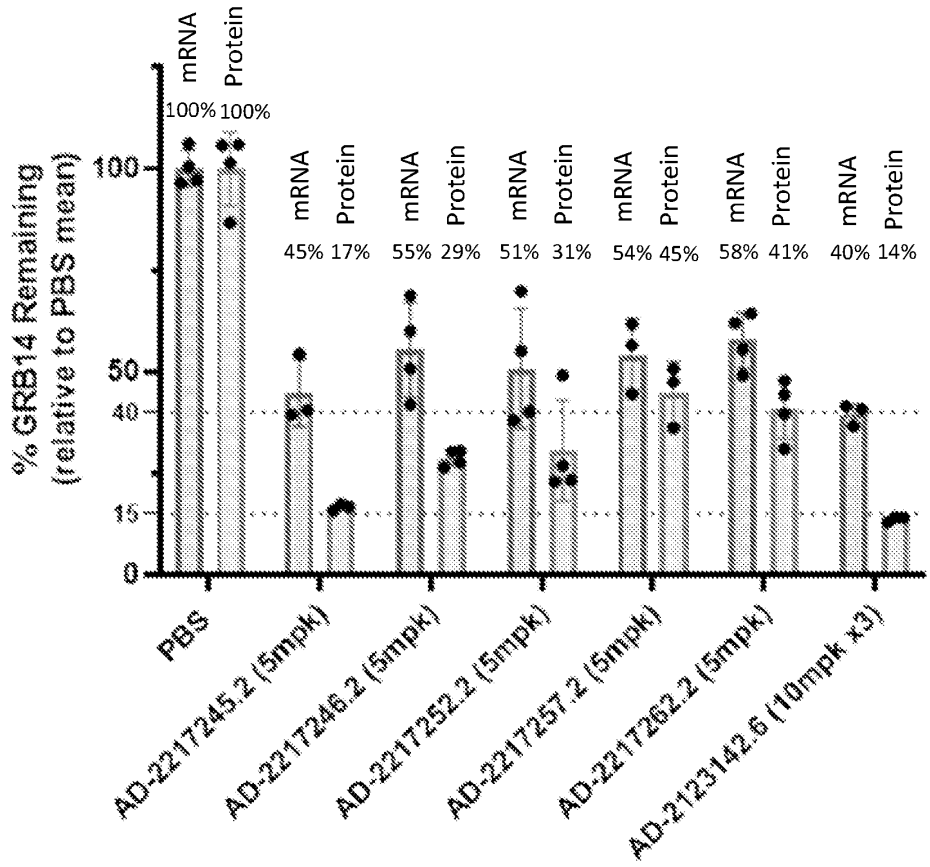

% GRB14 Remaining
(relative to PBS mean)
100
50
40
15
0
mRNA
Protein
100% 100%
45% 17%
55% 29%
51% 31%
54% 45%
58% 41%
40% 14%
PBS
AD-2217245.2 (5mpk)
AD-2217246.2 (5mpk)
AD-2217252.2 (5mpk)
AD-2217257.2 (5mpk)
AD-2217262.2 (5mpk)
AD-2123142.6 (10mpk x3)

FIG. 1

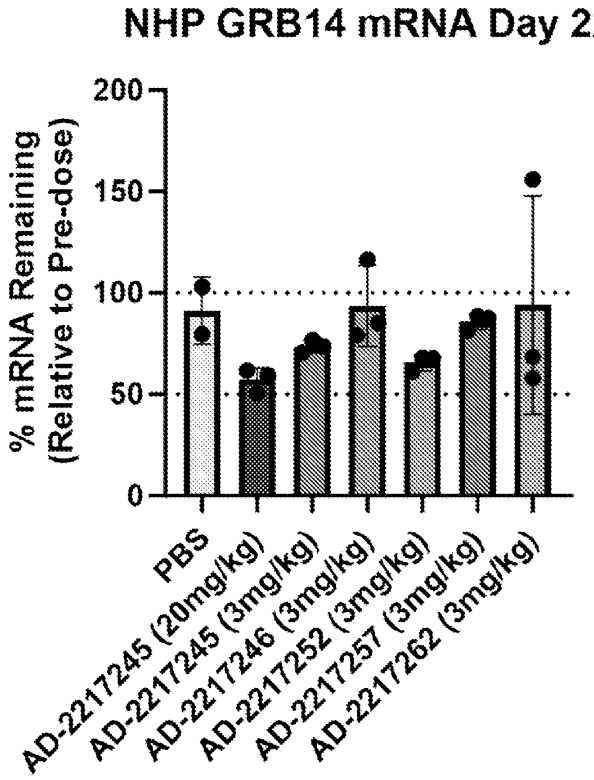

NHP GRB14 mRNA Day 22
% mRNA Remaining
(Relative to Pre-dose)
200
150
100
50
0
PBS
AD-2217245 (20mg/kg)
AD-2217245 (3mg/kg)
AD-2217246 (3mg/kg)
AD-2217252 (3mg/kg)
AD-2217257 (3mg/kg)
AD-2217262 (3mg/kg)

FIG. 2

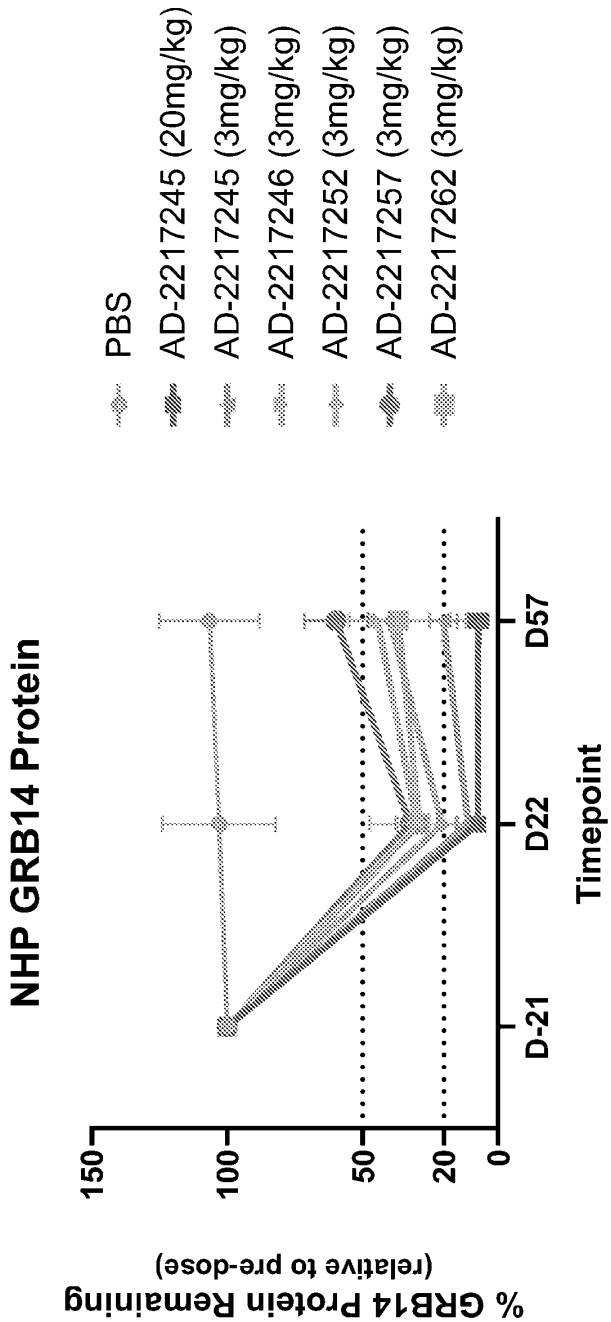

NHP GRB14 Protein
PBS
AD-2217245 (20mg/kg)
AD-2217245 (3mg/kg)
AD-2217246 (3mg/kg)
AD-2217252 (3mg/kg)
AD-2217257 (3mg/kg)
AD-2217262 (3mg/kg)
% GRB14 Protein Remaining
(relative to pre-dose)
150
100
50
20
0
D-21
D22
D57
Timepoint

FIG. 3

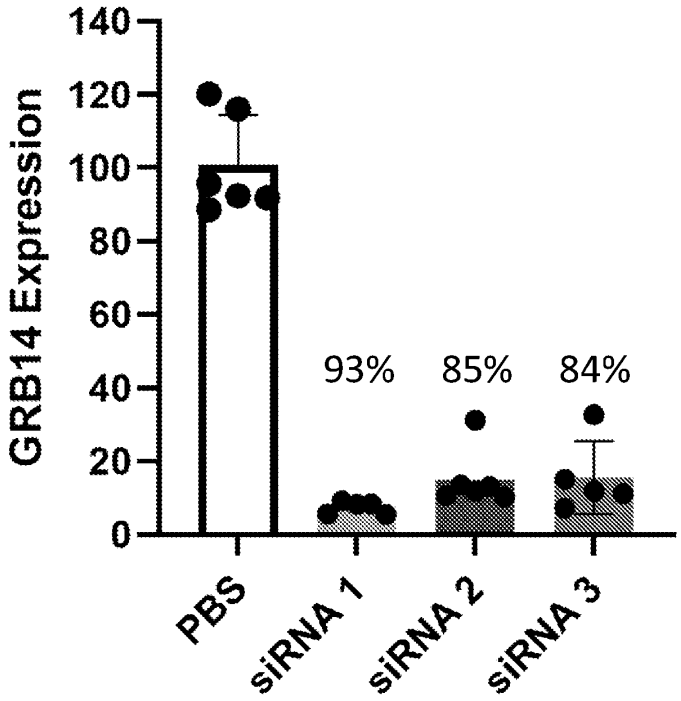

GRB14 Expression
140
120
100
80
60
40
20
0
93%
85%
84%
PBS
siRNA 1
siRNA 2
siRNA 3

FIG. 4A

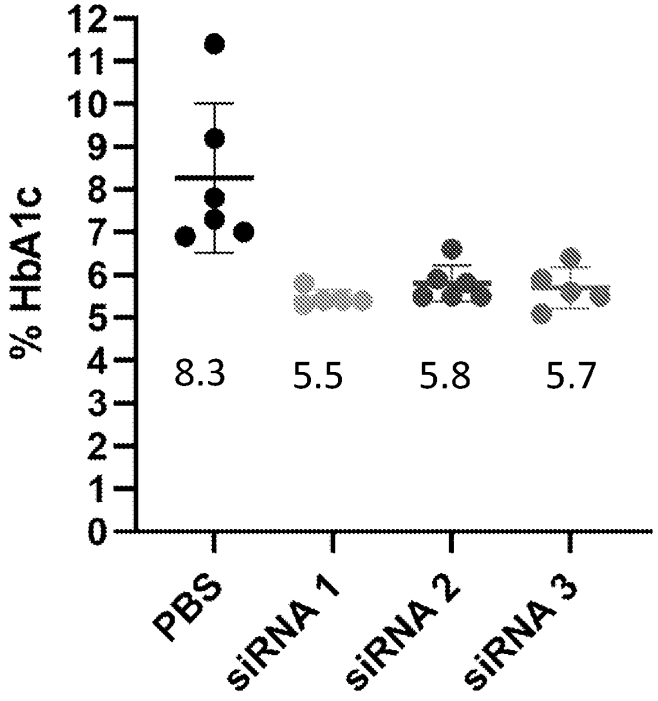

% HbA1c
12
11
10
9
8
7
6
5
4
3
2
1
0
8.3
5.5
5.8
5.7
PBS
siRNA 1
siRNA 2
siRNA 3

FIG. 4B

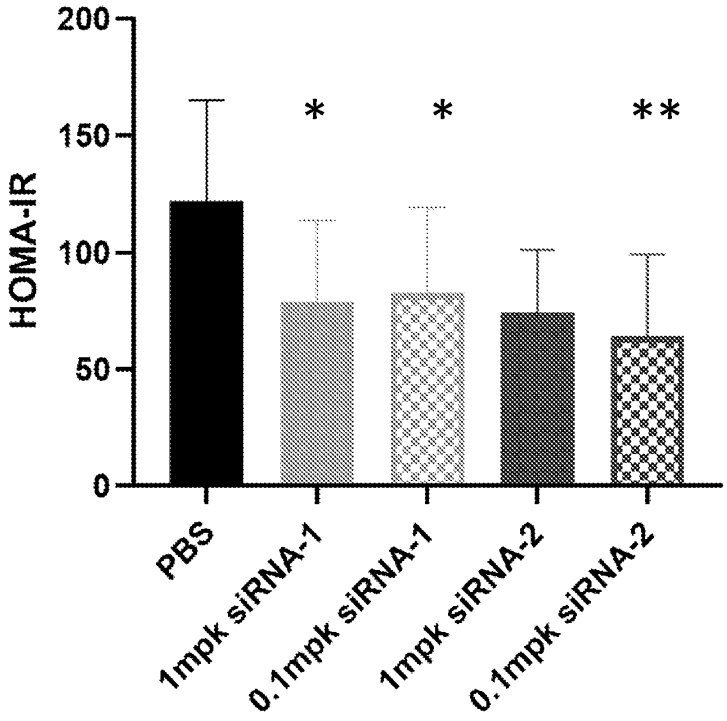

HOMA-IR
200
150
100
50
0
*
*
**
PBS
1mpk siRNA-1
0.1mpk siRNA-1
1mpk siRNA-2
0.1mpk siRNA-2

FIG. 5A

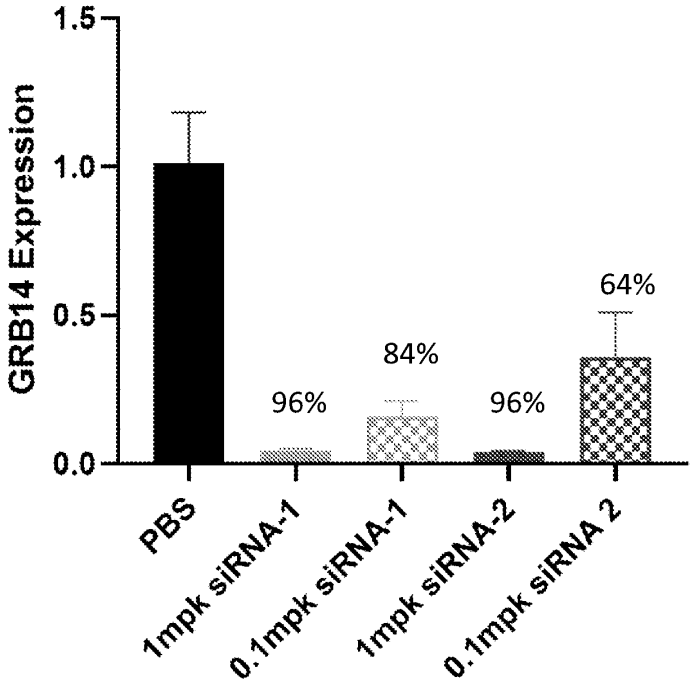

GRB14 Expression
1.5
1.0
0.5
0.0
96%
84%
96%
64%
PBS
1mpk siRNA-1
0.1mpk siRNA-1
1mpk siRNA-2
0.1mpk siRNA 2

FIG. 5B

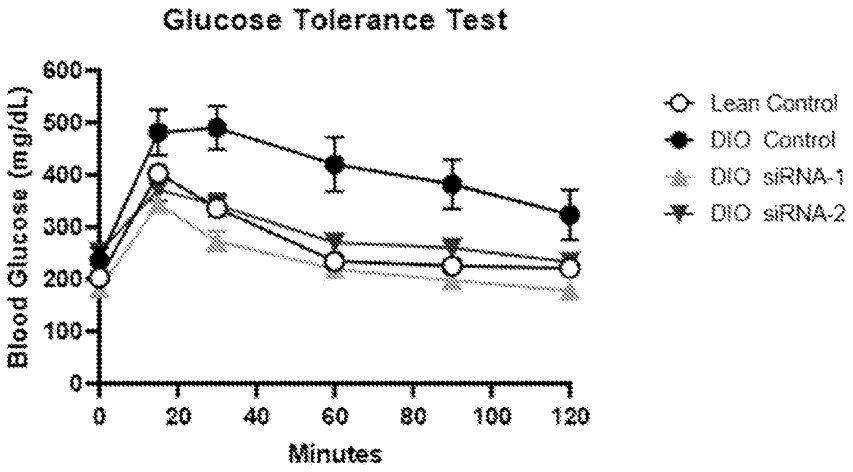


FIG. 6A

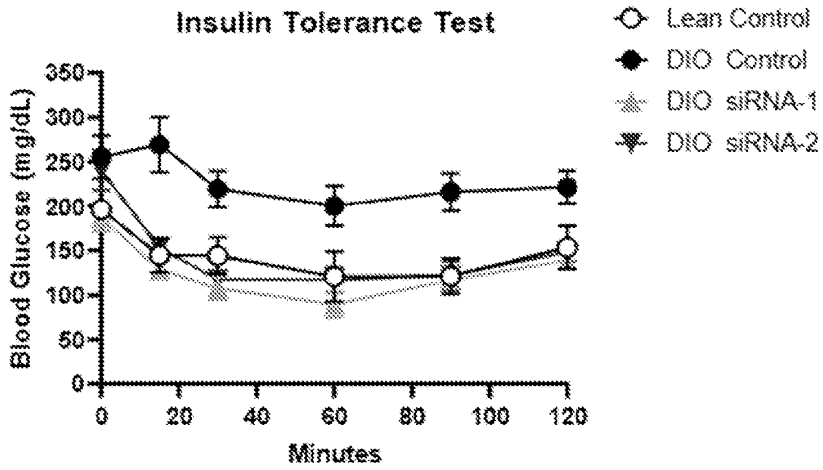


FIG. 6B

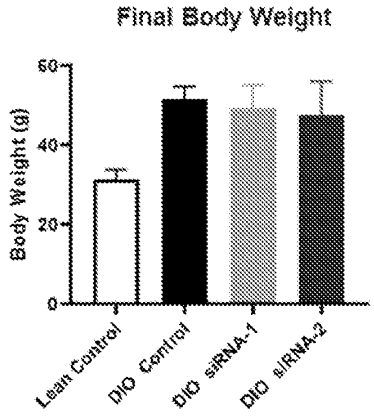


FIG. 6C