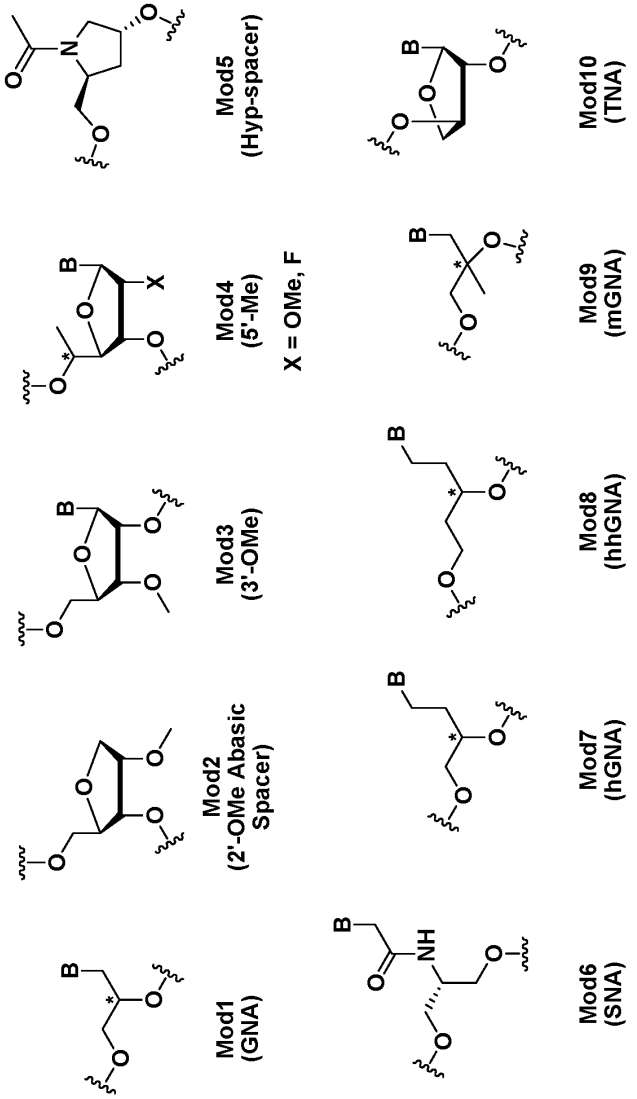

Mod1 (GNA)
Mod2 (2'-OMe Abasic Spacer)
Mod3 (3'-OMe)
Mod4 (5'-Me)
X = OMe, F
Mod5 (Hyp-spacer)
Mod6 (SNA)
Mod7 (hGNA)
Mod8 (hhGNA)
Mod9 (mGNA)
Mod10 (TNA)
h'GNA
*Both stereoisomers tested

FIG. 1

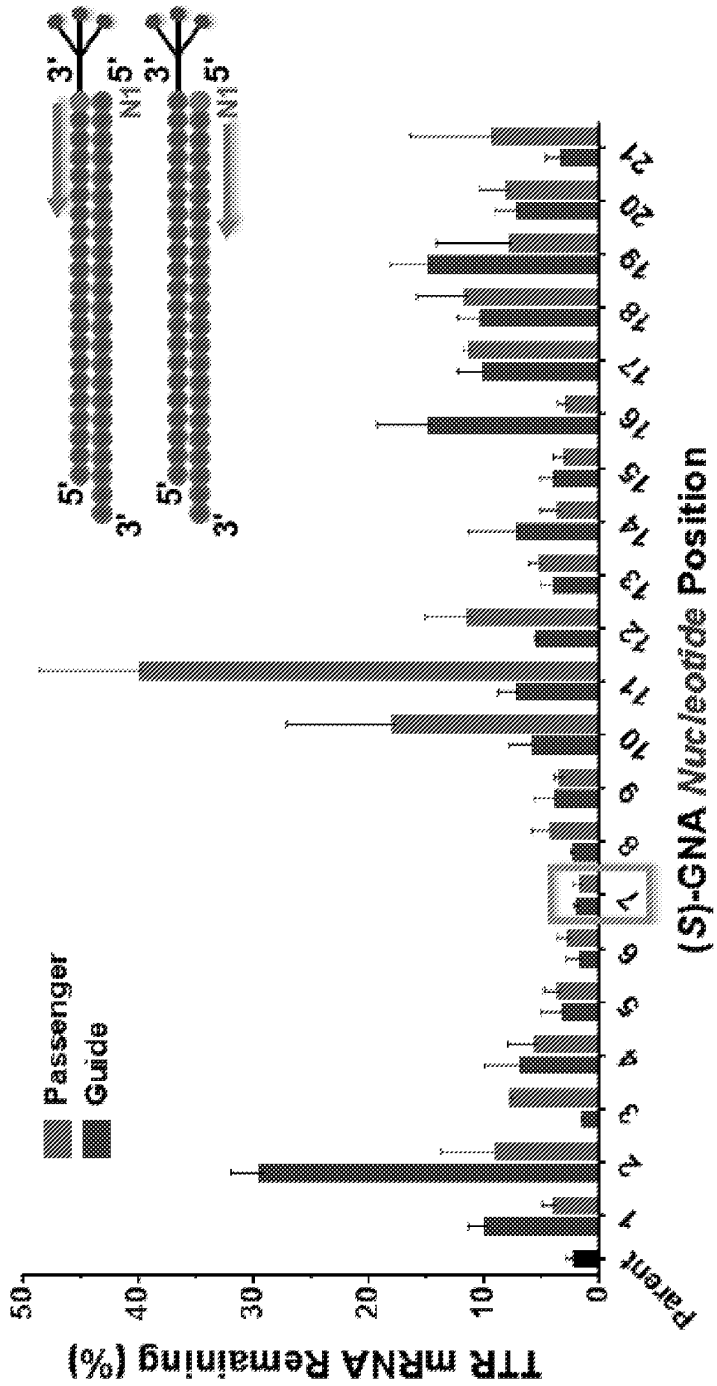

TTR mRNA Remaining (%)
50
40
30
20
10
0
Passenger
Guide
Parent
1
2
3
4
5
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
(S)-GNA Nucleotide Position
Relative to 5'-Guide
3'
5'
N1
5'
3'
3'
5'
N1

FIG. 2

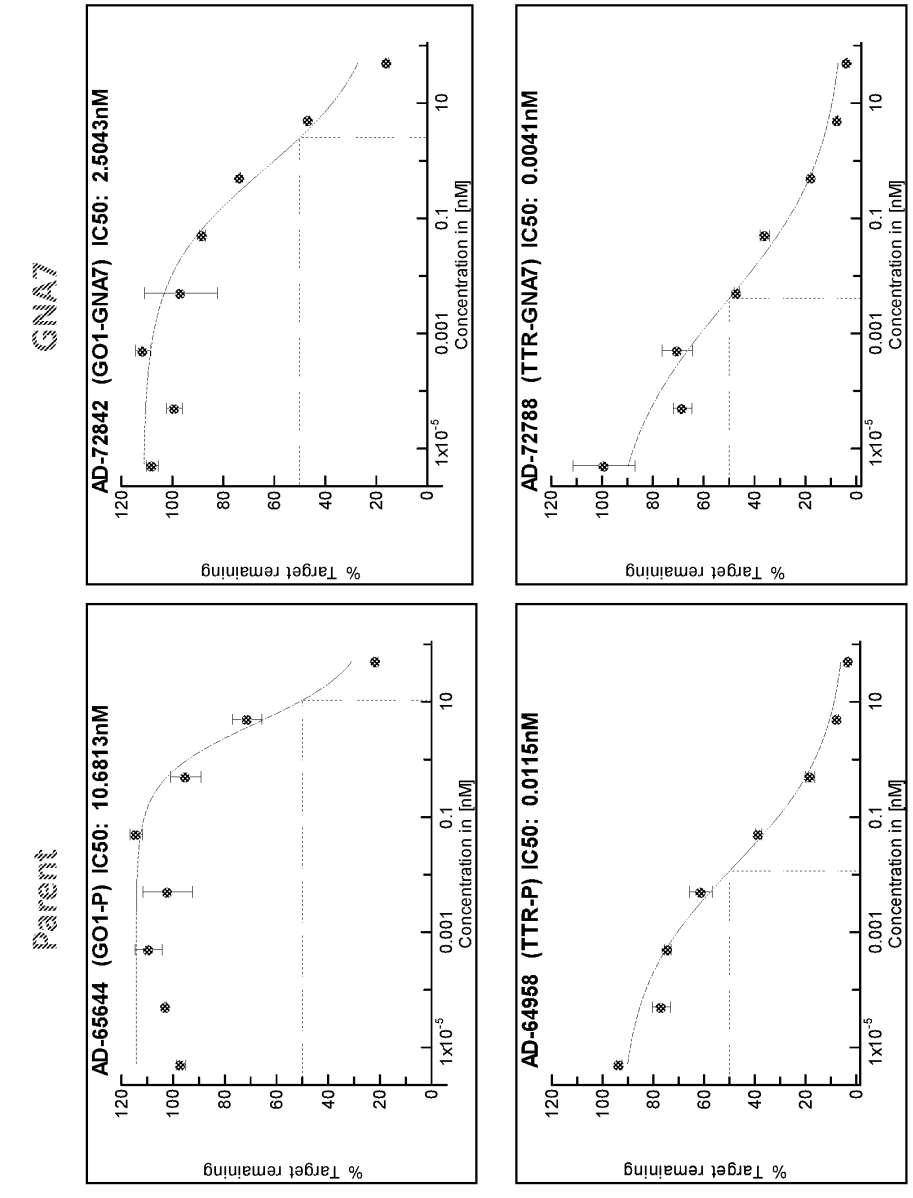

Parent
GNA7
GO1
TTR
AD-65644 (GO1-P) IC50: 10.6813nM
AD-72842 (GO1-GNA7) IC50: 2.5043nM
AD-64958 (TTR-P) IC50: 0.0115nM
AD-72788 (TTR-GNA7) IC50: 0.0041nM
% Target remaining
Concentration in [nM]
0
20
40
60
80
100
120
1x10-5
0.001
0.1
10

FIG. 3

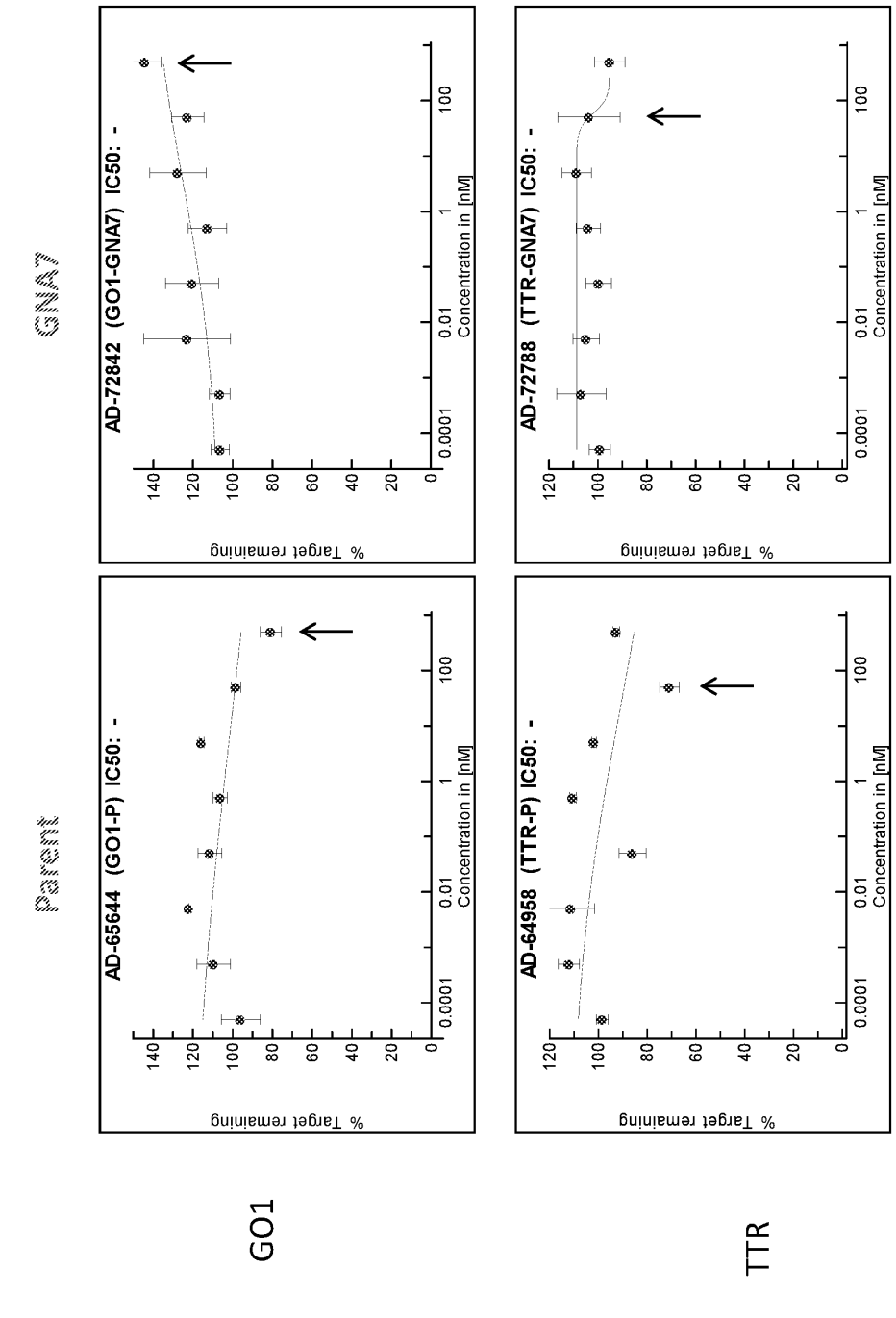

Parent
GNA7
GO1
TTR
AD-65644 (GO1-P) IC50: -
AD-72842 (GO1-GNA7) IC50: -
AD-64958 (TTR-P) IC50: -
AD-72788 (TTR-GNA7) IC50: -
% Target remaining
Concentration in [nM]
0.0001
0.01
1
100
0
20
40
60
80
100
120
140

FIG. 4

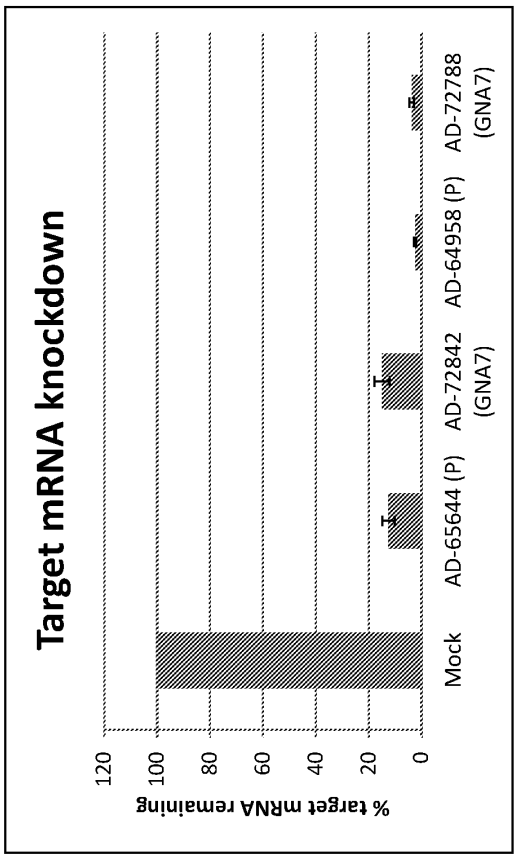
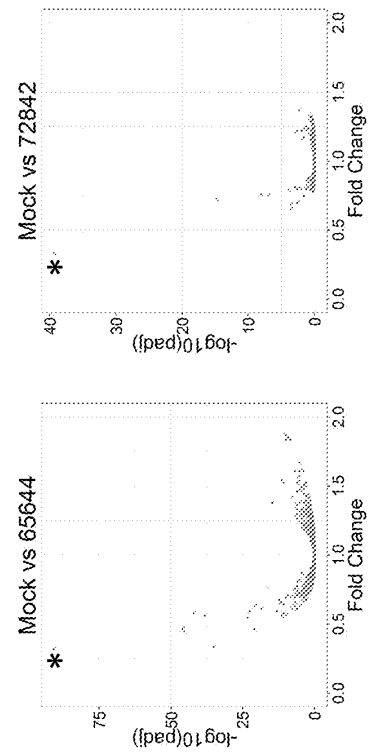

Target mRNA knockdown
% target mRNA remaining
120
100
80
60
40
20
0
Mock
AD-65644 (P)
AD-72842 (GNA7)
AD-64958 (P)
AD-72788 (GNA7)

FIG. 5



	DEG (down)	DEG (up)
AD-65644 (P)	190	160
AD-72842 (GNA7)	15	6
	-93%	-96%

	DEG (down)	DEG (up)
AD-64958 (P)	151	36
AD-72788 (GNA7)	1 (TTR)	5
	-100%	-86%

* on-target

FIG. 6

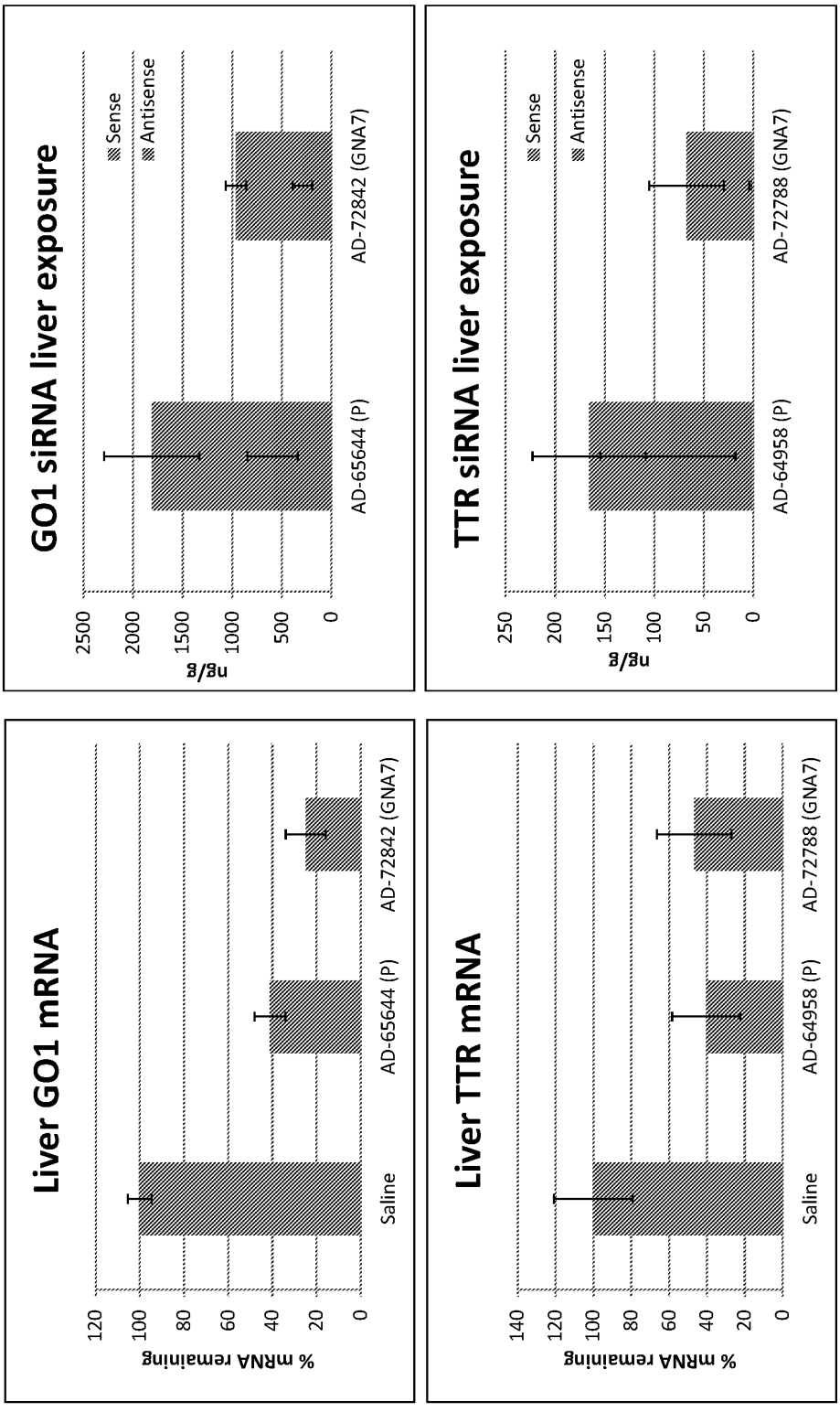

GO1 siRNA liver exposure
Sense
Antisense
ng/g
2500
2000
1500
1000
500
0
AD-65644 (P)
AD-72842 (GNA7)
TTR siRNA liver exposure
Sense
Antisense
ng/g
250
200
150
100
50
0
AD-64958 (P)
AD-72788 (GNA7)
Liver GO1 mRNA
% mRNA remaining
120
100
80
60
40
20
0
Saline
AD-65644 (P)
AD-72842 (GNA7)
Liver TTR mRNA
% mRNA remaining
140
120
100
80
60
40
20
0
Saline
AD-64958 (P)
AD-72788 (GNA7)

FIG. 7

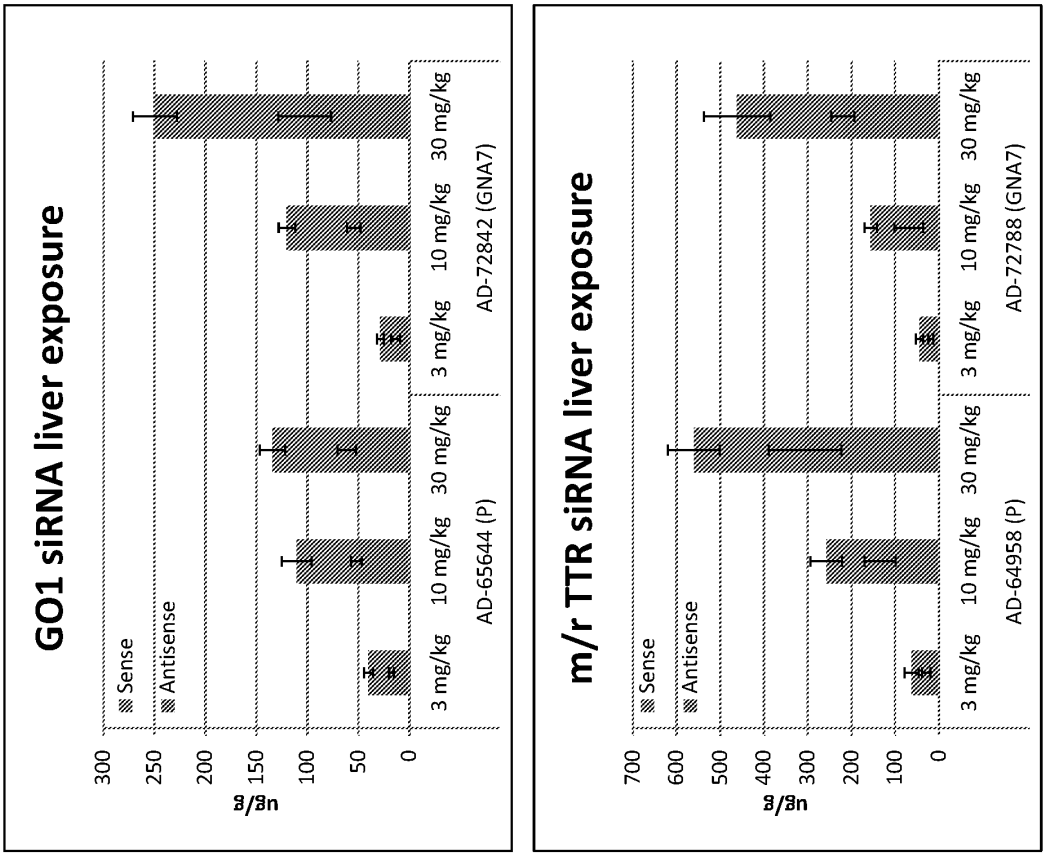

GO1 siRNA liver exposure
Sense
Antisense
ug/g
300
250
200
150
100
50
0
3 mg/kg
10 mg/kg
30 mg/kg
3 mg/kg
10 mg/kg
30 mg/kg
AD-65644 (P)
AD-72842 (GNA7)
m/r TTR siRNA liver exposure
Sense
Antisense
ug/g
700
600
500
400
300
200
100
0
3 mg/kg
10 mg/kg
30 mg/kg
3 mg/kg
10 mg/kg
30 mg/kg
AD-64958 (P)
AD-72788 (GNA7)

FIG. 8

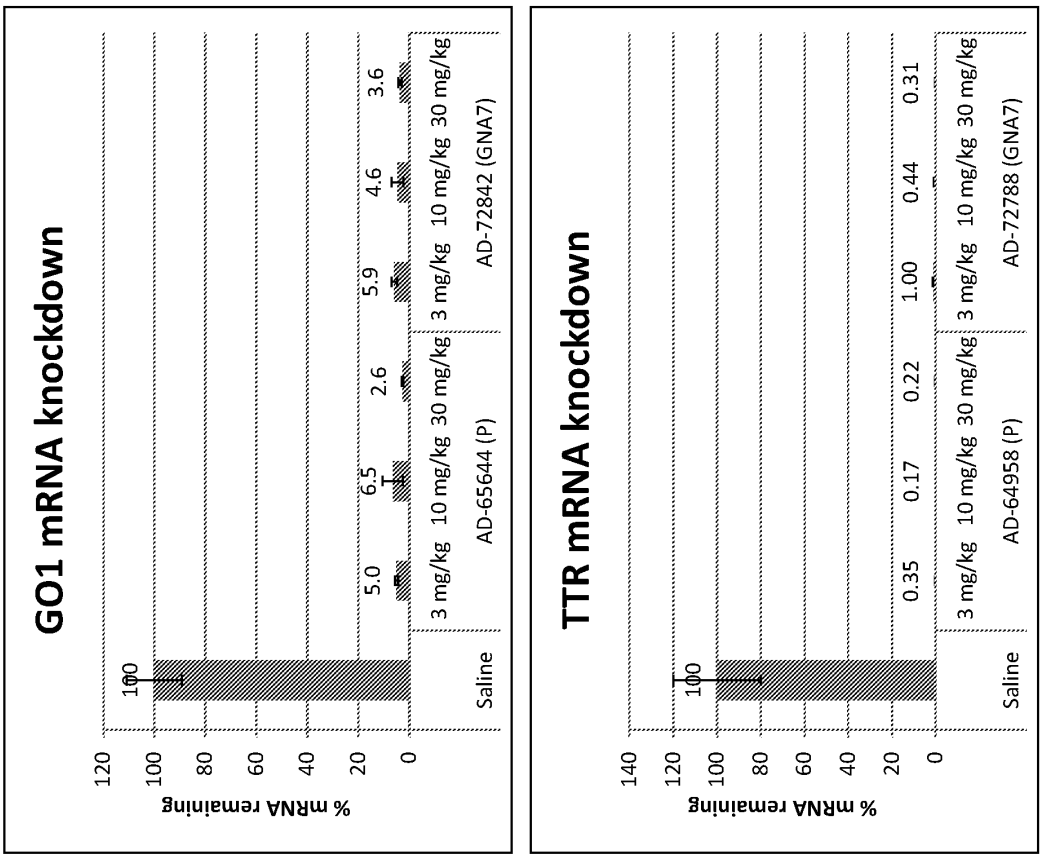

GO1 mRNA knockdown
% mRNA remaining
120
100
80
60
40
20
0
100
5.0
6.5
2.6
5.9
4.6
3.6
Saline
3 mg/kg
10 mg/kg
30 mg/kg
3 mg/kg
10 mg/kg
30 mg/kg
AD-65644 (P)
AD-72842 (GNA7)
TTR mRNA knockdown
% mRNA remaining
140
120
100
80
60
40
20
0
100
0.35
0.17
0.22
1.00
0.44
0.31
Saline
3 mg/kg
10 mg/kg
30 mg/kg
3 mg/kg
10 mg/kg
30 mg/kg
AD-64958 (P)
AD-72788 (GNA7)

FIG. 9

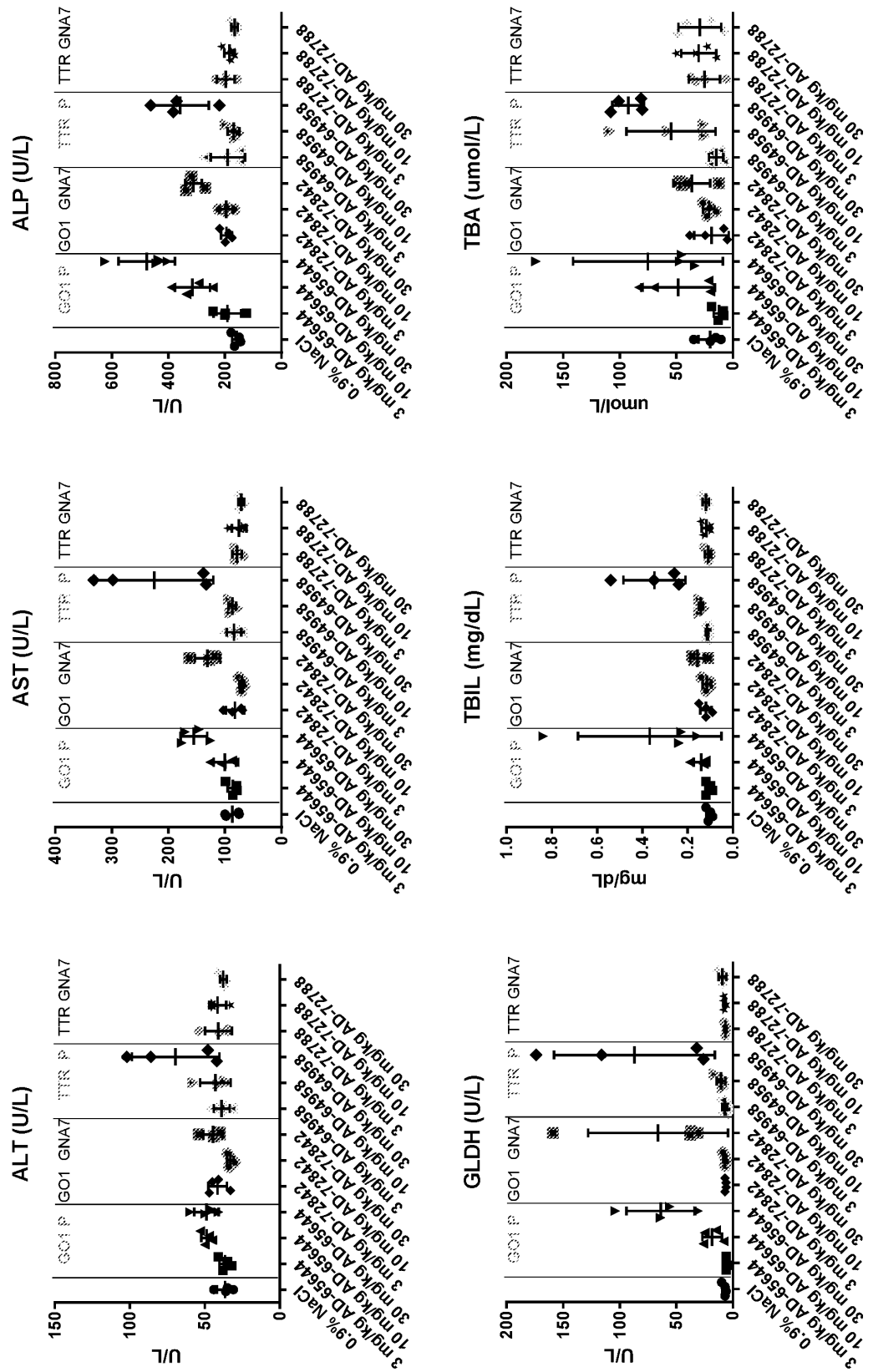

ALT (U/L)
AST (U/L)
ALP (U/L)
GLDH (U/L)
TBIL (mg/dL)
TBA (umol/L)
U/L
mg/dL
umol/L
GO1 P
GO1 GNA7
TTR P
TTR GNA7
0.9% NaCl
3 mg/kg AD-65644
10 mg/kg AD-65644
30 mg/kg AD-65644
3 mg/kg AD-72842
10 mg/kg AD-72842
30 mg/kg AD-72842
3 mg/kg AD-64958
10 mg/kg AD-64958
30 mg/kg AD-64958
3 mg/kg AD-72788
10 mg/kg AD-72788
30 mg/kg AD-72788

FIG. 10

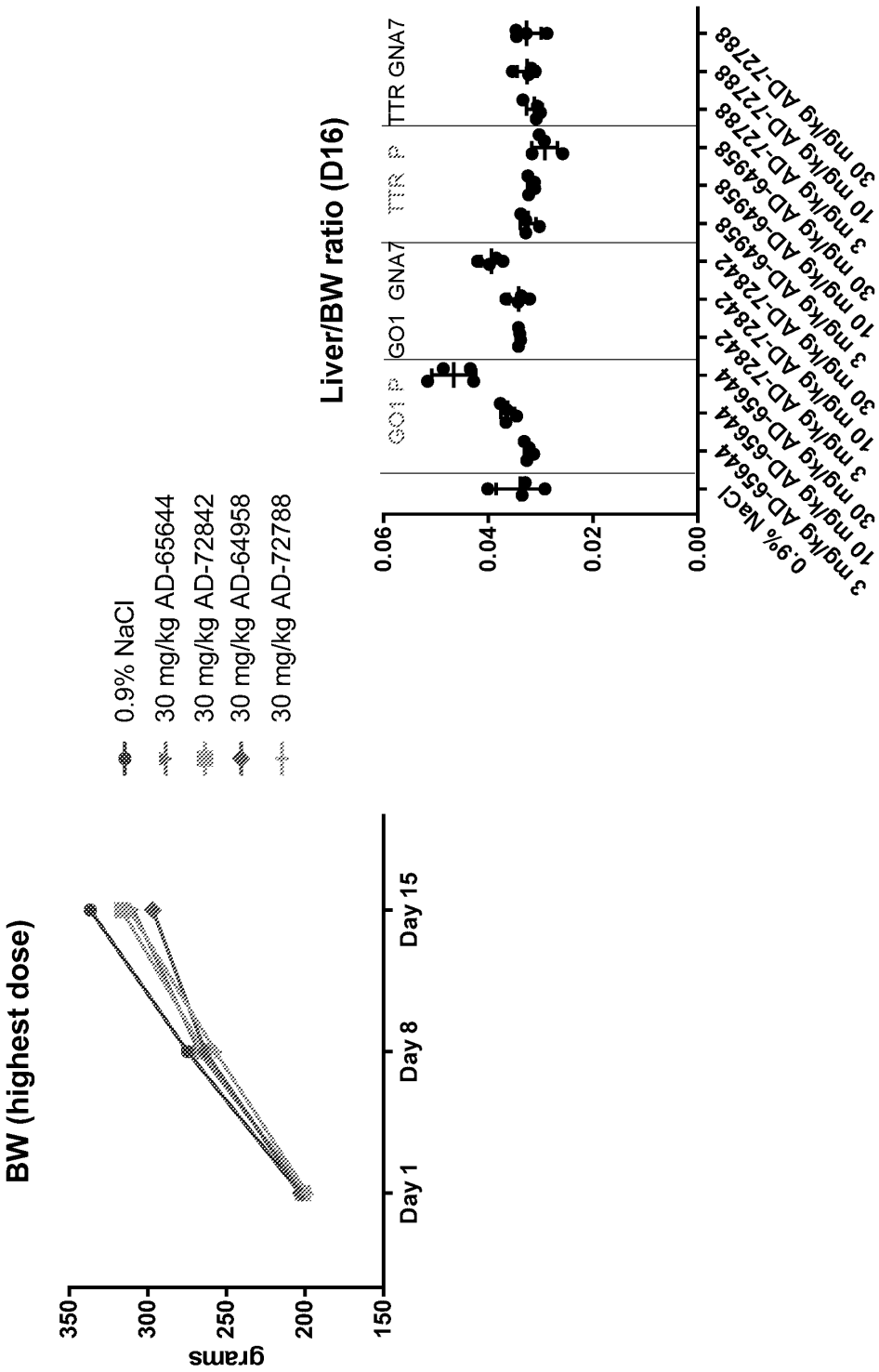

BW (highest dose)
grams
350
300
250
200
150
Day 1
Day 8
Day 15
0.9% NaCl
30 mg/kg AD-65644
30 mg/kg AD-72842
30 mg/kg AD-64958
30 mg/kg AD-72788
Liver/BW ratio (D16)
0.06
0.04
0.02
0.00
GO1 P
GO1 GNA7
TTR P
TTR GNA7
0.9% NaCl
3 mg/kg AD-65644
10 mg/kg AD-65644
30 mg/kg AD-65644
3 mg/kg AD-72842
10 mg/kg AD-72842
30 mg/kg AD-72842
3 mg/kg AD-64958
10 mg/kg AD-64958
30 mg/kg AD-64958
3 mg/kg AD-72788
10 mg/kg AD-72788
30 mg/kg AD-72788

FIG. 11

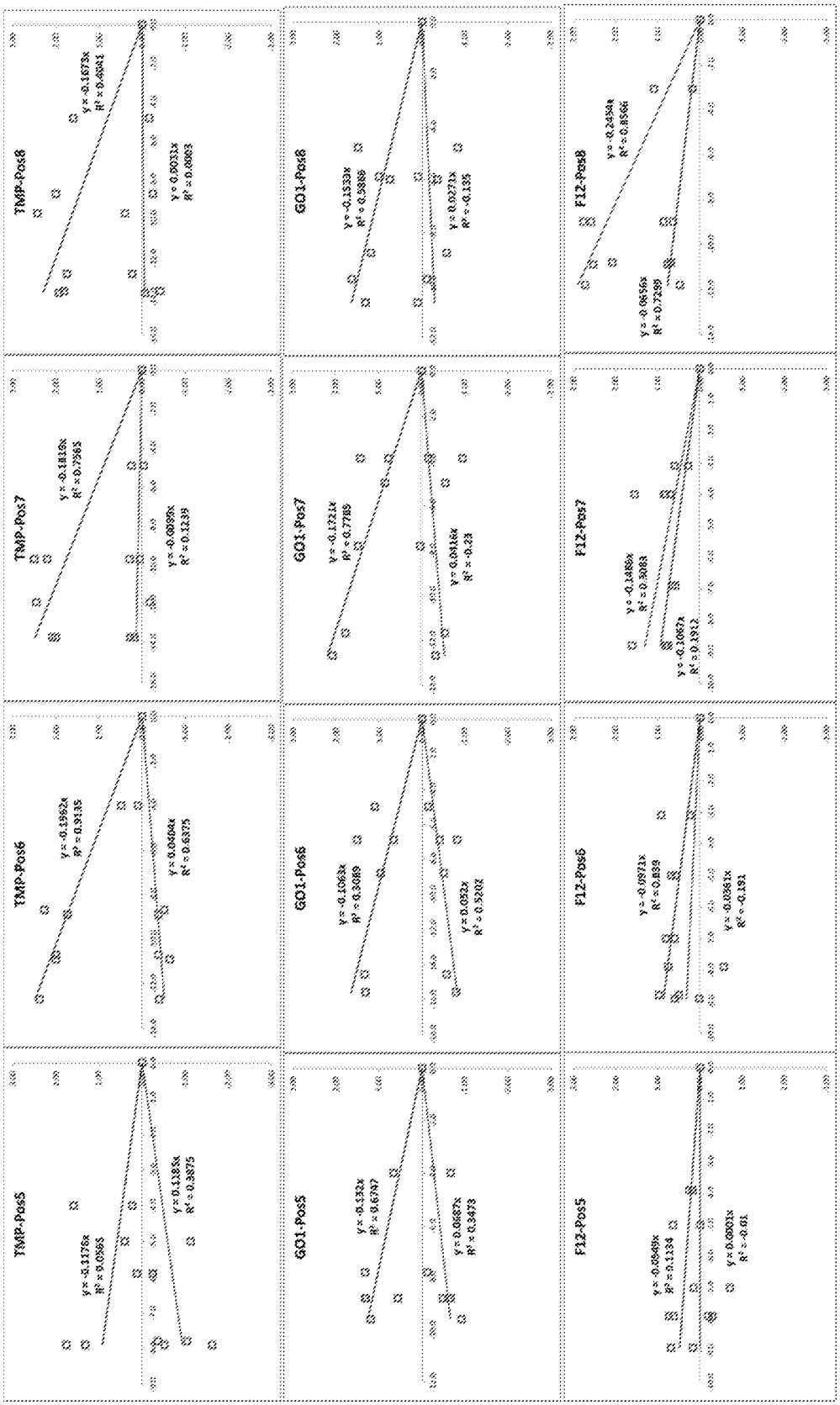

TMP-Pos8
GO1-Pos8
F12-Pos8
TMP-Pos7
GO1-Pos7
F12-Pos7
TMP-Pos6
GO1-Pos6
F12-Pos6
TMP-Pos5
GO1-Pos5
F12-Pos5

FIG. 12

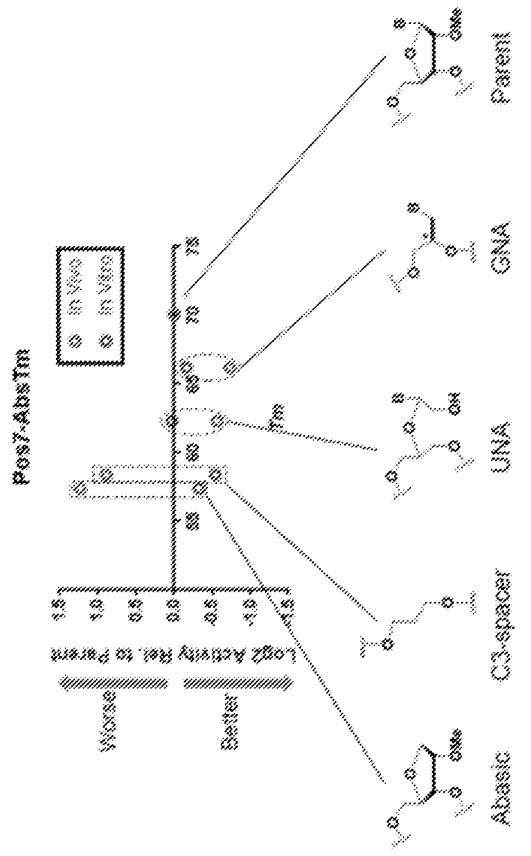

Pos7-AbsTm
In Vivo
In Vitro
Log2 Activity Rel. to Parent
Worse
Better
Parent
GNA
UNA
C3-spacer
Abasic

FIG. 13

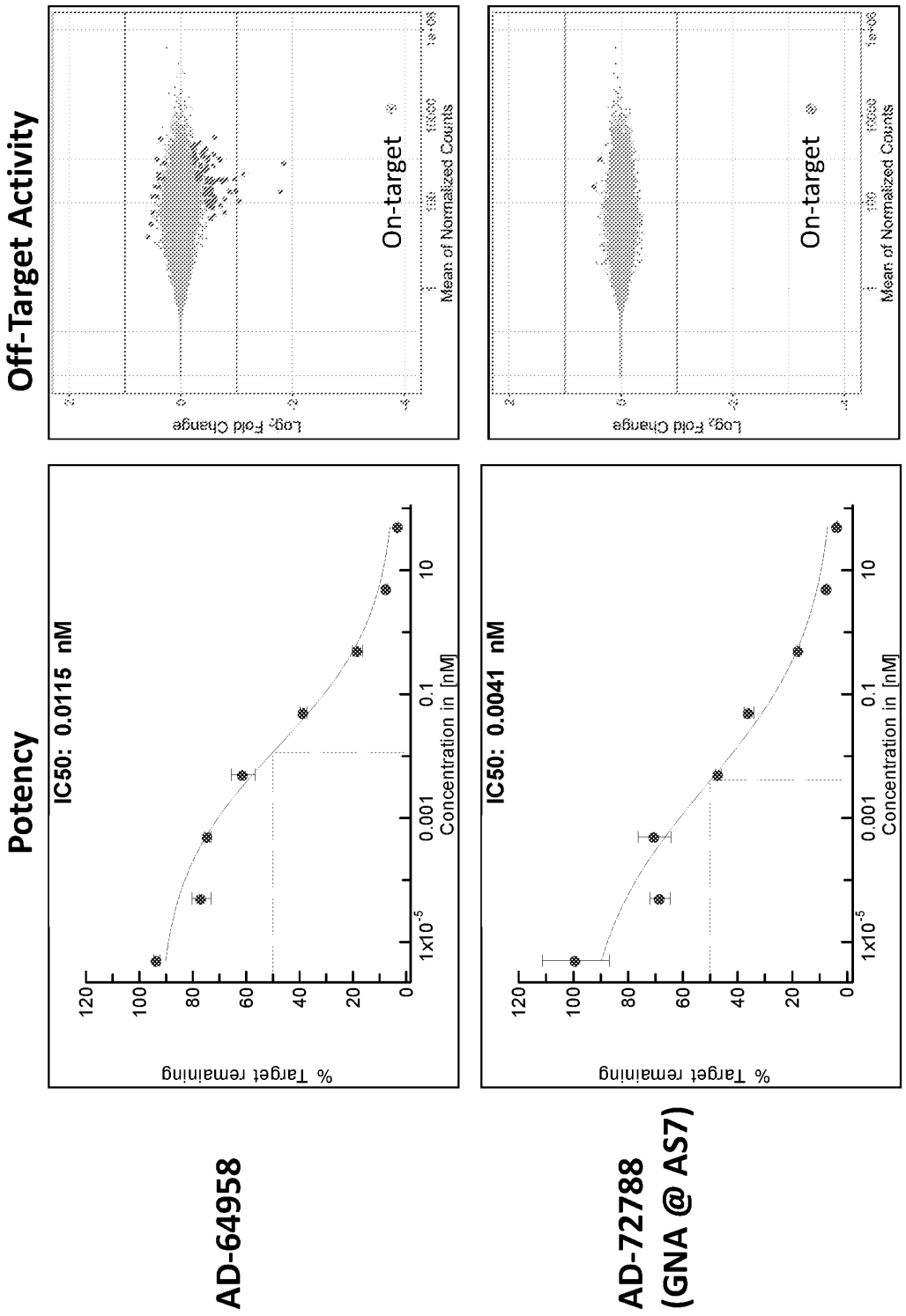

Potency
Off-Target Activity
AD-64958
AD-72788
(GNA @ AS7)
IC50: 0.0115 nM
IC50: 0.0041 nM
% Target remaining
Concentration in [nM]
log2 Fold Change
Mean of Normalized Counts
On-target

FIG. 14

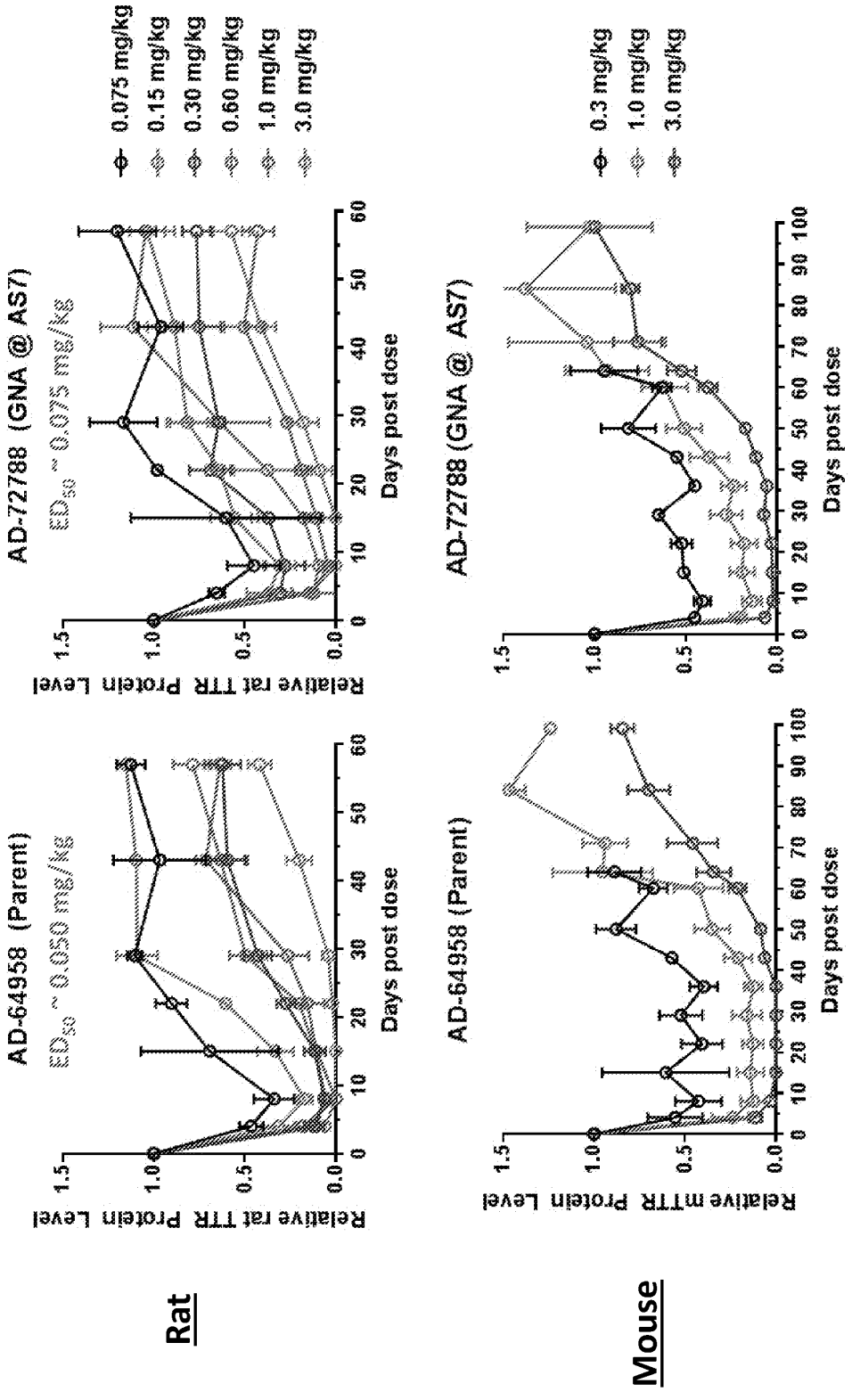
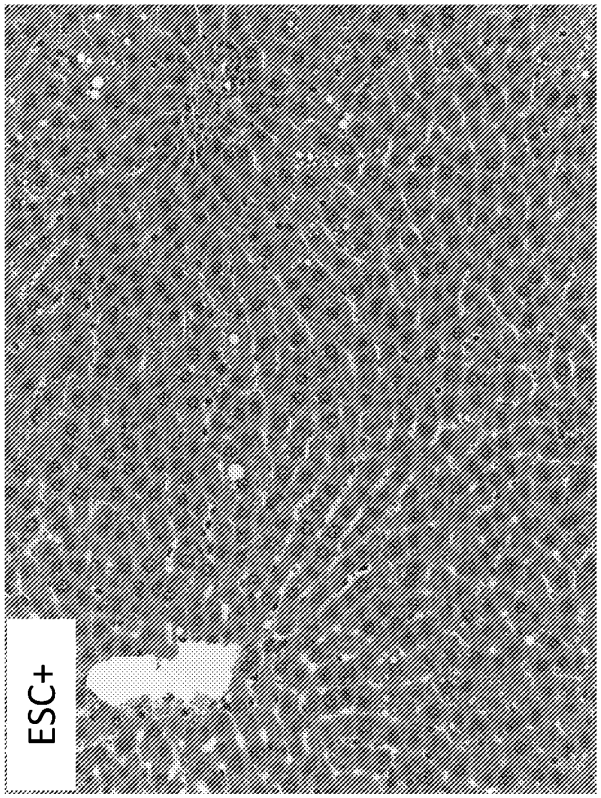

Rat
Mouse
AD-64958 (Parent)
AD-72788 (GNA @ AS7)
ED50 ~ 0.050 mg/kg
ED50 ~ 0.075 mg/kg
Relative rat TTR Protein Level
Relative mTTR Protein Level
Days post dose
0.075 mg/kg
0.15 mg/kg
0.30 mg/kg
0.60 mg/kg
1.0 mg/kg
3.0 mg/kg
0.3 mg/kg
1.0 mg/kg
3.0 mg/kg

FIG. 15



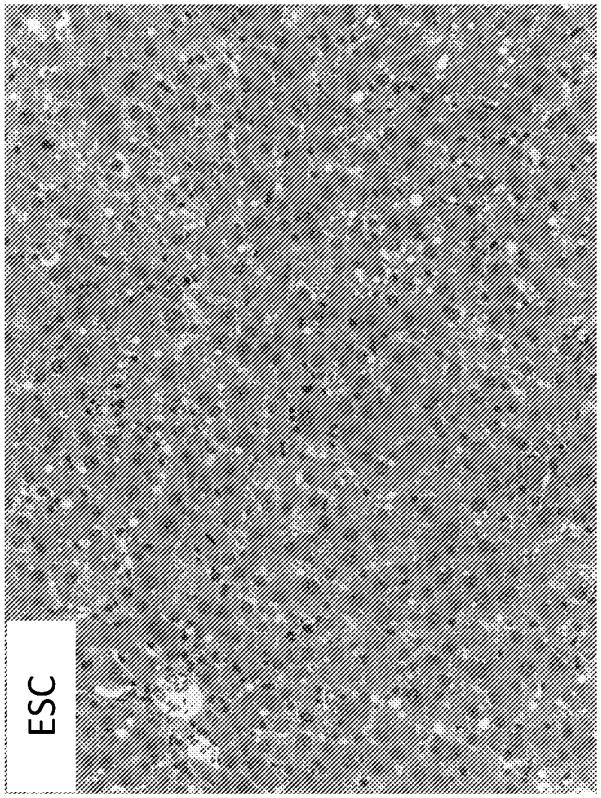


FIG. 16

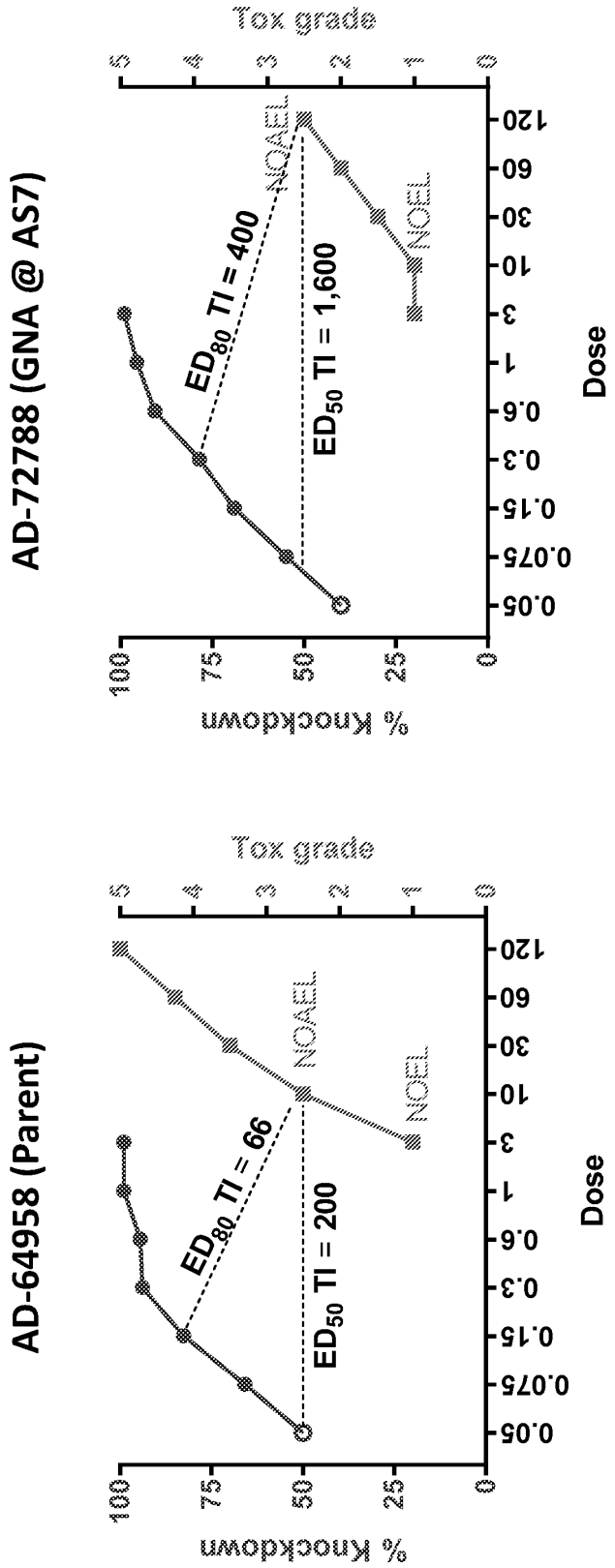

AD-72788 (GNA @ AS7)
% Knockdown
Tox grade
Dose
NOAEL
NOEL
ED80 TI = 400
ED50 TI = 1,600
AD-64958 (Parent)
% Knockdown
Tox grade
Dose
NOAEL
NOEL
ED80 TI = 66
ED50 TI = 200

FIG. 17

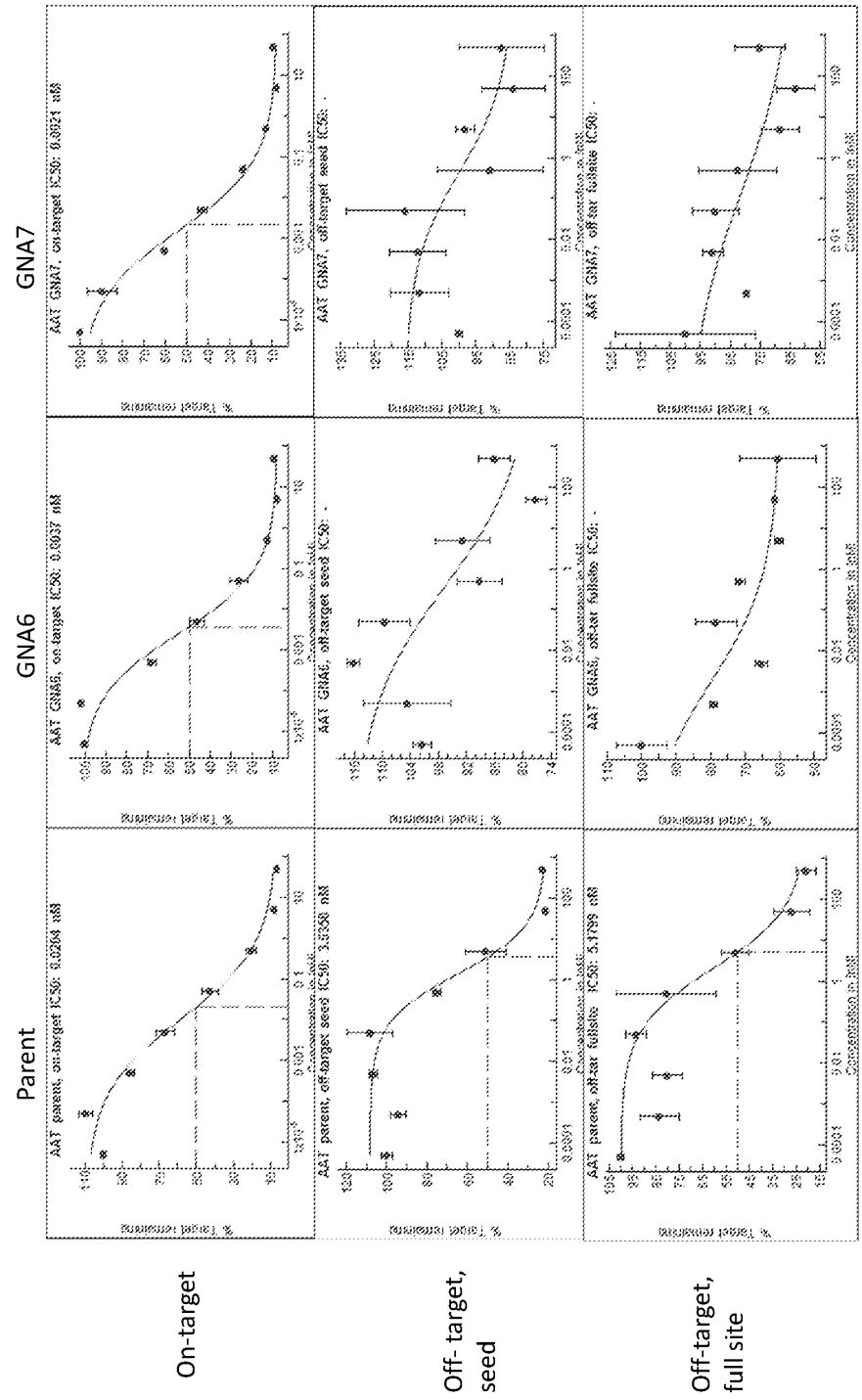

Parent
GNA6
GNA7
On-target
Off- target, seed
Off-target, full site

FIG. 18

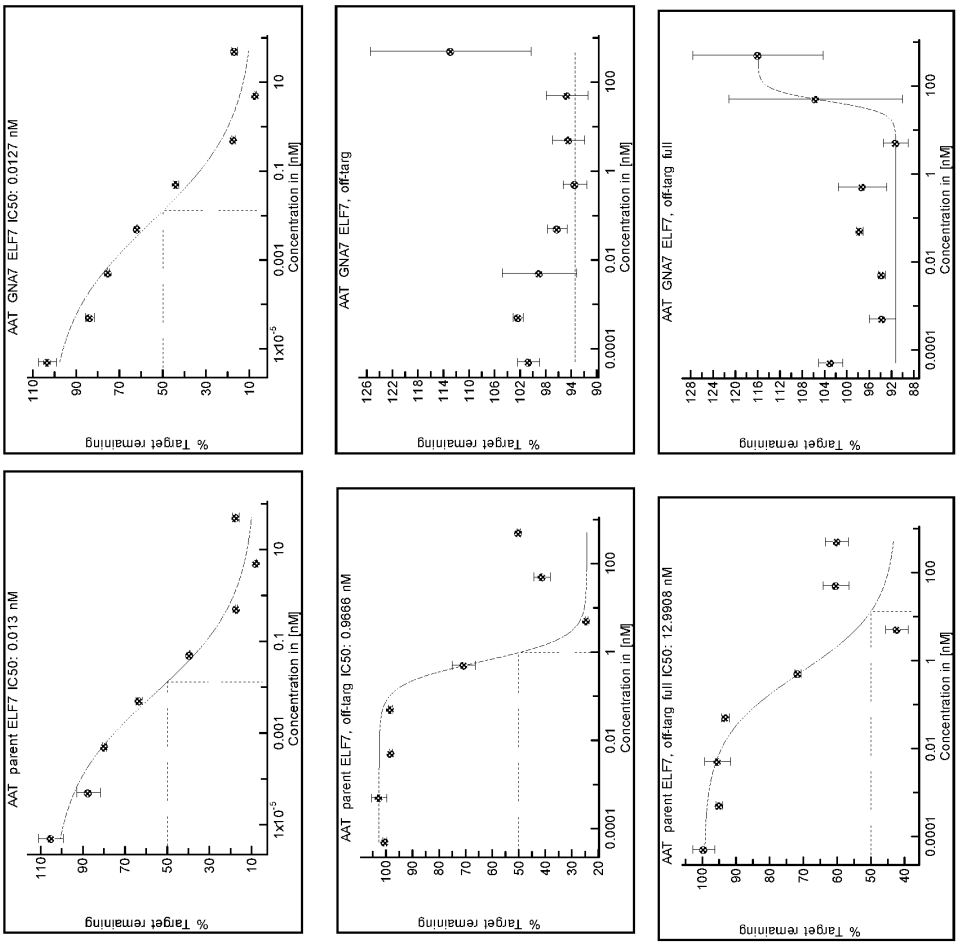
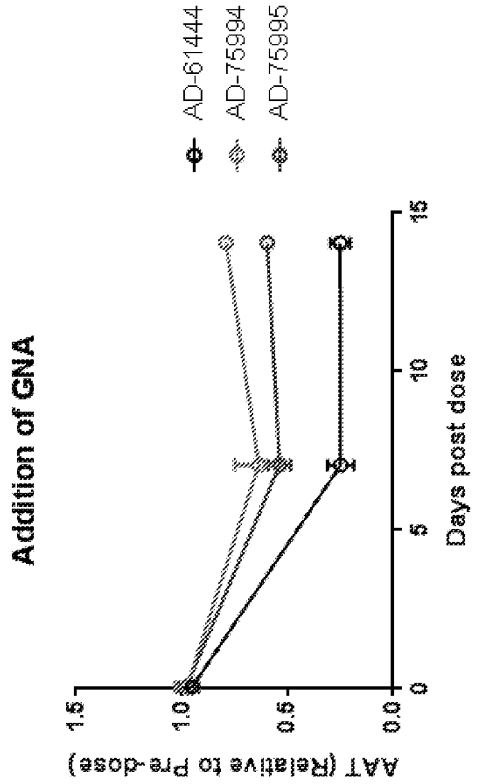

AAT GNA7 ELF7 IC50: 0.0127 nM
AAT parent ELF7 IC50: 0.013 nM
AAT GNA7 ELF7, off-targ
AAT parent ELF7, off-targ IC50: 0.9666 nM
AAT GNA7 ELF7, off-targ full
AAT parent ELF7, off-targ full IC50: 12.9908 nM
% Target remaining
Concentration in [nM]
On-target
Off- target, seed
Off-target, full site

FIG. 19


Addition of GNA
AD-61444
AD-75994
AD-75995
AAT (Relative to Pre-dose)
1.5
1.0
0.5
0.0
0
5
10
15
Days post dose

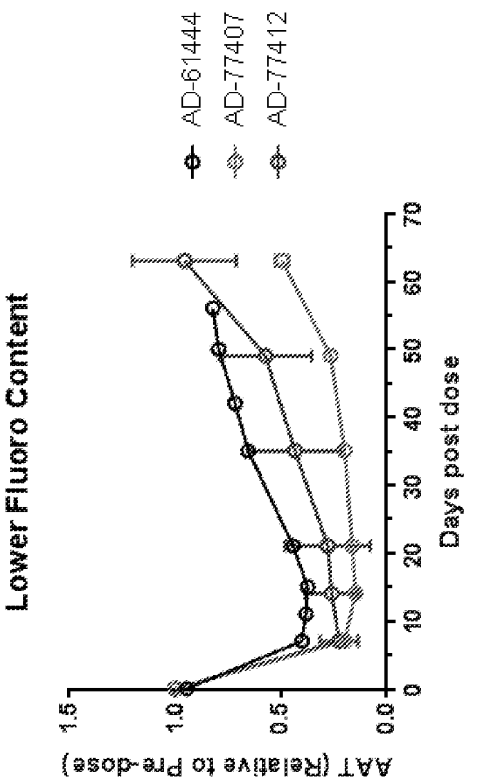

Lower Fluoro Content
AD-61444
AD-77407
AD-77412
AAT (Relative to Pre-dose)
1.5
1.0
0.5
0.0
0
10
20
30
40
50
60
70
Days post dose

FIG. 20

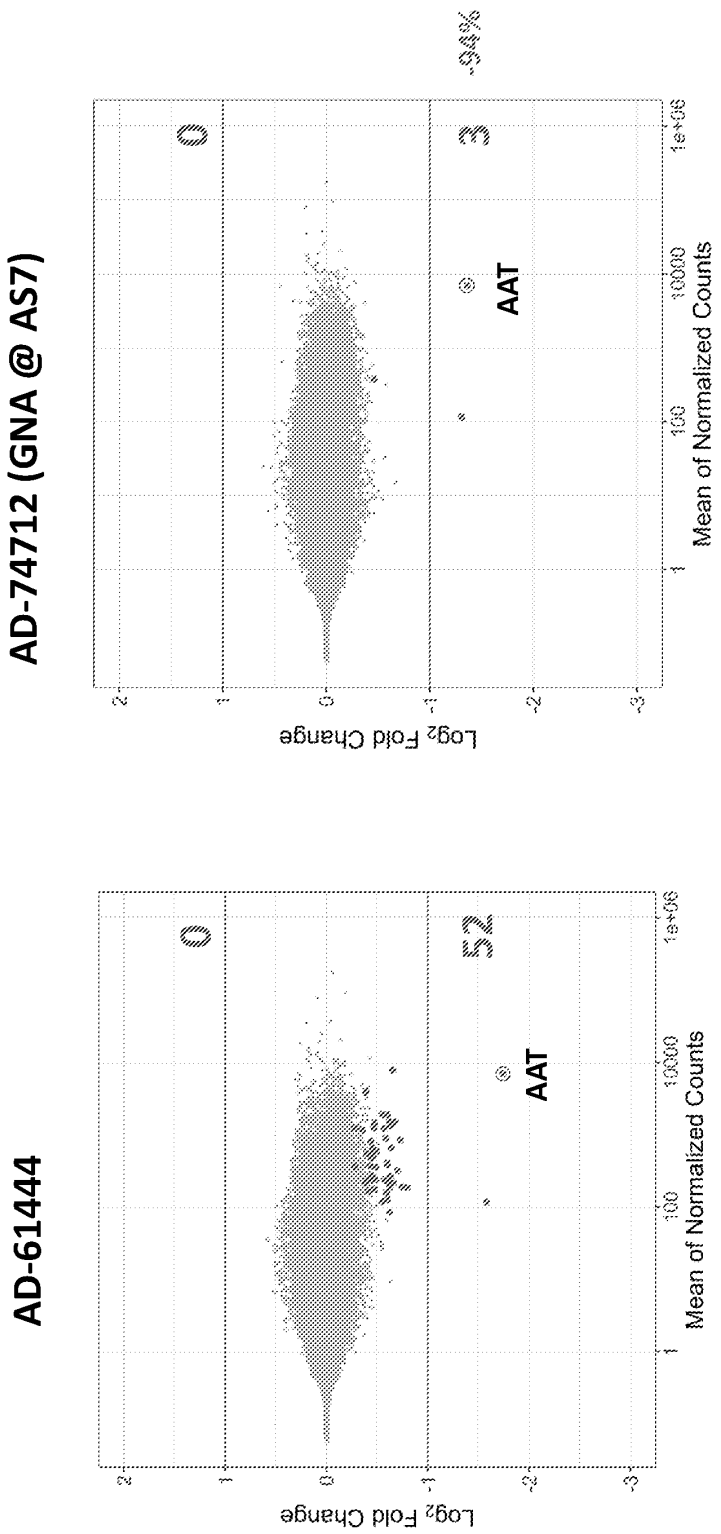

AD-74712 (GNA @ AS7)
0
3
-94%
AAT
Log2 Fold Change
Mean of Normalized Counts
AD-61444
0
52
AAT
Log2 Fold Change
Mean of Normalized Counts

FIG. 21

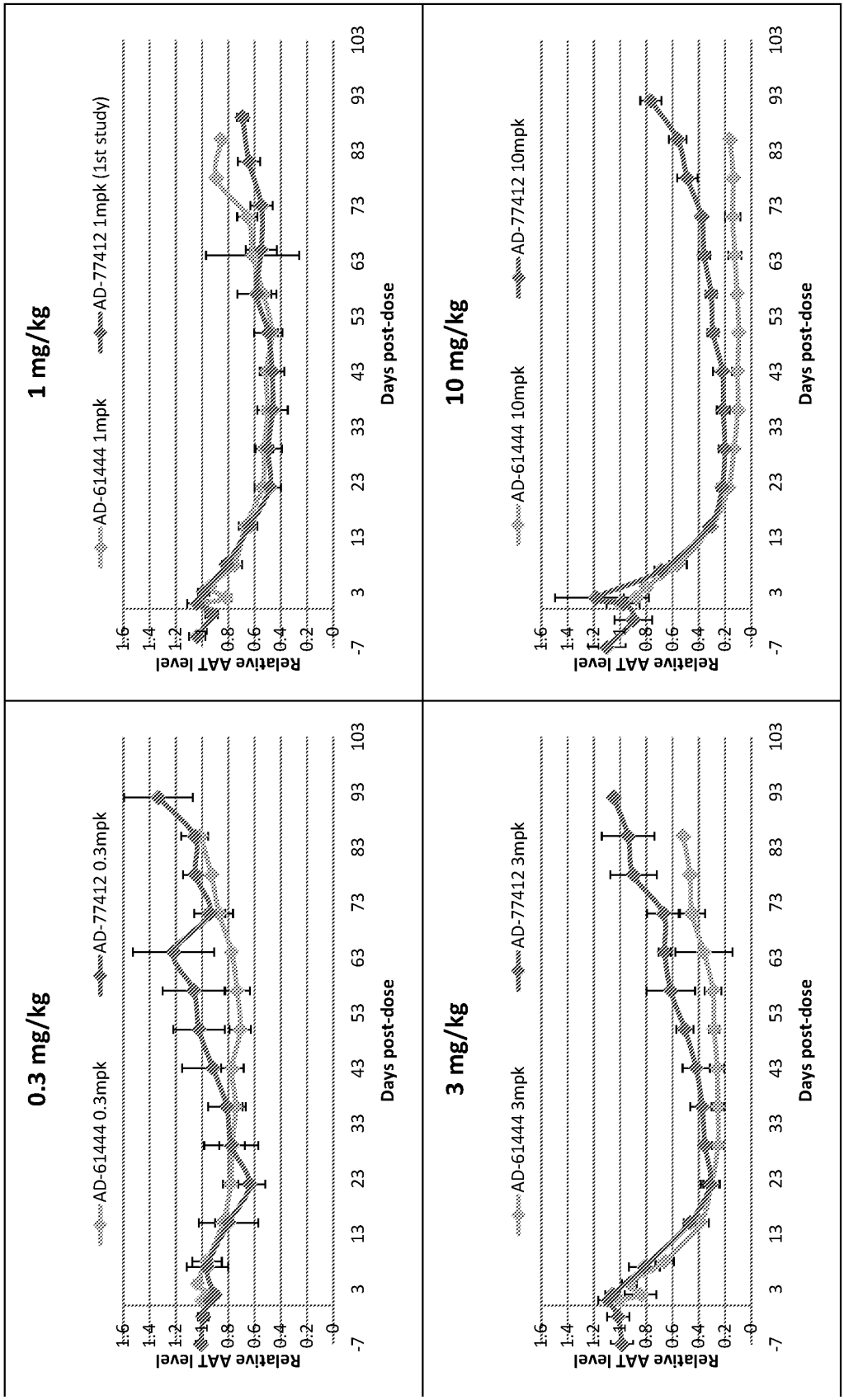

0.3 mg/kg
AD-61444 0.3mpk
AD-77412 0.3mpk
1 mg/kg
AD-61444 1mpk
AD-77412 1mpk (1st study)
3 mg/kg
AD-61444 3mpk
AD-77412 3mpk
10 mg/kg
AD-61444 10mpk
AD-77412 10mpk
Relative AAT level
Days post-dose
-7
3
13
23
33
43
53
63
73
83
93
103
0
0.2
0.4
0.6
0.8
1
1.2
1.4
1.6

FIG. 22

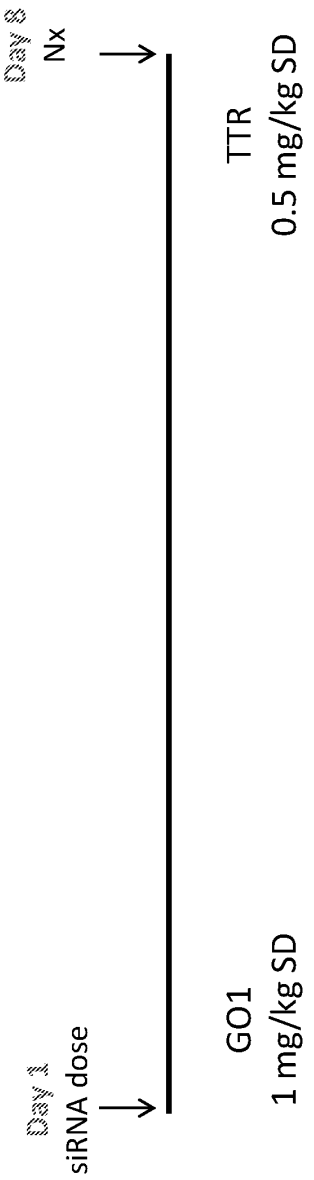


FIG. 23

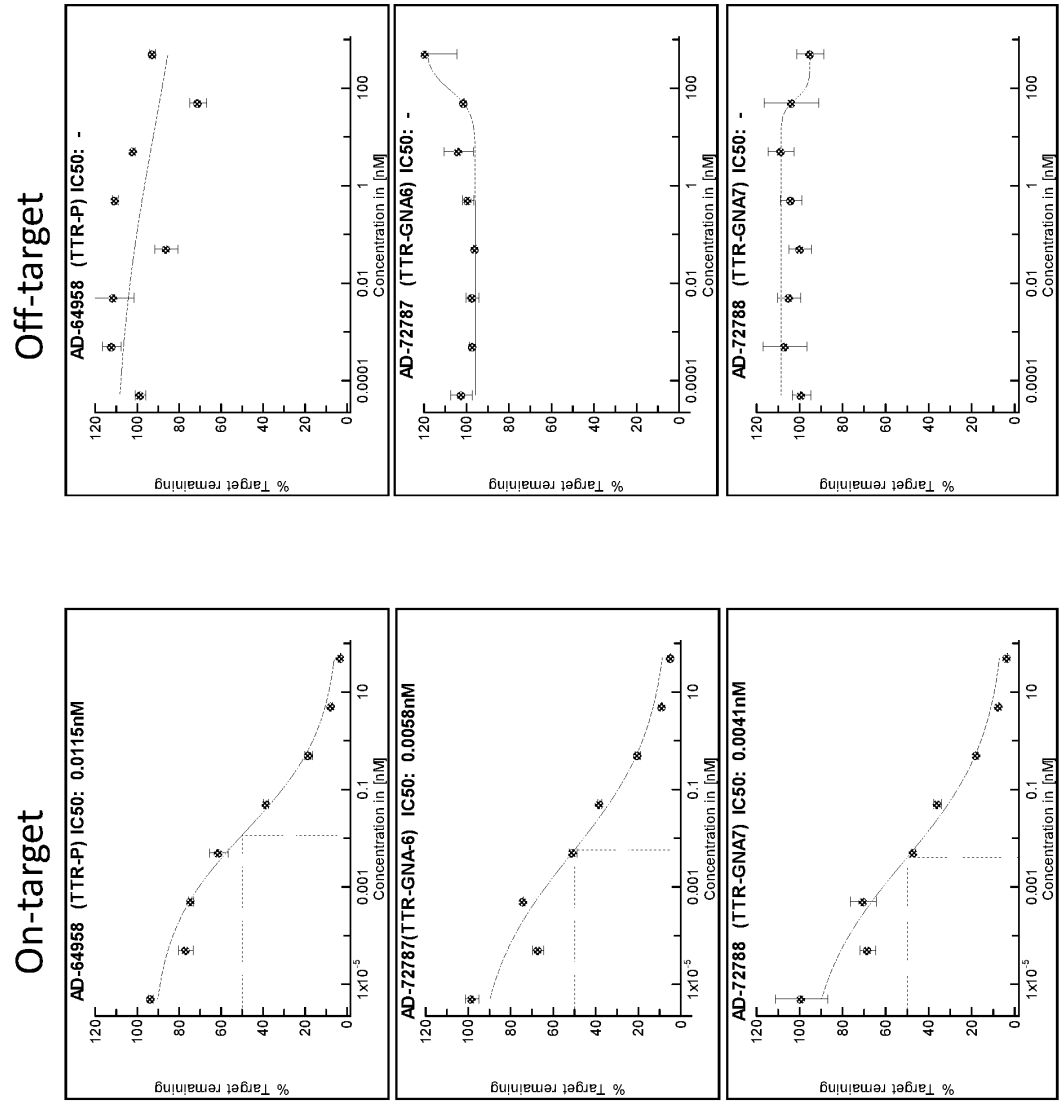

On-target
Off-target
AD-64958 (TTR-P) IC50: 0.0115nM
AD-72787(TTR-GNA-6) IC50: 0.0058nM
AD-72788 (TTR-GNA7) IC50: 0.0041nM
AD-64958 (TTR-P) IC50: -
AD-72787 (TTR-GNA6) IC50: -
AD-72788 (TTR-GNA7) IC50: -
% Target remaining
Concentration in [nM]

FIG. 24

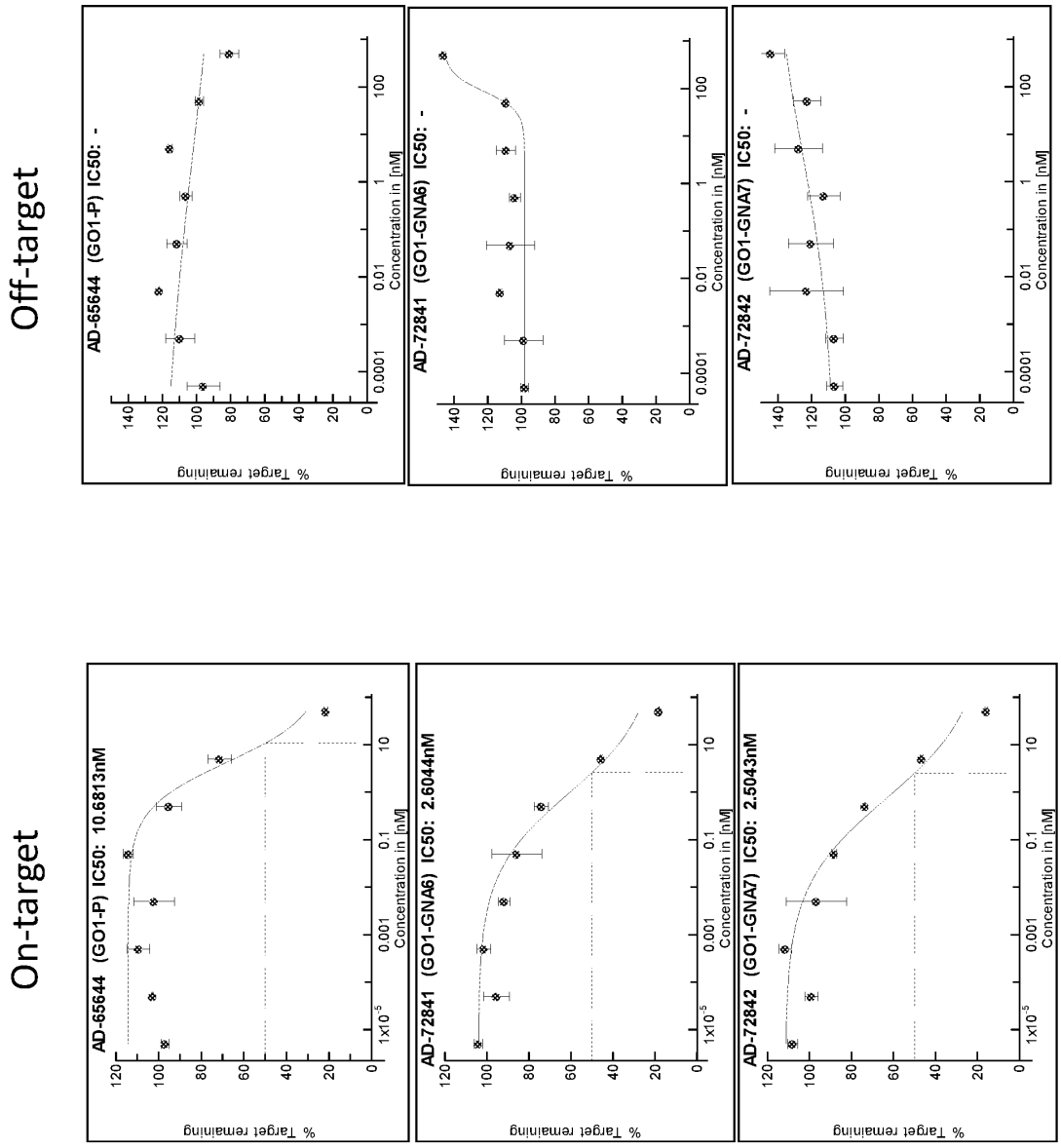

Off-target
AD-65644 (GO1-P) IC50: -
AD-72841 (GO1-GNA6) IC50: -
AD-72842 (GO1-GNA7) IC50: -
On-target
AD-65644 (GO1-P) IC50: 10.6813nM
AD-72841 (GO1-GNA6) IC50: 2.6044nM
AD-72842 (GO1-GNA7) IC50: 2.5043nM
% Target remaining
Concentration in [nM]

FIG. 25

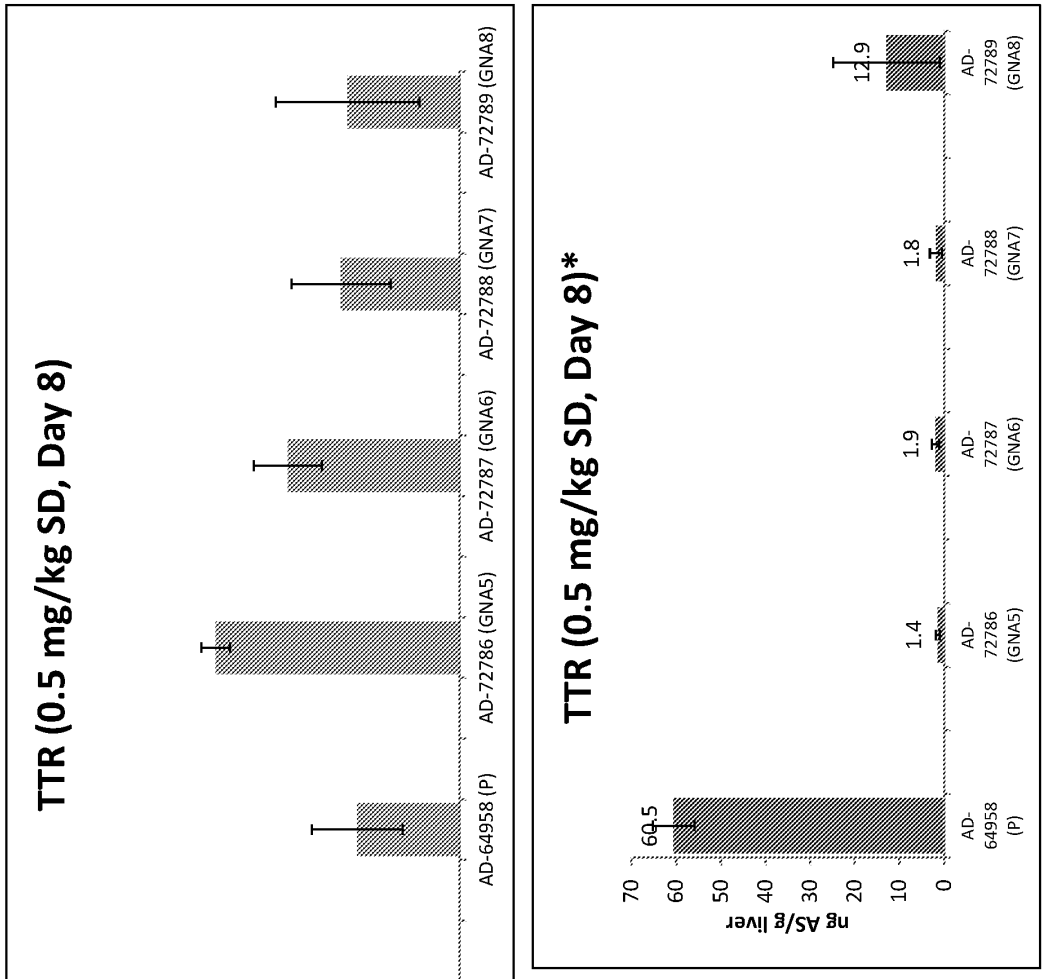

PD
TTR (0.5 mg/kg SD, Day 8)
% TTR mRNA remaining
0
20
40
60
80
100
120
140
Control
AD-64958 (P)
AD-72786 (GNA5)
AD-72787 (GNA6)
AD-72788 (GNA7)
AD-72789 (GNA8)
Test Article Accumulation in Liver
TTR (0.5 mg/kg SD, Day 8)*
ng AS/g liver
0
10
20
30
40
50
60
70
60.5
1.4
1.9
1.8
12.9
AD-64958 (P)
AD-72786 (GNA5)
AD-72787 (GNA6)
AD-72788 (GNA7)
AD-72789 (GNA8)

FIG. 26

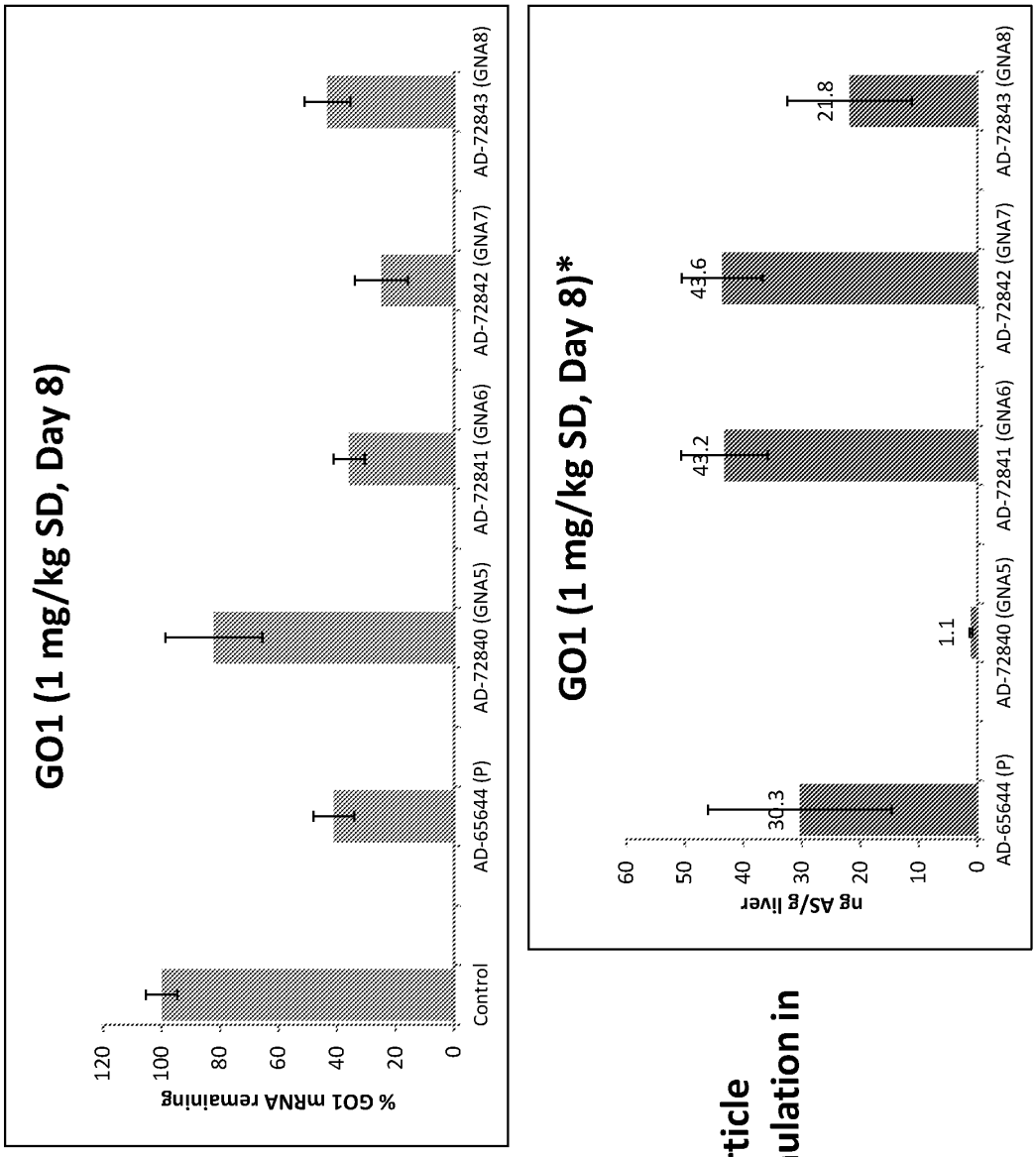

PD
GO1 (1 mg/kg SD, Day 8)
% GO1 mRNA remaining
120
100
80
60
40
20
0
Control
AD-65644 (P)
AD-72840 (GNA5)
AD-72841 (GNA6)
AD-72842 (GNA7)
AD-72843 (GNA8)
Test Article Accumulation in Liver
GO1 (1 mg/kg SD, Day 8)*
ng AS/g liver
60
50
40
30
20
10
0
30.3
1.1
43.2
43.6
21.8
AD-65644 (P)
AD-72840 (GNA5)
AD-72841 (GNA6)
AD-72842 (GNA7)
AD-72843 (GNA8)

FIG. 27

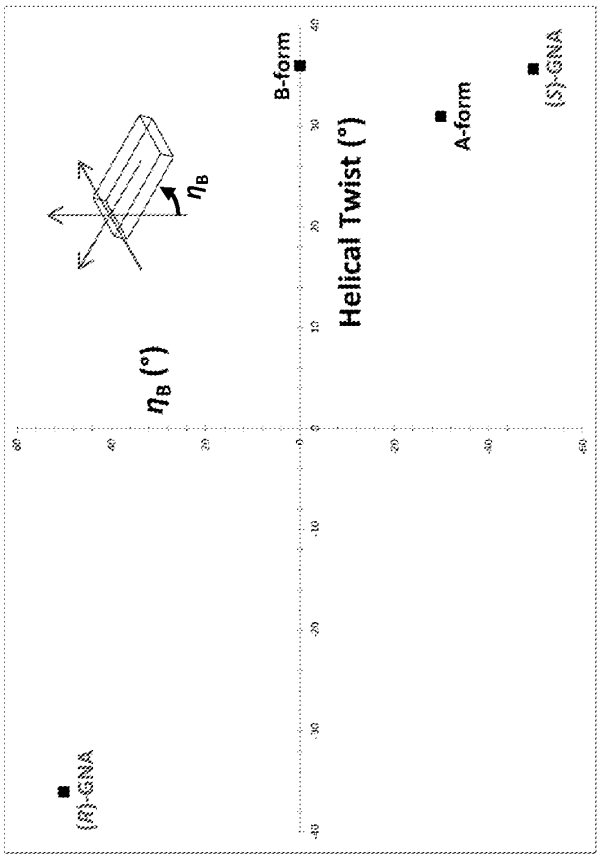

(R)-GNA
η_B (°)
η_B
B-form
A-form
(S)-GNA
Helical Twist (°)

FIG. 29

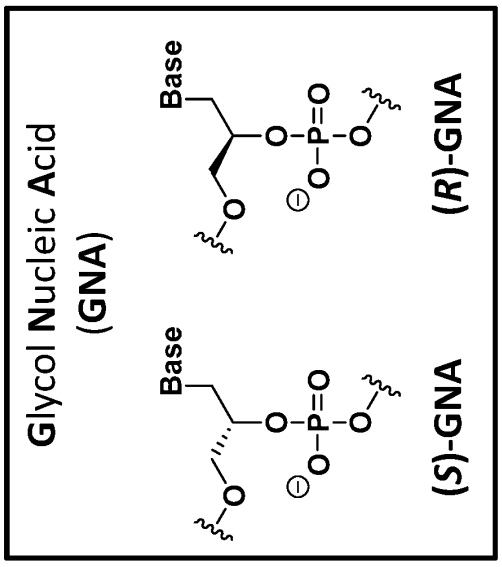

Glycol Nucleic Acid
(GNA)
Base
Base
(S)-GNA
(R)-GNA

FIG. 28

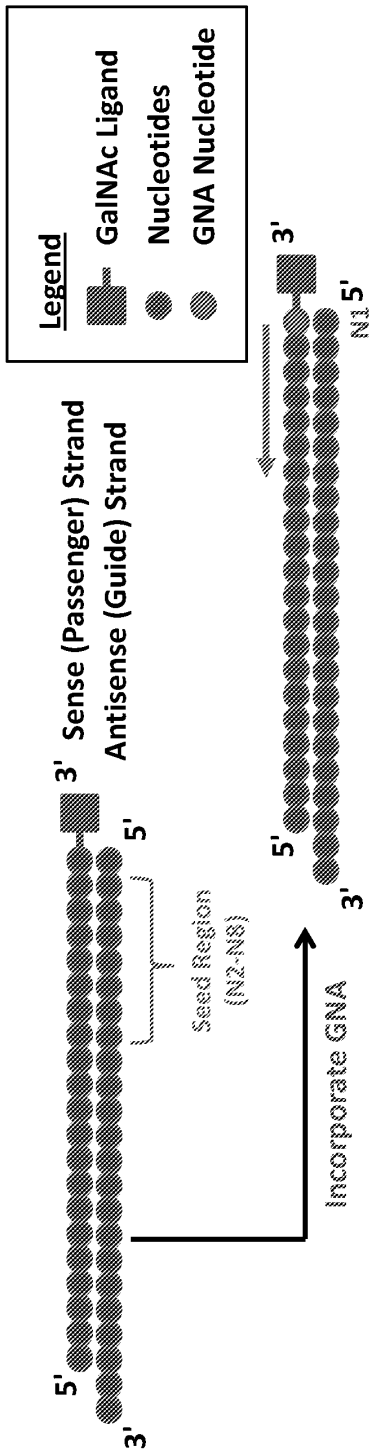

Legend
GalNAc Ligand
Nucleotides
GNA Nucleotide
Sense (Passenger) Strand
Antisense (Guide) Strand
3'
5'
Seed Region
(N2-N8)
Incorporate GNA
N1
5'
3'

FIG. 30

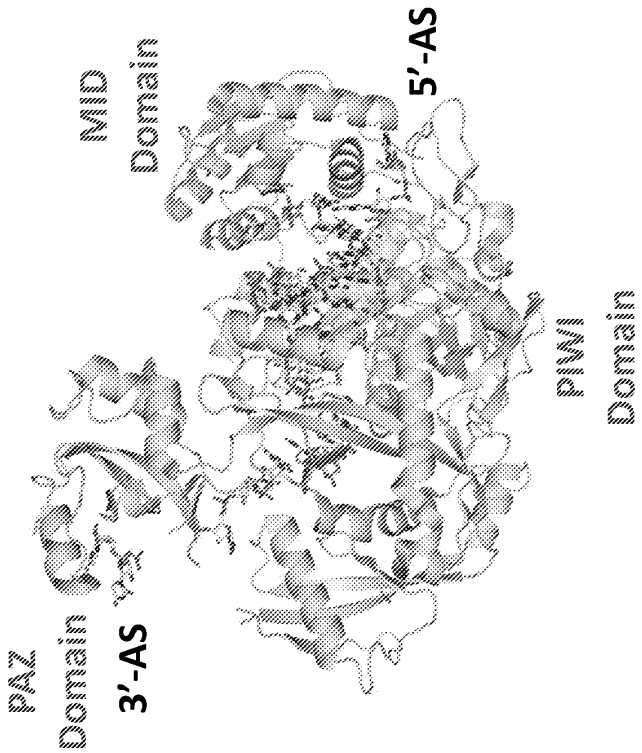


FIG. 31

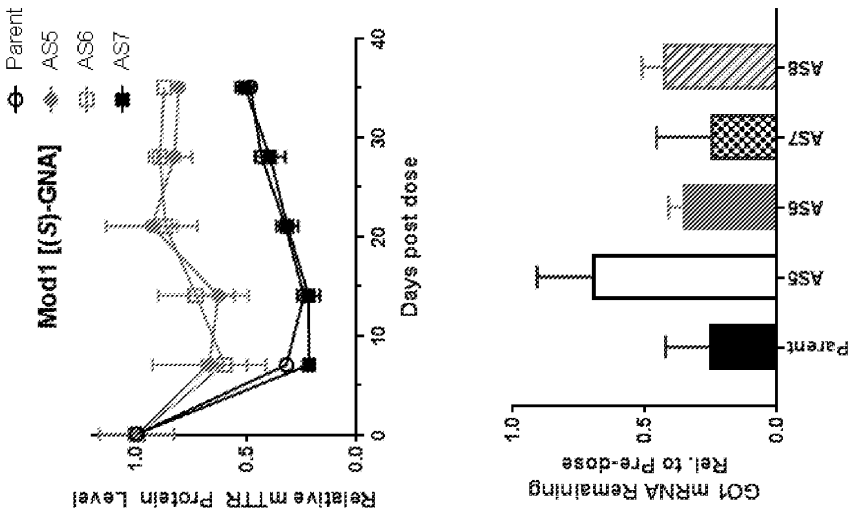


FIG. 32B

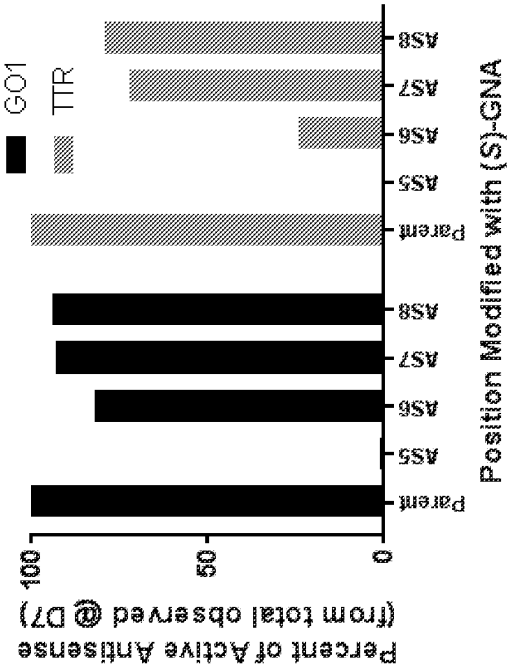


FIG. 32A

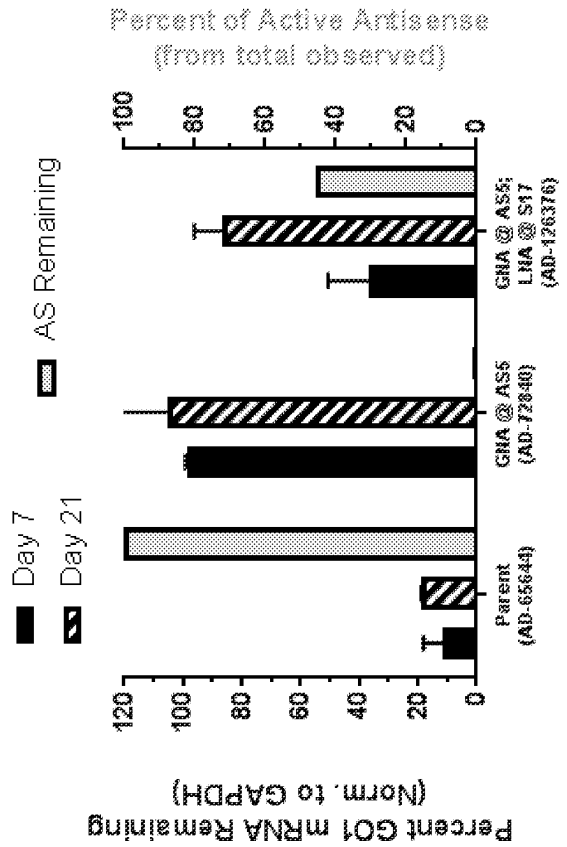

Percent of Active Antisense
(from total observed)
100
80
60
40
20
0
AS Remaining
Day 7
Day 21
GNA @ AS5;
LNA @ S17
(AD-126376)
GNA @ AS5
(AD-72640)
Parent
(AD-65644)
120
100
80
60
40
20
0
Percent GO1 mRNA Remaining
(Norm. to GAPDH)

FIG. 32C

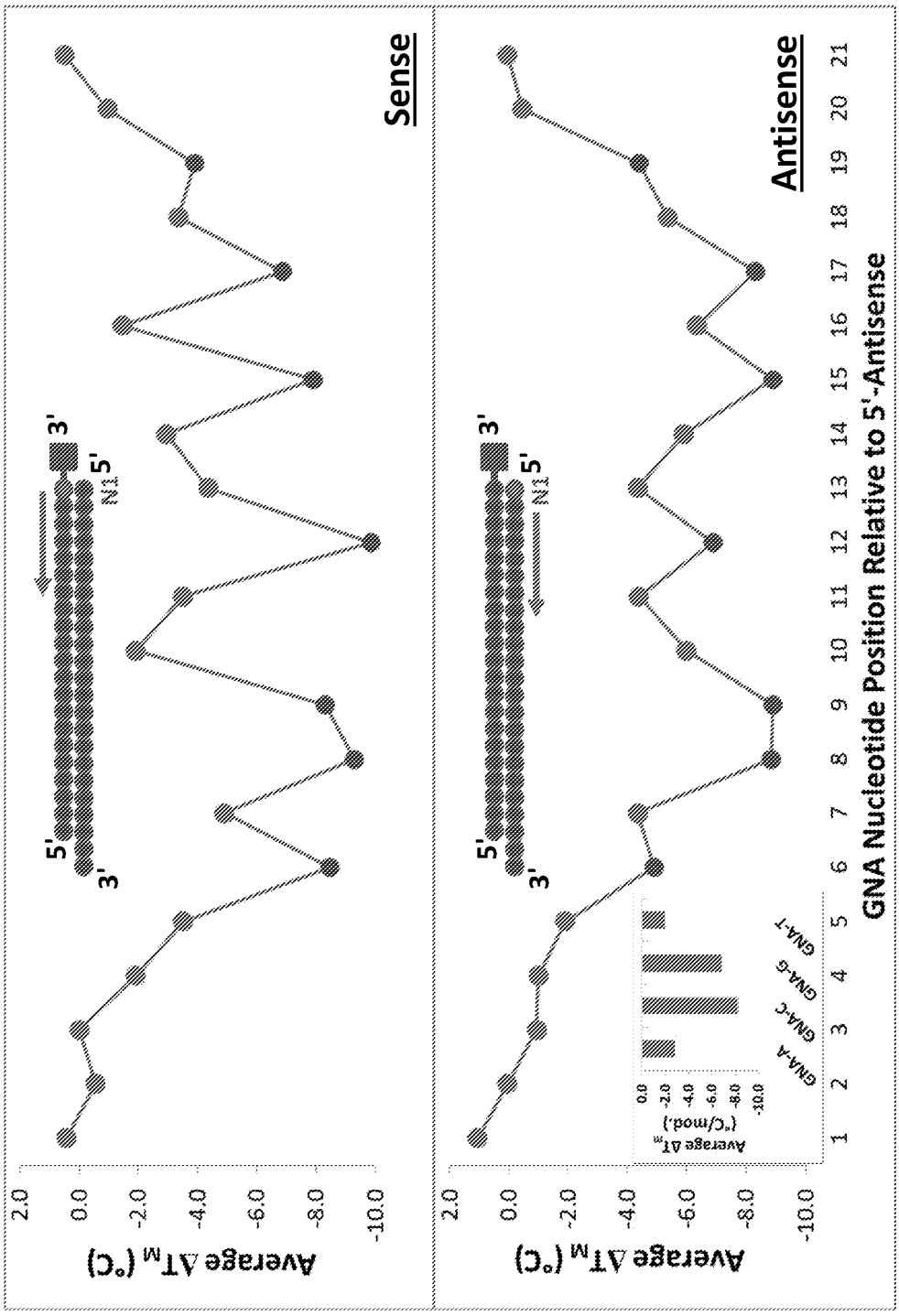
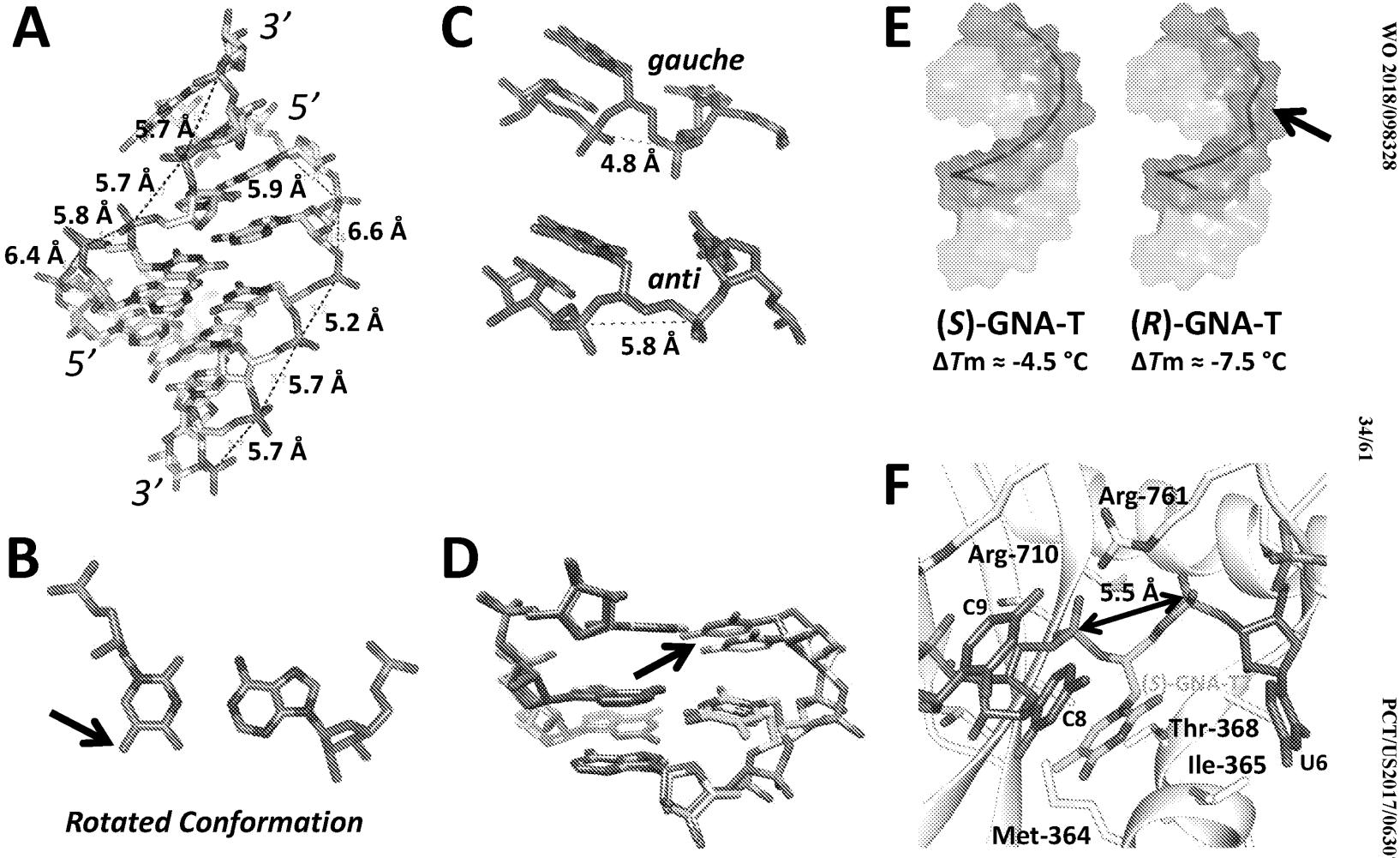

Sense
Antisense
Average ΔTM (°C)
2.0
0.0
-2.0
-4.0
-6.0
-8.0
-10.0
5'
3'
N1
GNA Nucleotide Position Relative to 5'-Antisense
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21
Average ΔTM (°C/mod.)
GNA-A
GNA-C
GNA-G
GNA-T

FIG. 33


A
3′
5′
5.7 Å
5.7 Å
5.9 Å
5.8 Å
6.6 Å
6.4 Å
5.2 Å
5′
5.7 Å
5.7 Å
3′
B
Rotated Conformation
C
gauche
4.8 Å
anti
5.8 Å
D
E
(S)-GNA-T
ΔTm ≈ -4.5 °C
(R)-GNA-T
ΔTm ≈ -7.5 °C
F
Arg-761
Arg-710
5.5 Å
C9
C8
Thr-368
Ile-365
U6
Met-364

FIGS. 34A-34F

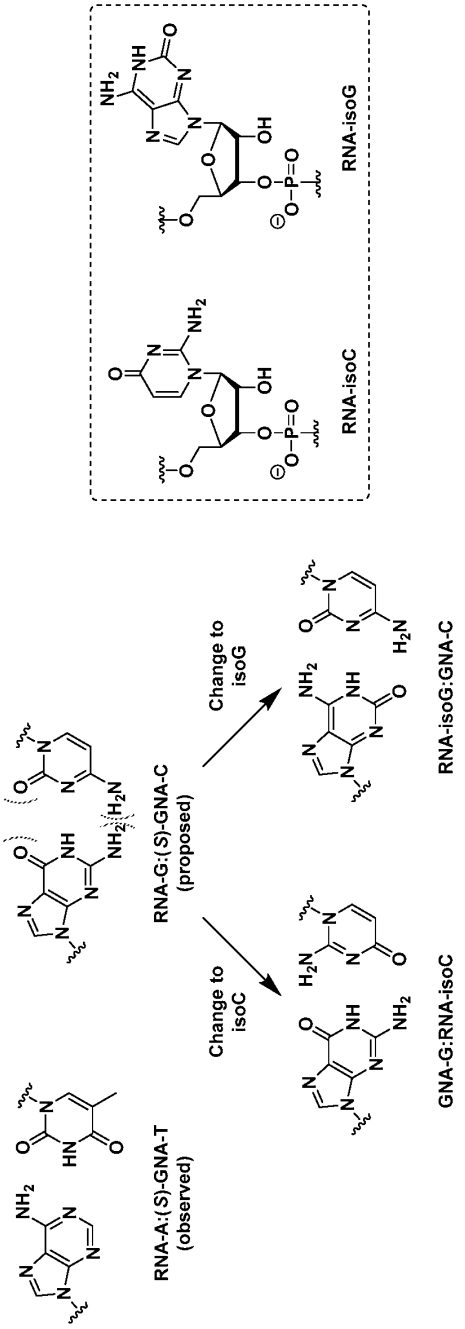

RNA-isoG
RNA-isoC
RNA-A:(S)-GNA-T
(observed)
RNA-G:(S)-GNA-C
(proposed)
Change to
isoG
Change to
isoC
RNA-isoG:GNA-C
GNA-G:RNA-isoC

FIG. 35

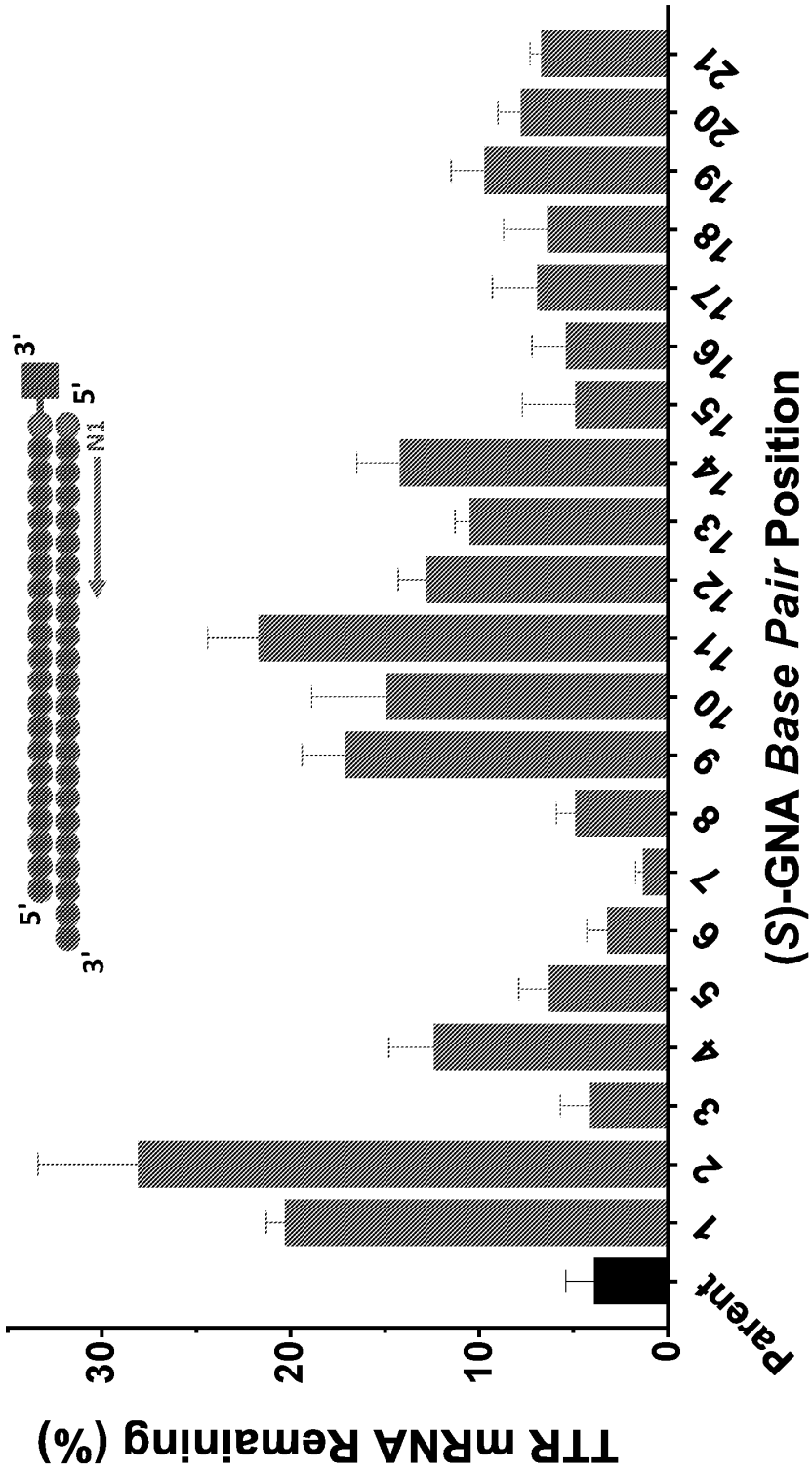
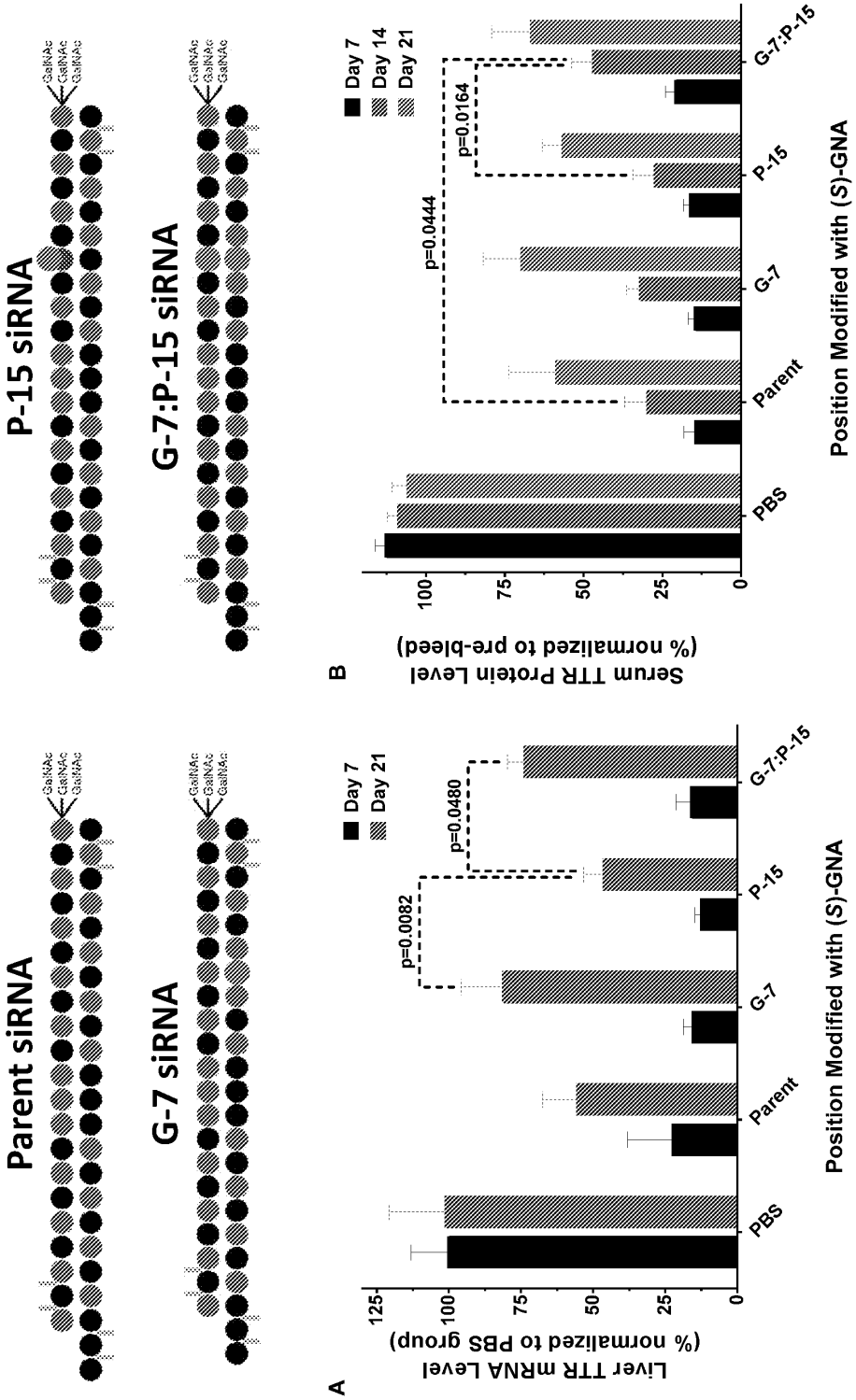
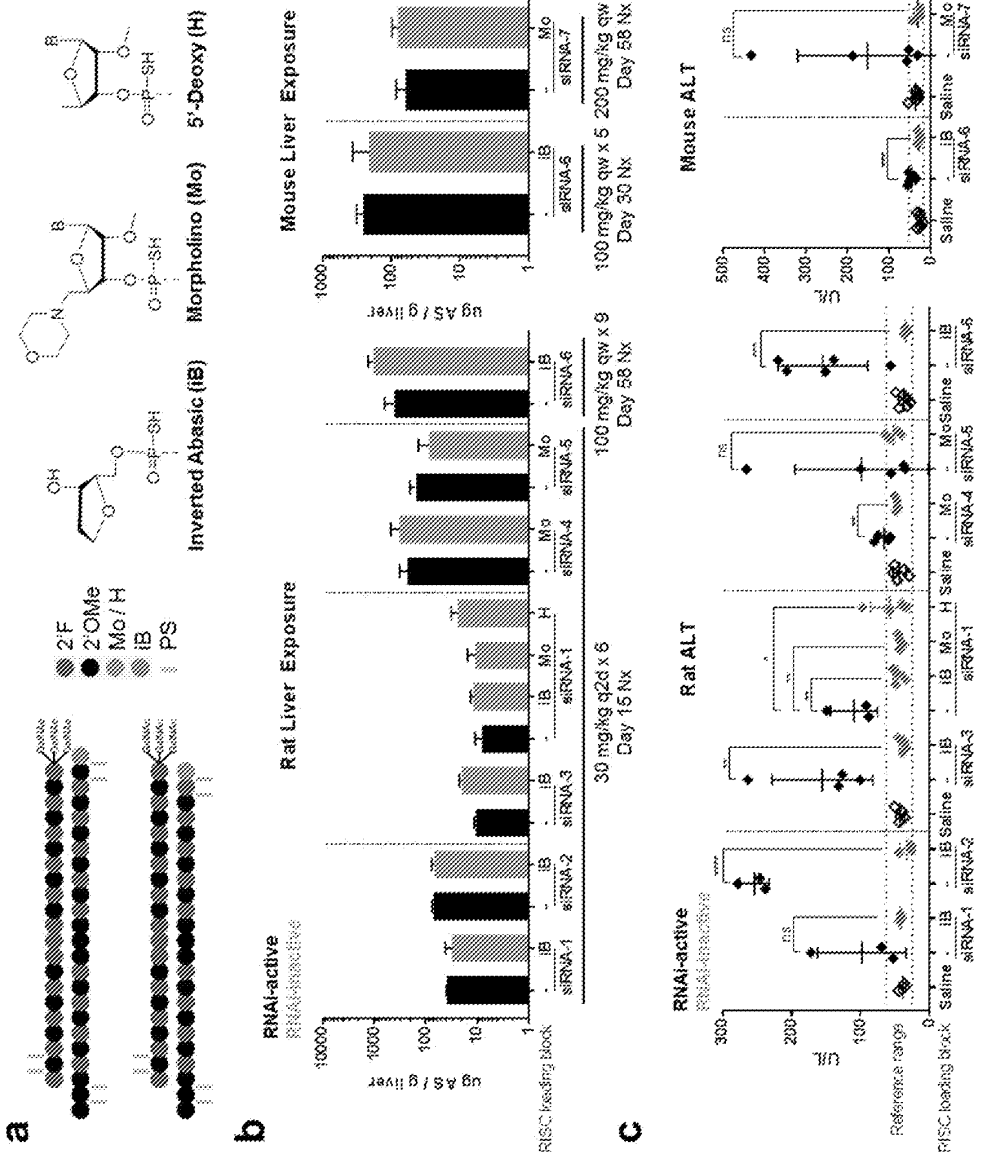

TTR mRNA Remaining (%)
0
10
20
30
Parent
1
2
3
4
5
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
(S)-GNA Base Pair Position
Relative to 5'-Guide
5'
3'
N1
3'
5'

FIG. 36


Parent siRNA
G-7 siRNA
P-15 siRNA
G-7:P-15 siRNA
GalNAc
A
Liver TTR mRNA Level
(% normalized to PBS group)
Day 7
Day 21
p=0.0082
p=0.0480
PBS
Parent
G-7
P-15
G-7:P-15
Position Modified with (S)-GNA
B
Serum TTR Protein Level
(% normalized to pre-bleed)
Day 7
Day 14
Day 21
p=0.0444
p=0.0164

FIGS. 37A-37B


a
2'F
2'OMe
Mo / H
iB
PS
Inverted Abasic (iB)
Morpholino (Mo)
5'-Deoxy (H)
b
RNAi-active
Rat Liver Exposure
Mouse Liver Exposure
ug AS / g liver
RISC loading block
Saline
siRNA-1
siRNA-2
siRNA-3
siRNA-4
siRNA-5
siRNA-6
siRNA-7
30 mg/kg q2d x 6
Day 15 Nx
100 mg/kg qw x 9
Day 58 Nx
100 mg/kg qw x 5
Day 30 Nx
200 mg/kg qw x 9
Day 58 Nx
c
RNAi-active
Rat ALT
Mouse ALT
U/L
Reference range

FIGS. 38A-38C

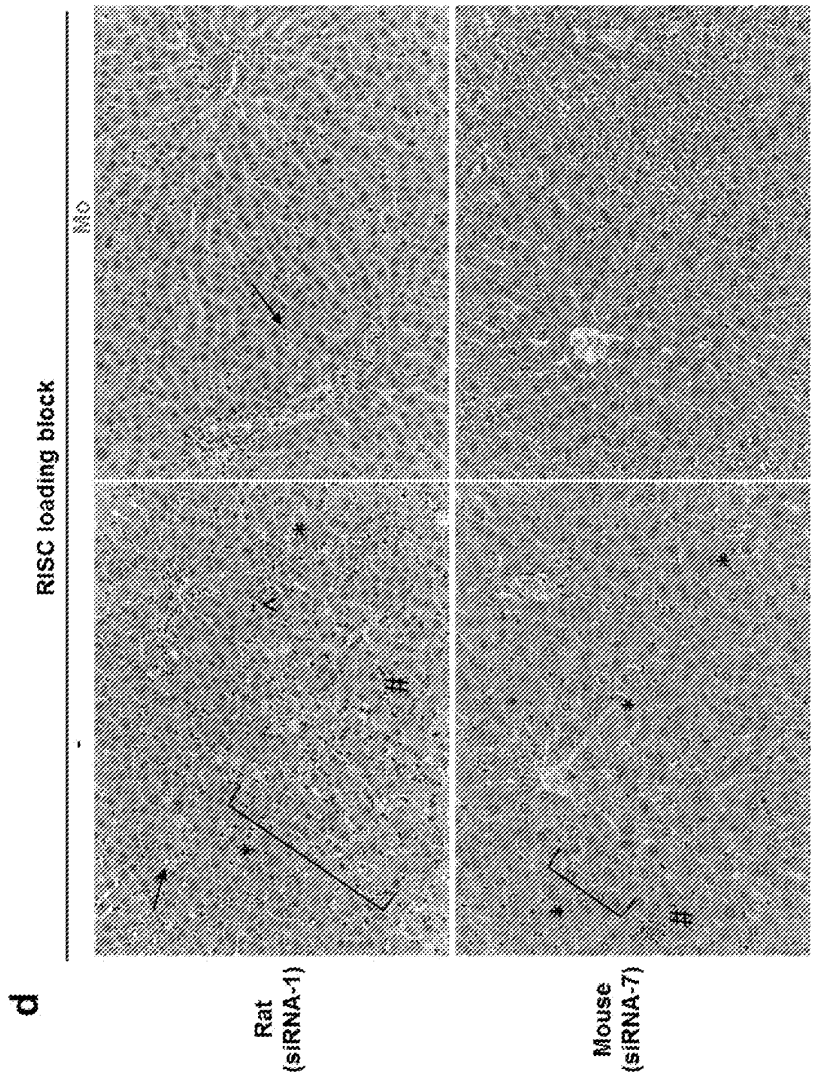
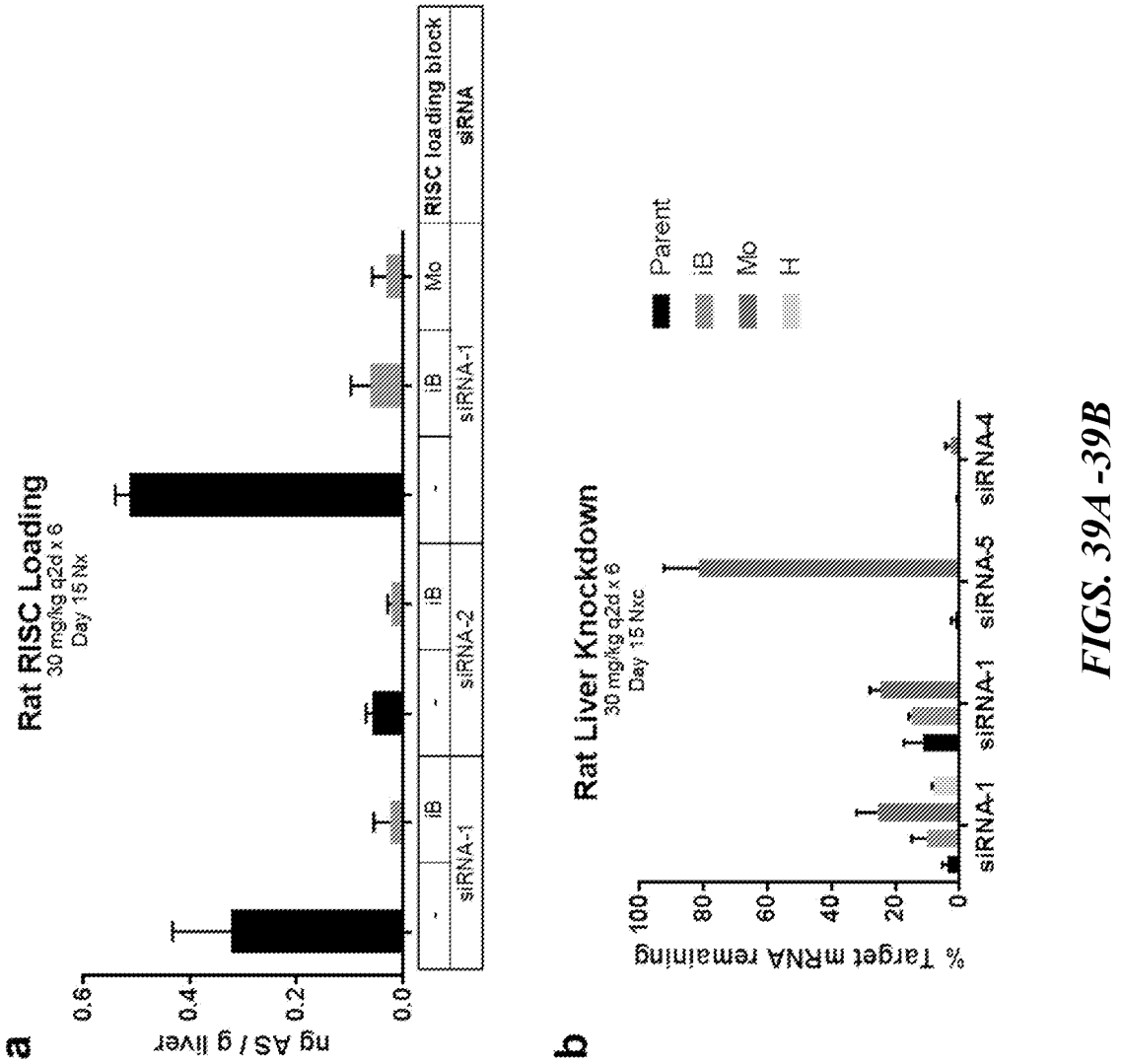


FIG. 38D


a
Rat RISC Loading
30 mg/kg q2d x 6
Day 15 Nx
ng AS / g liver
0.6
0.4
0.2
0.0
-
iB
-
iB
-
iB
Mo
RISC loading block
siRNA-1
siRNA-2
siRNA-1
siRNA
b
Rat Liver Knockdown
30 mg/kg q2d x 6
Day 15 Nxc
% Target mRNA remaining
100
80
60
40
20
0
Parent
iB
Mo
H
siRNA-1
siRNA-1
siRNA-5
siRNA-4

FIGS. 39A -39B

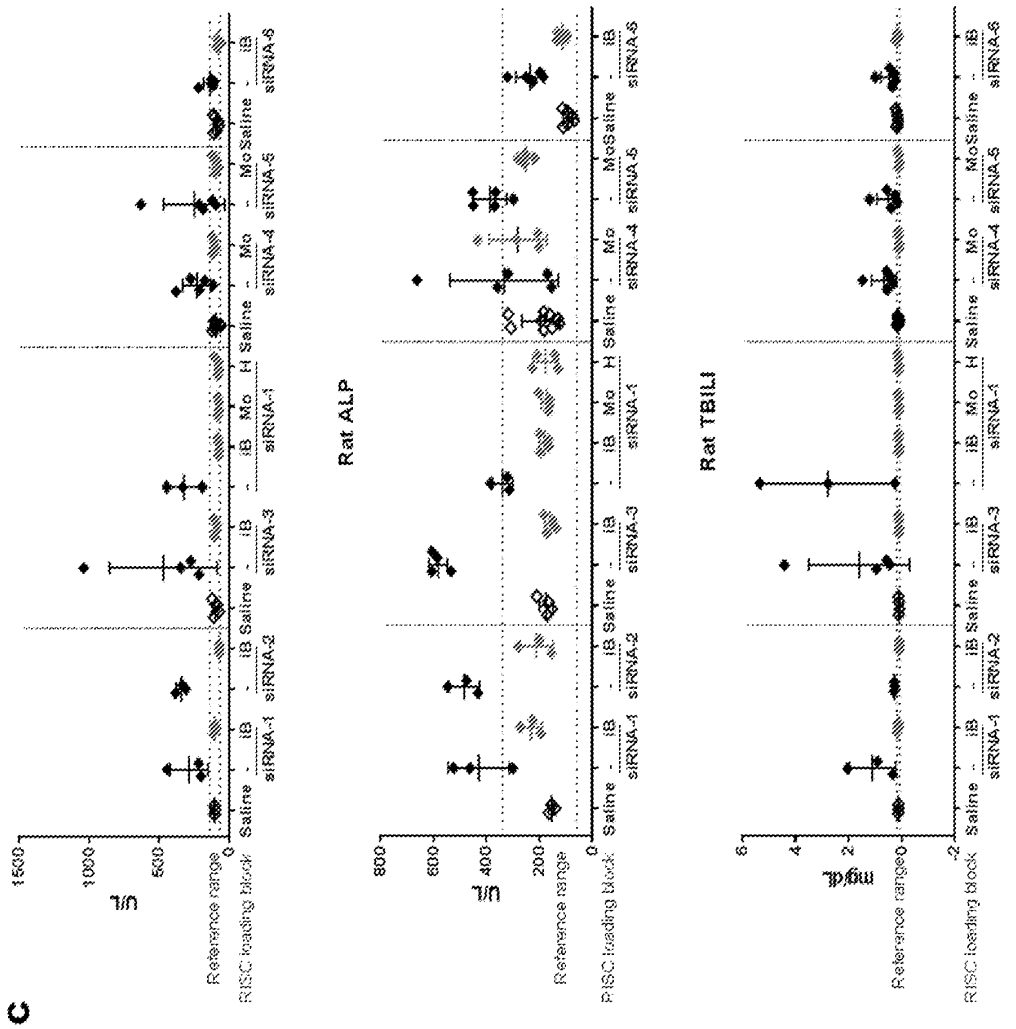
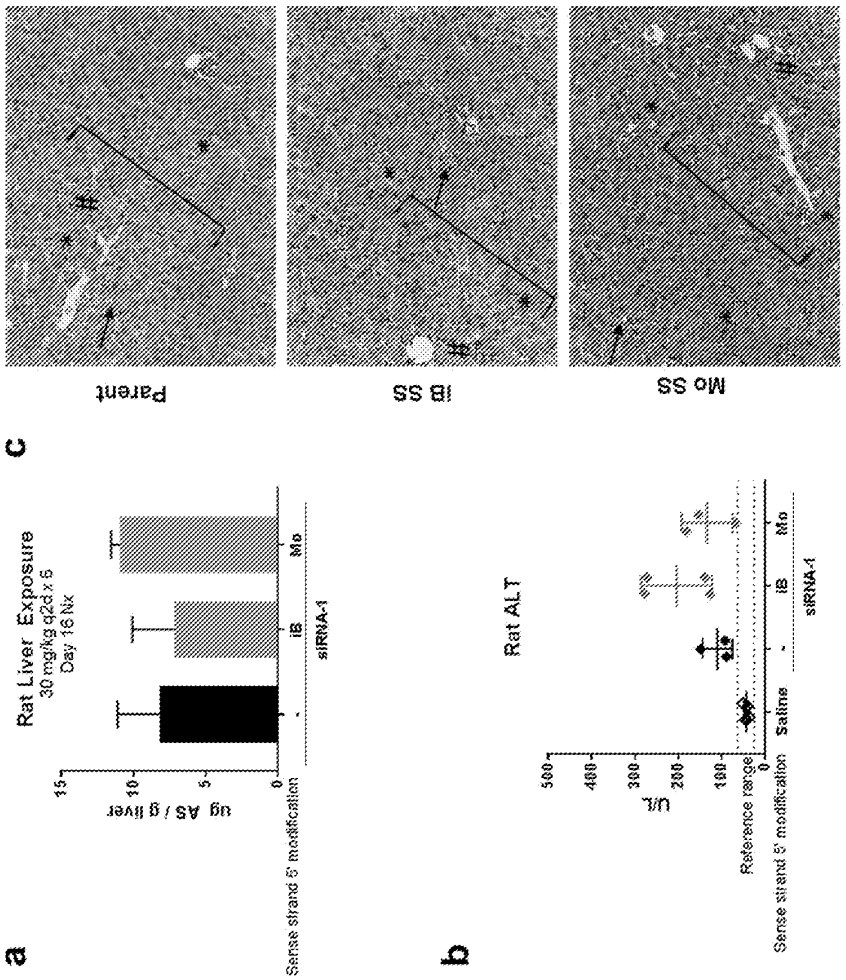
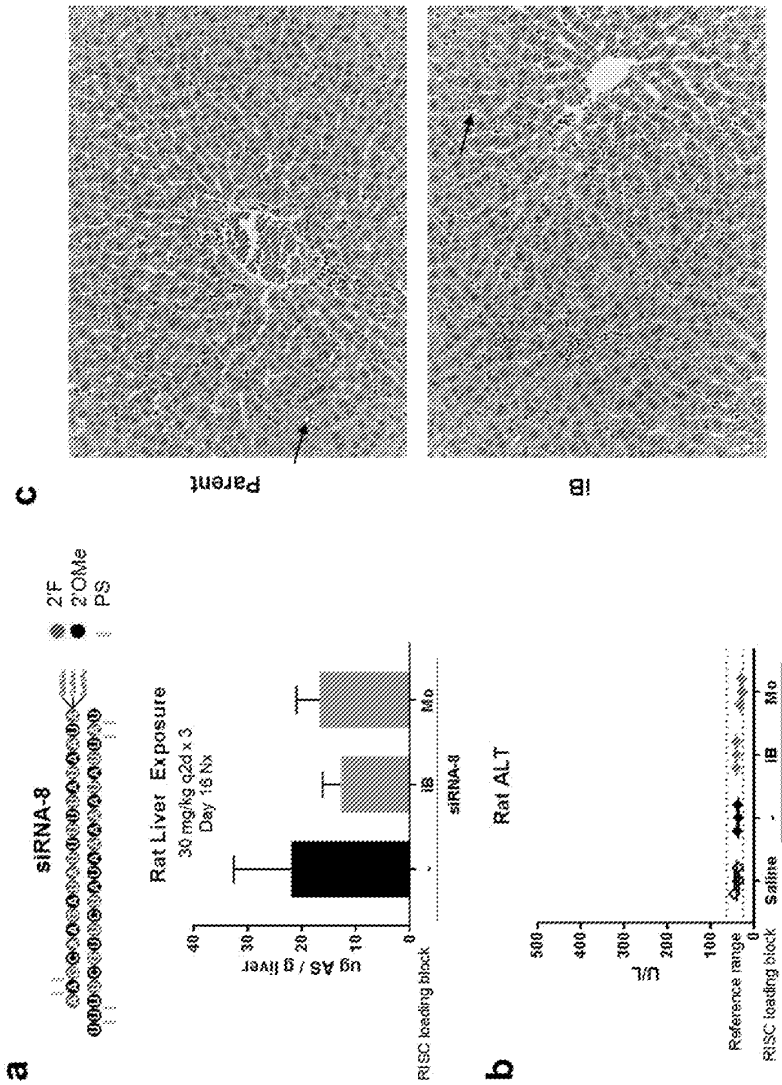

C
Rat AST
U/L
1500
1000
500
0
Reference range
RISC loading block
Saline
iB
siRNA-1
siRNA-2
siRNA-3
siRNA-4
siRNA-5
siRNA-6
Rat ALP
U/L
800
600
400
200
0
Rat TBILI
mg/dL
6
4
2
-2

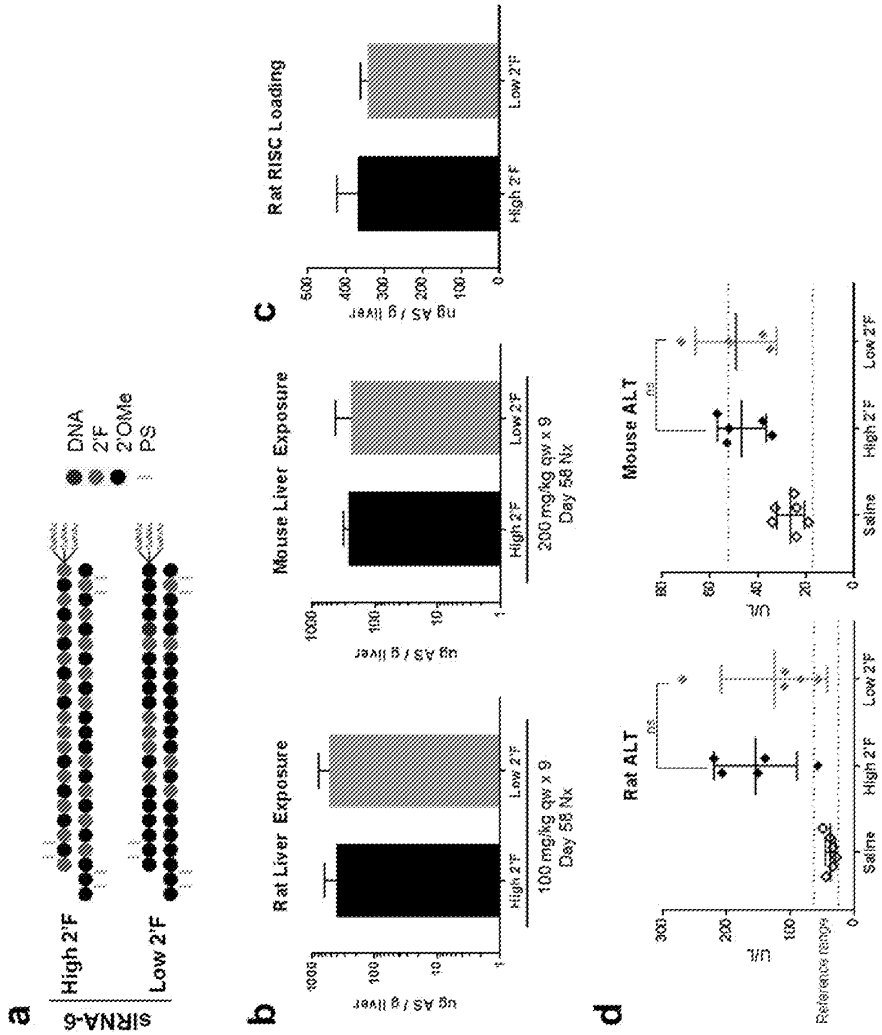
FIG. 39C


c
Parent
IB SS
Mo SS
a
Rat Liver Exposure
30 mg/kg q2d x 5
Day 16 Nx
ug AS / g liver
15
10
5
0
-
IB
Mo
Sense strand 5' modification
siRNA-1
b
Rat ALT
U/L
500
400
300
200
100
0
Reference range
Saline
-
IB
Mo
Sense strand 5' modification
siRNA-1

FIGS. 40A-40C


a
siRNA-8
2'F
2'OMe
PS
Rat Liver Exposure
30 mg/kg q2d x 3
Day 16 Nx
ug AS / g liver
40
30
20
10
0
RISC loading block
-
IB
Mo
siRNA-8
b
Rat ALT
U/L
500
400
300
200
100
0
Reference range
RISC loading block
Saline
-
IB
Mo
siRNA-8
c
Parent
IB

FIGS. 41A-41C


a
siRNA-6
High 2'F
Low 2'F
DNA
2'F
2'OMe
PS
b
Rat Liver Exposure
ug AS / g liver
100 mg/kg qw x 9
Day 58 Nx
High 2'F
Low 2'F
Mouse Liver Exposure
ug AS / g liver
200 mg/kg qw x 9
Day 58 Nx
c
Rat RISC Loading
ng AS / g liver
d
Rat ALT
U/L
Reference range
Saline
High 2'F
Low 2'F
Mouse ALT
U/L

FIGS. 42A-42D

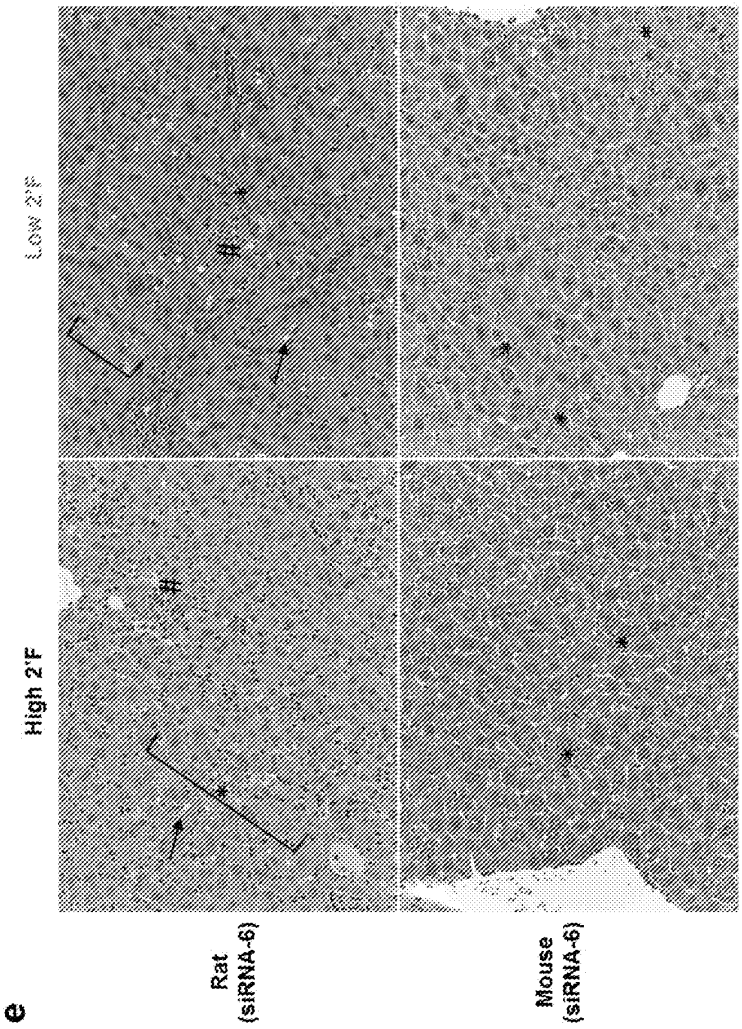


FIG. 42E

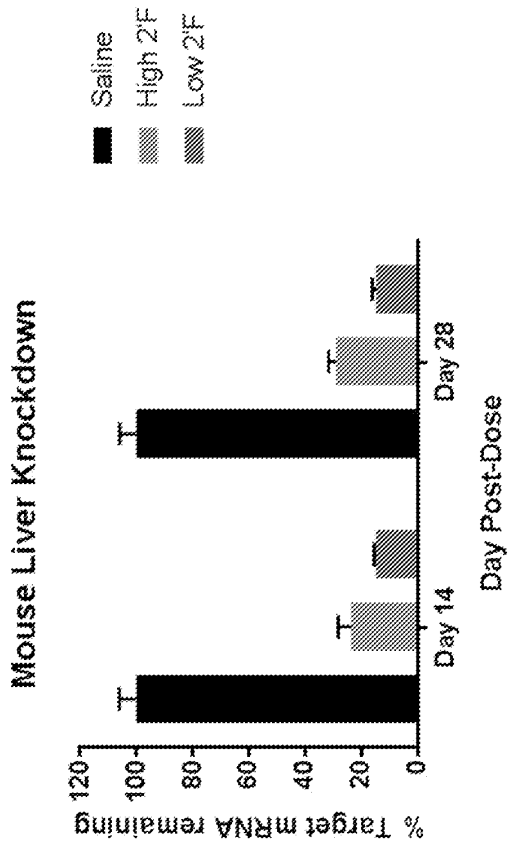
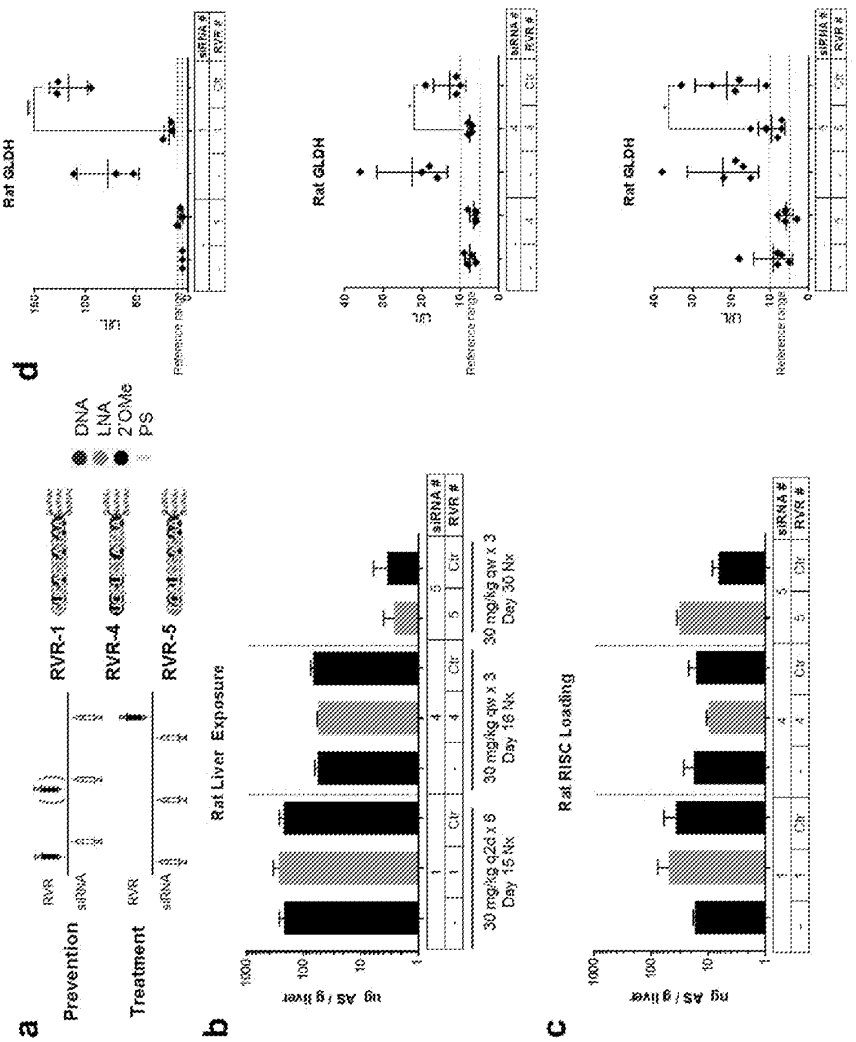

Mouse Liver Knockdown
Saline
High 2'F
Low 2'F
% Target mRNA remaining
120
100
80
60
40
20
0
Day 14
Day 28
Day Post-Dose

FIG. 43


a
Prevention
Treatment
RVR
siRNA
RVR-1
RVR-4
RVR-5
DNA
LNA
2'OMe
PS
b
Rat Liver Exposure
ng AS / g liver
30 mg/kg q2d x 6
Day 15 Nx
30 mg/kg qw x 3
Day 16 Nx
30 mg/kg qw x 3
Day 30 Nx
siRNA #
RVR #
Ctr
c
Rat RISC Loading
ng AS / g liver
d
Rat GLDH
Reference range

FIGS. 44A-44D

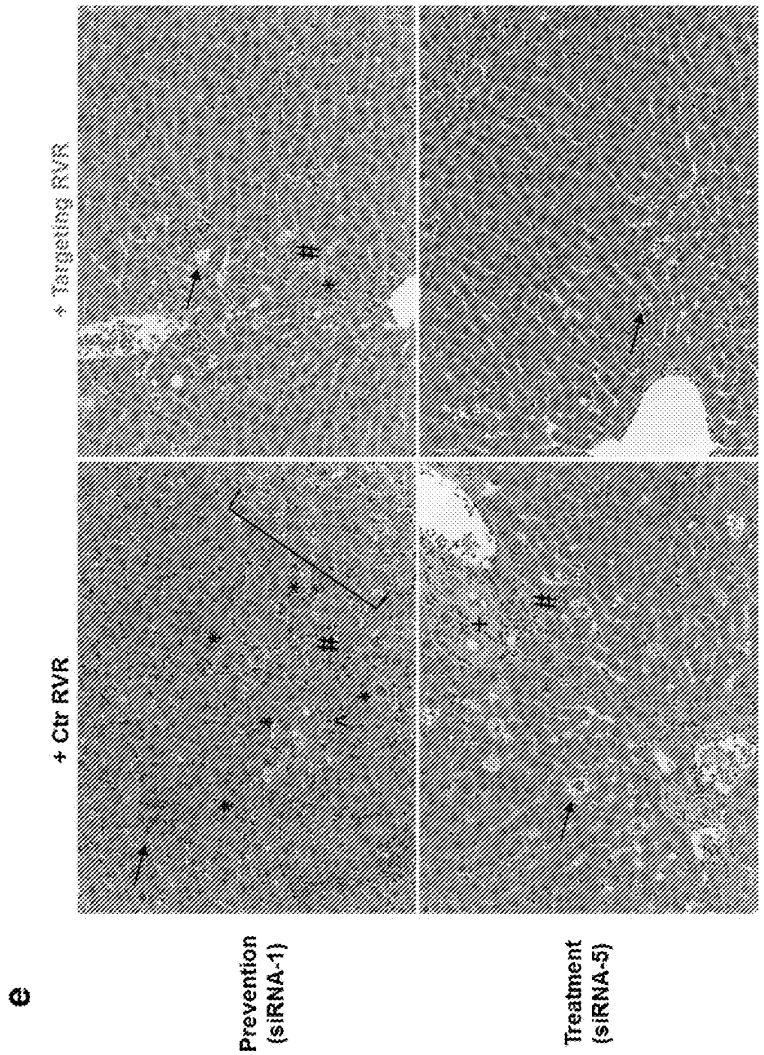

e
Prevention (siRNA-1)
Treatment (siRNA-5)
+ Ctr RVR
+ Targeting RVR

FIG. 44E

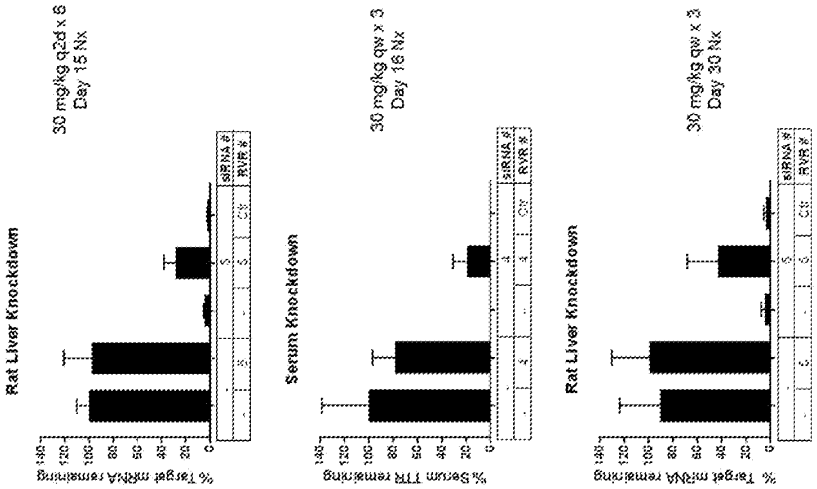
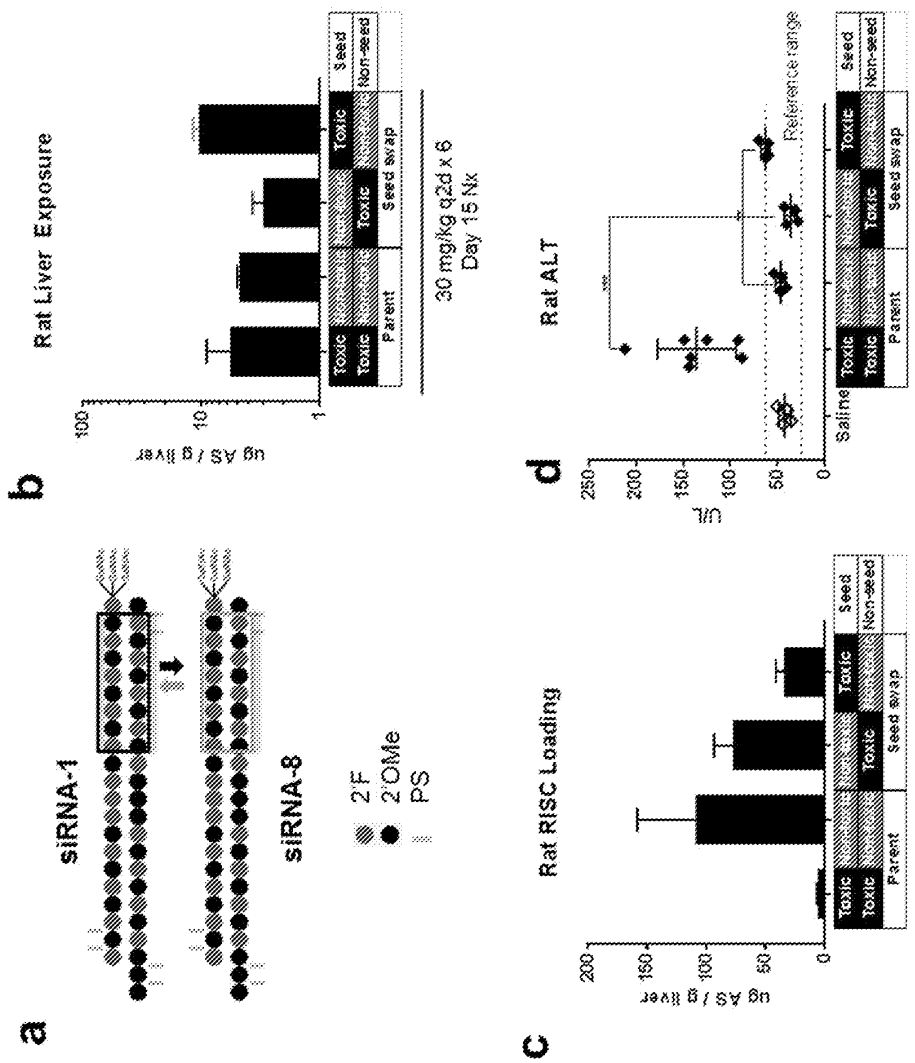

Rat Liver Knockdown
30 mg/kg q2d x 6
Day 15 Nx
% Target mRNA remaining
siRNA #
RVR #
Serum Knockdown
30 mg/kg qw x 3
Day 16 Nx
% Serum TTR remaining
siRNA #
RVR #
Rat Liver Knockdown
30 mg/kg qw x 3
Day 30 Nx
% Target mRNA remaining
siRNA #
RVR #

FIG. 45


a
siRNA-1
siRNA-8
2'F
2'OMe
PS
b
Rat Liver Exposure
ug AS / g liver
100
10
1
Toxic
Toxic
Toxic
Toxic
Seed
Non-seed
Parent
Seed swap
30 mg/kg q2d x 6
Day 15 Nk
c
Rat RISC Loading
ug AS / g liver
200
150
100
50
0
d
Rat ALT
U/L
250
200
150
100
50
0
Reference range
Saline

FIGS. 46A-46D

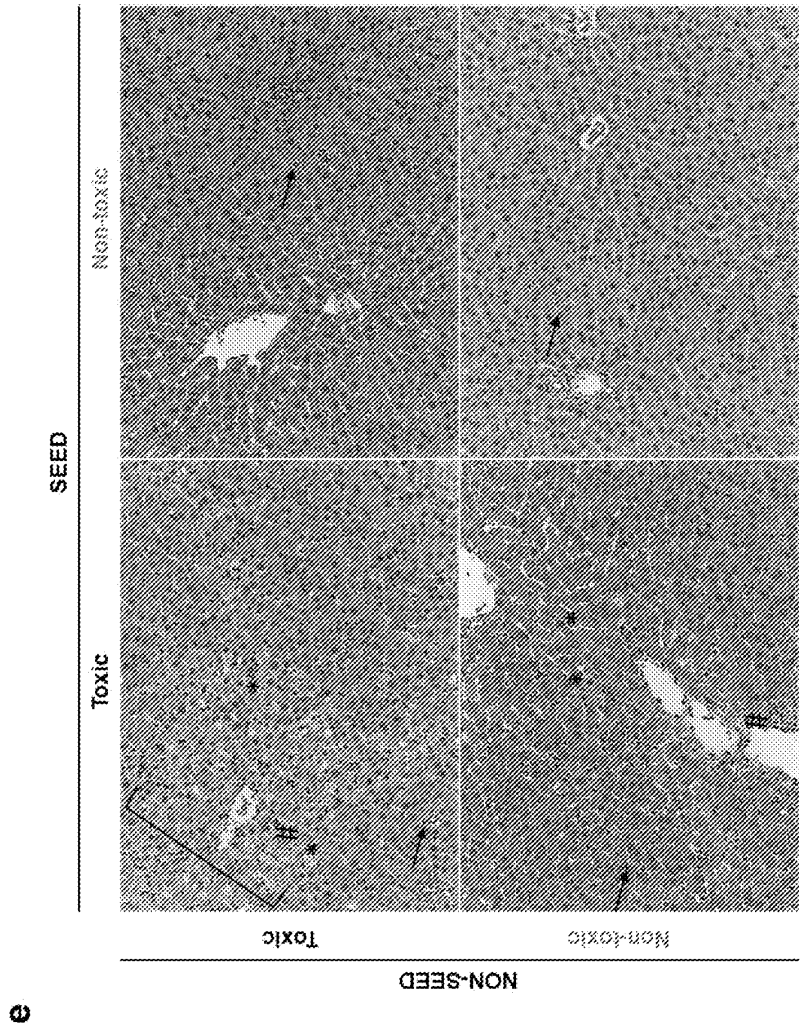


FIG. 46E

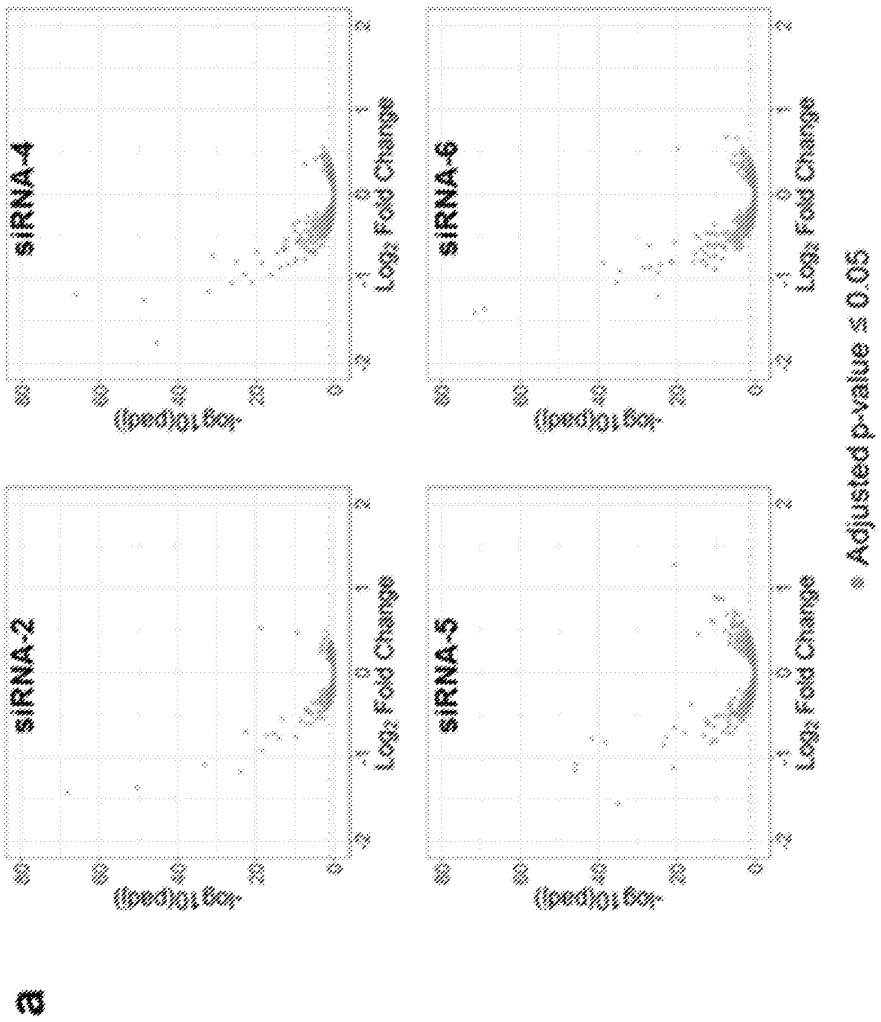

a
siRNA-4
siRNA-6
siRNA-2
siRNA-5
Log2 Fold Change
-log10(padj)
* Adjusted p-value ≤ 0.05

FIG. 47A

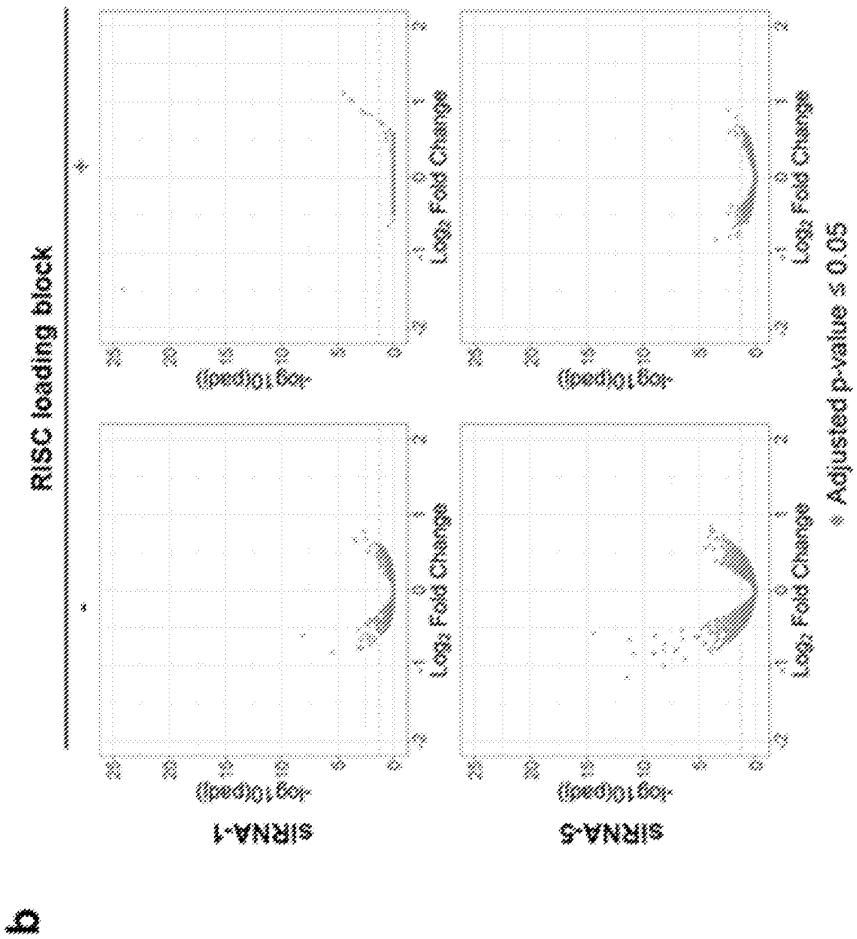
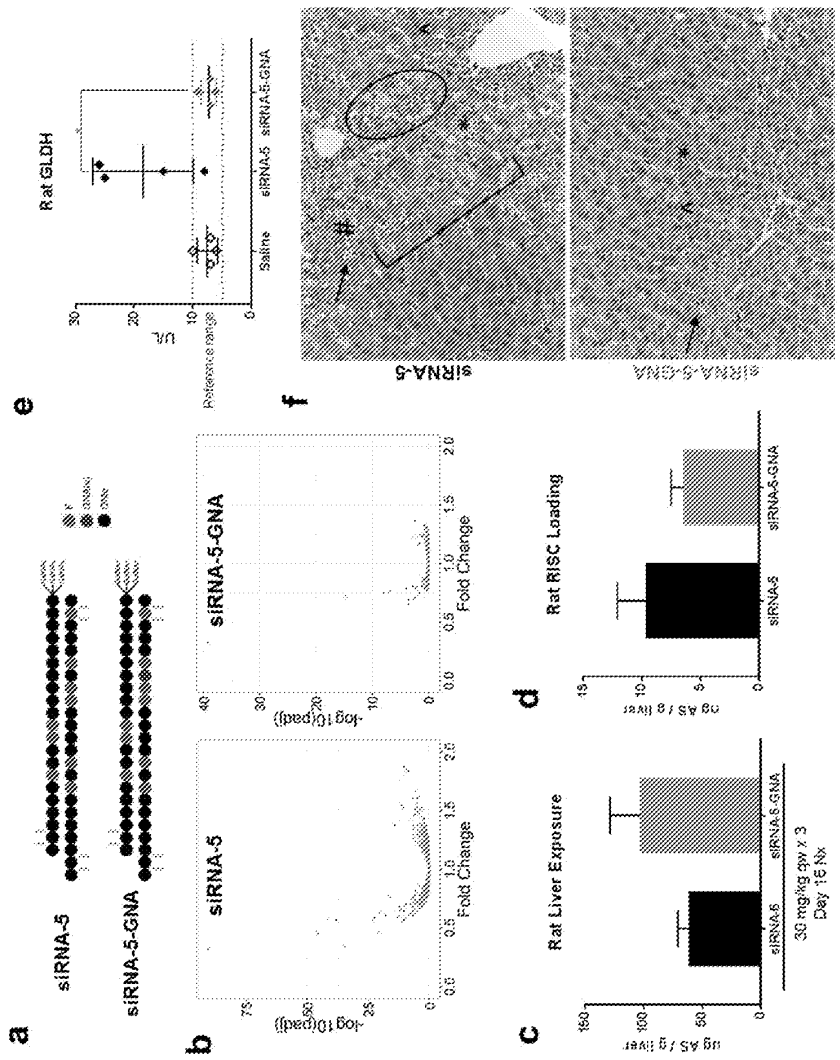
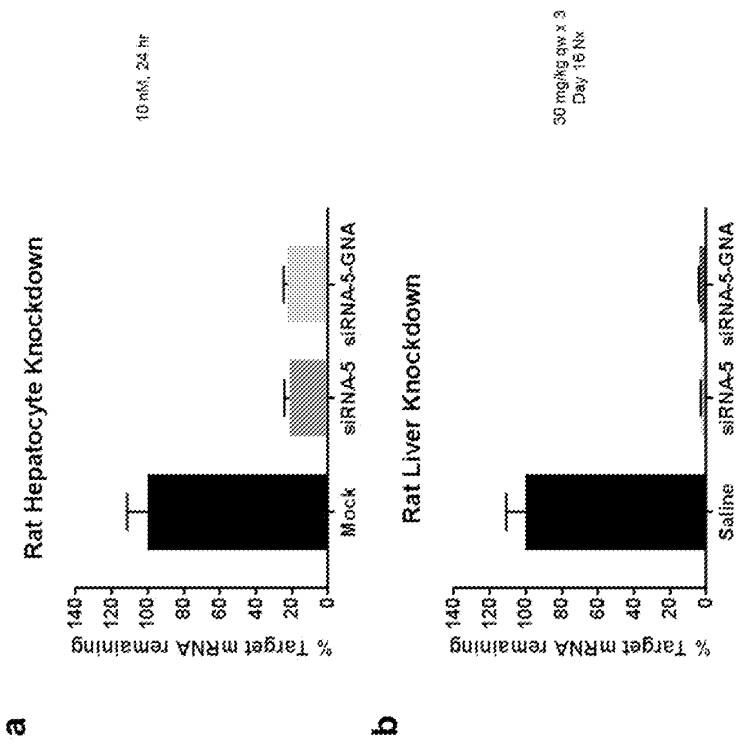

b
siRNA-1
siRNA-5
RISC loading block
Log2 Fold Change
-log10(padj)
* Adjusted p-value ≤ 0.05

FIG. 47B


a
siRNA-5
siRNA-5-GNA
b
siRNA-5
siRNA-5-GNA
-log 10(pad))
Fold Change
c
Rat Liver Exposure
ug AS / g liver
siRNA-5
siRNA-5-GNA
30 mg/kg qw x 3
Day 15 Nx
d
Rat RISC Loading
ng AS / g liver
siRNA-5
siRNA-5-GNA
e
Rat GLDH
ULN
Potential range
Saline
siRNA-5
siRNA-5-GNA
f
siRNA-5
siRNA-5-GNA

FIGS. 48A-48F


a
Rat Hepatocyte Knockdown
10 nM, 24 hr
% Target mRNA remaining
140
120
100
80
60
40
20
0
Mock
siRNA-5
siRNA-5-GNA
b
Rat Liver Knockdown
30 mg/kg qw x 3
Day 16 Nx
% Target mRNA remaining
140
120
100
80
60
40
20
0
Saline
siRNA-5
siRNA-5-GNA

FIGS. 49A-49B

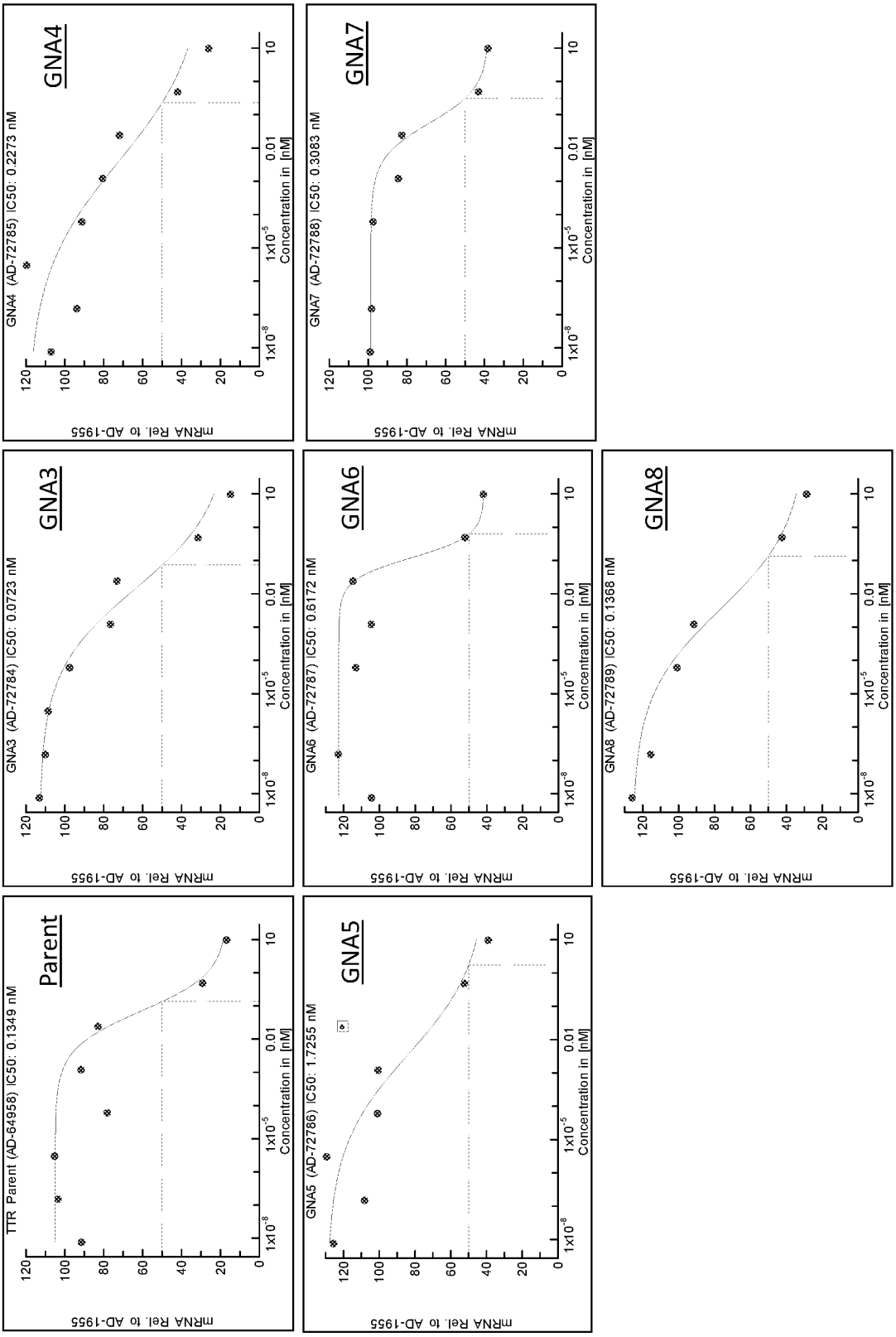

Parent
TTR Parent (AD-64958) IC50: 0.1349 nM
GNA3
GNA3 (AD-72784) IC50: 0.0723 nM
GNA4
GNA4 (AD-72785) IC50: 0.2273 nM
GNA5
GNA5 (AD-72786) IC50: 1.7255 nM
GNA6
GNA6 (AD-72787) IC50: 0.6172 nM
GNA7
GNA7 (AD-72788) IC50: 0.3083 nM
GNA8
GNA8 (AD-72789) IC50: 0.1368 nM
mRNA Rel. to AD-1955
Concentration in [nM]

FIG. 50

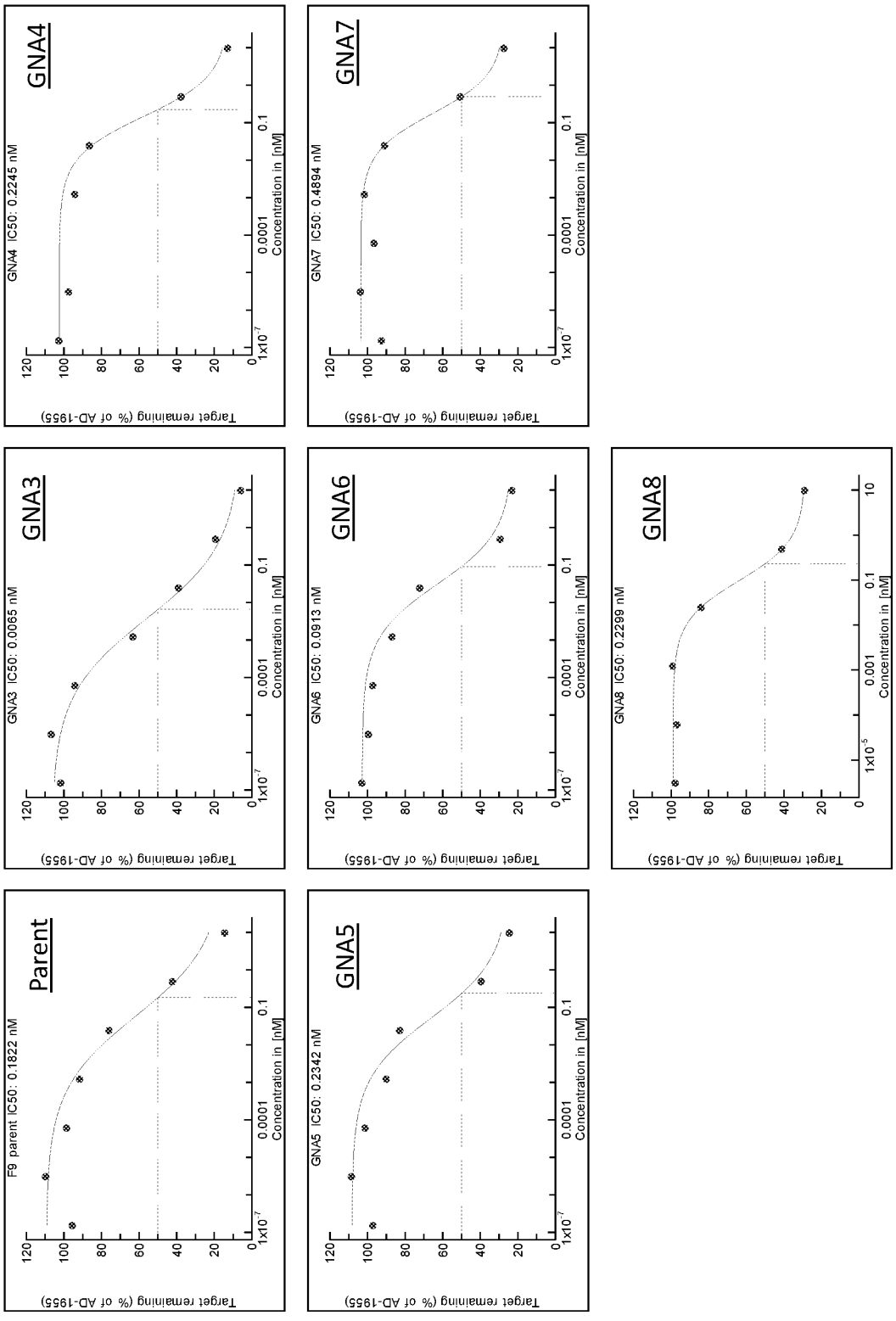

Parent
F9 parent IC50: 0.1822 nM
GNA3
GNA3 IC50: 0.0065 nM
GNA4
GNA4 IC50: 0.2245 nM
GNA5
GNA5 IC50: 0.2342 nM
GNA6
GNA6 IC50: 0.0913 nM
GNA7
GNA7 IC50: 0.4894 nM
GNA8
GNA8 IC50: 0.2299 nM
Target remaining (% of AD-1955)
Concentration in [nM]

FIG. 51

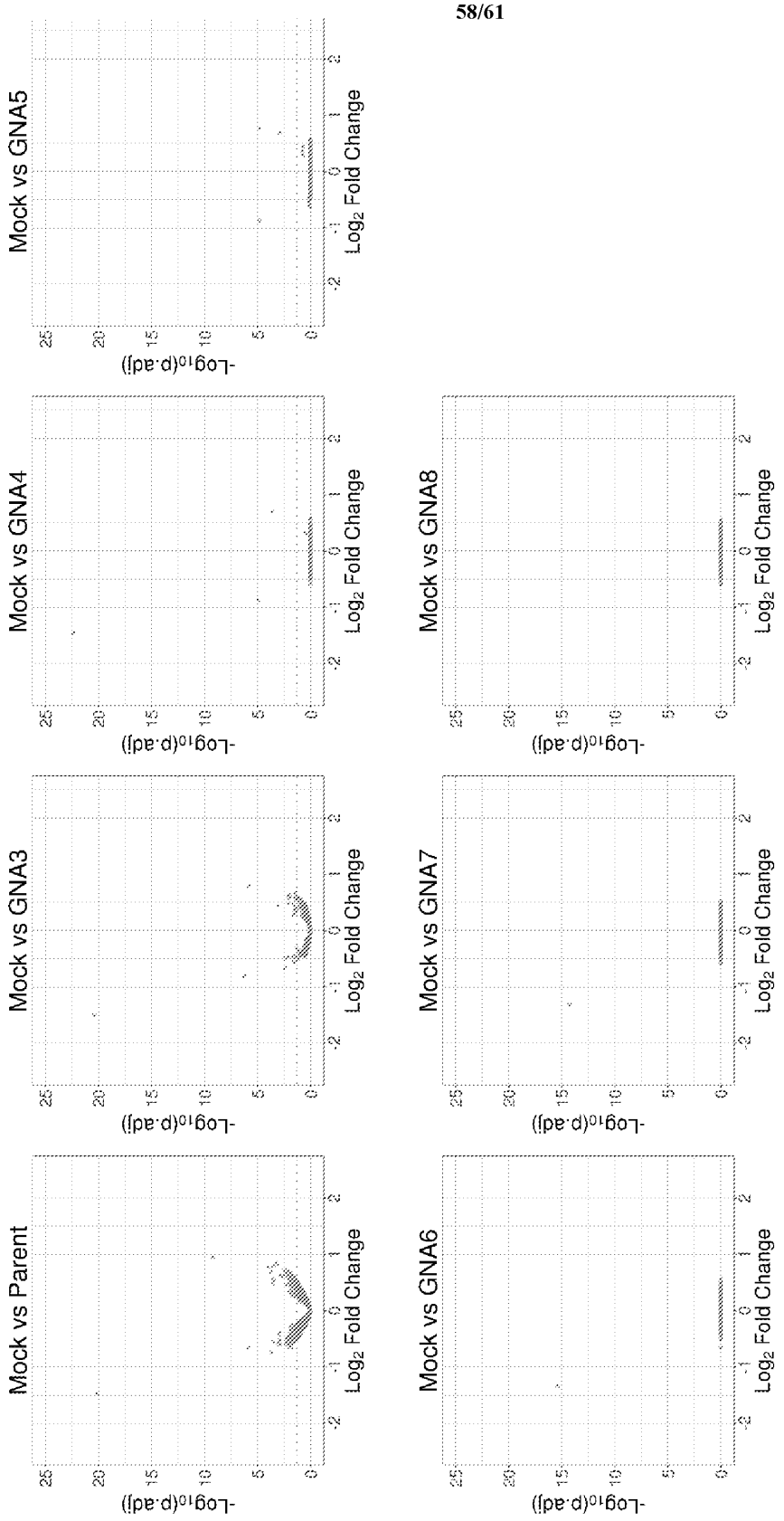

Mock vs Parent
Mock vs GNA3
Mock vs GNA4
Mock vs GNA5
Mock vs GNA6
Mock vs GNA7
Mock vs GNA8
-Log10(p.adj)
Log2 Fold Change

FIG. 52

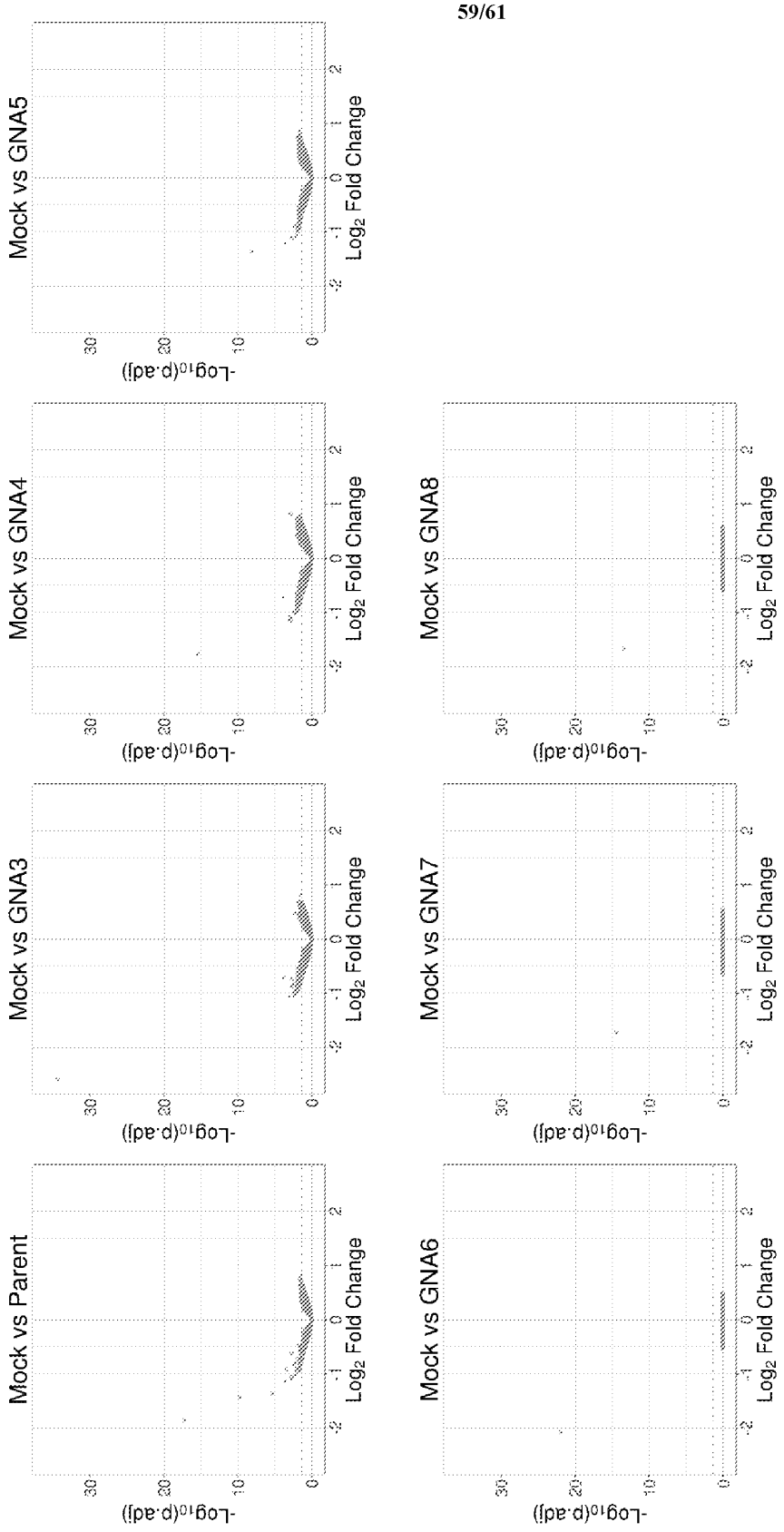

Mock vs Parent
Mock vs GNA3
Mock vs GNA4
Mock vs GNA5
Mock vs GNA6
Mock vs GNA7
Mock vs GNA8
-Log10(p.adj)
Log2 Fold Change

FIG. 53

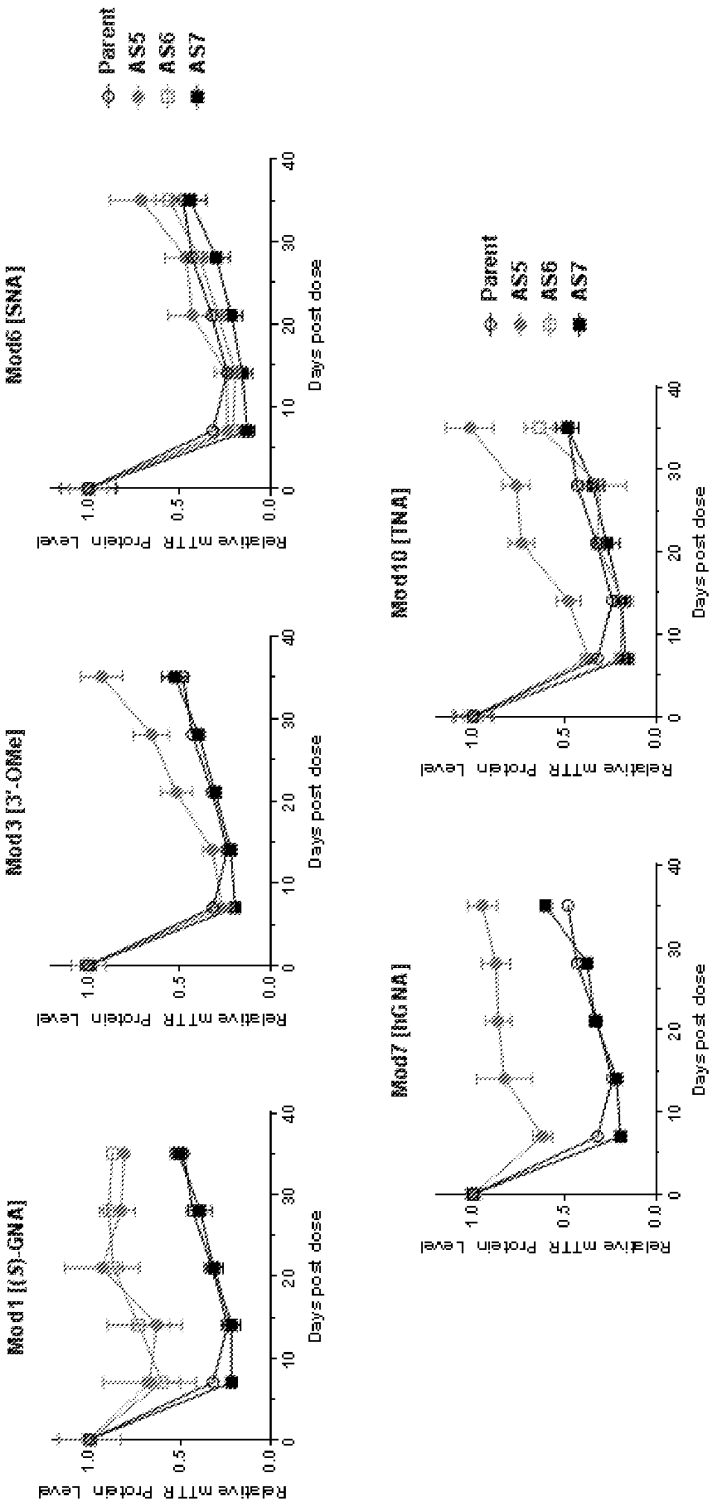

Parent
AS5
AS6
AS7
Mod1 [(S)-GNA]
Mod3 [3'-OMe]
Mod6 [SNA]
Mod7 [hGNA]
Mod10 [TNA]
Relative mTTR Protein Level
Days post dose
0.0
0.5
1.0
0
10
20
30
40

FIG. 54

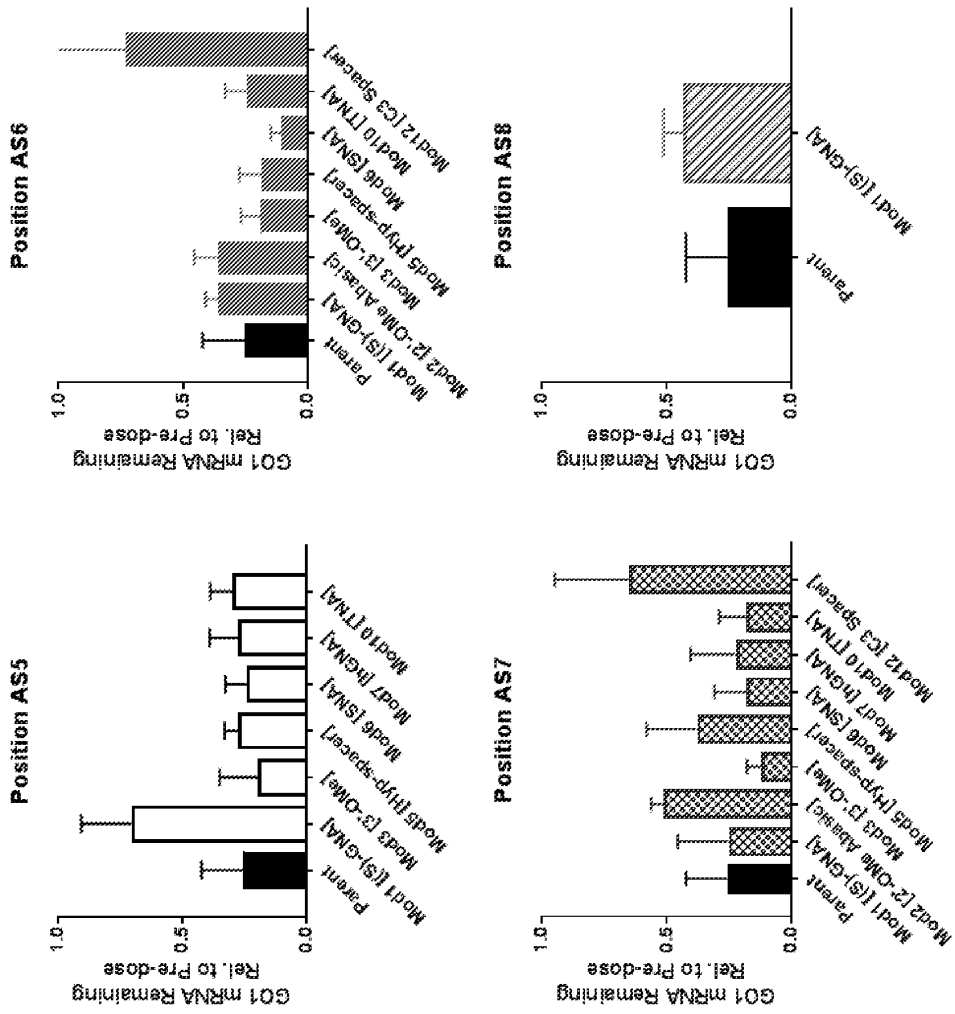

Position AS6
G01 mRNA Remaining
Rel. to Pre-dose
1.0
0.5
0.0
Parent
Mod1 [(S)-GNA]
Mod2 [2'-OMe Abasic]
Mod3 [3'-OMe]
Mod5 [Hyp-spacer]
Mod8 [SNA]
Mod10 [TNA]
Mod12 [C3 Spacer]
Position AS8
G01 mRNA Remaining
Rel. to Pre-dose
1.0
0.5
0.0
Parent
Mod1 [(S)-GNA]
Position AS5
G01 mRNA Remaining
Rel. to Pre-dose
1.0
0.5
0.0
Parent
Mod1 [(S)-GNA]
Mod3 [3'-OMe]
Mod5 [Hyp-spacer]
Mod8 [SNA]
Mod7 [hGNA]
Mod10 [TNA]
Position AS7
G01 mRNA Remaining
Rel. to Pre-dose
1.0
0.5
0.0
Parent
Mod1 [(S)-GNA]
Mod2 [2'-OMe Abasic]
Mod3 [3'-OMe]
Mod5 [Hyp-spacer]
Mod8 [SNA]
Mod7 [hGNA]
Mod10 [TNA]
Mod12 [C3 Spacer]

FIG. 55