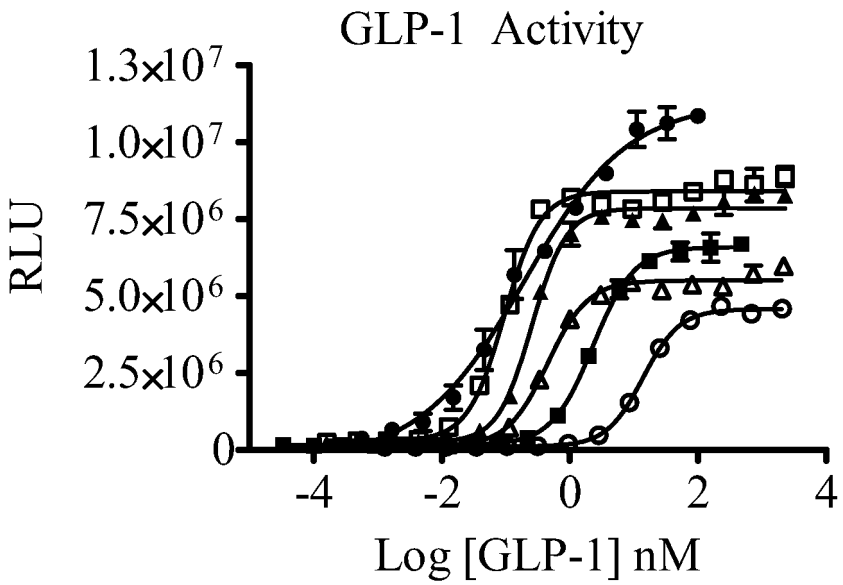

GLP-1 Activity
RLU
1.3x10^7
1.0x10^7
7.5x10^6
5.0x10^6
2.5x10^6
0
-4
-2
0
2
4
Log [GLP-1] nM

FIG 1A

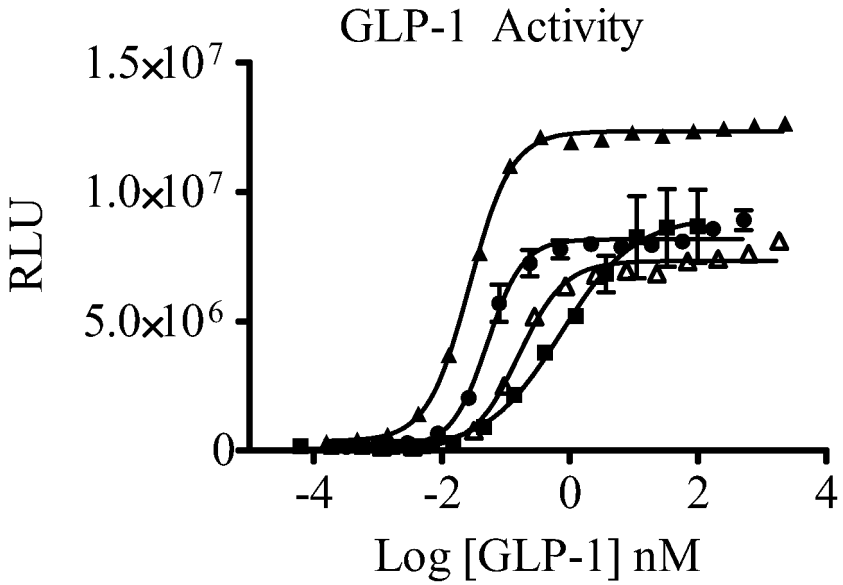

GLP-1 Activity
RLU
1.5x10^7
1.0x10^7
5.0x10^6
0
-4
-2
0
2
4
Log [GLP-1] nM

FIG 1B

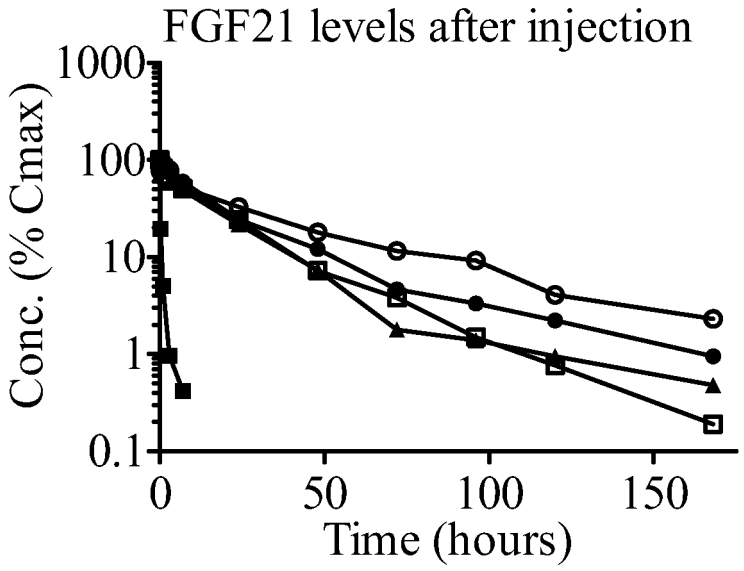

FGF21 levels after injection
Conc. (% Cmax)
1000
100
10
1
0.1
0
50
100
150
Time (hours)

FIG 2

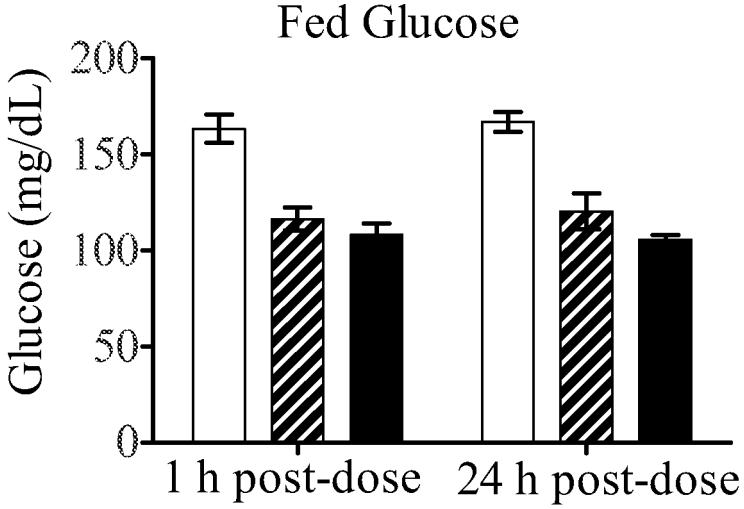

Fed Glucose
Glucose (mg/dL)
200
150
100
50
0
1 h post-dose
24 h post-dose

FIG 3A

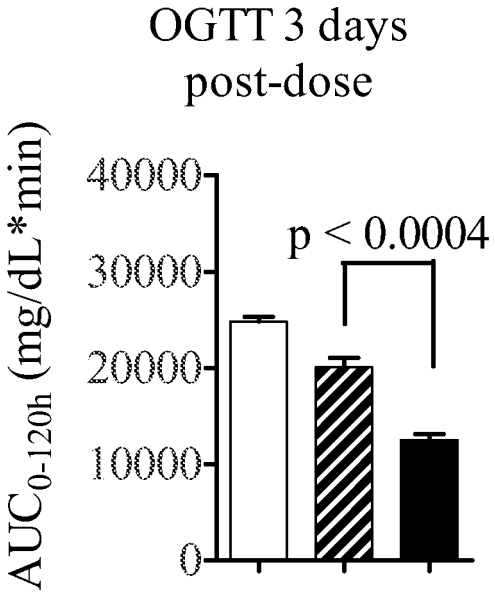

OGTT 3 days
post-dose
AUC0-120h (mg/dL*min)
40000
30000
20000
10000
0
p < 0.0004

FIG 3B

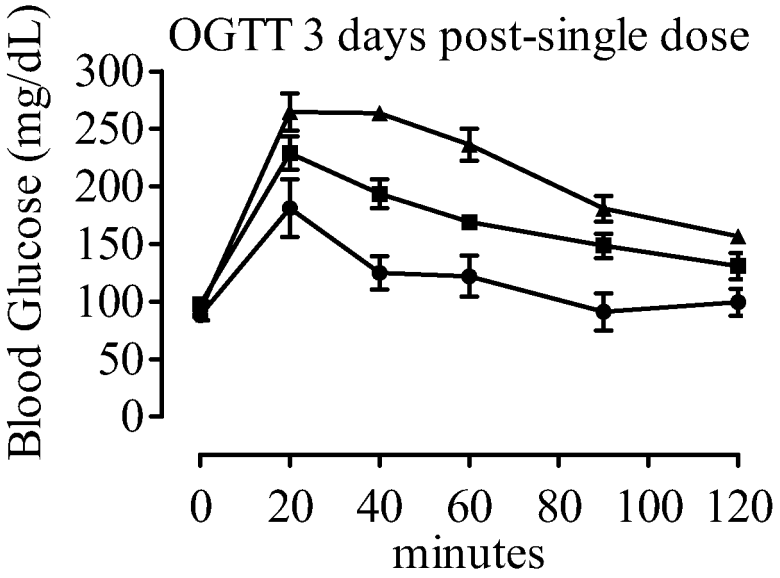

OGTT 3 days post-single dose
Blood Glucose (mg/dL)
300
250
200
150
100
50
0
0 20 40 60 80 100 120
minutes

FIG 3C

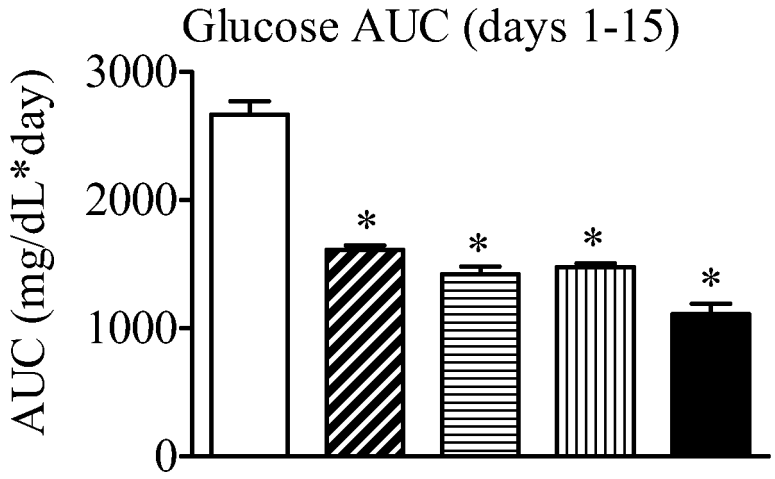


FIG 4

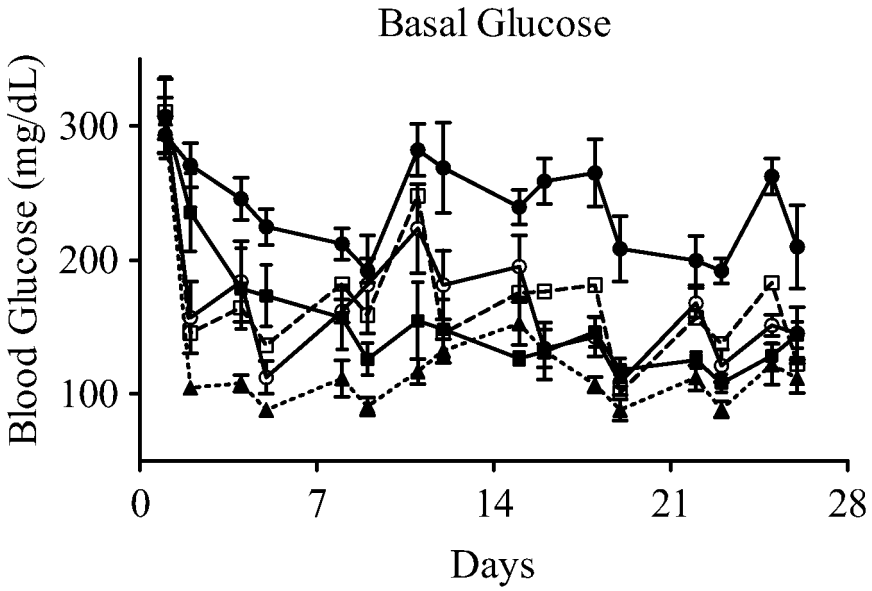

Basal Glucose
Blood Glucose (mg/dL)
300
200
100
0
7
14
21
28
Days

FIG 5A

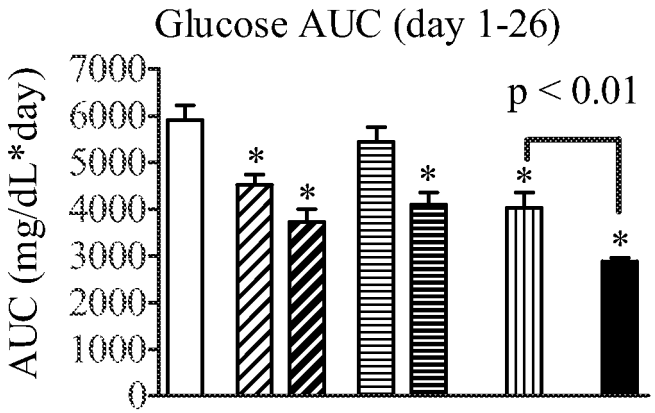

Glucose AUC (day 1-26)
AUC (mg/dL*day)
7000
6000
5000
4000
3000
2000
1000
0
p < 0.01
*

FIG 5B

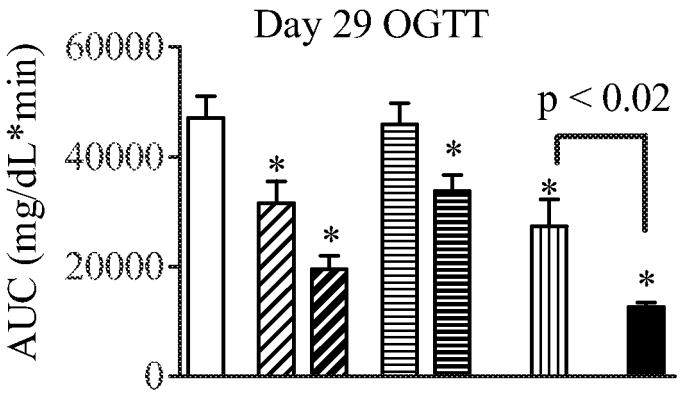

Day 29 OGTT
AUC (mg/dL*min)
60000
40000
20000
0
p < 0.02
*
*
*
*
*

FIG 5C

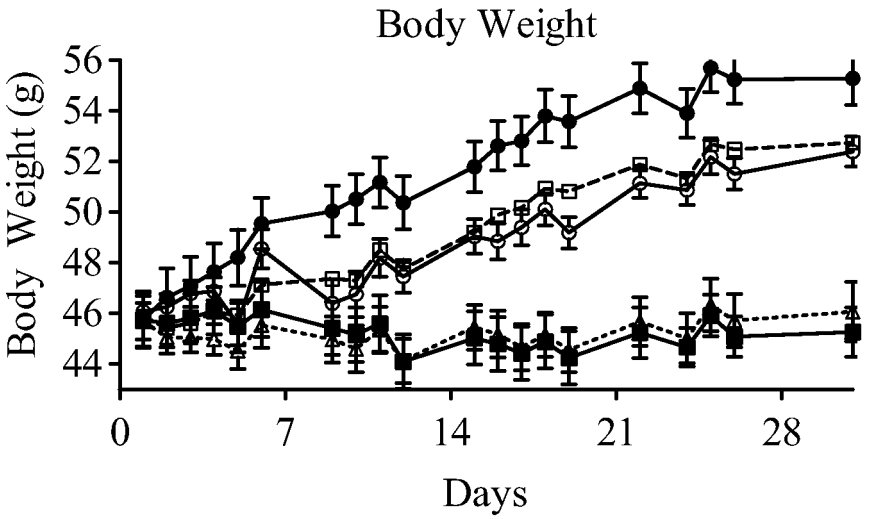

Body Weight
Body Weight (g)
56
54
52
50
48
46
44
0
7
14
21
28
Days

FIG 5D

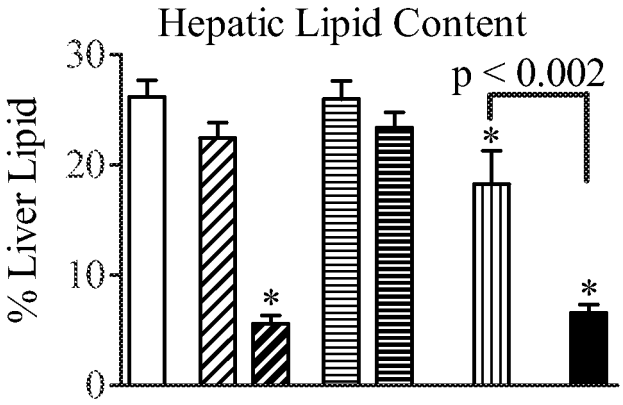

Hepatic Lipid Content
p < 0.002
% Liver Lipid
30
20
10
0
*
*
*

FIG 5E

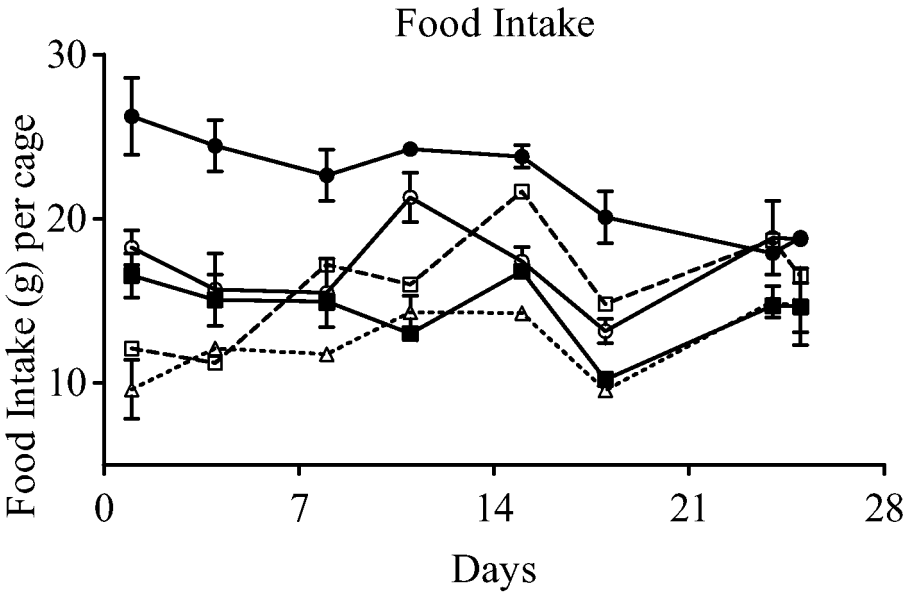

Food Intake
Food Intake (g) per cage
30
20
10
0
7
14
21
28
Days

FIG 5F

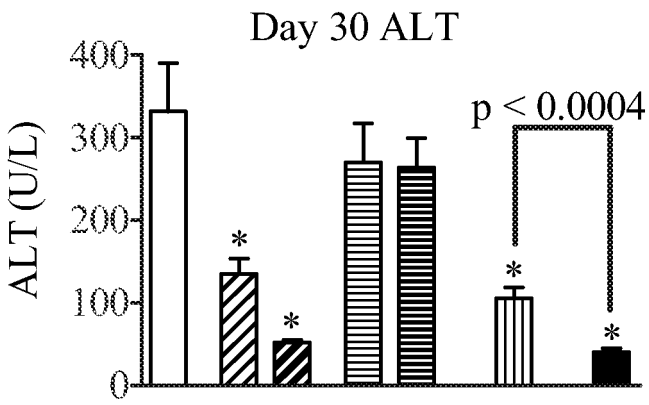


FIG 5G

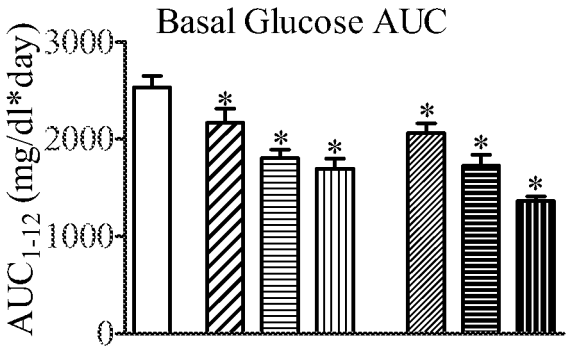


FIG 6A

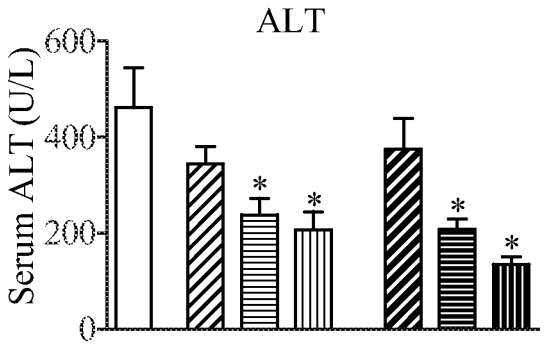


FIG 6B

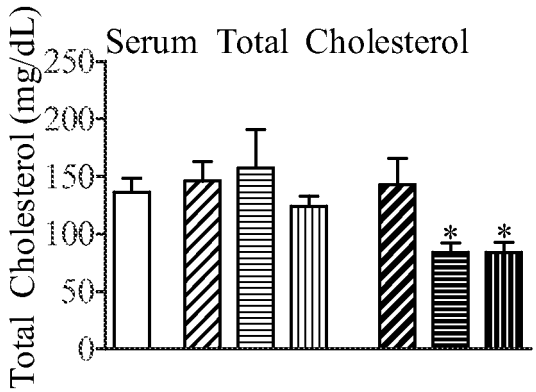


FIG 6C

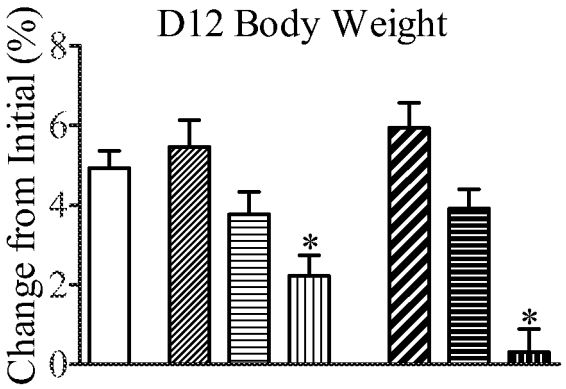


FIG 6D

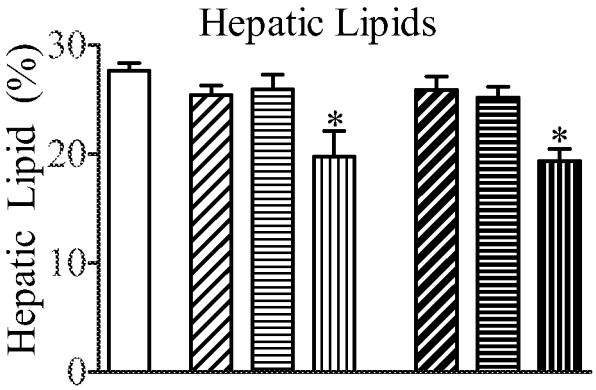


FIG 6E

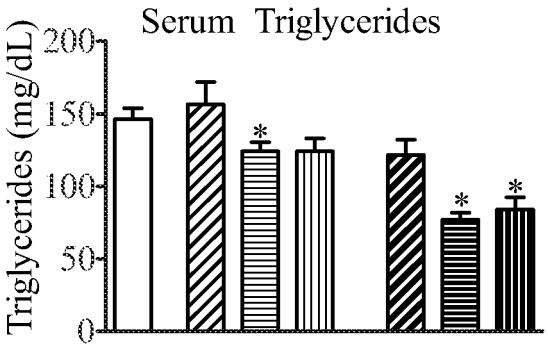


FIG 6F

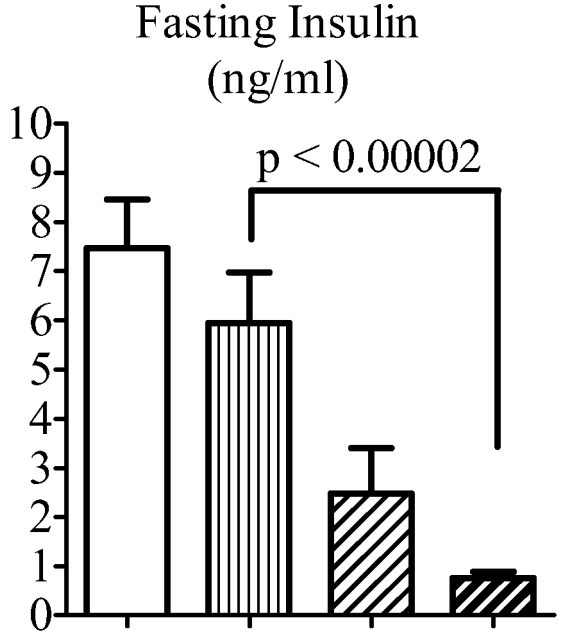

Fasting Insulin
(ng/ml)
10
9
8
7
6
5
4
3
2
1
0
p < 0.00002

FIG 7A

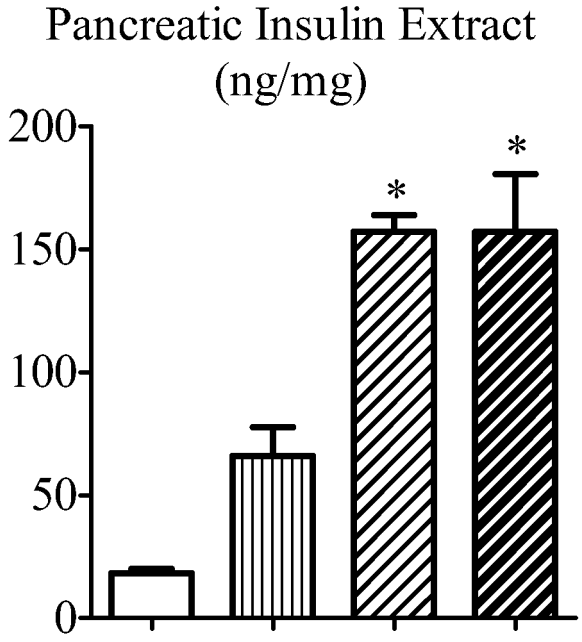

Pancreatic Insulin Extract
(ng/mg)
200
150
100
50
0
*
*

FIG 7B

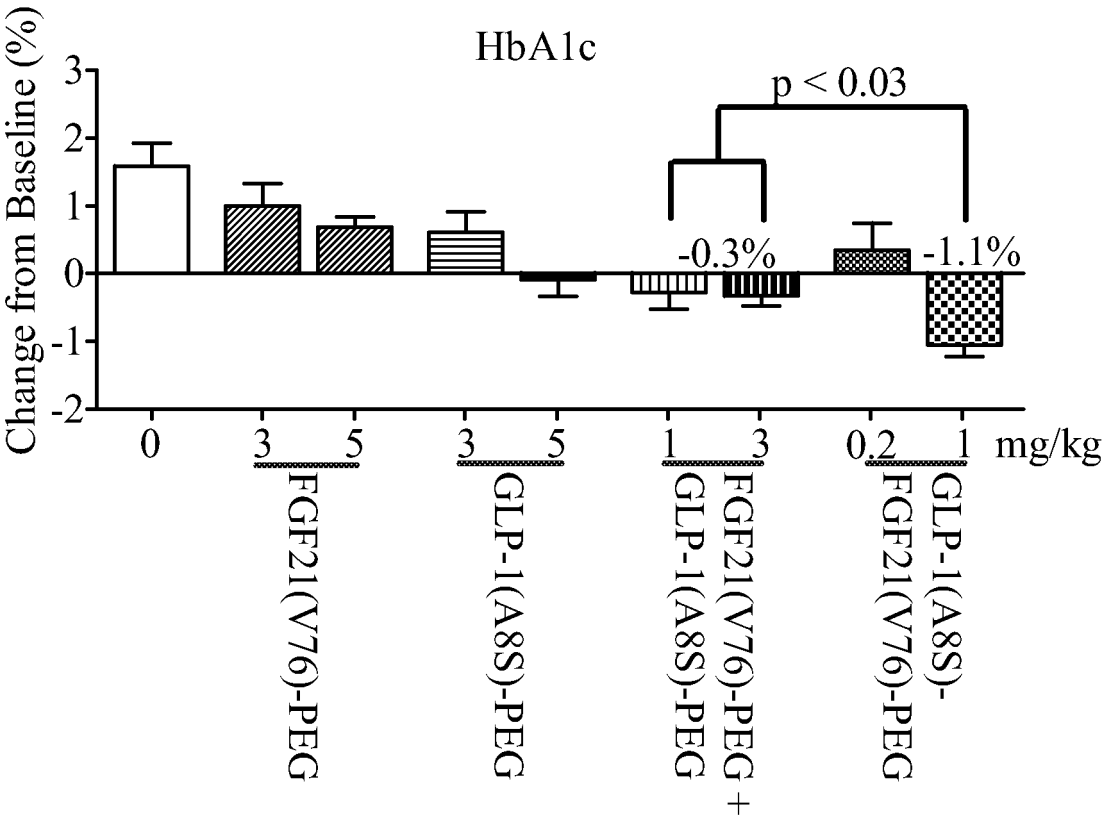

HbA1c
Change from Baseline (%)
3
2
1
0
-1
-2
p < 0.03
-0.3%
-1.1%
0
3
5
3
5
1
3
0.2
1
mg/kg
FGF21(V76)-PEG
GLP-1(A8S)-PEG
GLP-1(A8S)-PEG
FGF21(V76)-PEG +
FGF21(V76)-PEG
GLP-1(A8S)-

FIG 8A

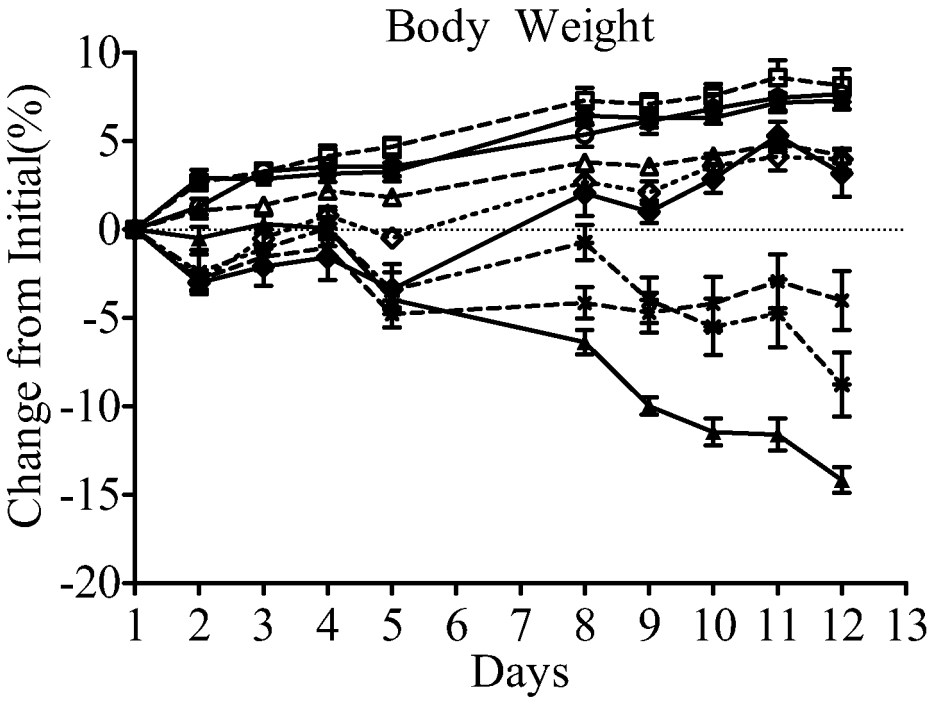

Body Weight
Change from Initial(%)
10
5
0
-5
-10
-15
-20
1 2 3 4 5 6 7 8 9 10 11 12 13
Days

FIG 8B

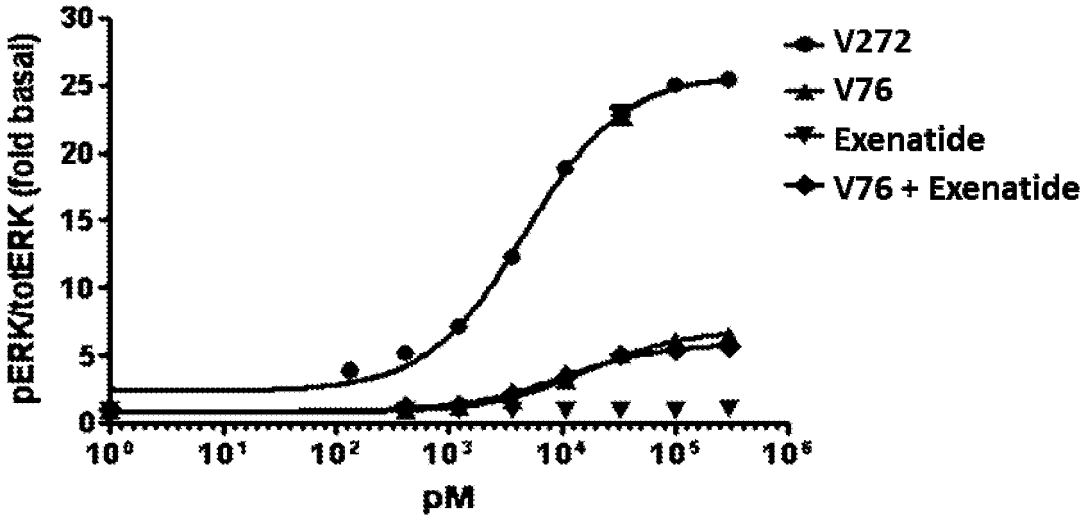

pERK/totERK (fold basal)
30
25
20
15
10
5
0
10^0
10^1
10^2
10^3
10^4
10^5
10^6
pM
V272
V76
Exenatide
V76 + Exenatide

FIG 9

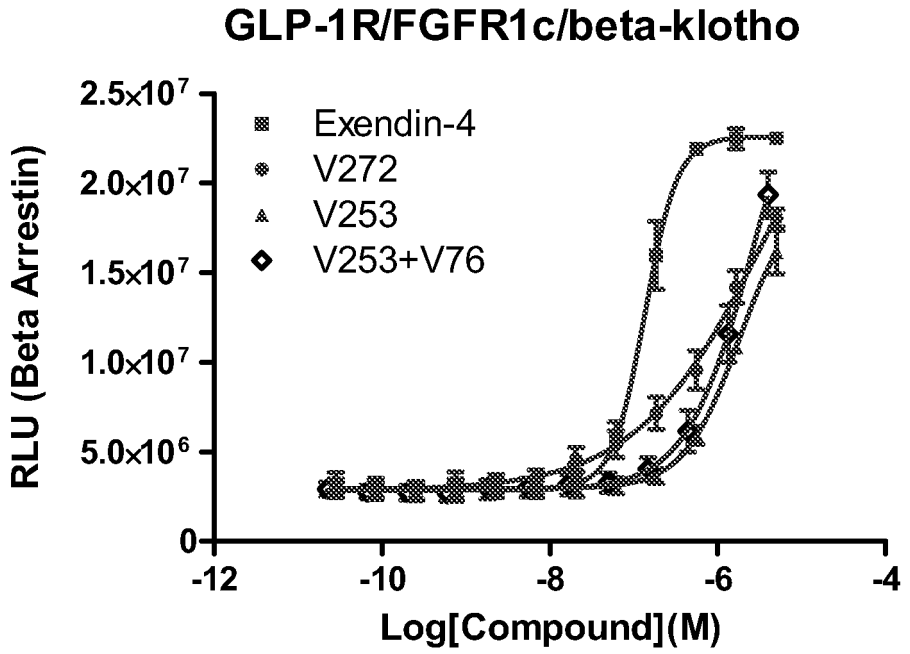


FIG 10A

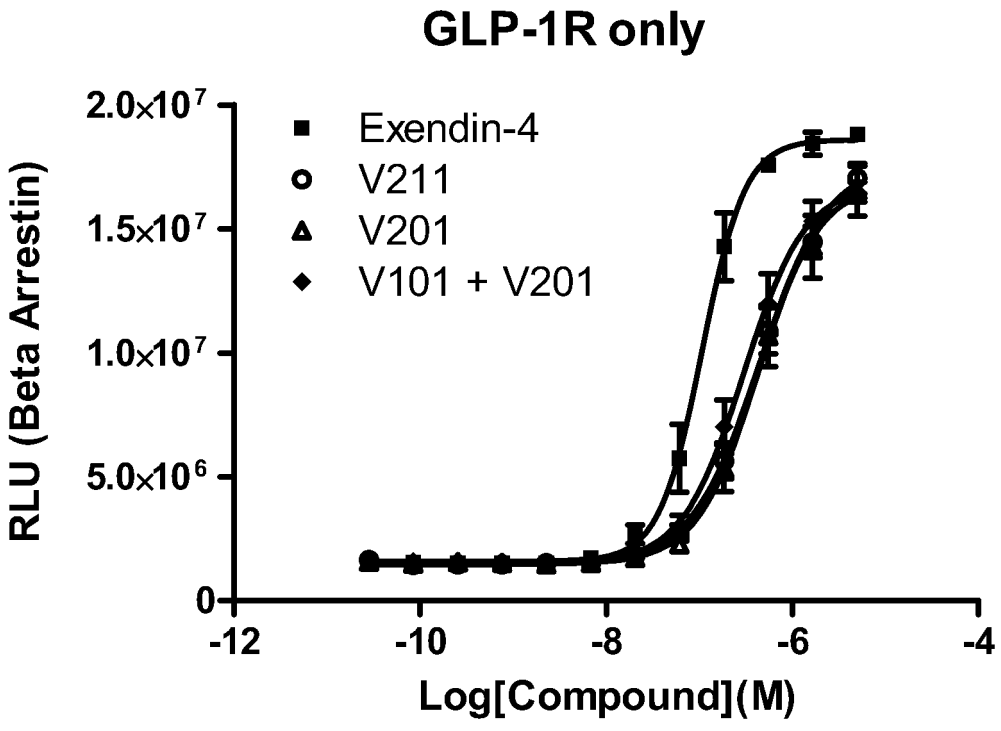


FIG 10B

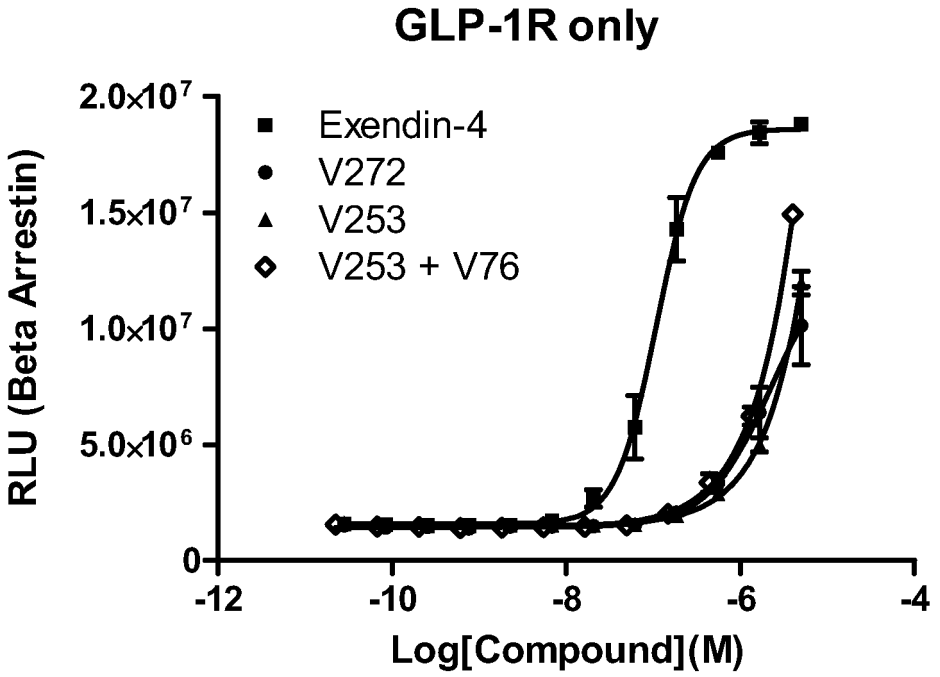


FIG 10C

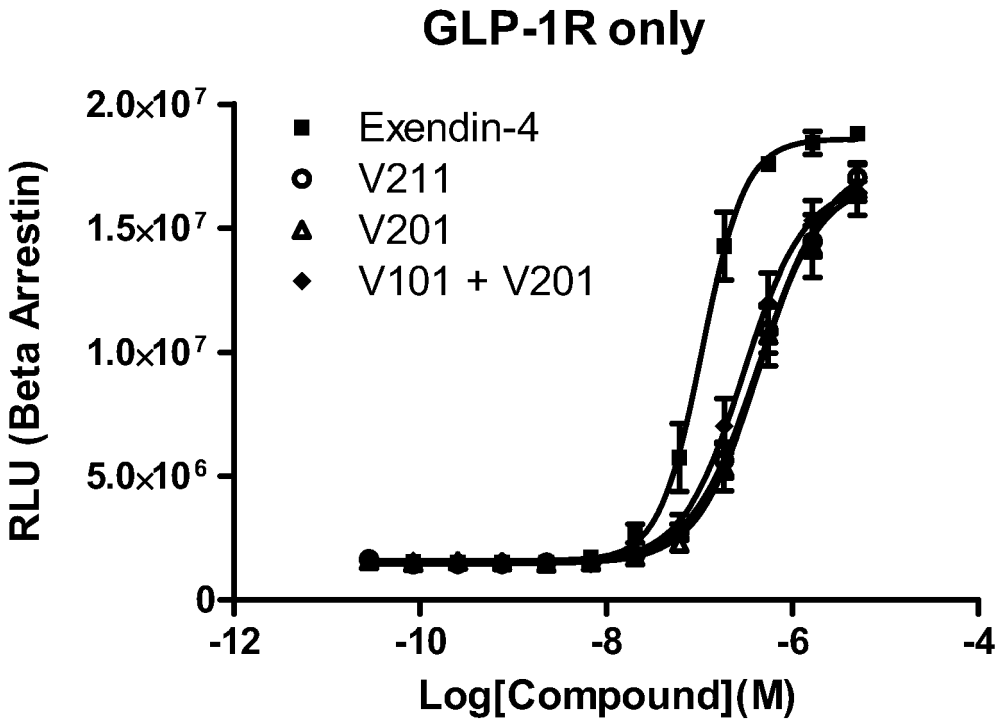


FIG 10D