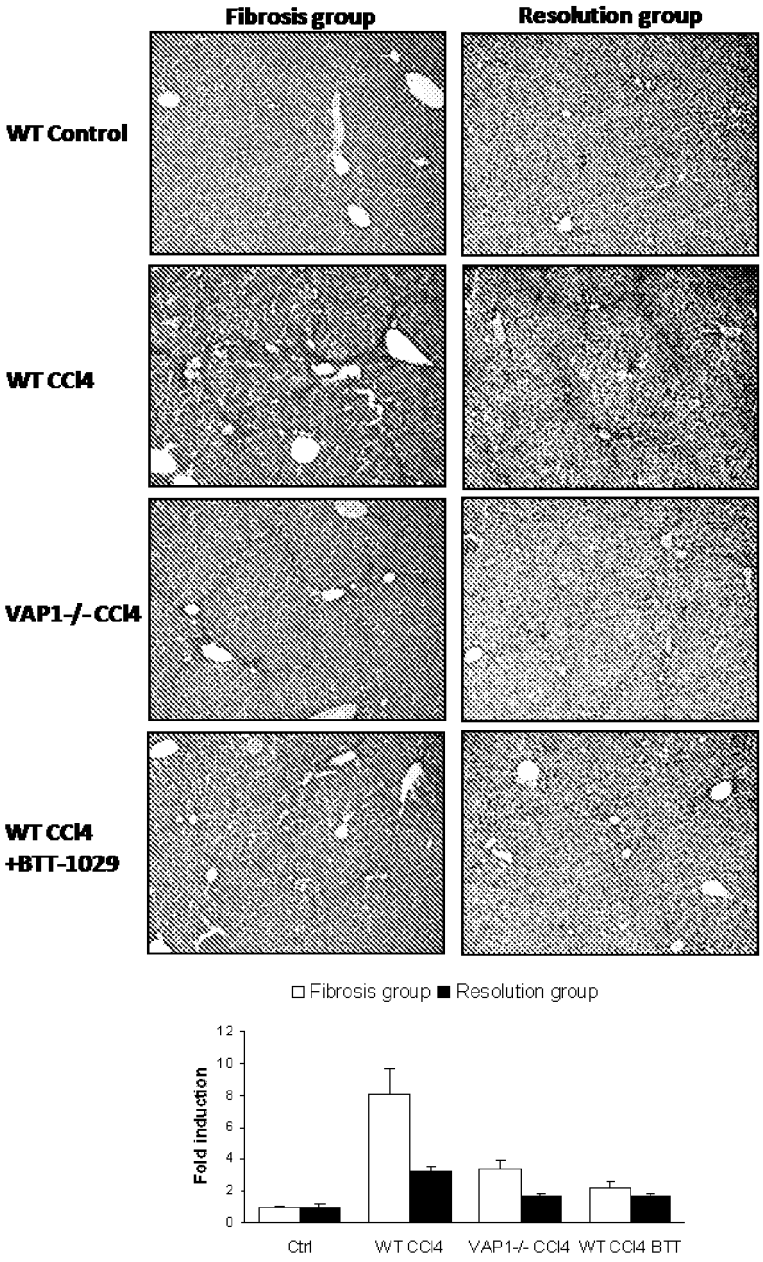

Fibrosis group
Resolution group
WT Control
WT CCl4
VAP1-/- CCl4
WT CCl4
+BTT-1029
Fibrosis group
Resolution group
Fold induction
0
2
4
6
8
10
12
Ctrl
WT CCl4
VAP1-/- CCl4
WT CCl4 BTT

FIG. 1

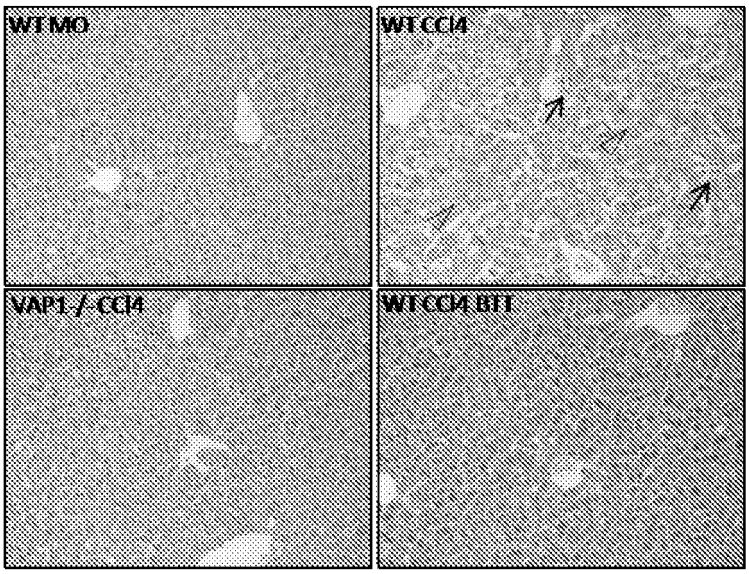


FIG. 2

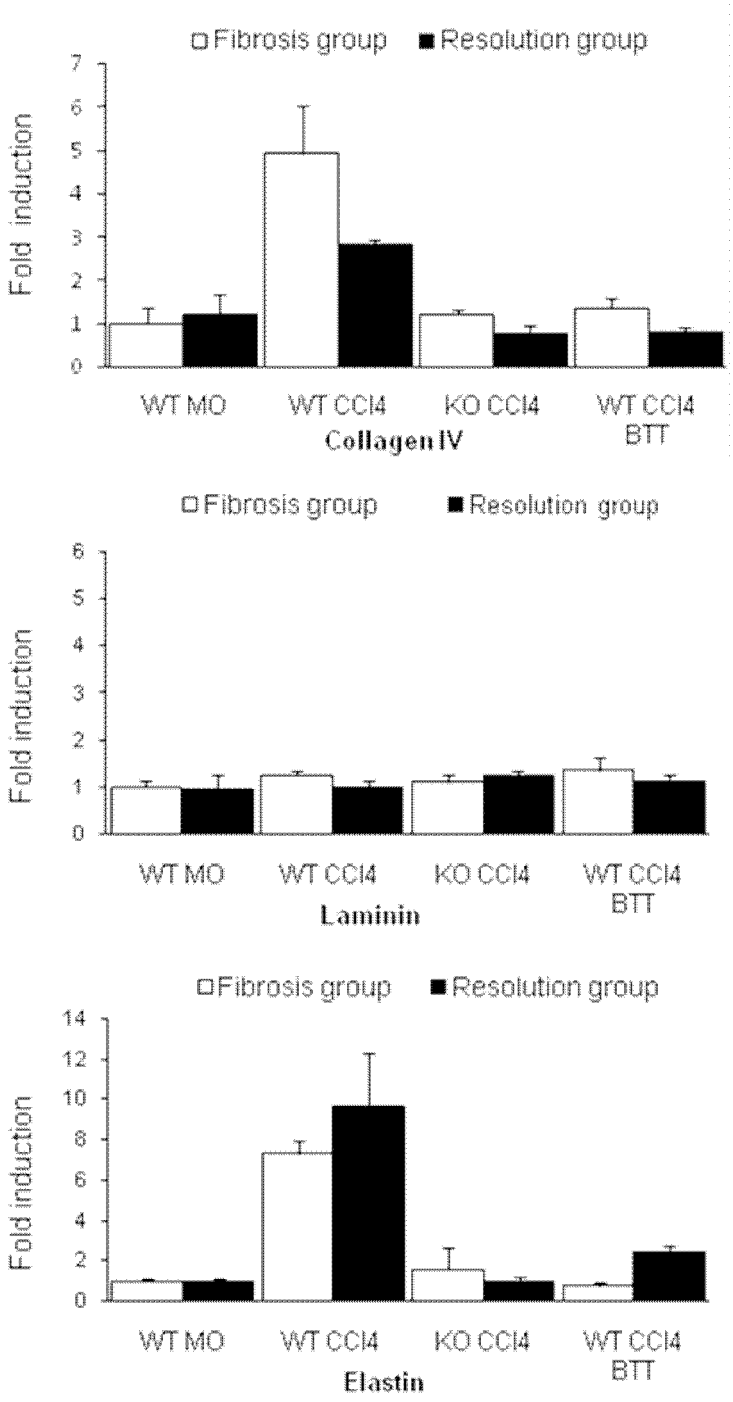

Fibrosis group
Resolution group
Fold induction
0
1
2
3
4
5
6
7
WT MO
WT CCl4
KO CCl4
WT CCl4 BTT
Collagen IV
Fibrosis group
Resolution group
Fold induction
0
1
2
3
4
5
6
WT MO
WT CCl4
KO CCl4
WT CCl4 BTT
Laminin
Fibrosis group
Resolution group
Fold induction
0
2
4
6
8
10
12
14
WT MO
WT CCl4
KO CCl4
WT CCl4 BTT
Elastin

FIG. 3

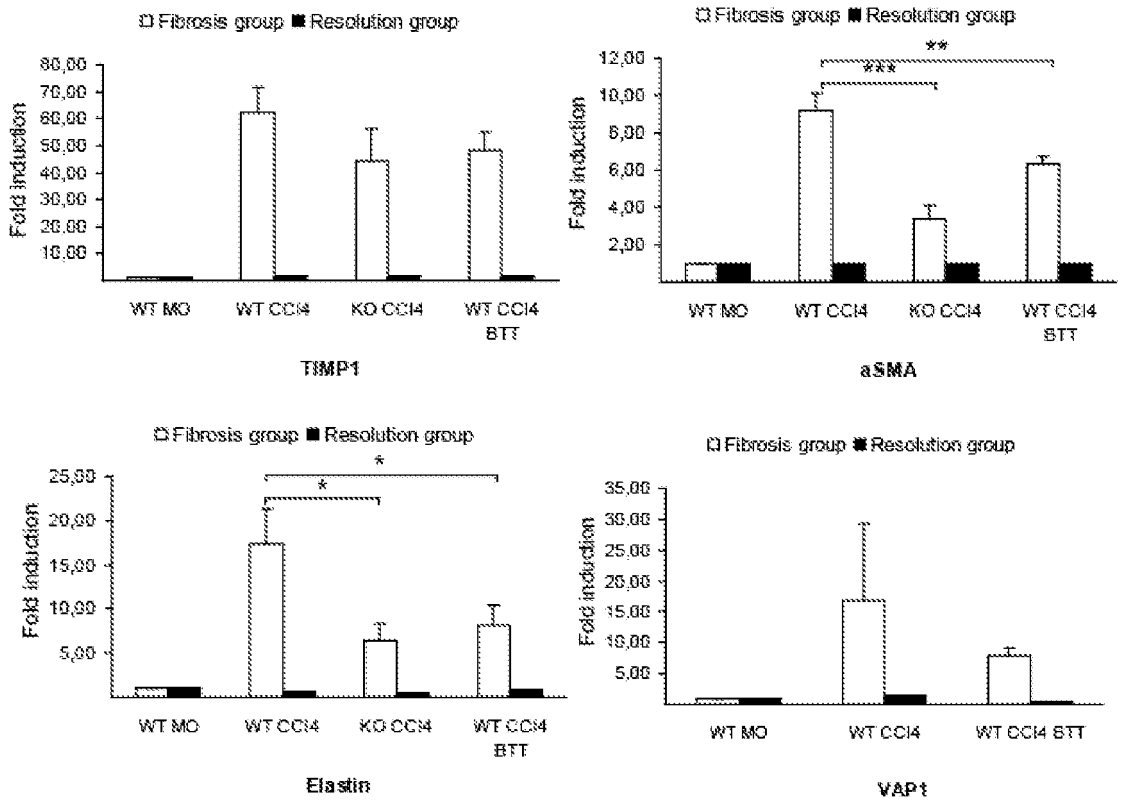

Fibrosis group
Resolution group
Fold induction
WT MO
WT CCl4
KO CCl4
WT CCl4 BTT
TIMP1
aSMA
Elastin
VAP1

FIG. 4

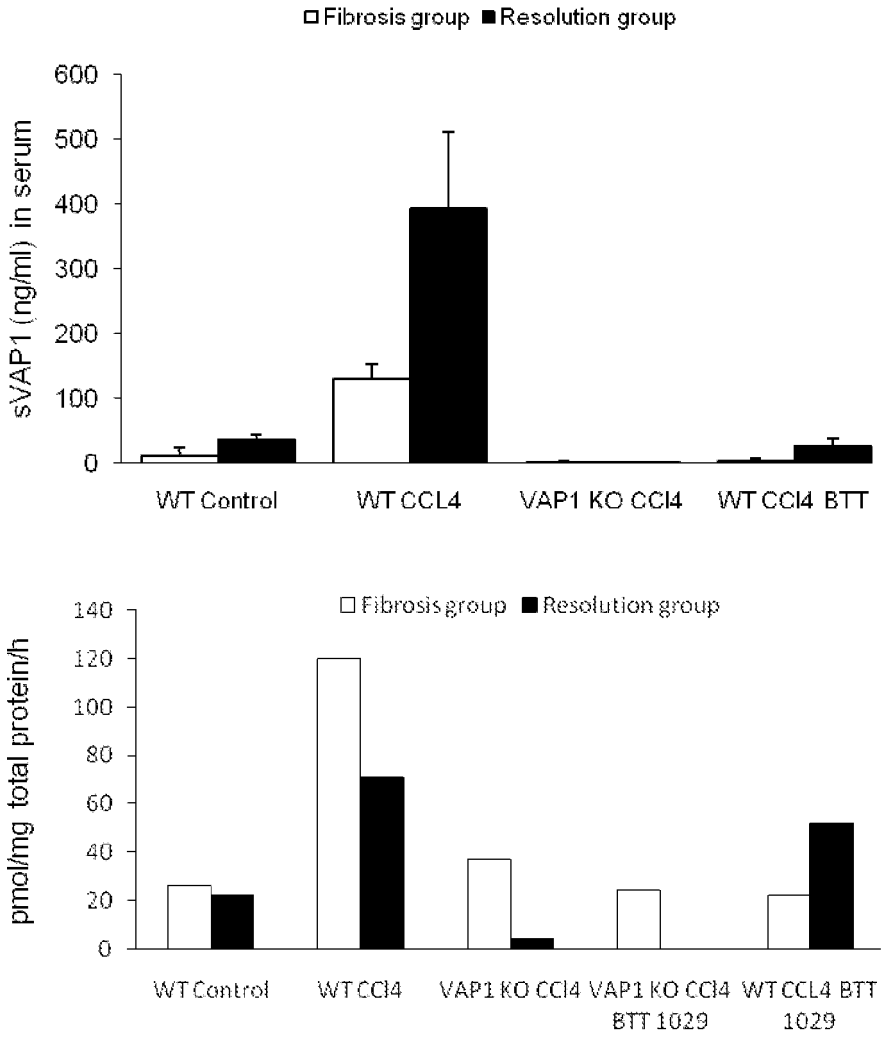

Fibrosis group
Resolution group
sVAP1 (ng/ml) in serum
600
500
400
300
200
100
0
WT Control
WT CCL4
VAP1 KO CCl4
WT CCl4 BTT
Fibrosis group
Resolution group
pmol/mg total protein/h
140
120
100
80
60
40
20
0
WT Control
WT CCl4
VAP1 KO CCl4
VAP1 KO CCl4 BTT 1029
WT CCL4 BTT 1029

FIG. 5

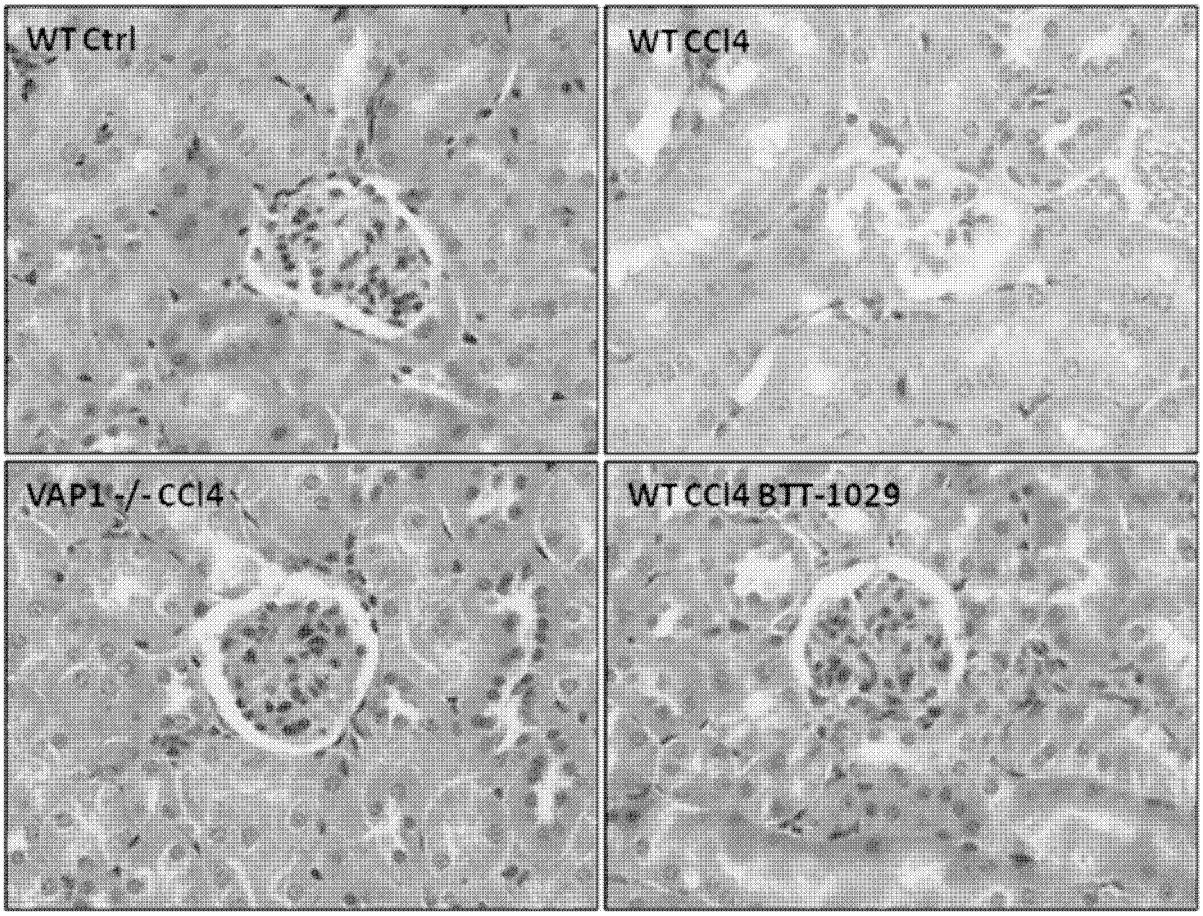
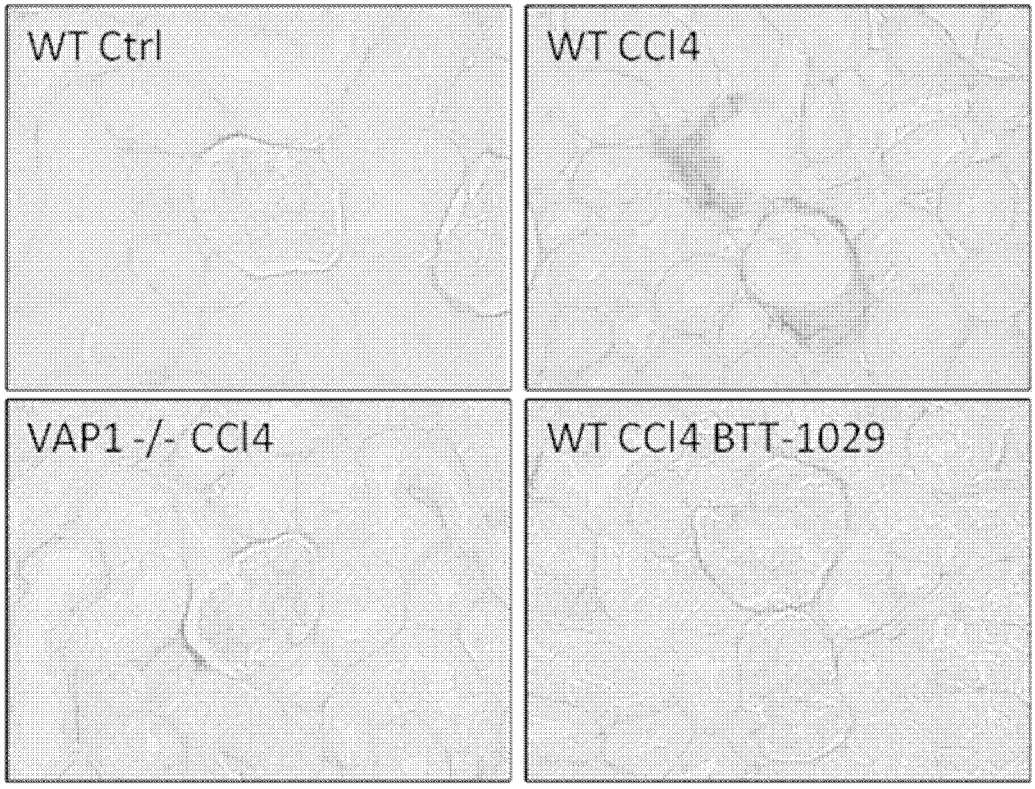


FIG. 6


WT Ctrl
WT CCl4
VAP1 -/- CCl4
WT CCl4 BTT-1029

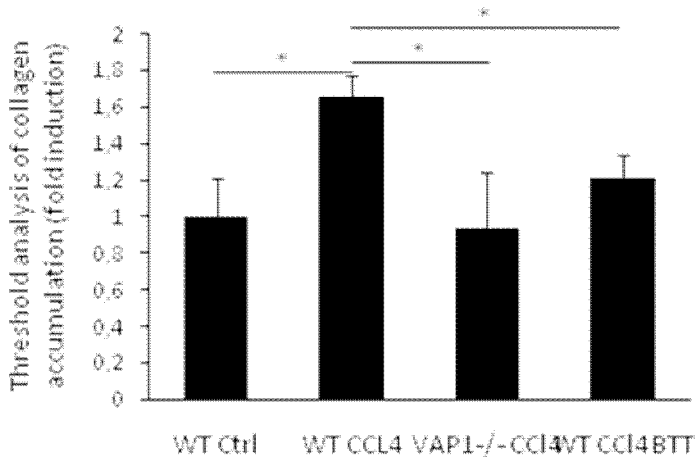

Threshold analysis of collagen accumulation (fold induction)
2
1,8
1,6
1,4
1,2
1
0,8
0,6
0,4
0,2
0
*
*
*
WT Ctrl
WT CCL4
VAP1-/-CCl4
WT CCl4 BTT

FIG. 7

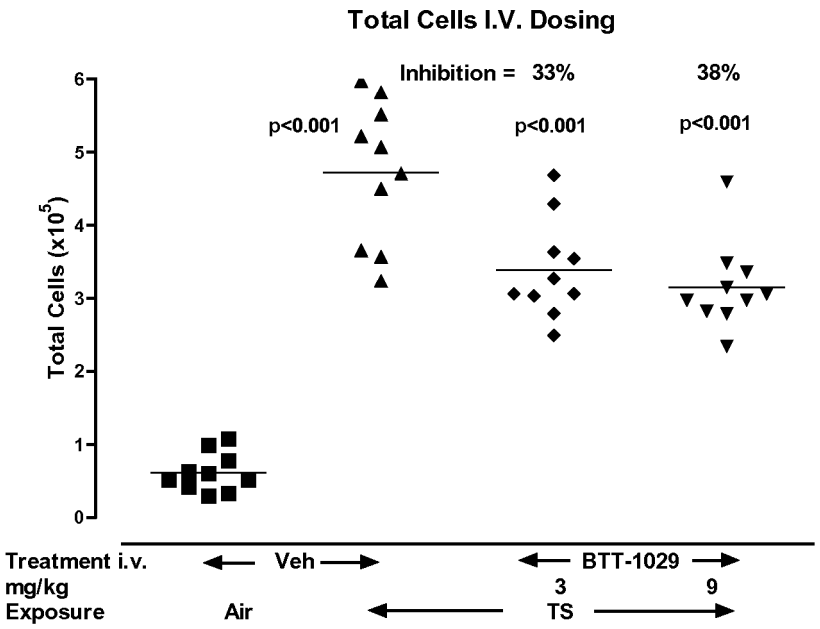

Total Cells I.V. Dosing
Inhibition = 33% 38%
p<0.001
p<0.001
p<0.001
Total Cells (x10^5)
0
1
2
3
4
5
6
Treatment i.v.
Veh
BTT-1029
mg/kg
3
9
Exposure
Air
TS

FIG. 8

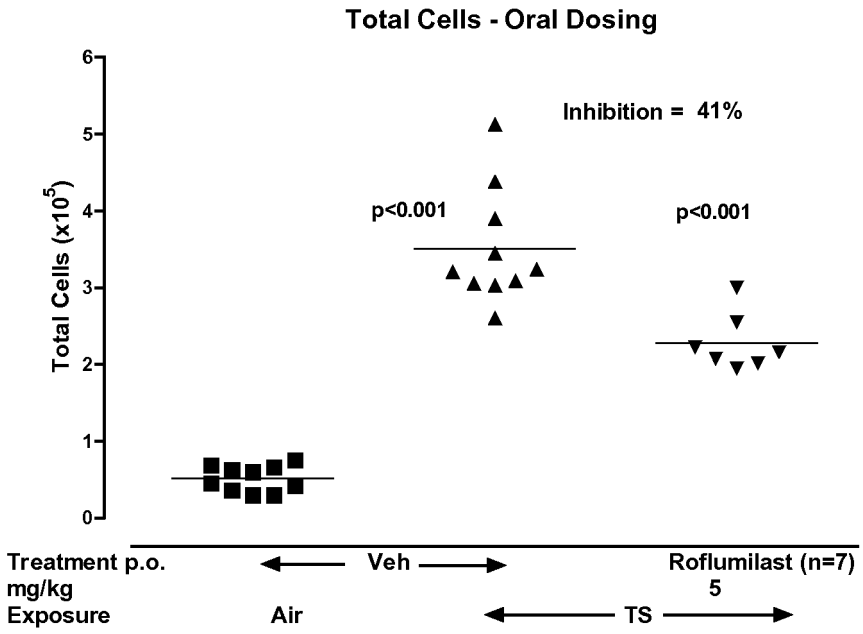

Total Cells - Oral Dosing
Total Cells (x10^5)
6
5
4
3
2
1
0
Inhibition = 41%
p<0.001
p<0.001
Treatment p.o.
mg/kg
Exposure
Air
Veh
TS
Roflumilast (n=7)
5

FIG. 9

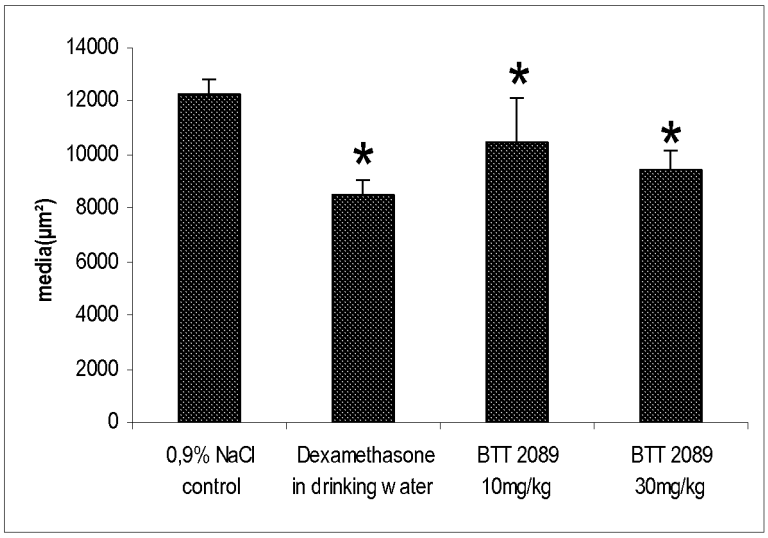

14000
12000
10000
8000
6000
4000
2000
0
media(µm²)
*
*
*
0,9% NaCl control
Dexamethasone in drinking w ater
BTT 2089 10mg/kg
BTT 2089 30mg/kg

FIG. 10

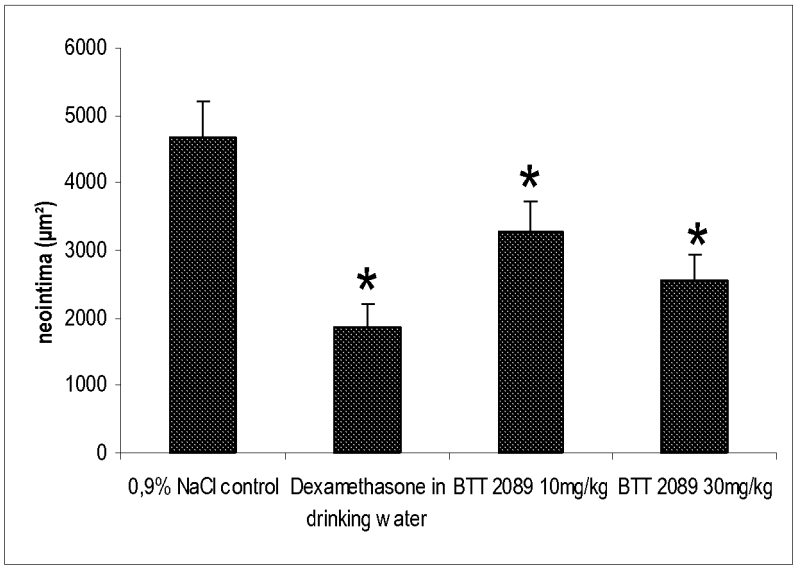

6000
5000
4000
3000
2000
1000
0
neointima (µm²)
*
*
*
0,9% NaCl control
Dexamethasone in drinking water
BTT 2089 10mg/kg
BTT 2089 30mg/kg

FIG. 11

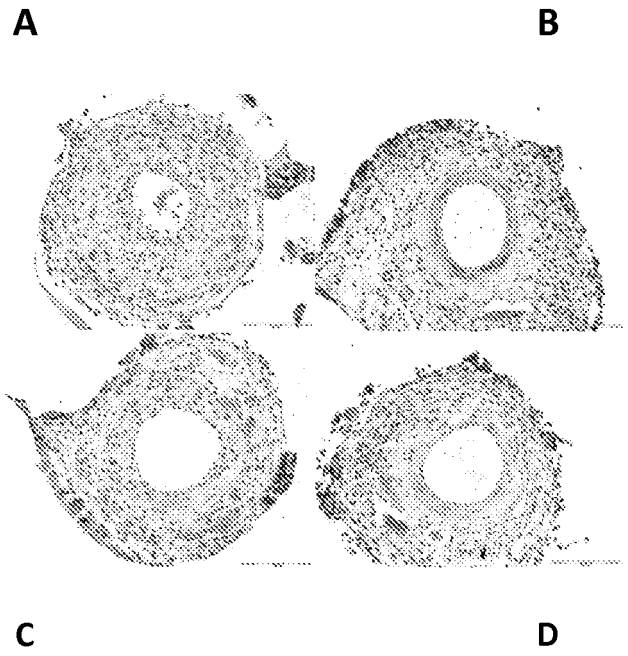
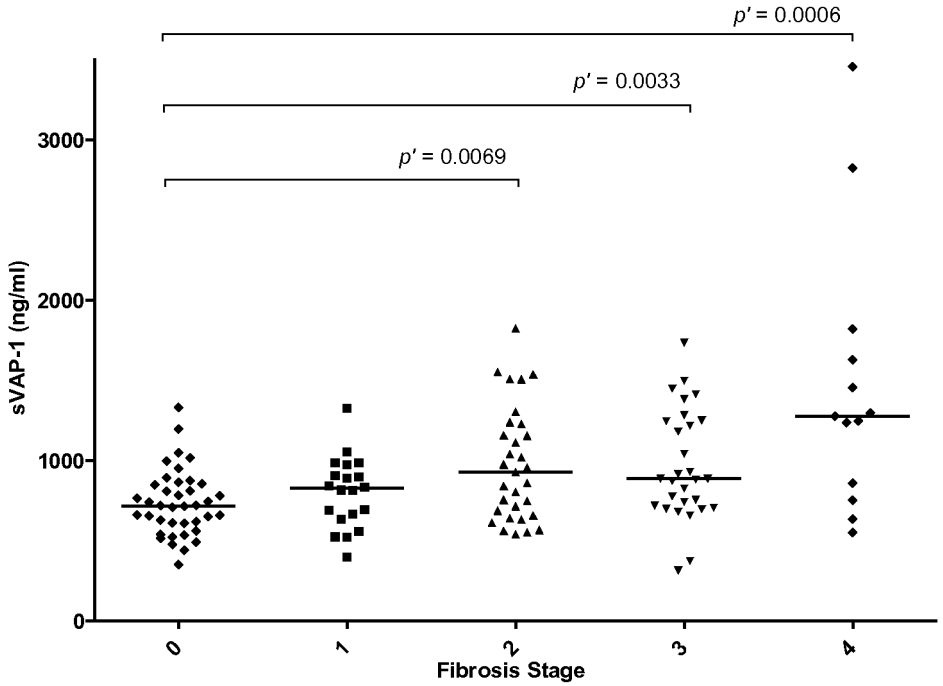


FIG. 12



Kruskal-Wallis test $p = 0.0003$
Cuzick trend test $p = <0.0001$
p' values calculated by Mann Whitney test with Bonferroni correction

FIG. 13

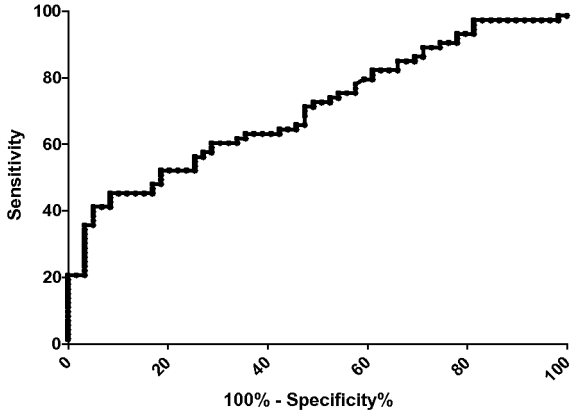


FIG. 14A

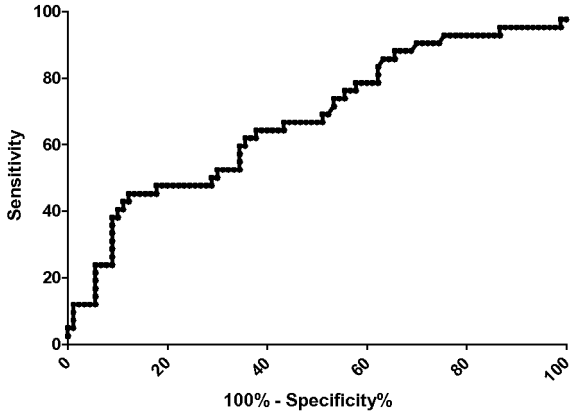


FIG. 14B

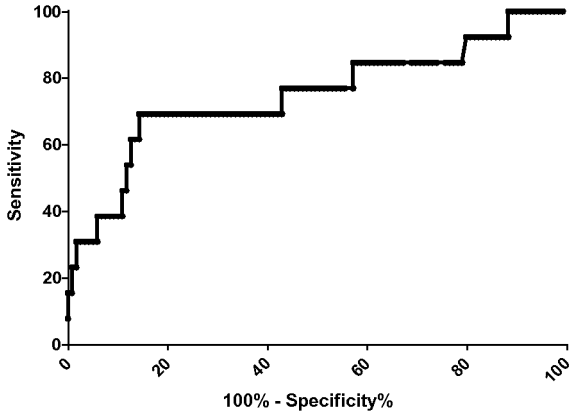


FIG. 14C

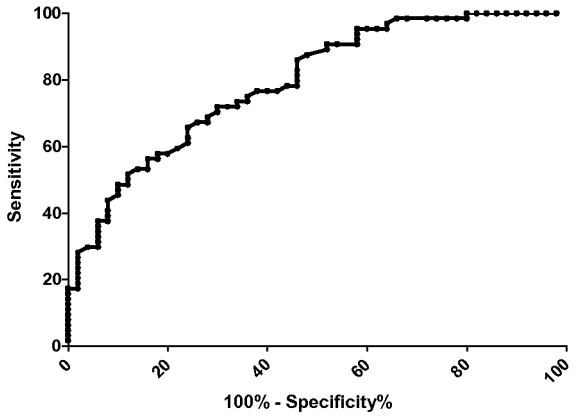


FIG. 15A

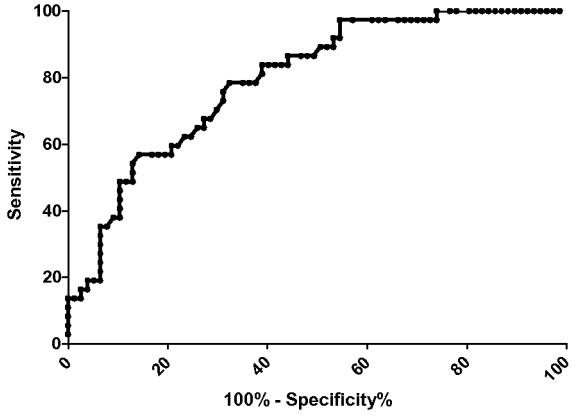


FIG. 15B

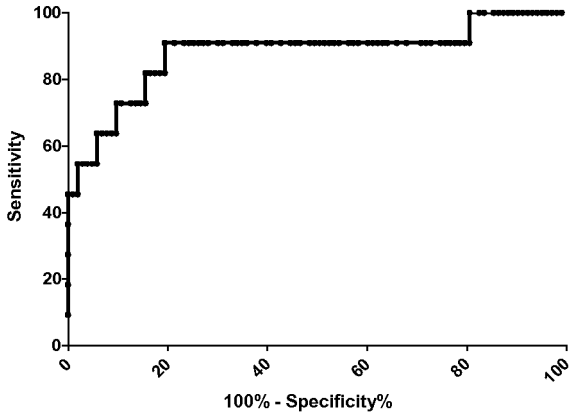


FIG. 15C

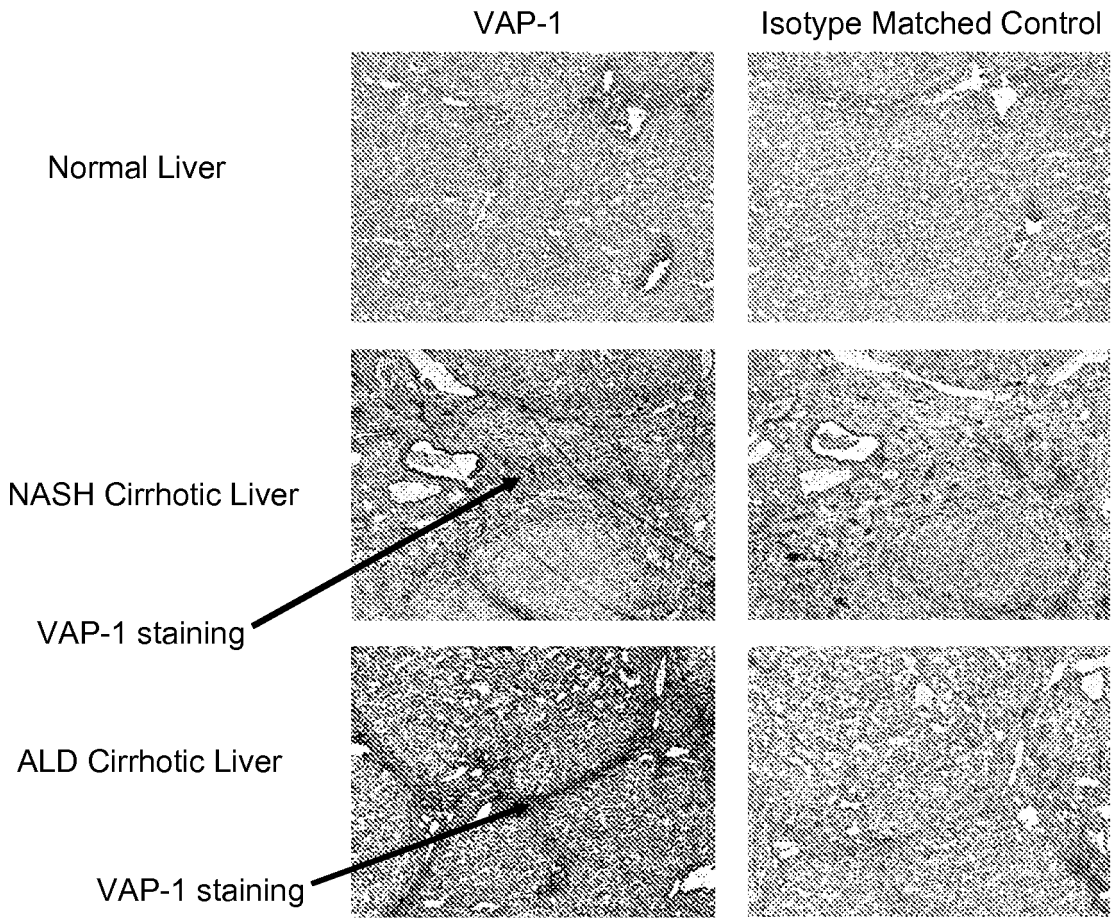

VAP-1
Isotype Matched Control
Normal Liver
NASH Cirrhotic Liver
VAP-1 staining
ALD Cirrhotic Liver
VAP-1 staining

FIG. 16

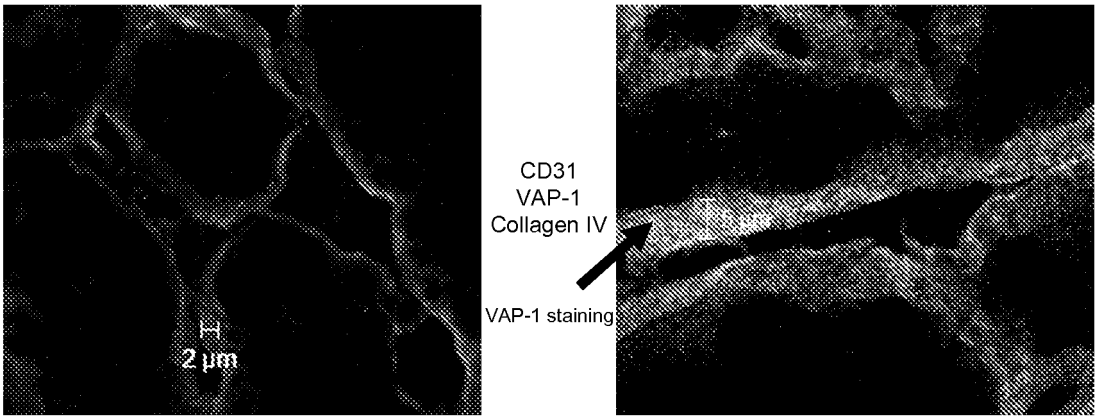

2 µm
CD31
VAP-1
Collagen IV
VAP-1 staining
Normal Liver (sinusoids) x 100
NASH Cirrhotic Liver (sinusoids) x 100

FIG. 17

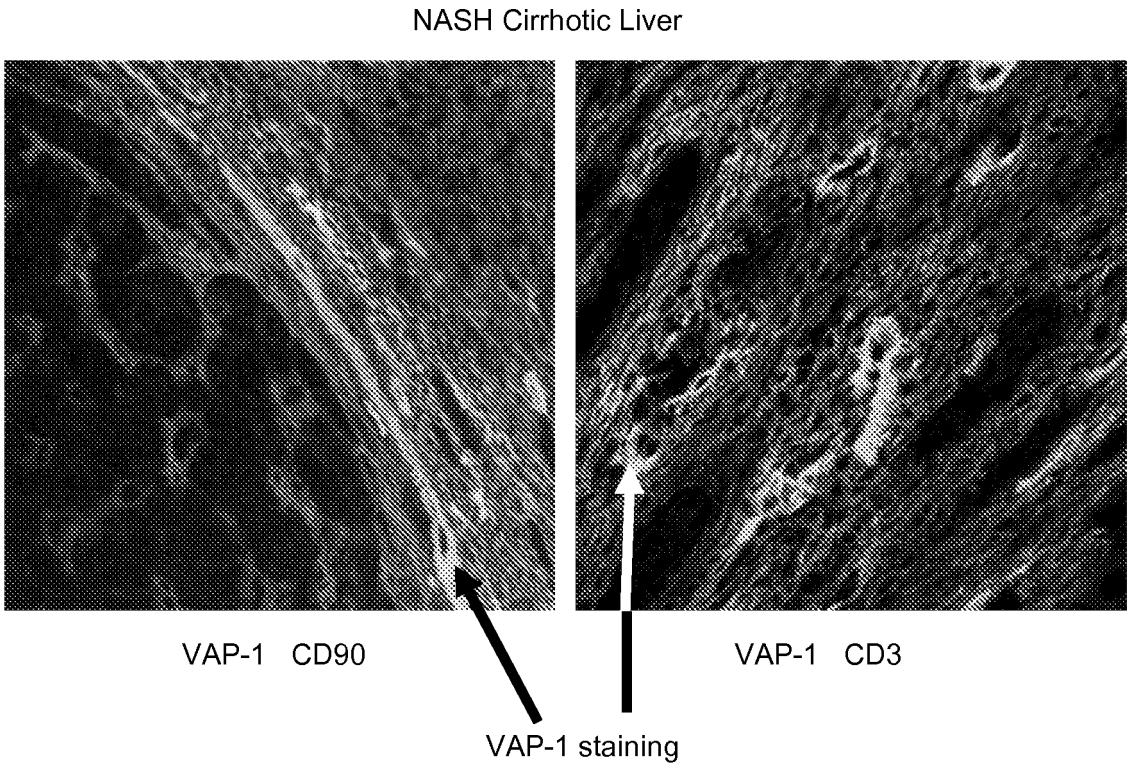


FIG. 18

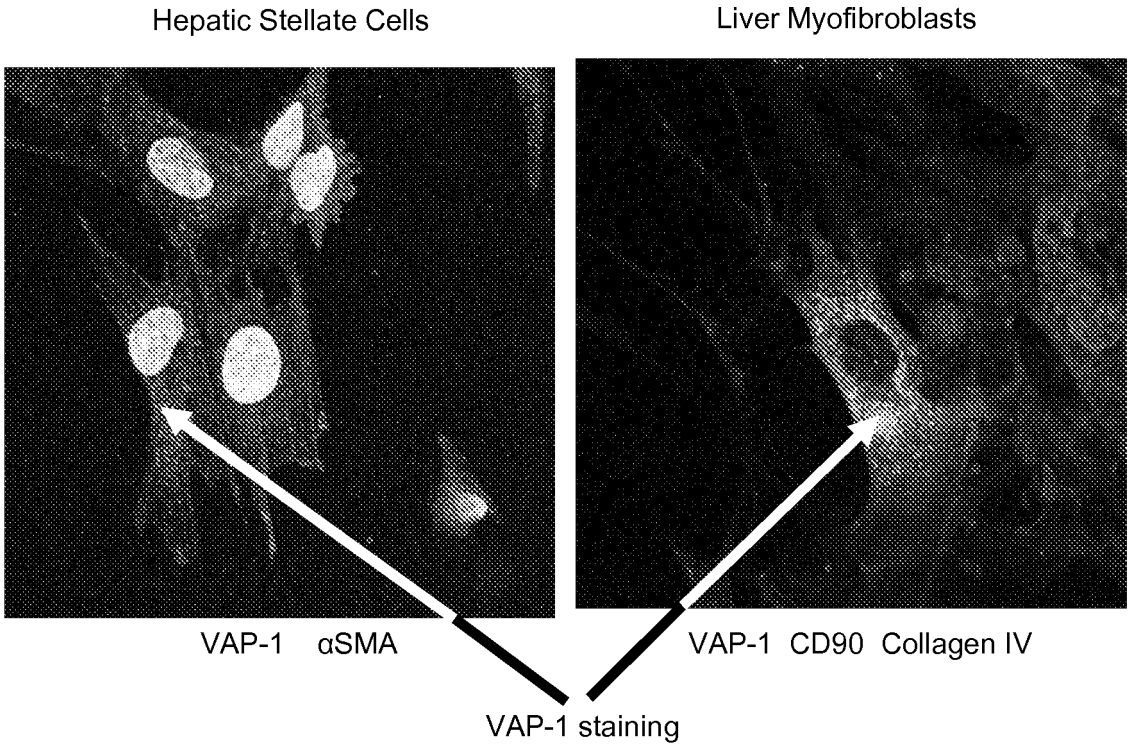

Hepatic Stellate Cells
Liver Myofibroblasts
VAP-1 αSMA
VAP-1 CD90 Collagen IV
VAP-1 staining

FIG. 19