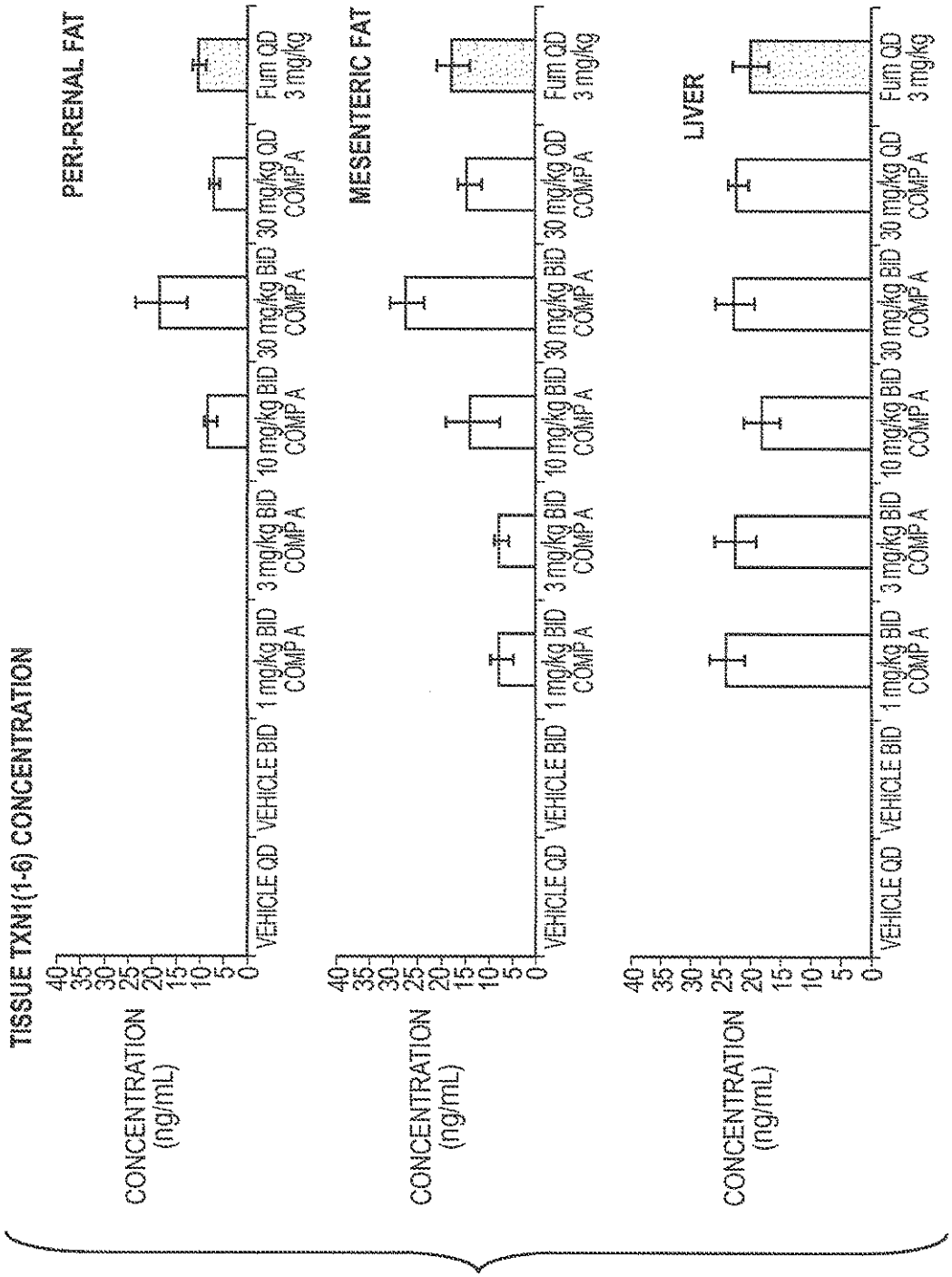
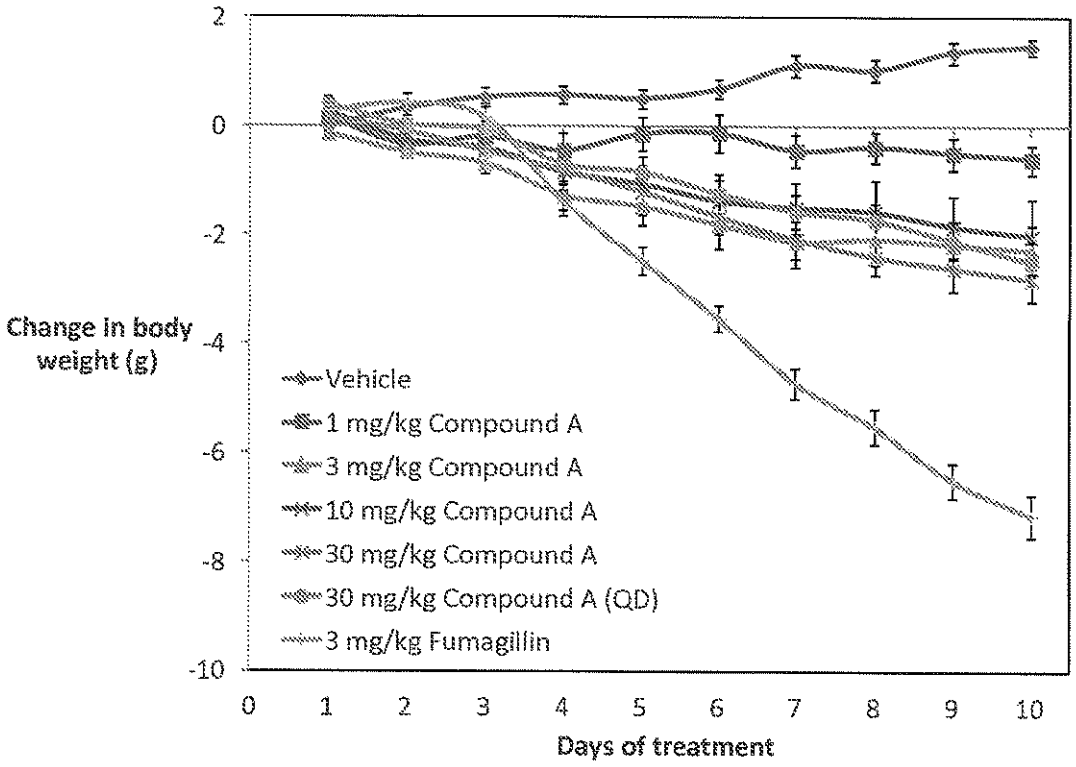

TISSUE TXN1(1-6) CONCENTRATION
PERI-RENAL FAT
MESENTERIC FAT
LIVER
CONCENTRATION (ng/mL)
0
5
10
15
20
25
30
35
40
VEHICLE QD
VEHICLE BID
1 mg/kg BID COMP A
3 mg/kg BID COMP A
10 mg/kg BID COMP A
30 mg/kg BID COMP A
30 mg/kg QD COMP A
Fum QD 3 mg/kg

FIG. 1

Figure 2


Change in body weight (g)
2
0
-2
-4
-6
-8
-10
0
1
2
3
4
5
6
7
8
9
10
Days of treatment
Vehicle
1 mg/kg Compound A
3 mg/kg Compound A
10 mg/kg Compound A
30 mg/kg Compound A
30 mg/kg Compound A (QD)
3 mg/kg Fumagillin

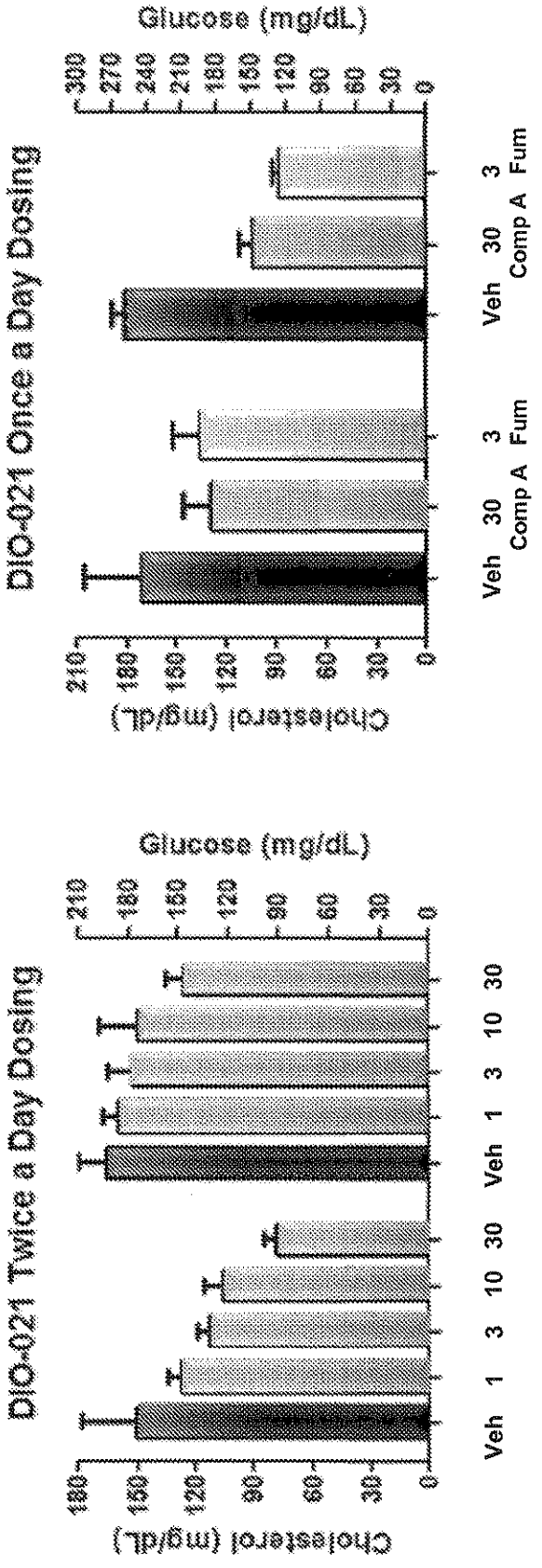

DIO-021 Once a Day Dosing
Glucose (mg/dL)
300
270
240
210
180
150
120
90
60
30
0
Cholesterol (mg/dL)
210
180
150
120
90
60
30
0
Veh
30
Comp A
3
Fum
DIO-021 Twice a Day Dosing
Glucose (mg/dL)
210
180
150
120
90
60
30
0
Cholesterol (mg/dL)
180
150
120
90
60
30
0
Veh
1
3
10
30
Veh
1
3
10
30

Figure 3

Figure 4

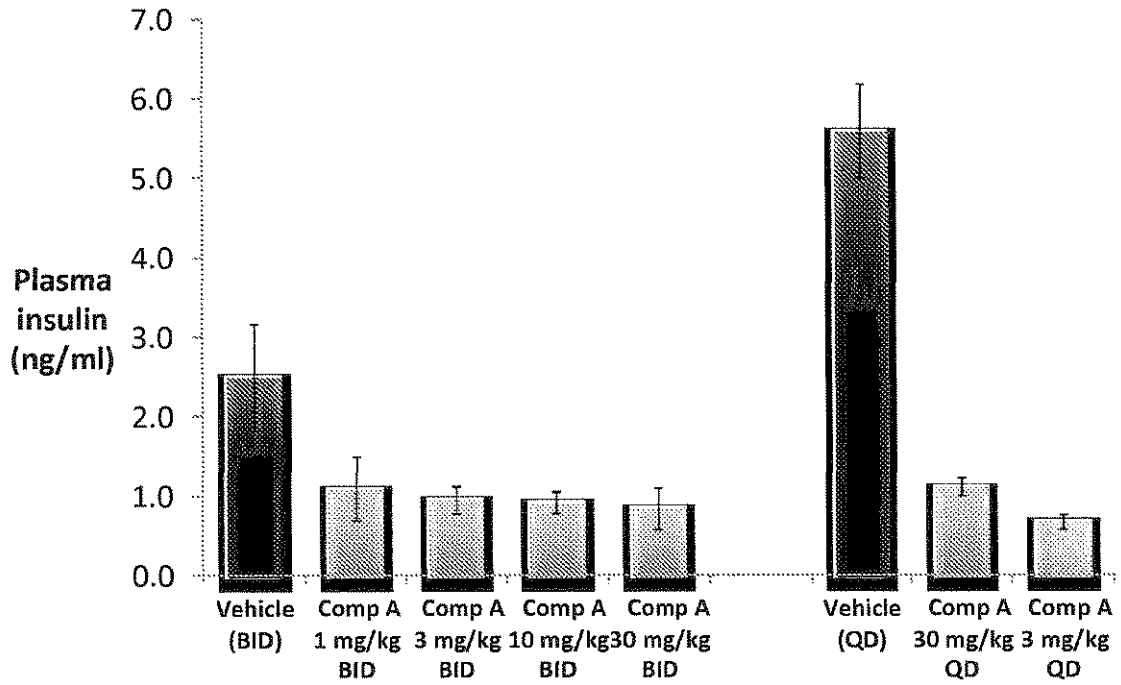
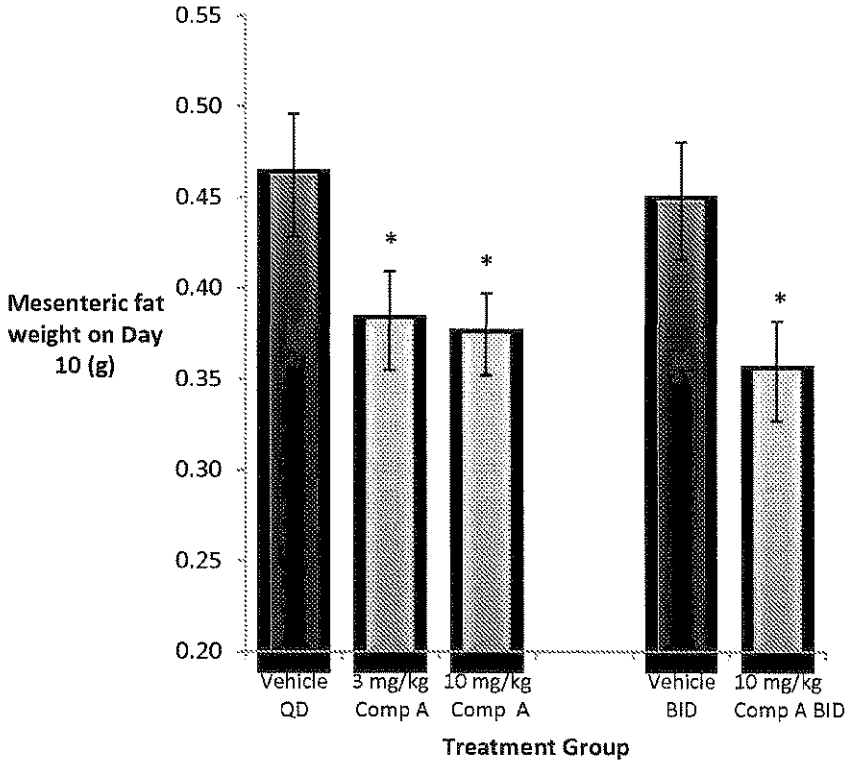

Plasma insulin (ng/ml)
7.0
6.0
5.0
4.0
3.0
2.0
1.0
0.0
Vehicle (BID)
Comp A 1 mg/kg BID
Comp A 3 mg/kg BID
Comp A 10 mg/kg BID
Comp A 30 mg/kg BID
Vehicle (QD)
Comp A 30 mg/kg QD
Comp A 3 mg/kg QD

Figure 5


0.55
0.50
0.45
0.40
0.35
0.30
0.25
0.20
Mesenteric fat weight on Day 10 (g)
*
*
*
Vehicle QD
3 mg/kg Comp A
10 mg/kg Comp A
Vehicle BID
10 mg/kg Comp A BID
Treatment Group

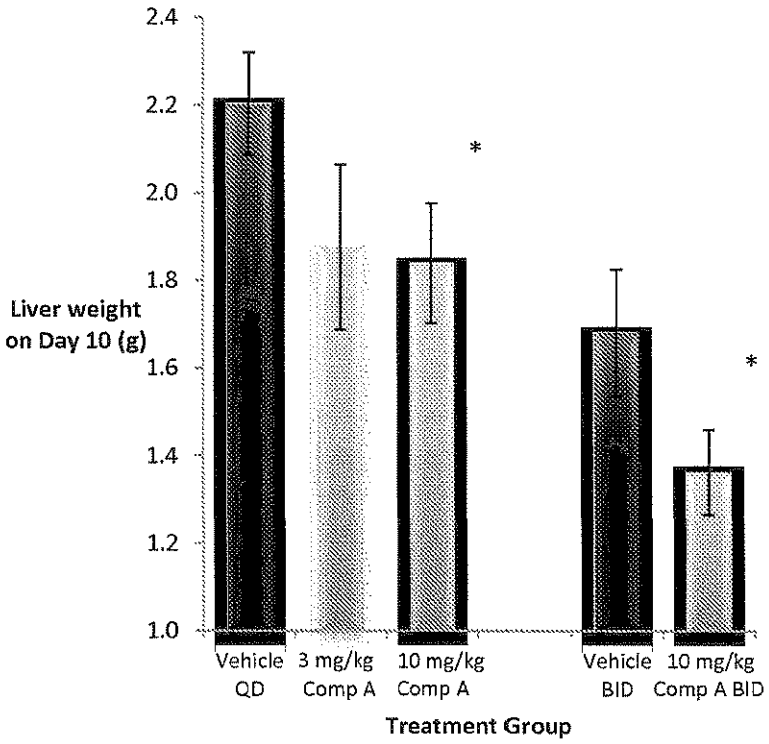

2.4
2.2
2.0
1.8
1.6
1.4
1.2
1.0
Liver weight on Day 10 (g)
*
*
Vehicle QD
3 mg/kg Comp A
10 mg/kg Comp A
Vehicle BID
10 mg/kg Comp A BID
Treatment Group

Figure 6

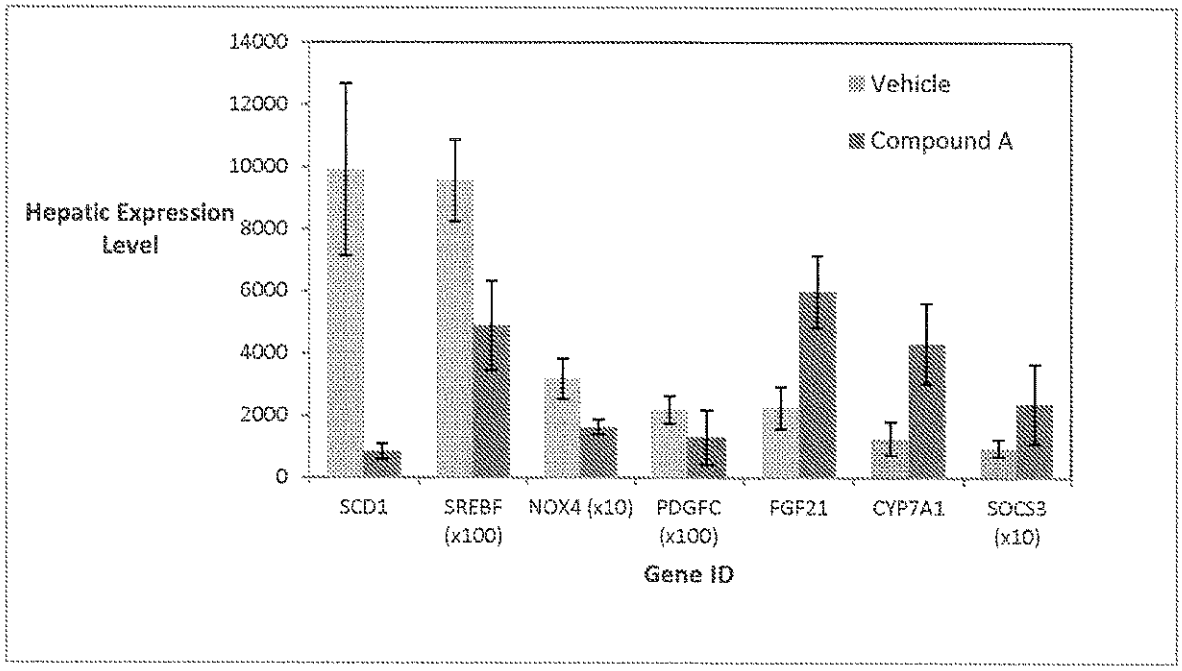

14000
12000
10000
8000
6000
4000
2000
0
Hepatic Expression Level
Vehicle
Compound A
SCD1
SREBF (x100)
NOX4 (x10)
PDGFC (x100)
FGF21
CYP7A1
SOCS3 (x10)
Gene ID

Figure 7

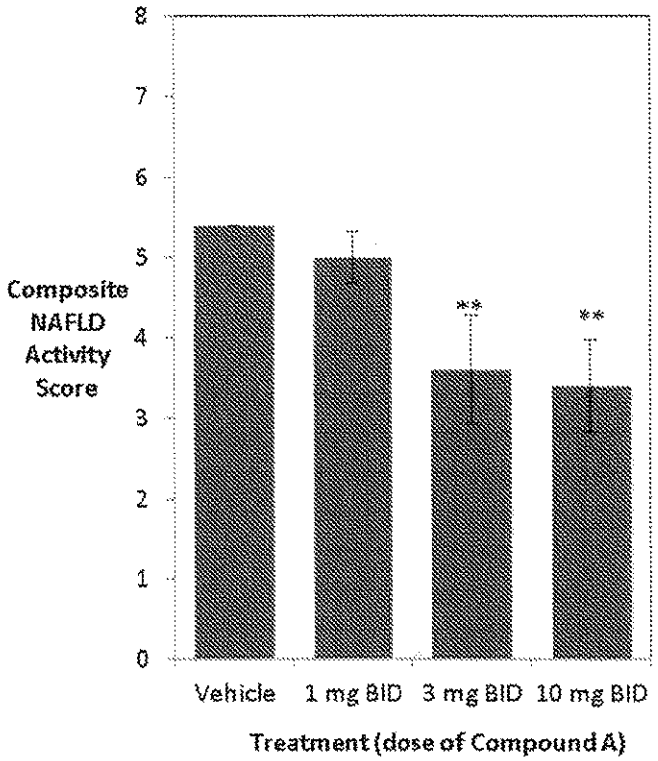

Composite NAFLD Activity Score
8
7
6
5
4
3
2
1
0
**
**
Vehicle
1 mg BID
3 mg BID
10 mg BID
Treatment (dose of Compound A)

Figure 8

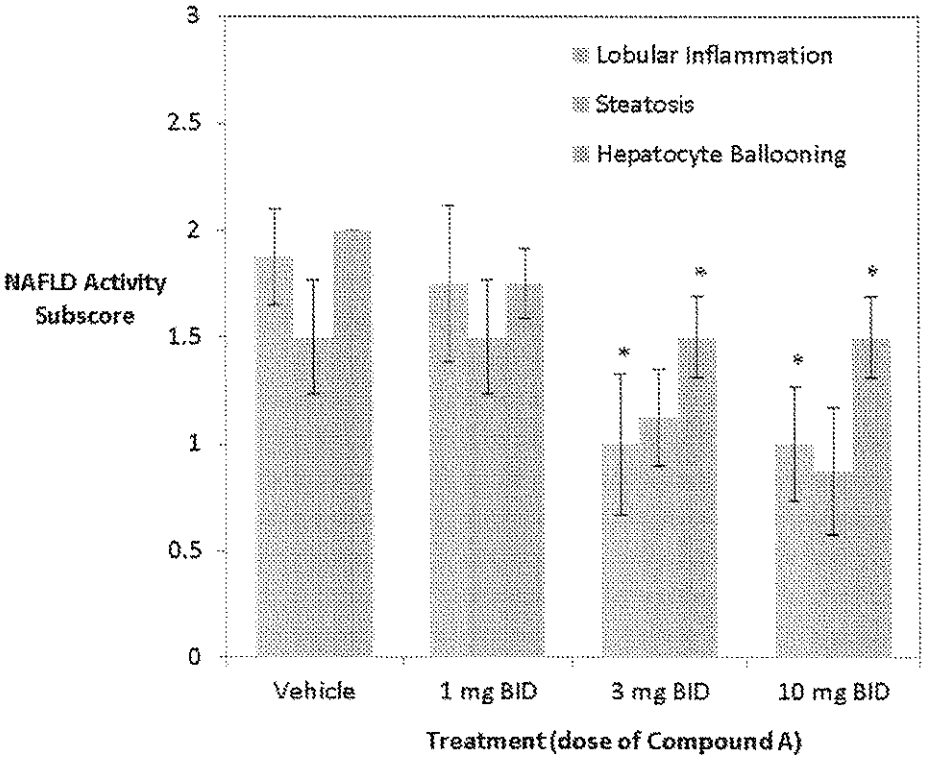

3
2.5
2
1.5
1
0.5
0
NAFLD Activity Subscore
Lobular Inflammation
Steatosis
Hepatocyte Ballooning
*
*
*
*
Vehicle
1 mg BID
3 mg BID
10 mg BID
Treatment (dose of Compound A)

Figure 9

Whole Blood HbA_{1c}

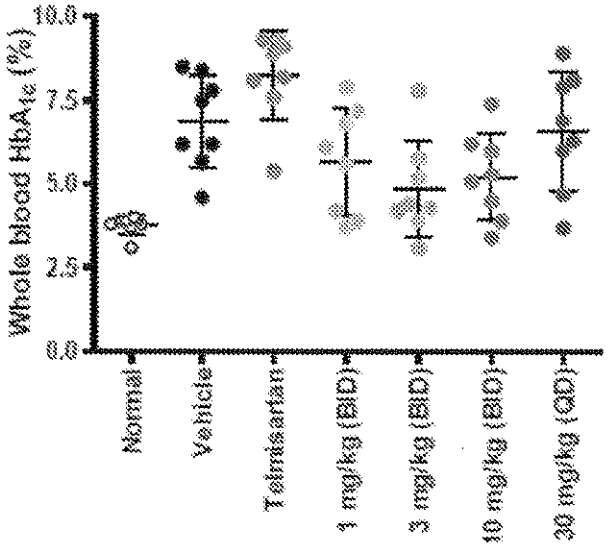


Figure 10

Plasma Total Cholesterol

