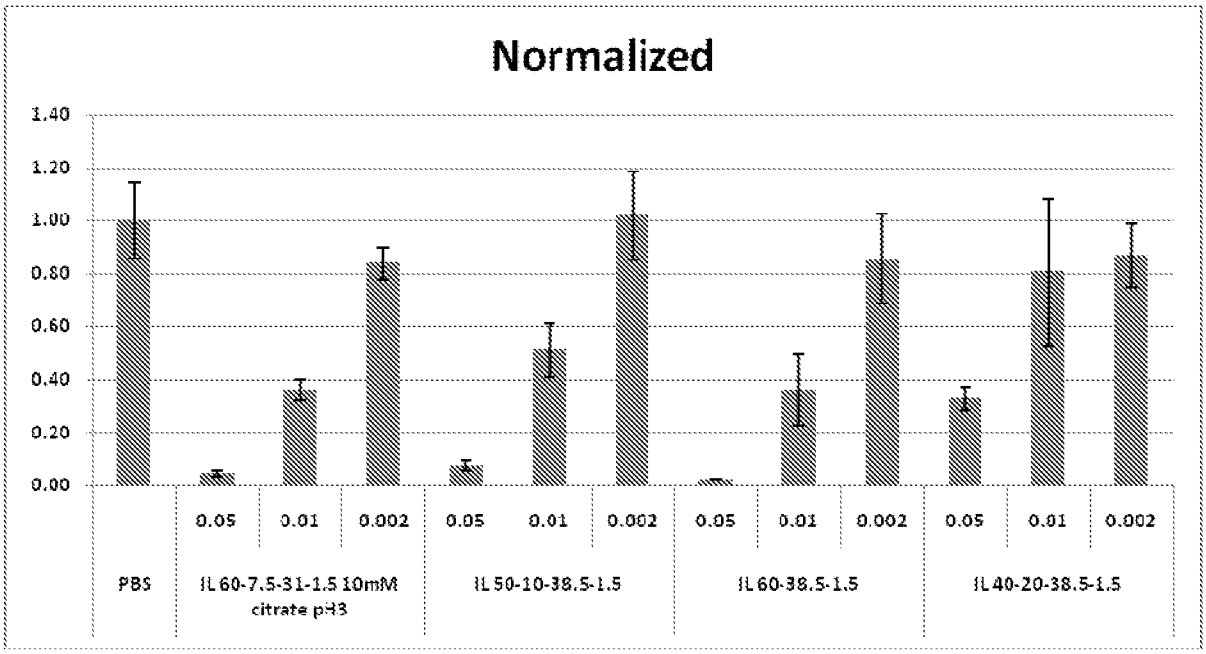

Normalized
1.40
1.20
1.00
0.80
0.60
0.40
0.20
0.00
0.05
0.01
0.002
PBS
IL60-7.5-31-1.5 10mM citrate pH3
IL50-10-38.5-1.5
IL60-38.5-1.5
IL40-20-38.5-1.5

FIG. 3

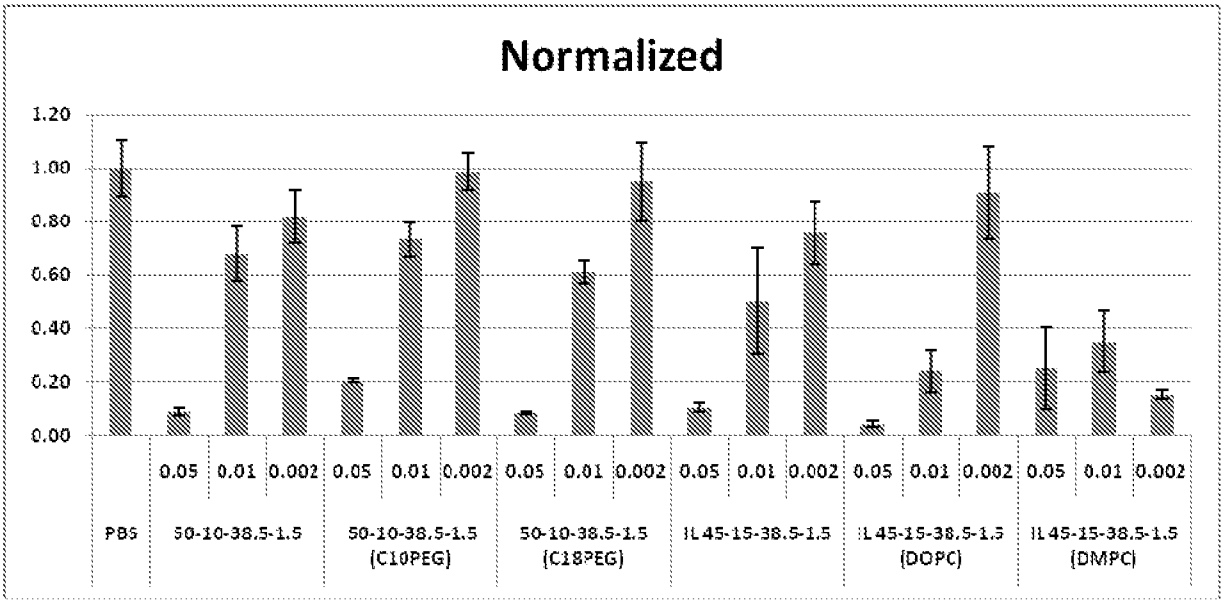

Normalized
1.20
1.00
0.80
0.60
0.40
0.20
0.00
0.05
0.01
0.002
PBS
50-10-38.5-1.5
50-10-38.5-1.5 (C10PEG)
50-10-38.5-1.5 (C18PEG)
IL45-15-38.5-1.5
IL45-15-38.5-1.5 (DOPC)
IL45-15-38.5-1.5 (DMPC)

FIG. 4

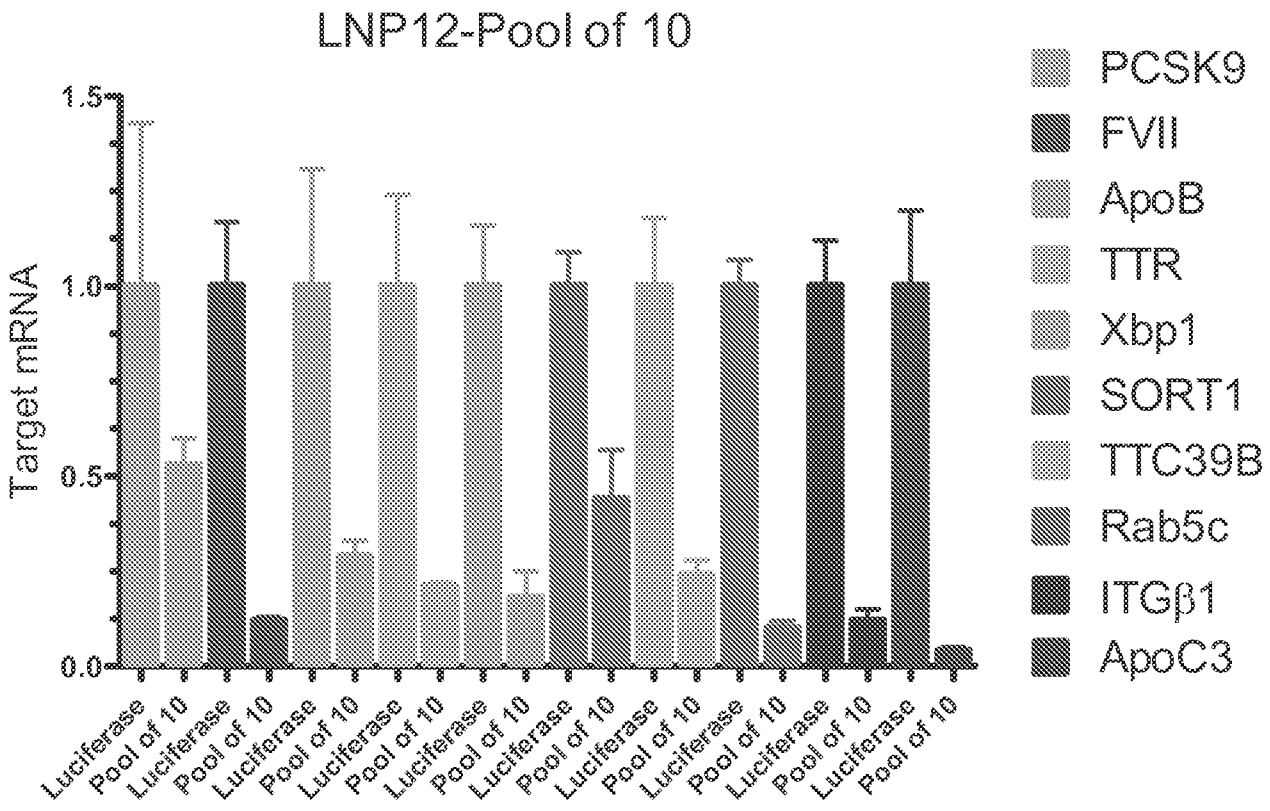

LNP12-Pool of 10
Target mRNA
1.5
1.0
0.5
0.0
Luciferase
Pool of 10
PCSK9
FVII
ApoB
TTR
Xbp1
SORT1
TTC39B
Rab5c
ITGβ1
ApoC3

FIG. 5

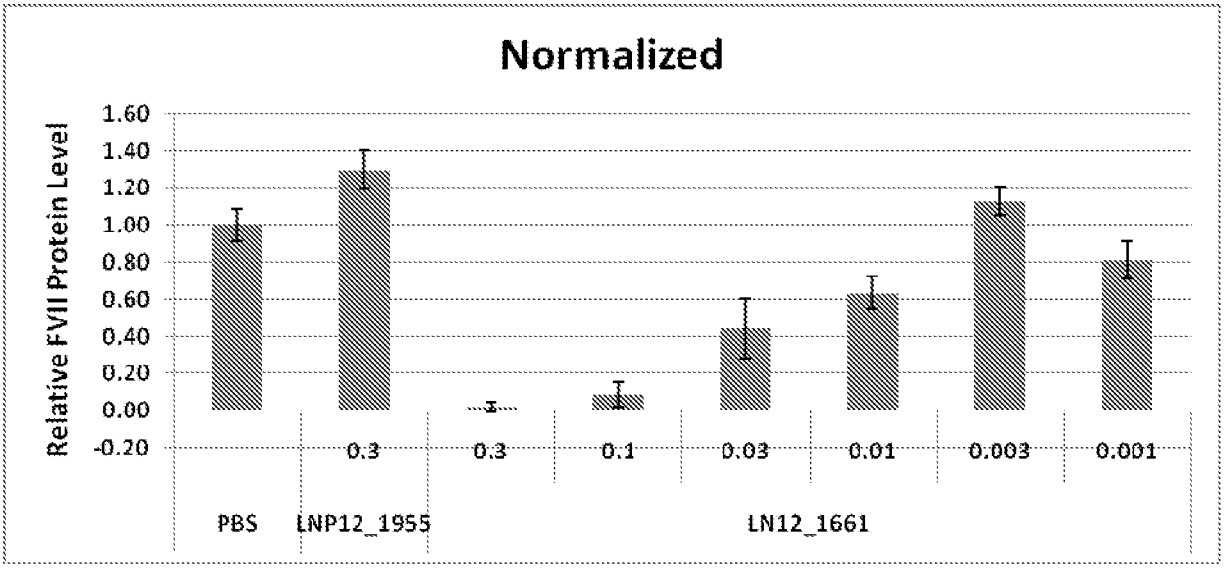

Normalized
Relative FVII Protein Level
1.60
1.40
1.20
1.00
0.80
0.60
0.40
0.20
0.00
-0.20
0.3
0.3
0.1
0.03
0.01
0.003
0.001
PBS
LNP12_1955
LN12_1661

FIG. 6

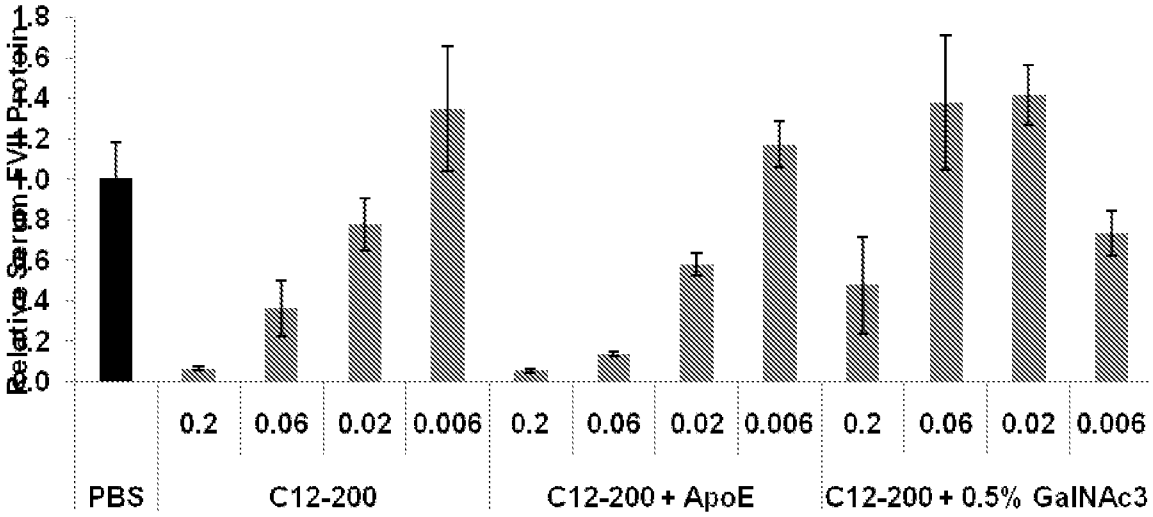

Relative Serum FVII Protein
1.8
1.6
1.4
1.2
1.0
0.8
0.6
0.4
0.2
0.0
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006
PBS
C12-200
C12-200 + ApoE
C12-200 + 0.5% GalNAc3

FIG. 7

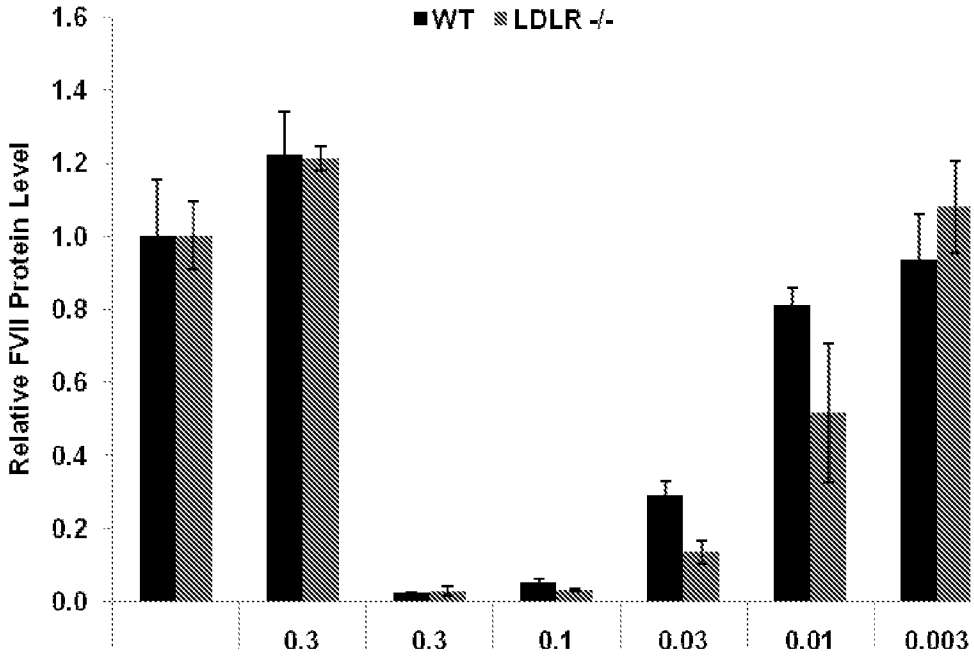

WT
LDLR -/-
Relative FVII Protein Level
1.6
1.4
1.2
1.0
0.8
0.6
0.4
0.2
0.0
0.3
0.3
0.1
0.03
0.01
0.003

FIG. 8

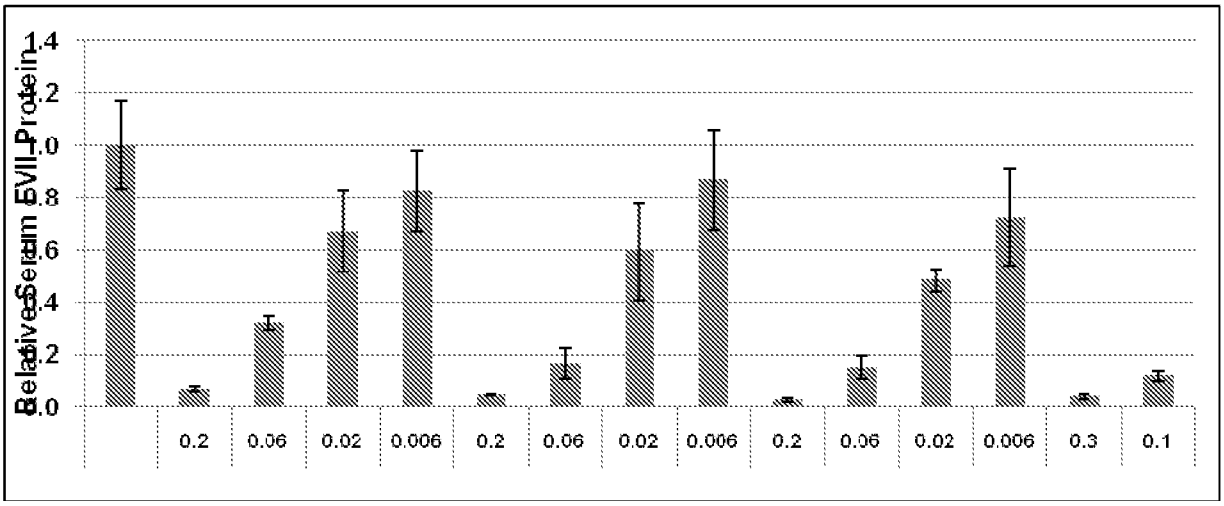

Relative Serum FVII Protein
1.4
1.2
1.0
0.8
0.6
0.4
0.2
0.0
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006
0.3
0.1

FIG. 9a

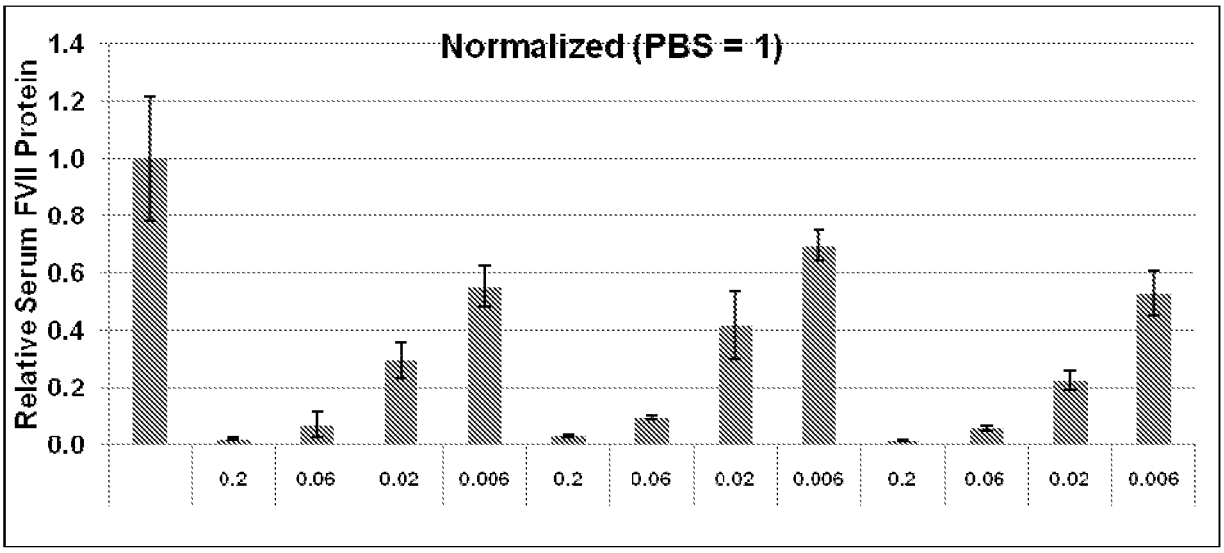

Normalized (PBS = 1)
Relative Serum FVII Protein
1.4
1.2
1.0
0.8
0.6
0.4
0.2
0.0
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006
0.2
0.06
0.02
0.006

FIG. 9b

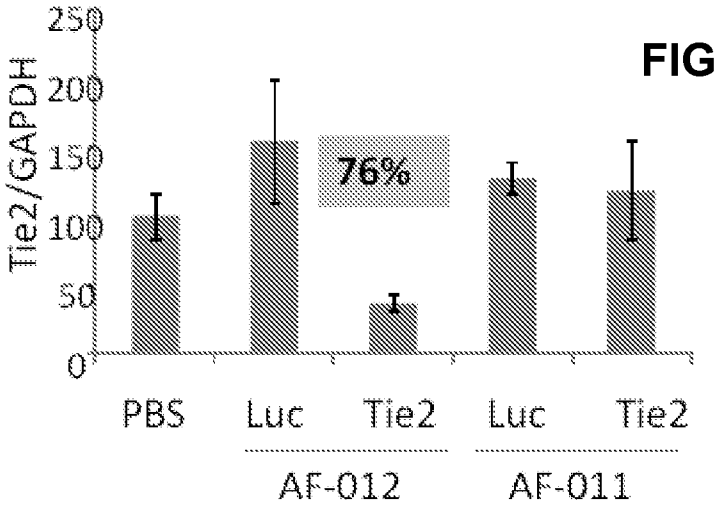


FIG. 10A. Heart

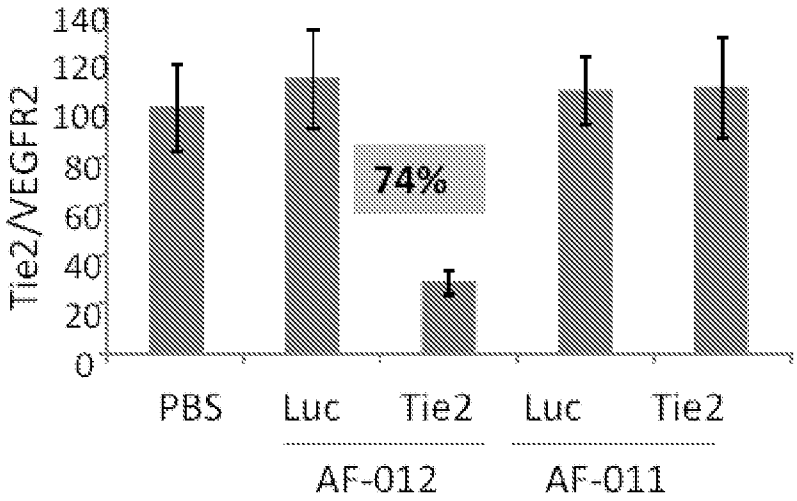


FIG. 10B

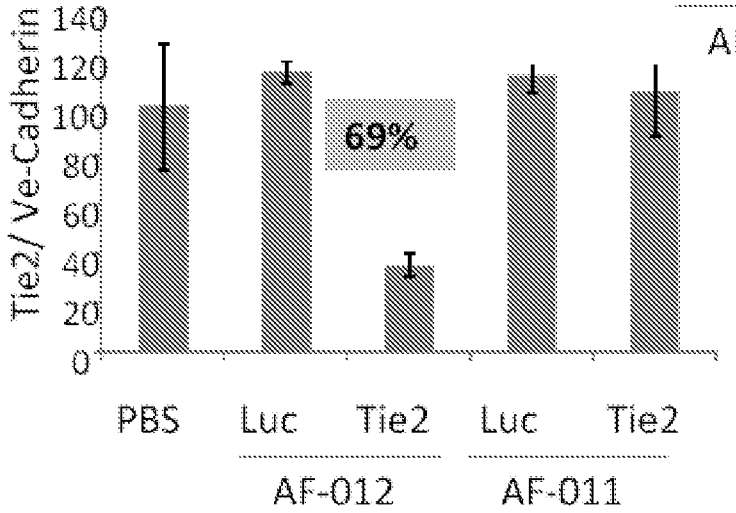


FIG. 10C

FIG. 11A. Liver

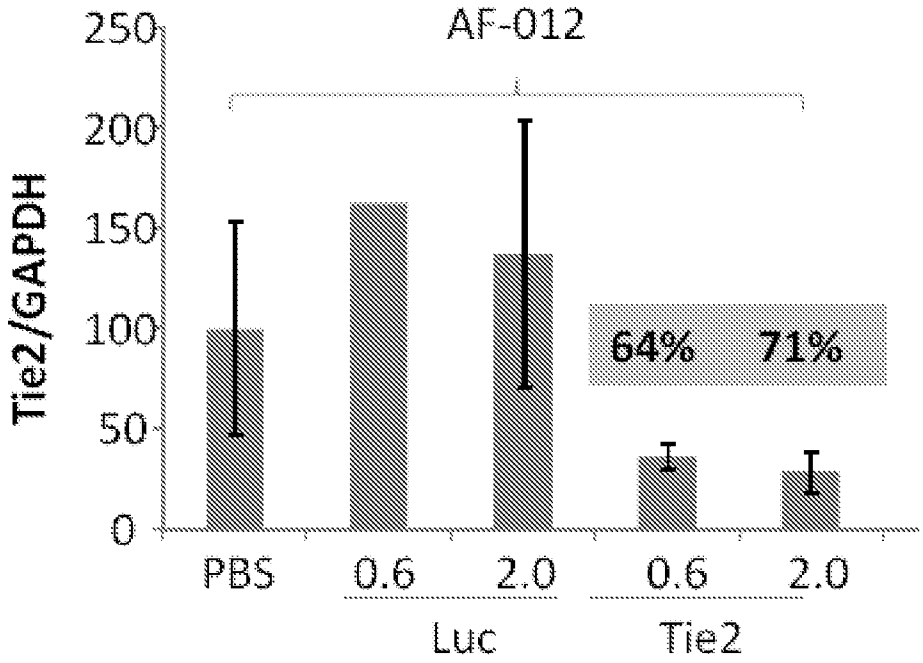


FIG. 11B

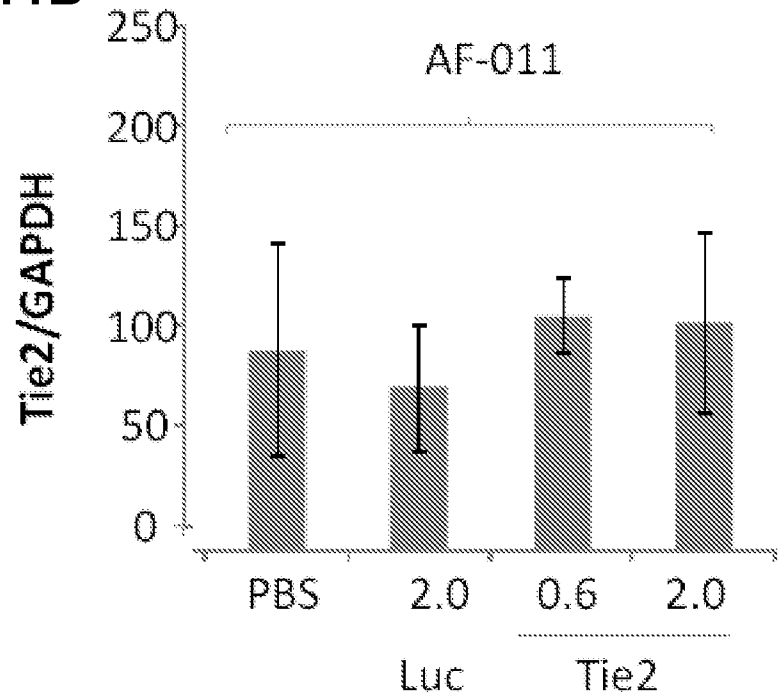


FIG. 12A. Liver

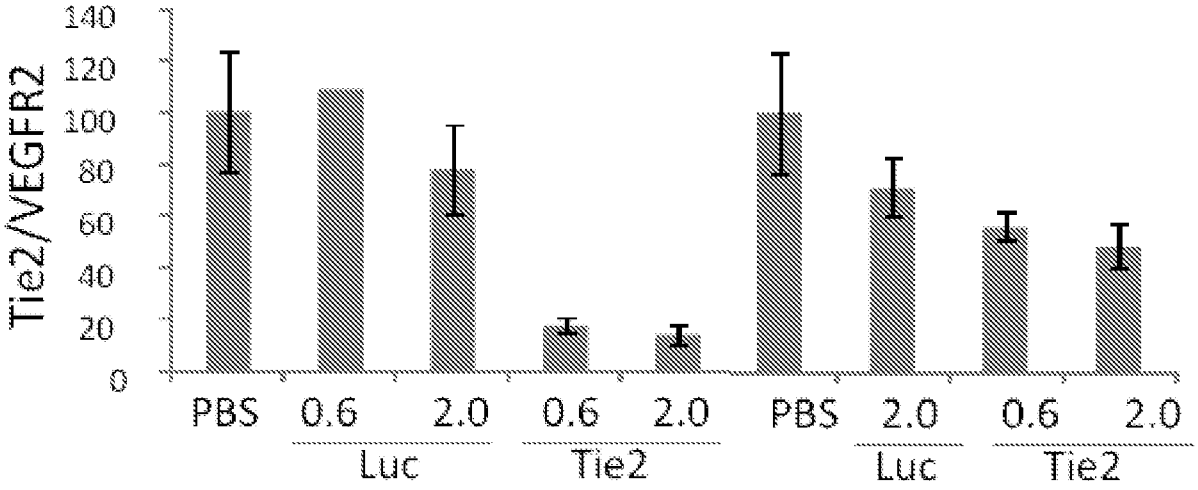


FIG. 12B. Liver

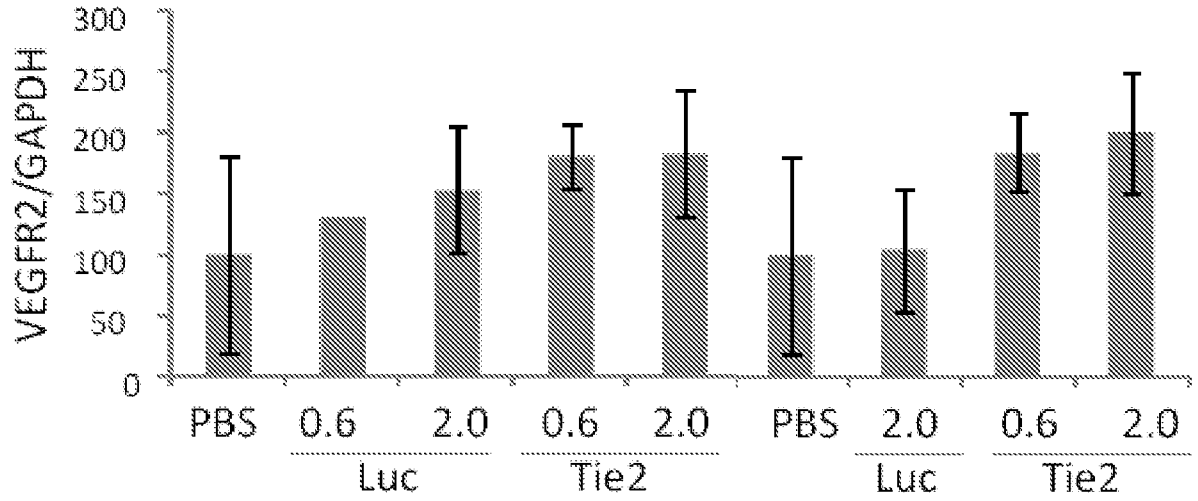


FIG. 13A. Lung

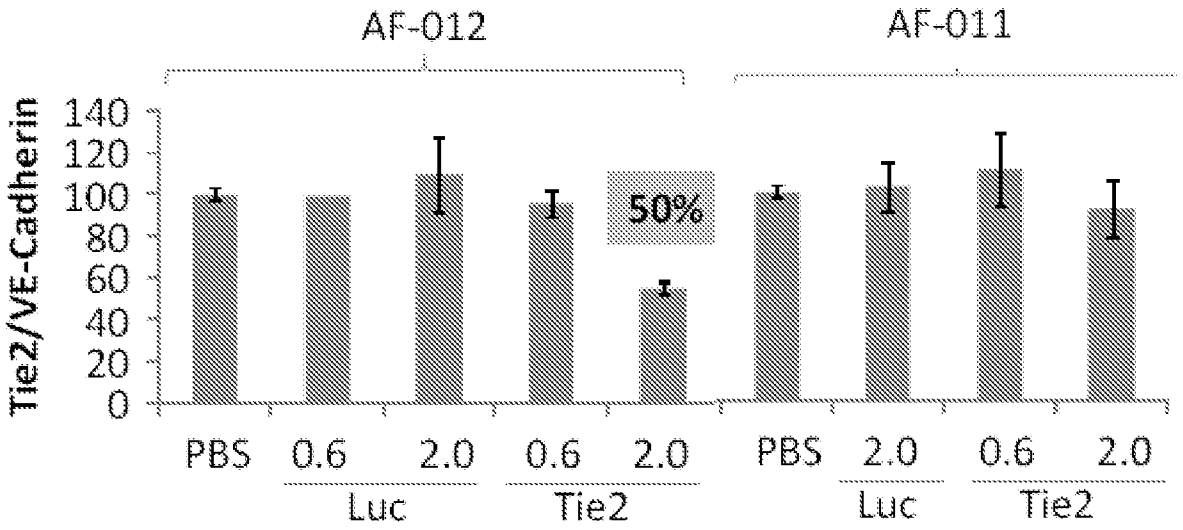


FIG. 13B

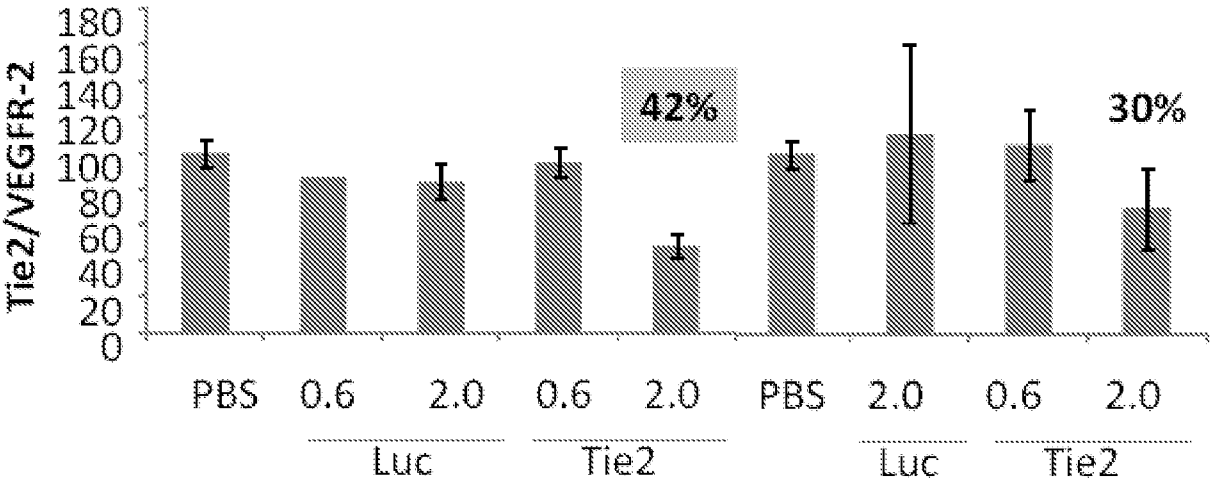


FIG. 14A

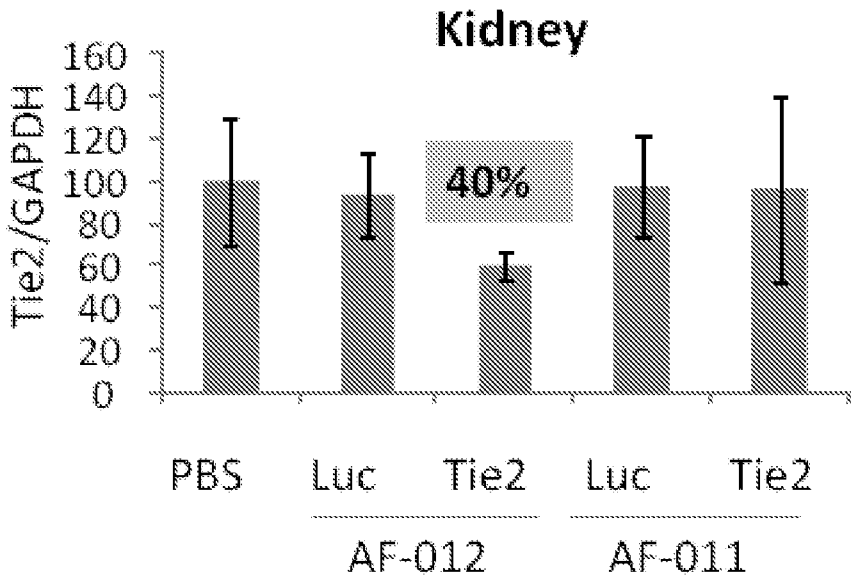


FIG. 14B

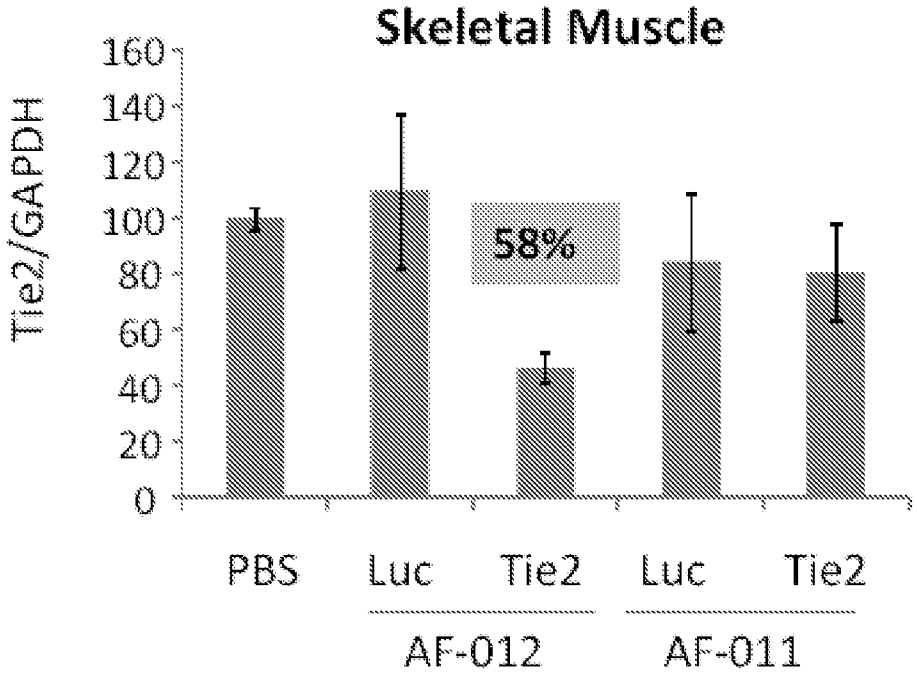


FIG. 15

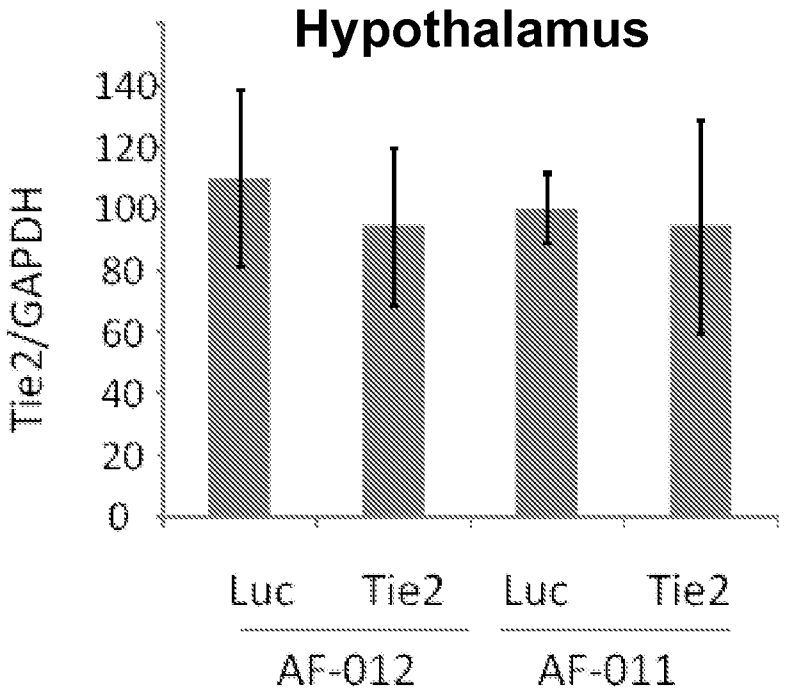

Hypothalamus
Tie2/GAPDH
140
120
100
80
60
40
20
0
Luc
Tie2
Luc
Tie2
AF-012
AF-011

FIG. 16A

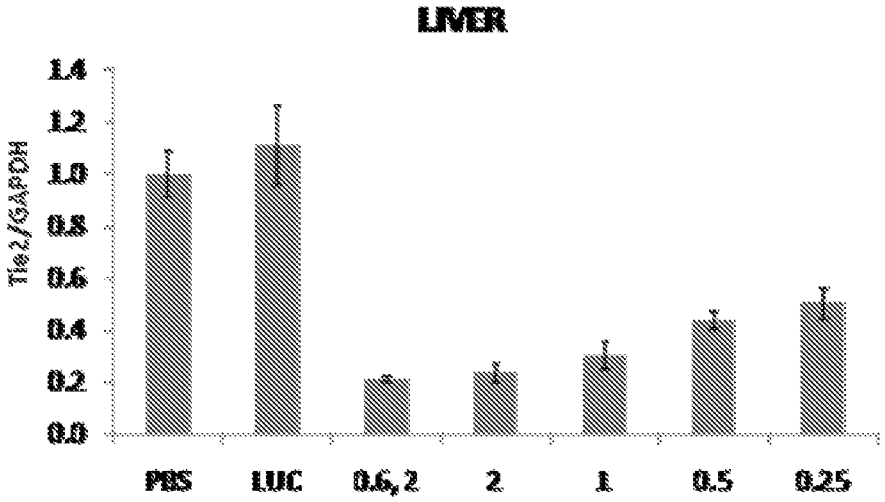


FIG. 16B

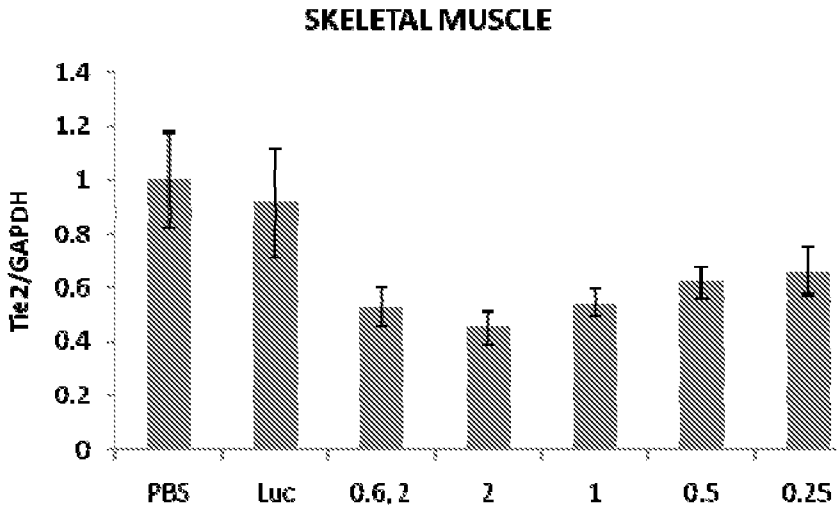


FIG. 17A

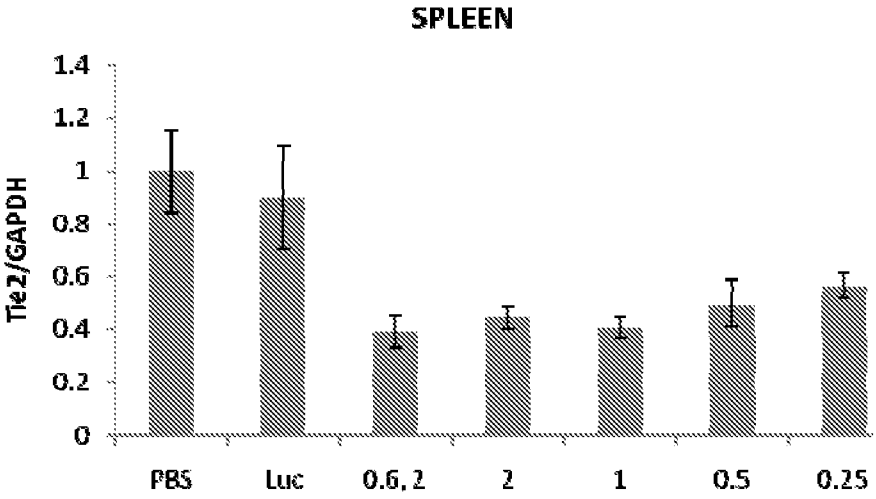


FIG. 17B

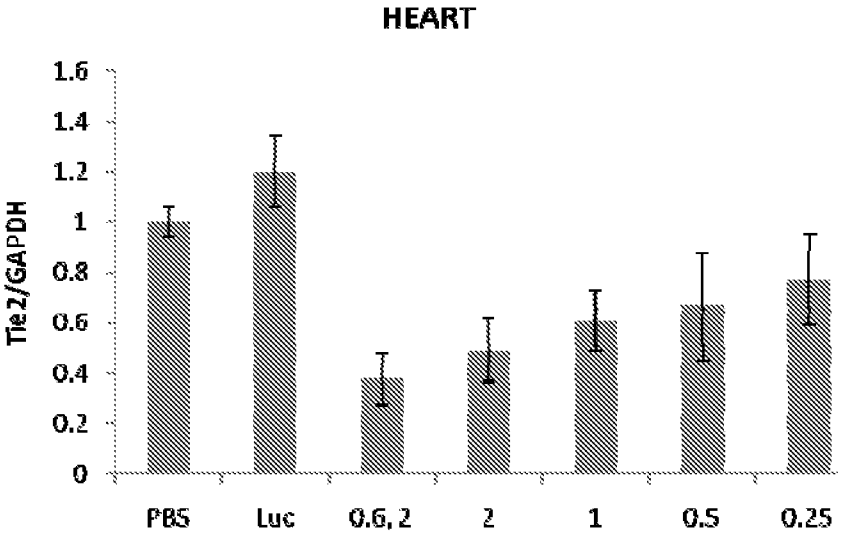


FIG. 18A

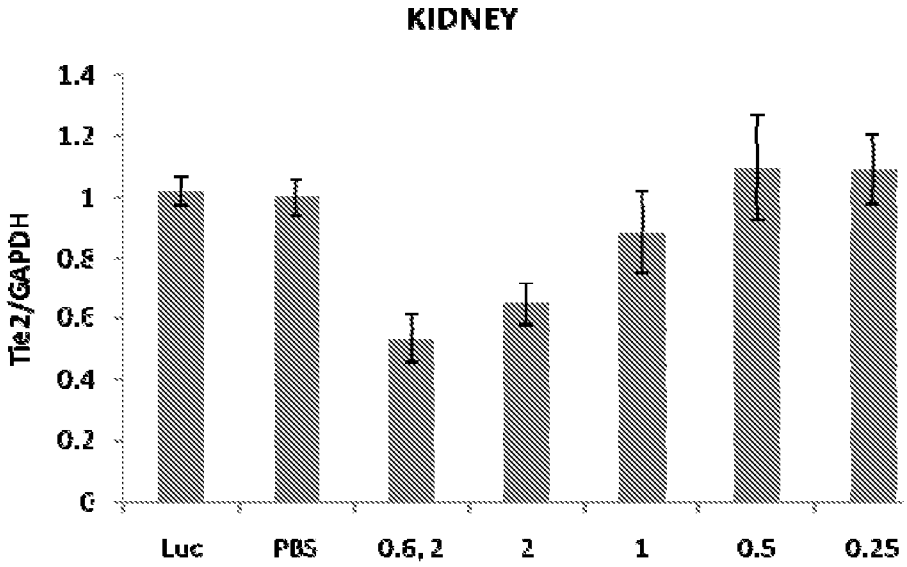


FIG. 18B

