Figure 1A

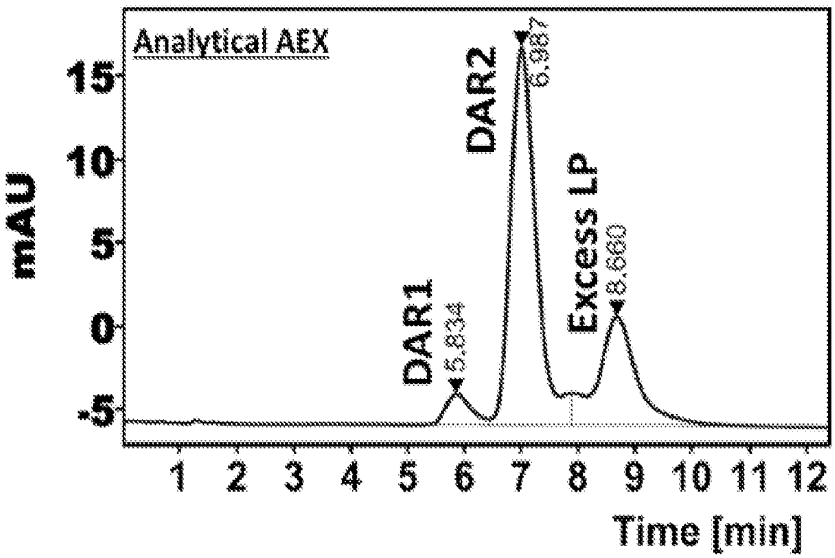


Figure 1B

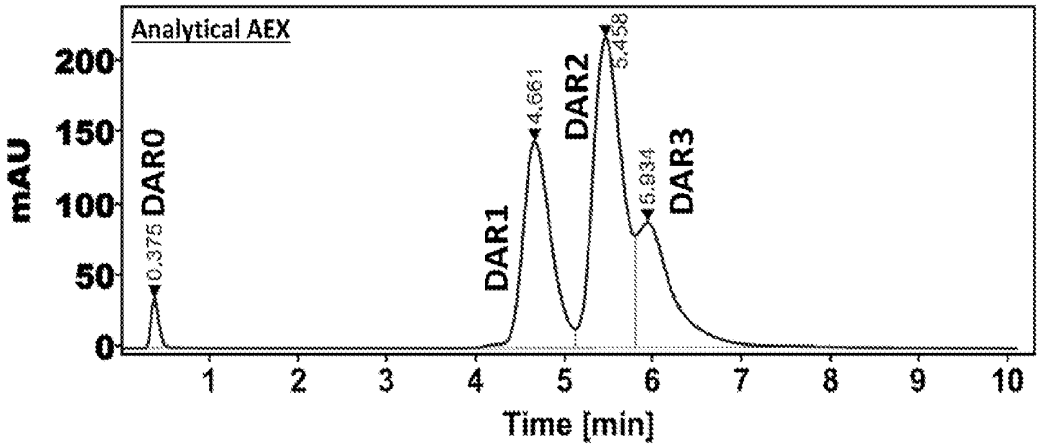


Figure 1C

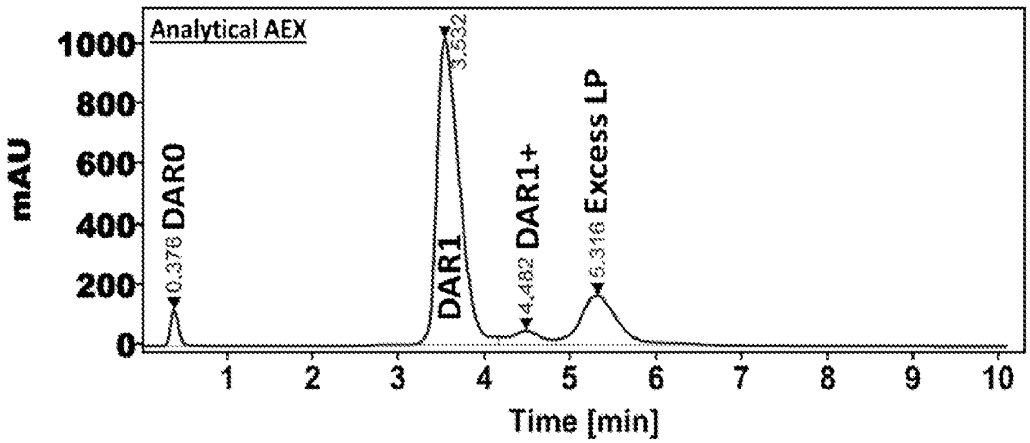


Figure 1D

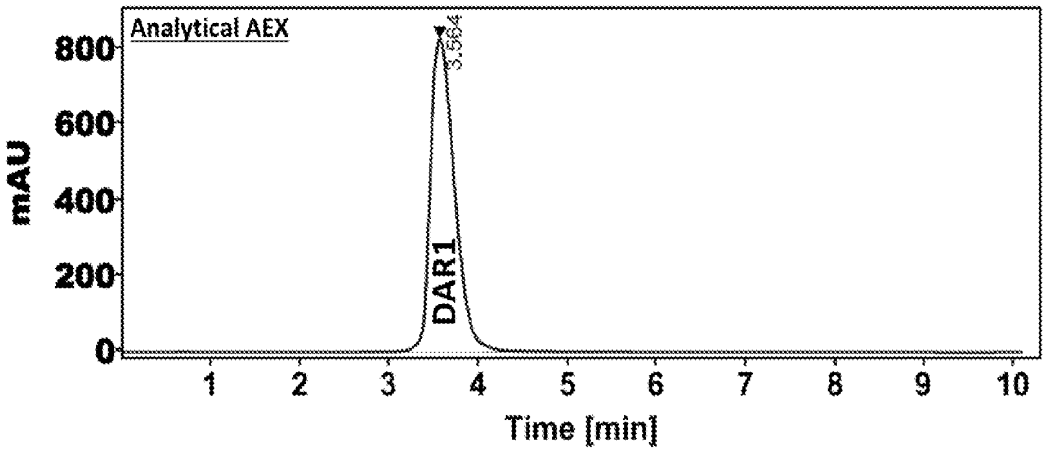


Figure 1E

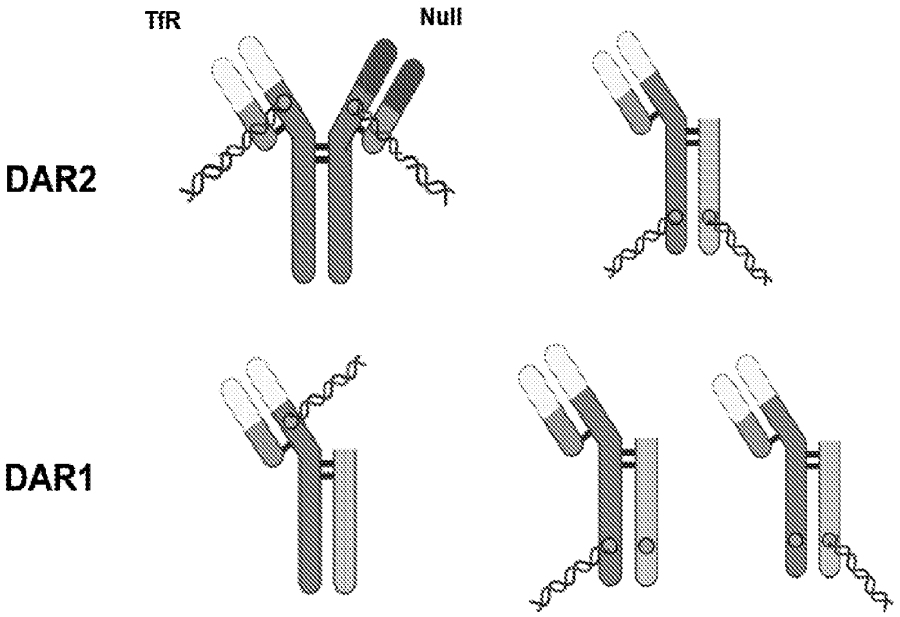


Figure 2

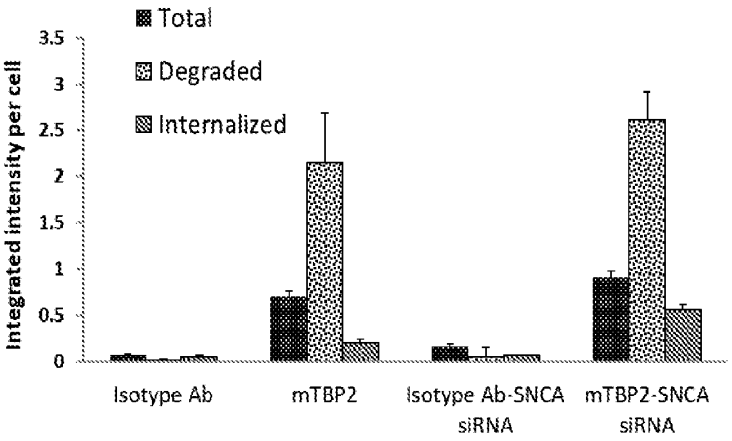


Figure 3

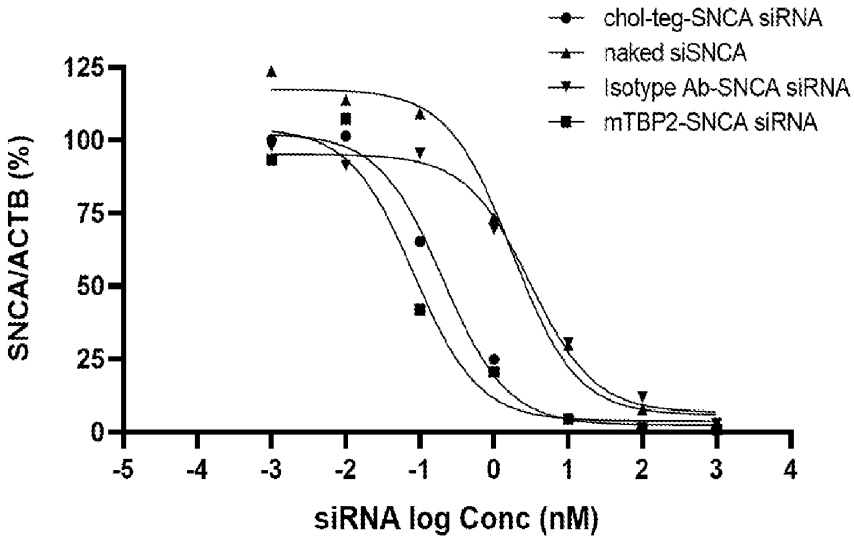


Figure 4

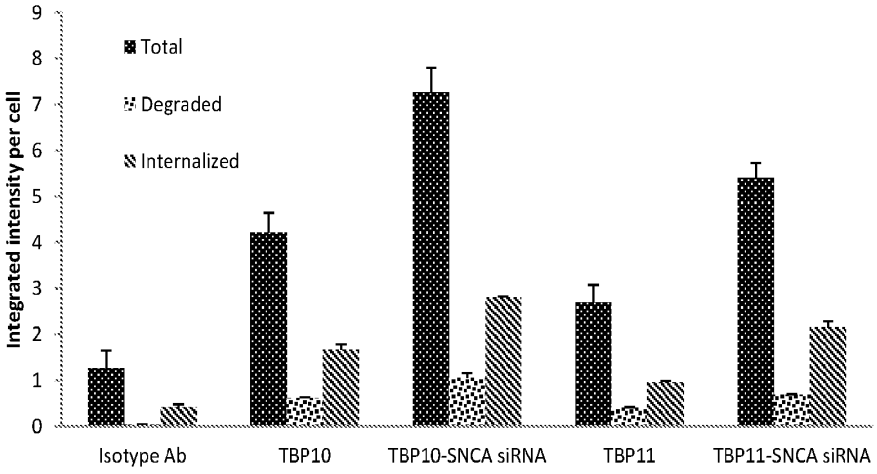


Figure 5

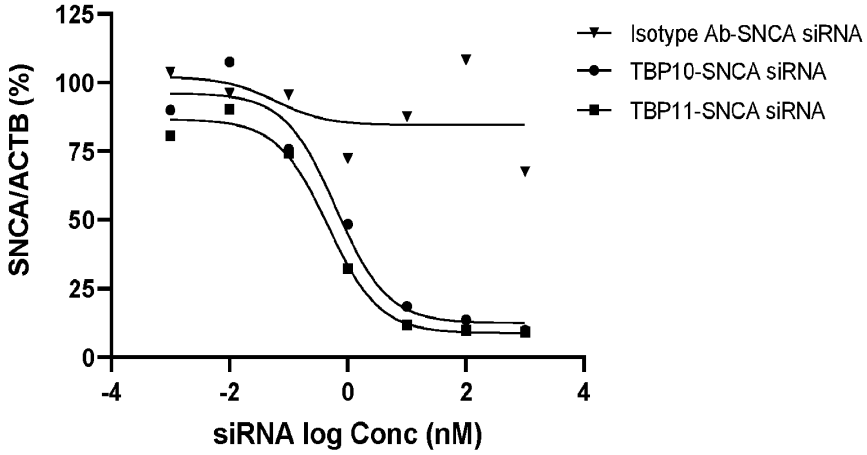


Figure 6A

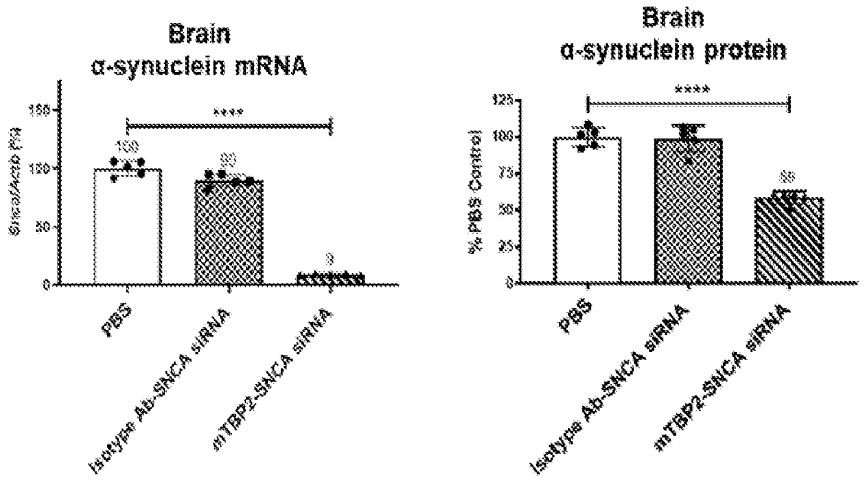
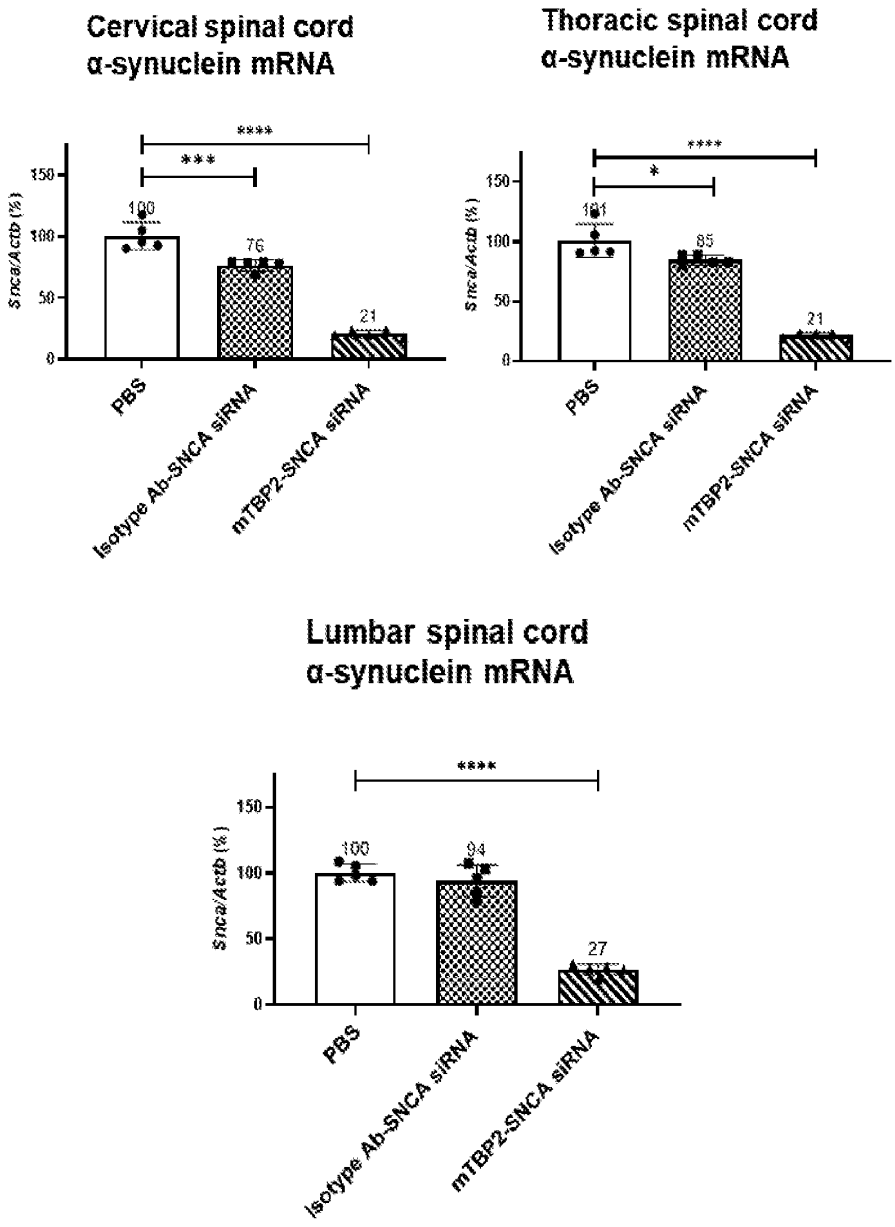


Figure 6B


Cervical spinal cord
α-synuclein mRNA
Thoracic spinal cord
α-synuclein mRNA
Lumbar spinal cord
α-synuclein mRNA
Snca/Actb (%)
PBS
Isotype Ab-SNCA siRNA
mTBP2-SNCA siRNA
100
76
21
101
85
21
100
94
27
****
***
*

7/22

Figure 6C

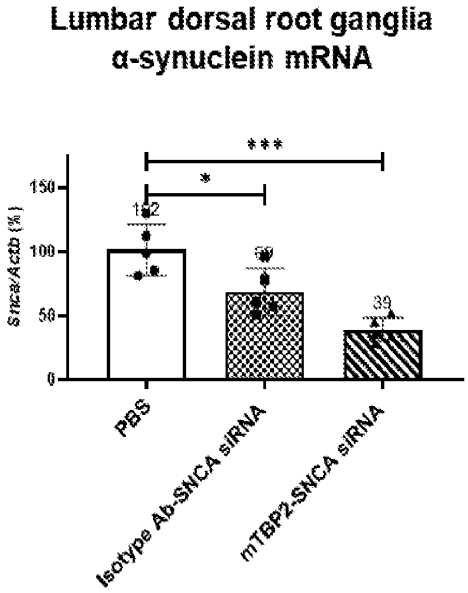


Figure 7A

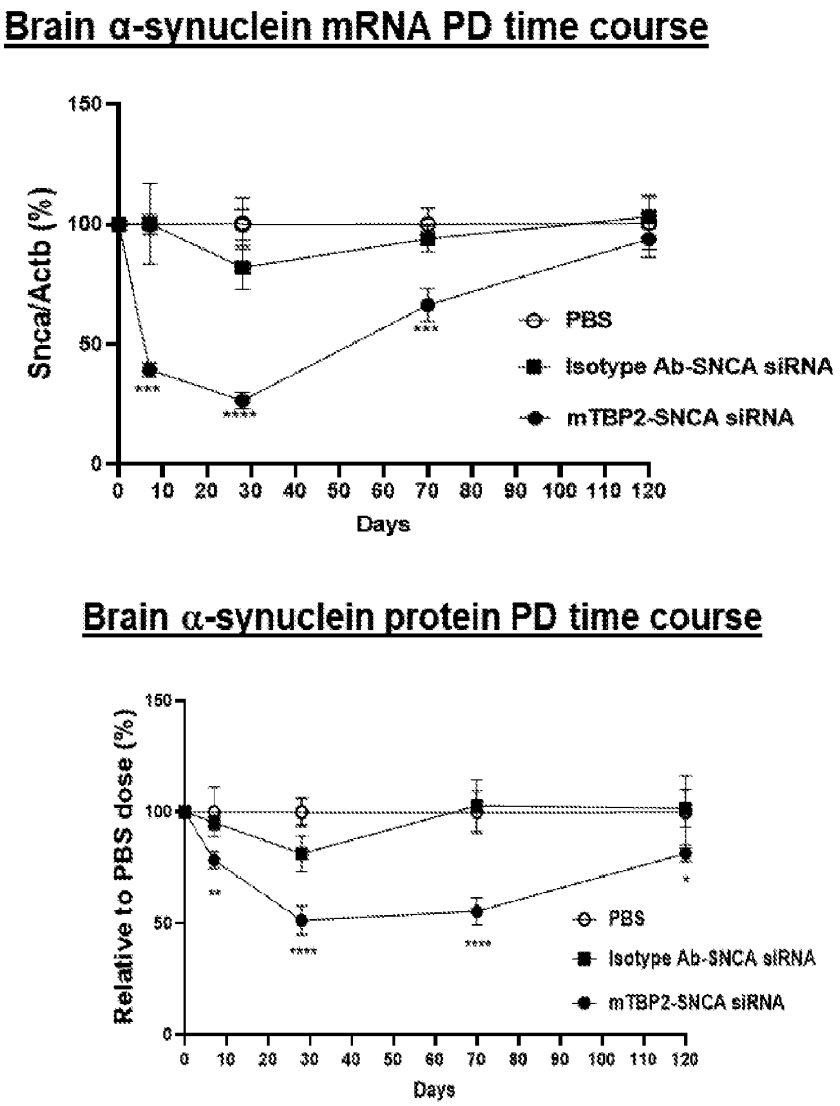
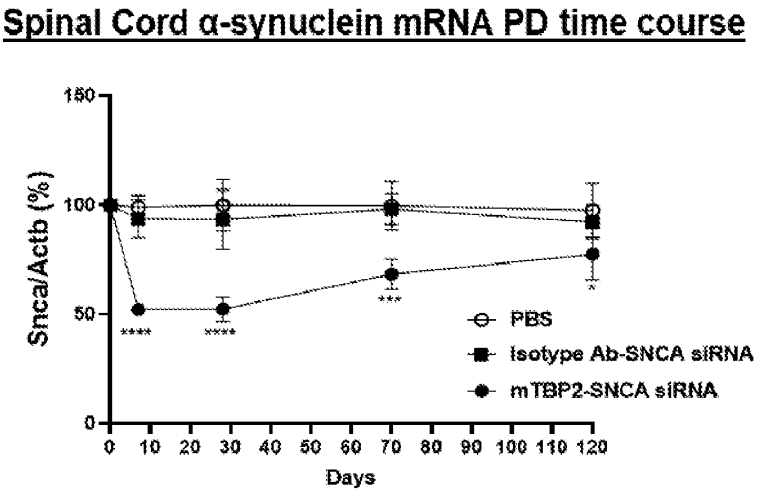
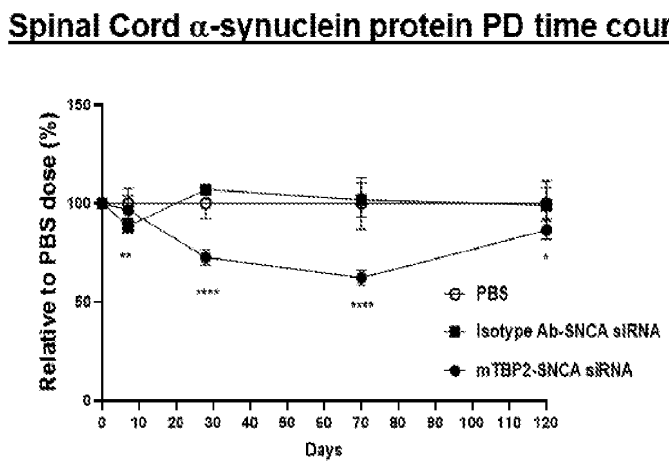

Brain α-synuclein mRNA PD time course
Snca/Actb (%)
150
100
50
0
0 10 20 30 40 50 60 70 80 90 100 110 120
Days
PBS
Isotype Ab-SNCA siRNA
mTBP2-SNCA siRNA
***
****
***
Brain α-synuclein protein PD time course
Relative to PBS dose (%)
150
100
50
0
0 10 20 30 40 50 60 70 80 90 100 110 120
Days
PBS
Isotype Ab-SNCA siRNA
mTBP2-SNCA siRNA
**
****
****
*

Figure 7B


Spinal Cord α-synuclein mRNA PD time course
Snca/Actb (%)
150
100
50
0
0 10 20 30 40 50 60 70 80 90 100 110 120
Days
PBS
Isotype Ab-SNCA siRNA
mTBP2-SNCA siRNA


Spinal Cord α-synuclein protein PD time course
Relative to PBS dose (%)
150
100
50
0
0 10 20 30 40 50 60 70 80 90 100 110 120
Days
PBS
Isotype Ab-SNCA siRNA
mTBP2-SNCA siRNA

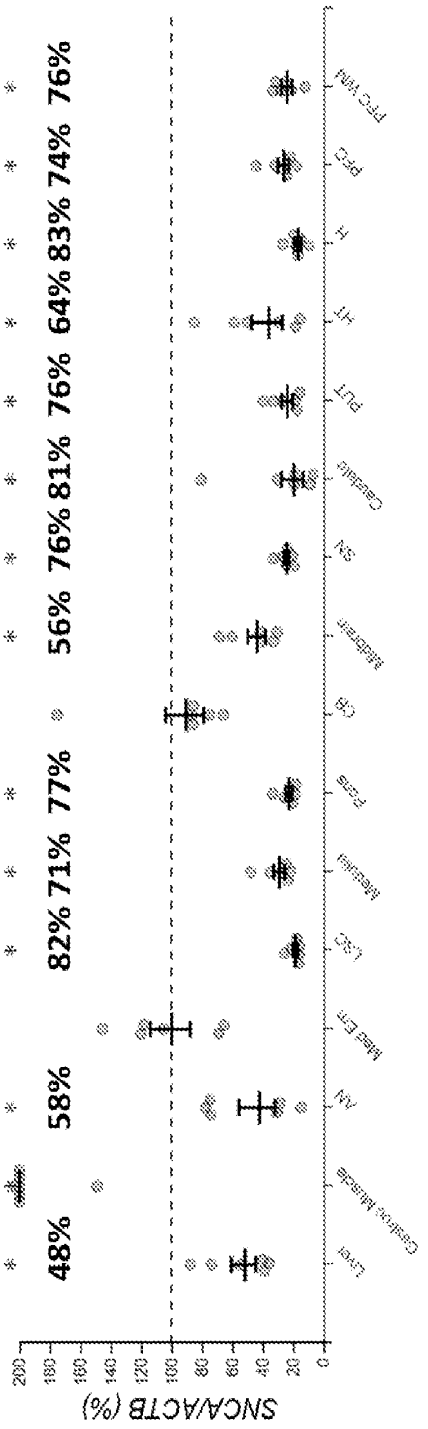

SNCA/ACTB (%)
200
180
160
140
120
100
80
60
40
20
0
48%
58%
82%
71%
77%
56%
76%
81%
76%
64%
83%
74%
76%
Liver
Cerebellar tissue
Ad
Mast Cn
Ute
Mesothel
Pros
Ov
Melanom
SM
Cervix
PLT
HT
Fi
PPC
PPC Vas

Figure 8A

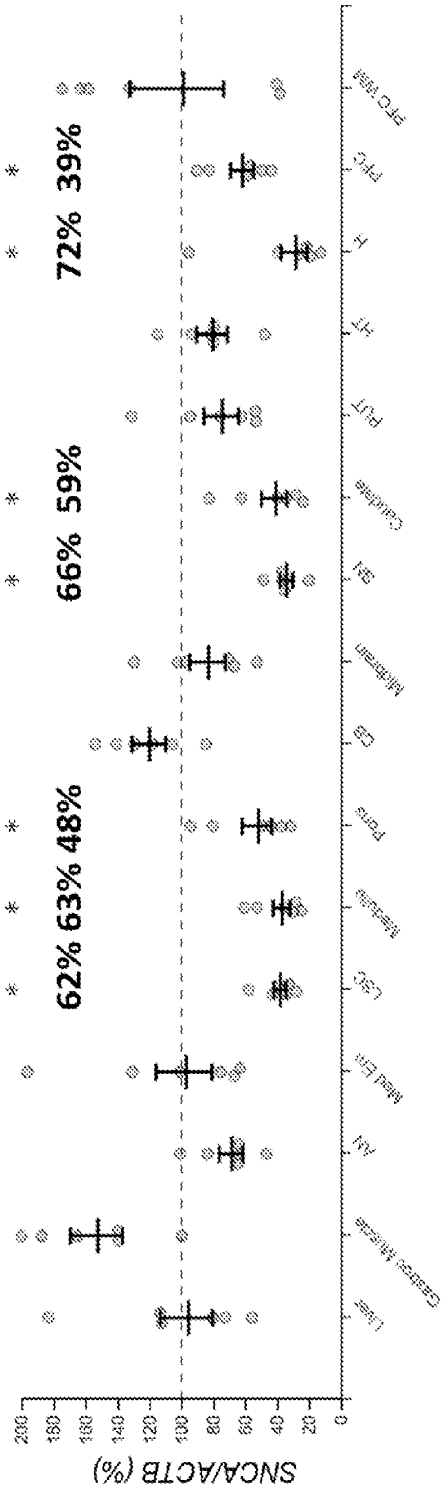

SNCA/ACTB (%)
200
160
140
120
100
80
60
40
20
0
Liver
Gastroc Muscle
TA
Med Gm
Lat Gm
Cortex
Pons
CB
Midbrain
SN
Caudate
Put
HT
Hip
PFC
PFC Dors
72% 39%
66% 59%
62% 63% 48%

Figure 8B

Figure 8C

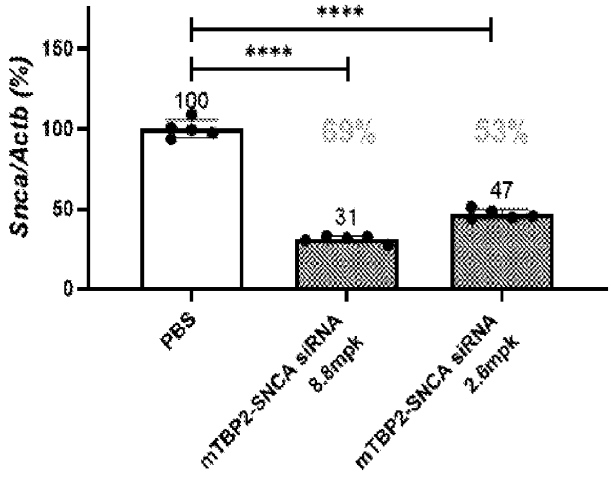


Figure 9A

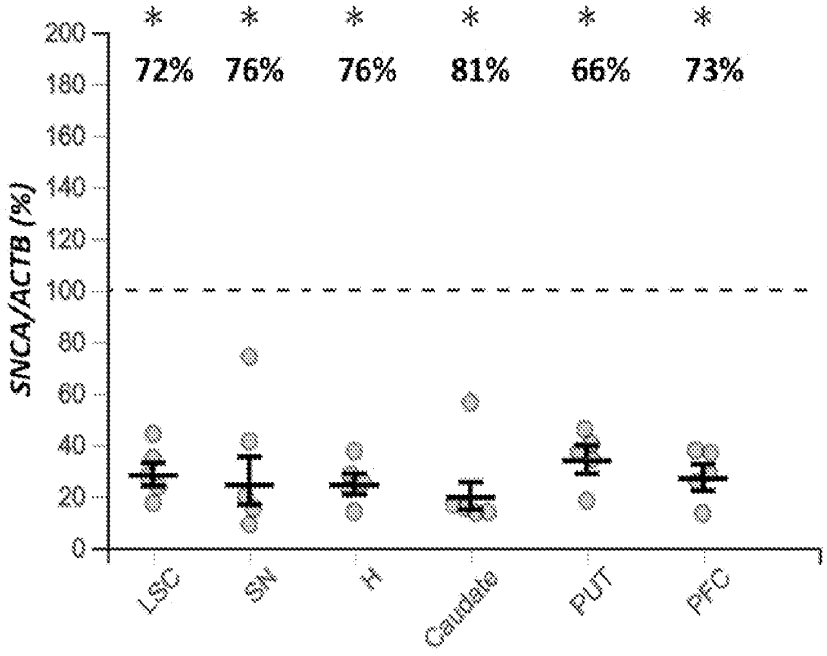


Figure 9B

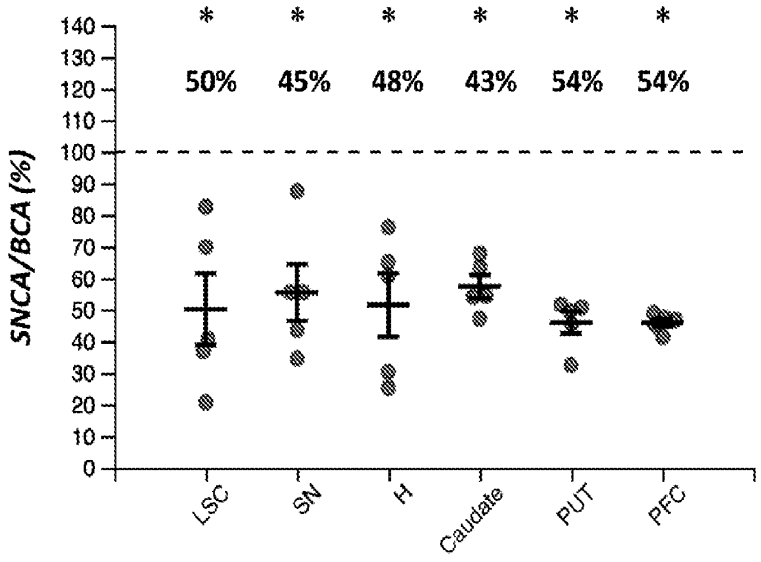


Figure 9C

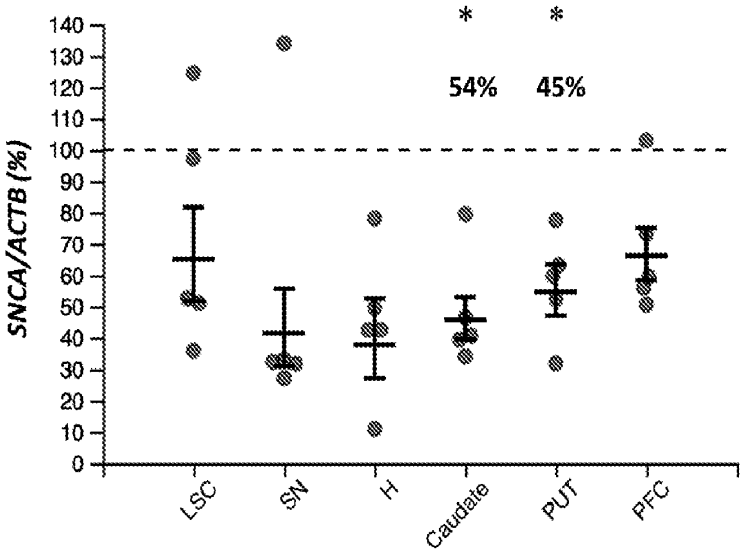


Figure 9D

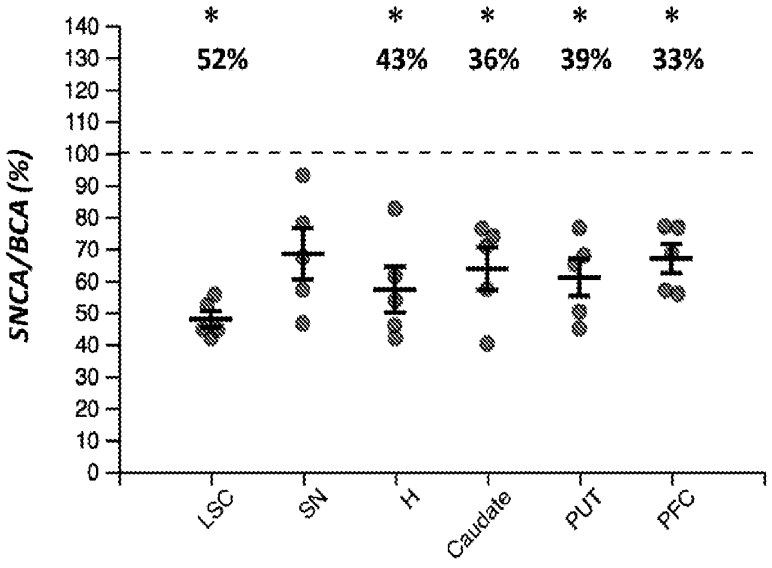


Figure 9E

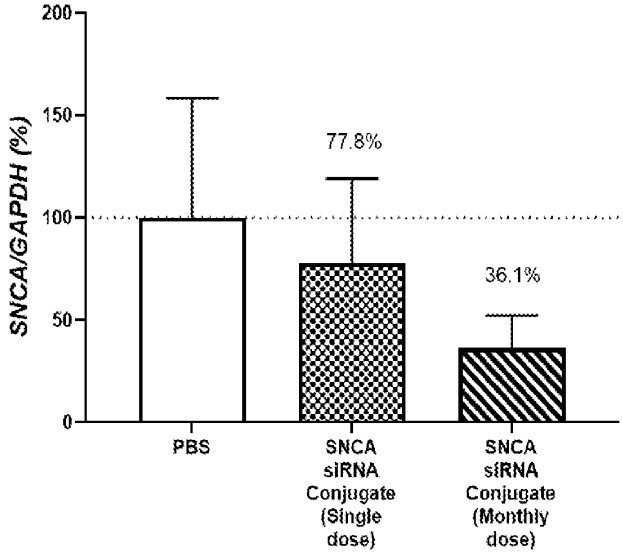


Figure 10A

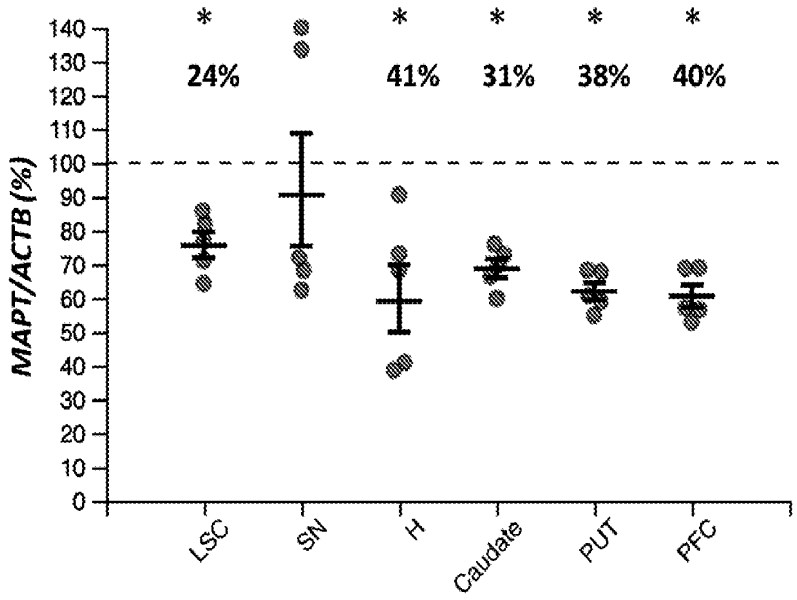


Figure 10B

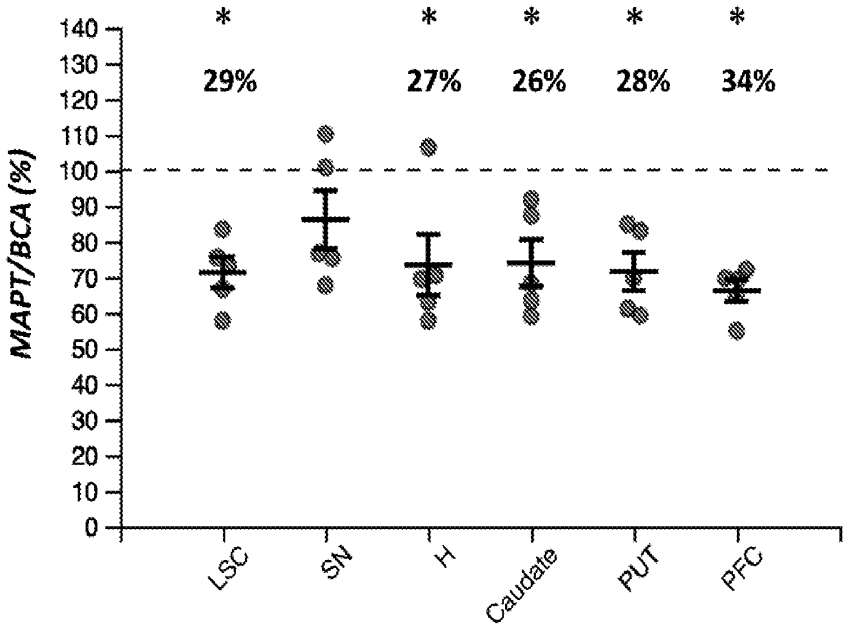


Figure 11A

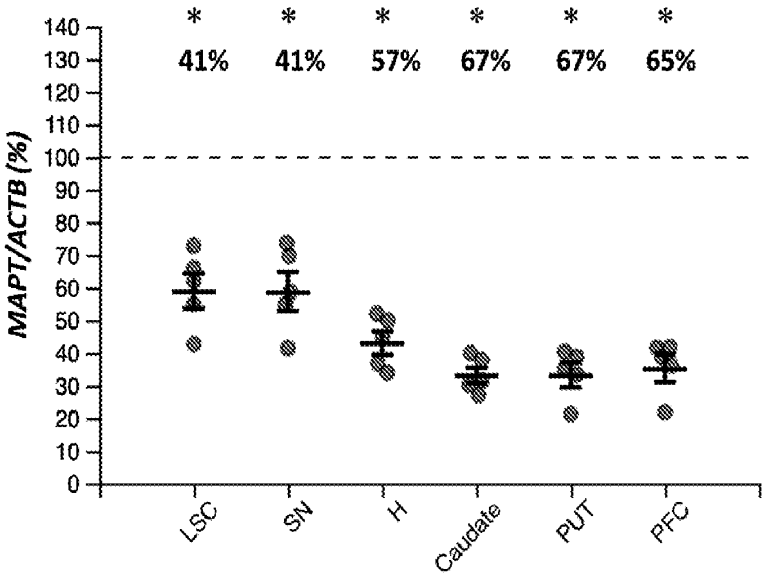


Figure 11B

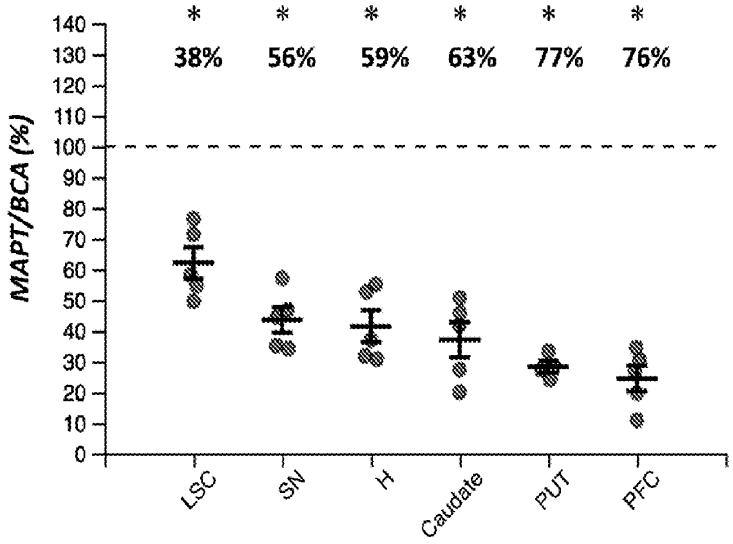


Figure 12A

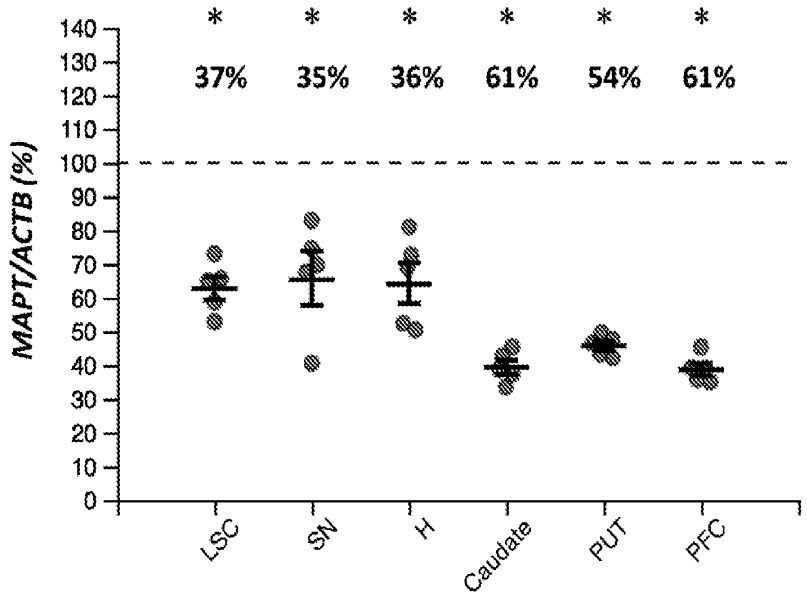


Figure 12B

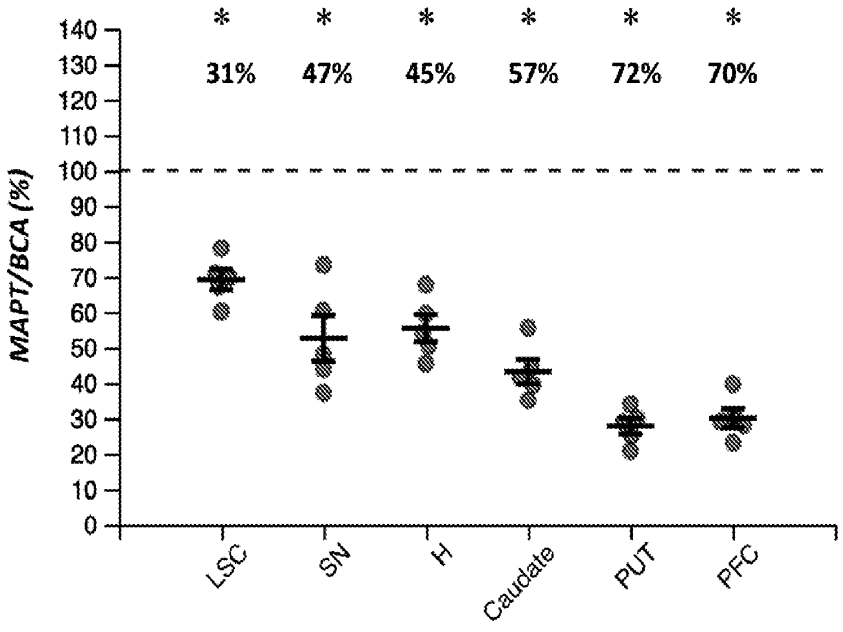


Figure 13A

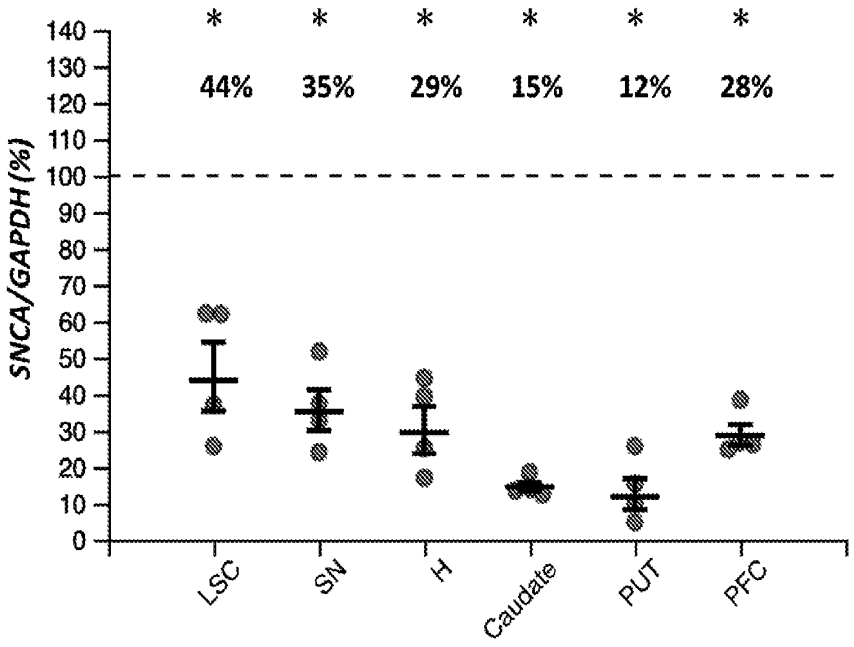


Figure 13B

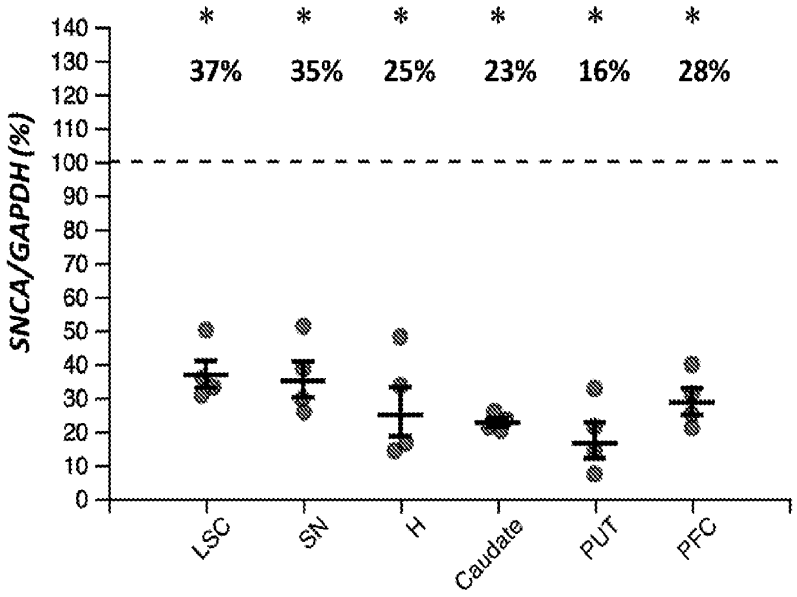


Figure 13C

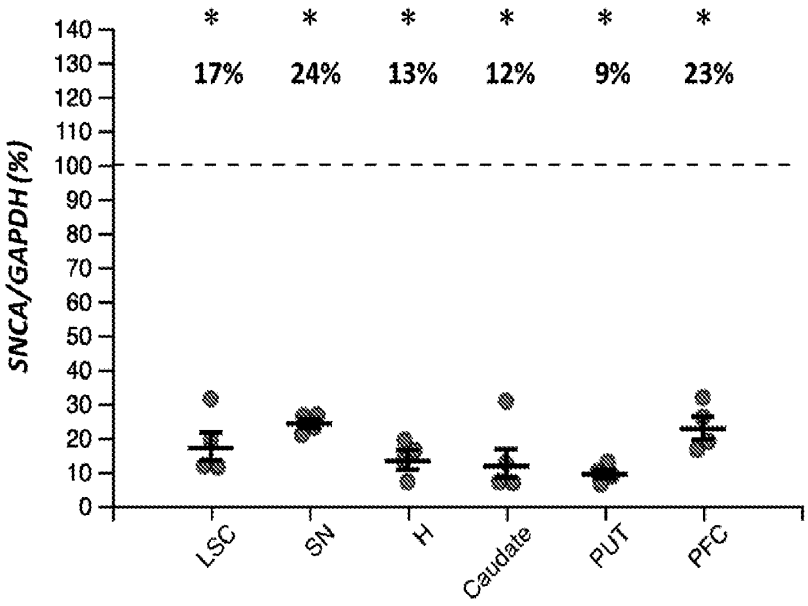


Figure 13D

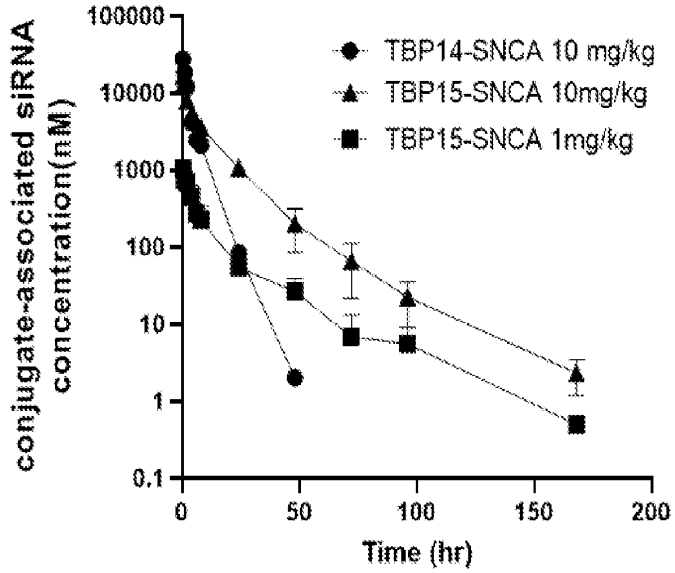


Figure 13E

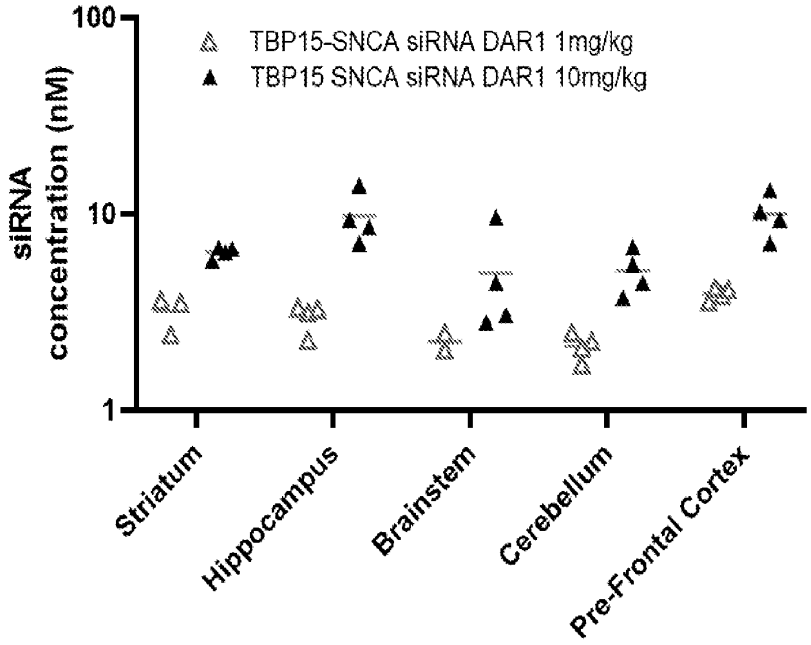


Figure 14A

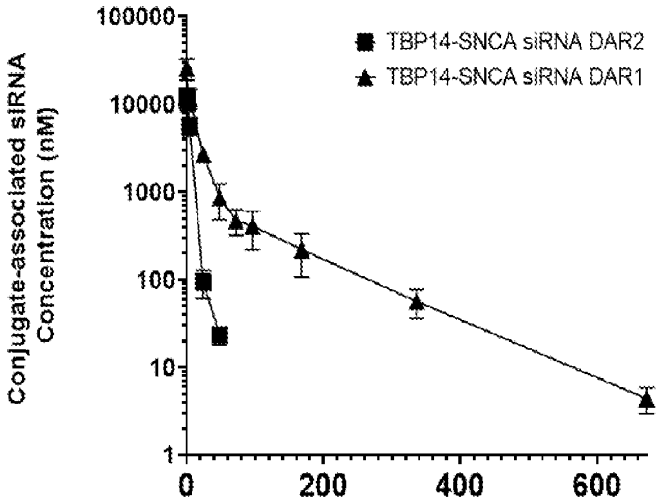


Figure 14B

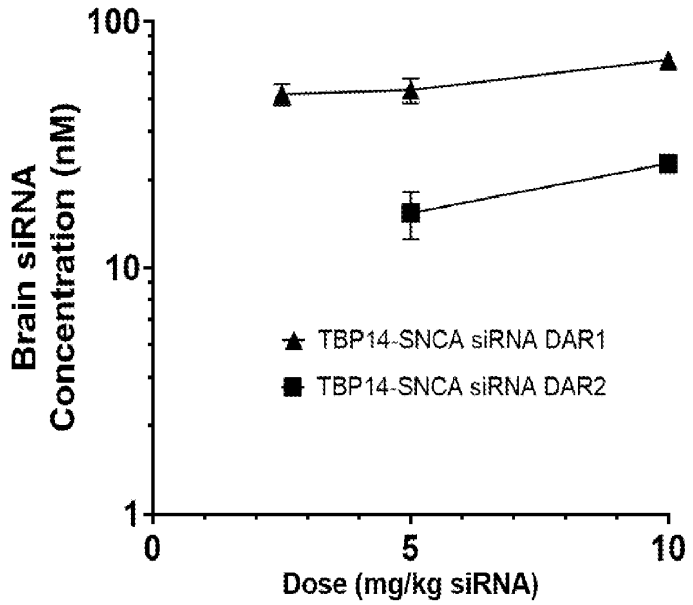


Figure 14C

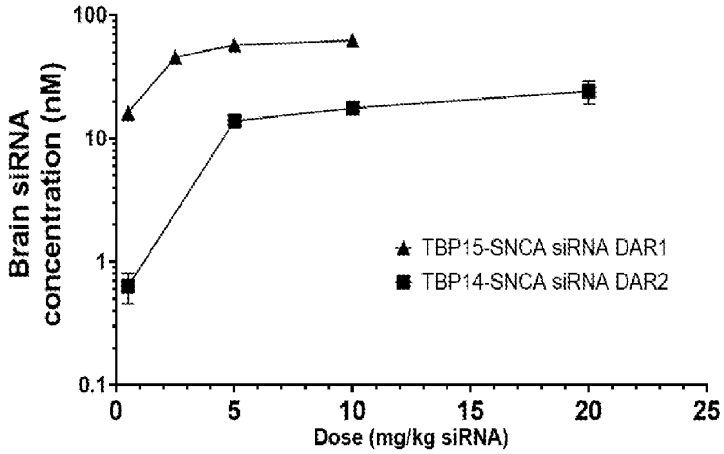
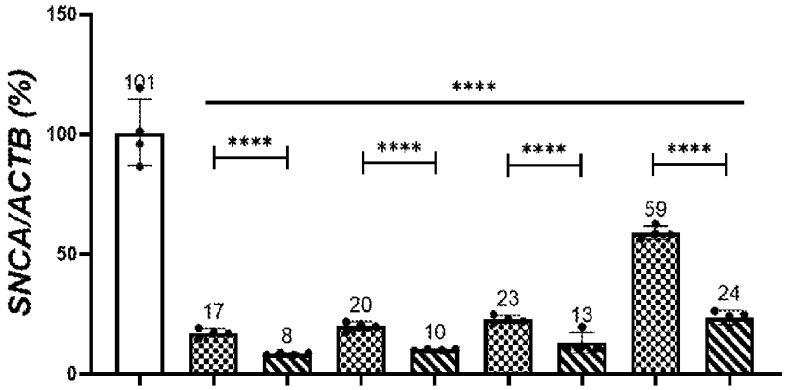


Figure 14D



DAR	2	1	2	1	2	1	2	1
mAb (mg/kg)	66	66	33	33	16.5	16.5	1.65	3.3
siRNA (mg/kg)	20	10	10	5	5	2.5	0.5	0.5

Figure 14E

