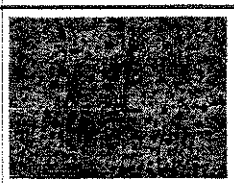
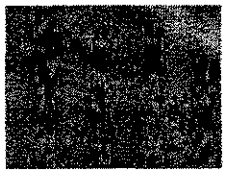
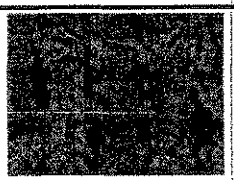
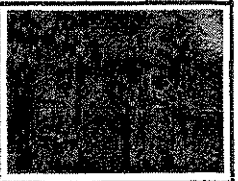
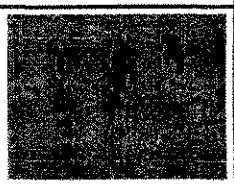
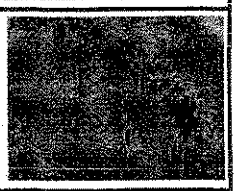
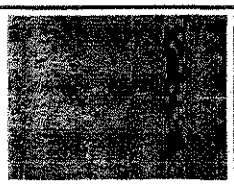
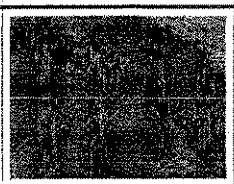
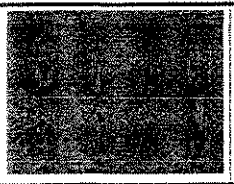
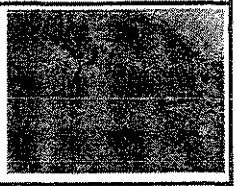
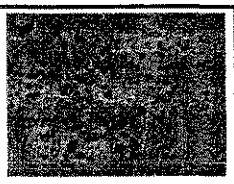
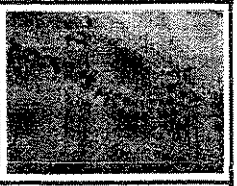
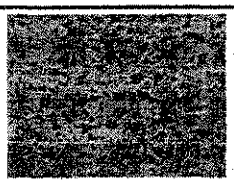
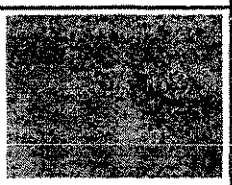
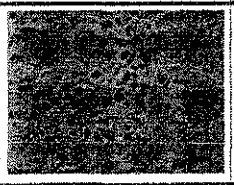
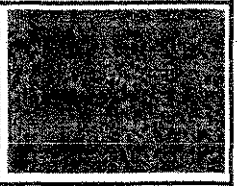
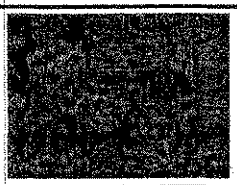
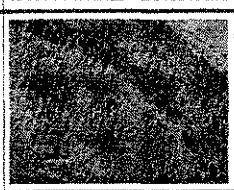
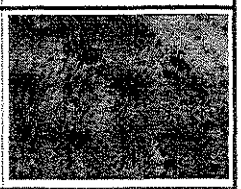
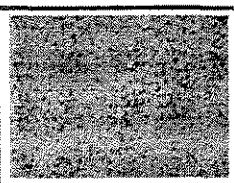
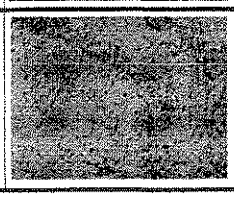
Clone	Cortex	Hippocampus (CA1)	Clone	Cortex	Hippocampus (CA1)
6C10F9 C12A11			6C10E5 E9C12		
6H1A11 C11			6H1G6E 6		
2B6A10 C11			2B6G7A 12		
3A8A12 G7			3A8E12 H8		
7C2(1) F10C10 D3			7C2(2) B9F11D 5		
Control AT100 antibody			***Control*** PG5 antibody		
Control No primary antibody					

FIGURE 1-1

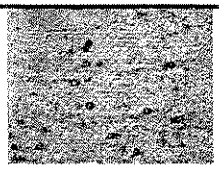
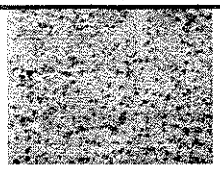
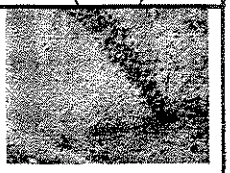
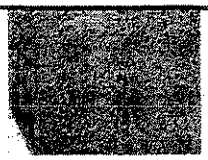
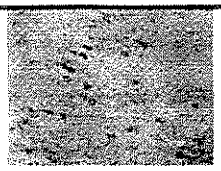
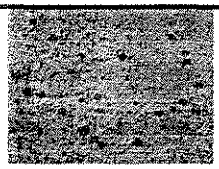
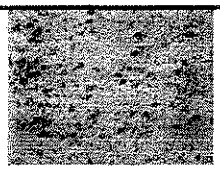
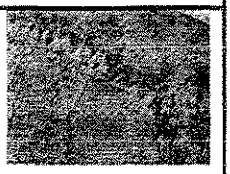
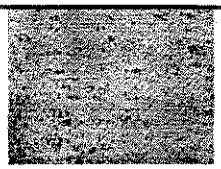
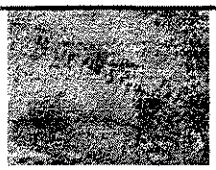
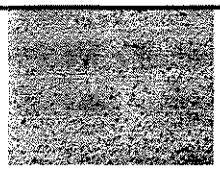
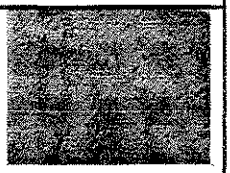
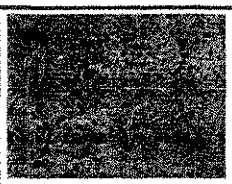
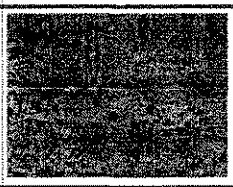
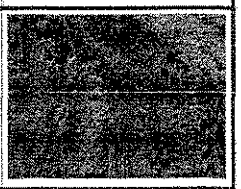
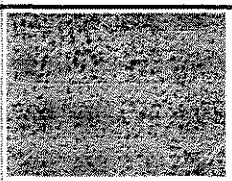
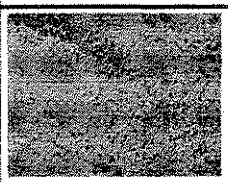
Clone	Cortex	Hippocampus (CA1)	Clone	Cortex	Hippocampus (CA1)
A4-2A1-18			A6-2G5-08		
A4-4A6-48			A6-2G5-30		
A4-2A1-40			A6-2G5-41		
A6-1D2-12			A4-4A6-18		
Control AT100 antibody			***Control*** PG5 antibody		
Control No primary antibody					

FIGURE 1-2

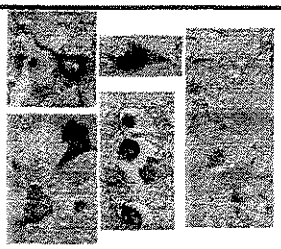
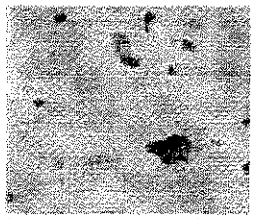
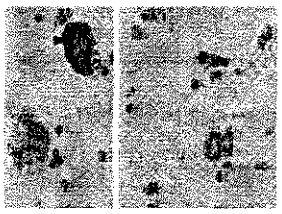
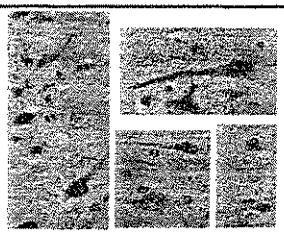
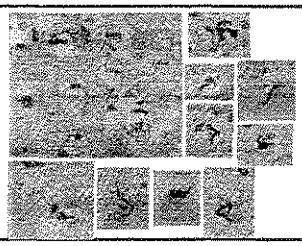
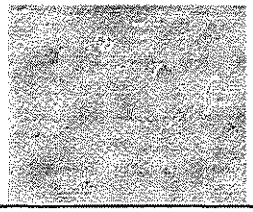
Diagnosis	Region	ACI-3A8-Ab1 TAUPIR staining
AD	Hippocampus	
FAD	Anterior hippocampus	
AGD	Anterior hippocampus	
PSP	Pallidum	
FTDP-17	Frontal cortex	
CBD	Frontal cortex	
Healthy control	Hippocampus	

FIGURE 2

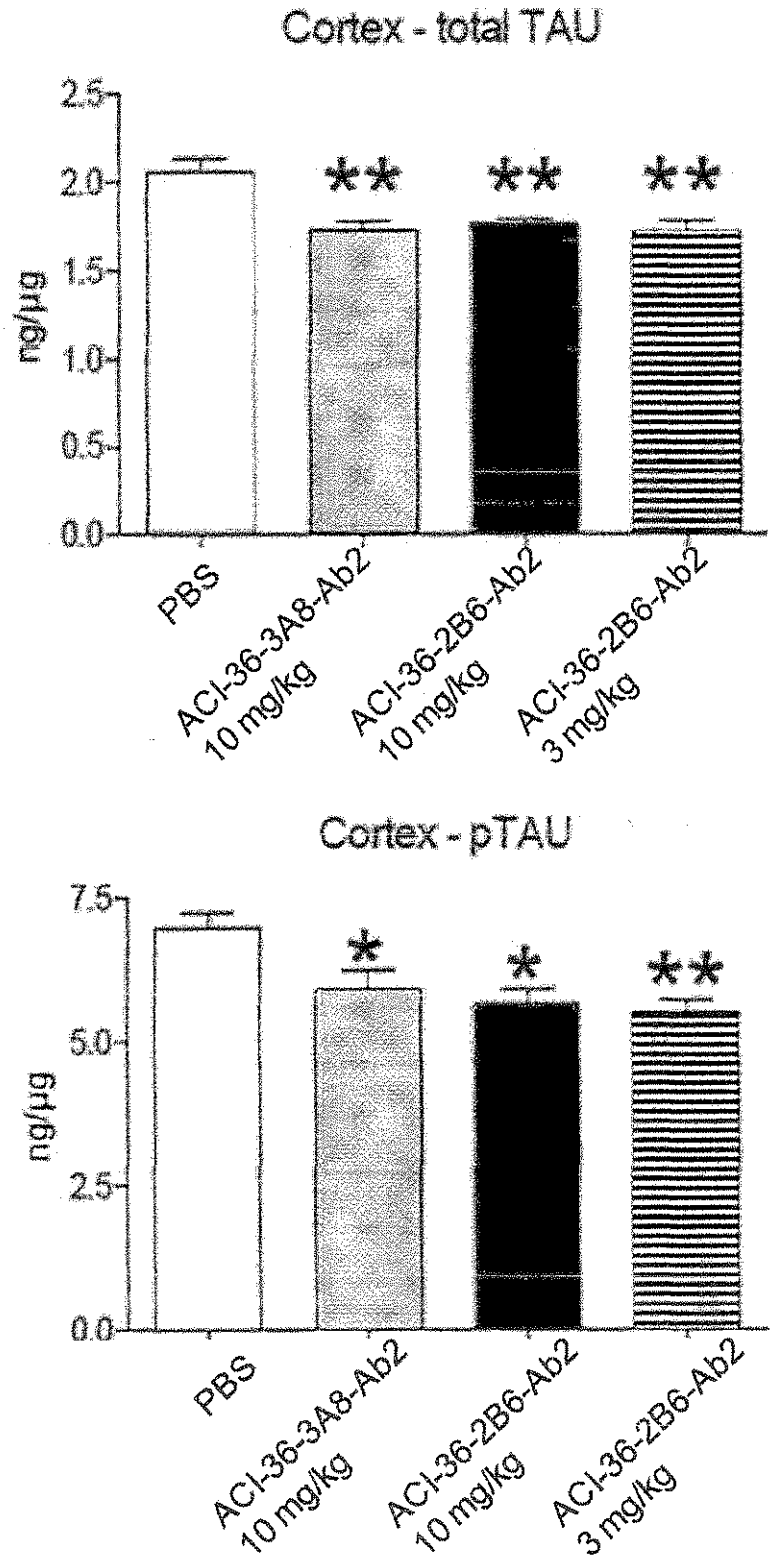

Cortex - total TAU
ng/µg
2.5
2.0
1.5
1.0
0.5
0.0
**
**
**
PBS
ACI-36-3A8-Ab2 10 mg/kg
ACI-36-2B6-Ab2 10 mg/Kg
ACI-36-2B6-Ab2 3 mg/kg
Cortex - pTAU
ng/µg
7.5
5.0
2.5
0.0
*
*
**
PBS
ACI-36-3A8-Ab2 10 mg/kg
ACI-36-2B6-Ab2 10 mg/kg
ACI-36-2B6-Ab2 3 mg/kg

FIGURE 3

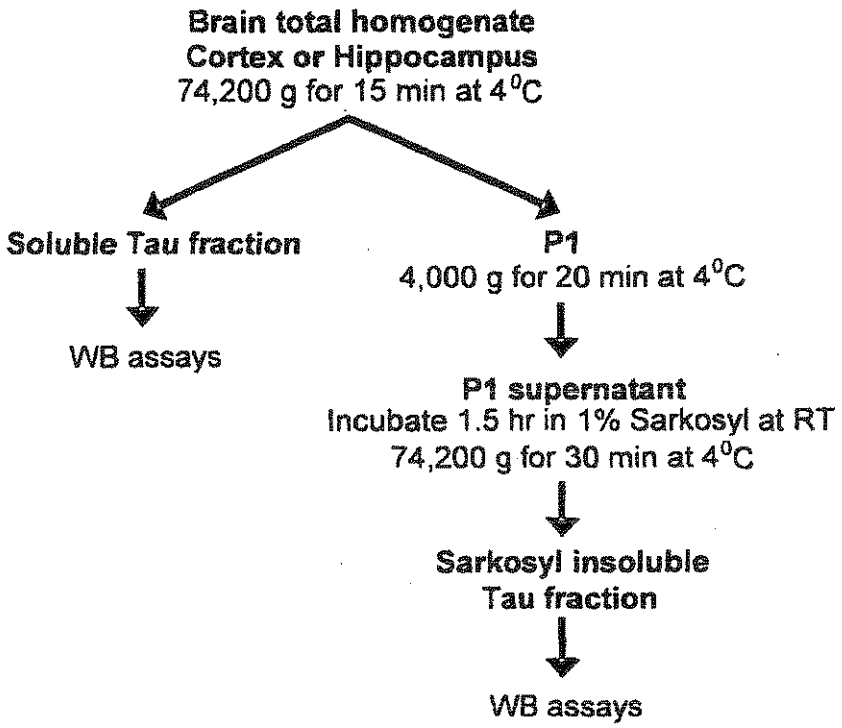


FIGURE 4-1

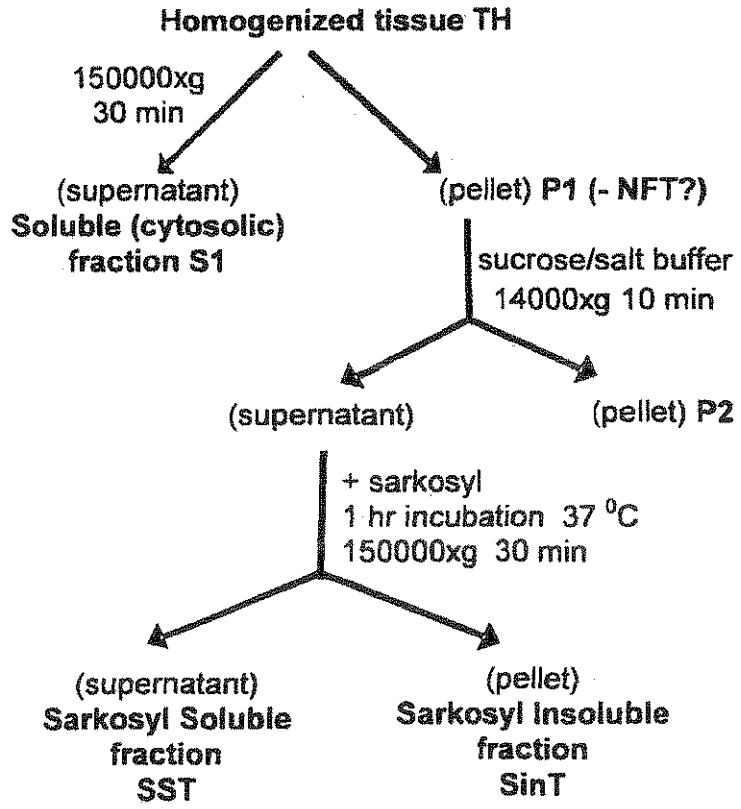


FIGURE 4-2

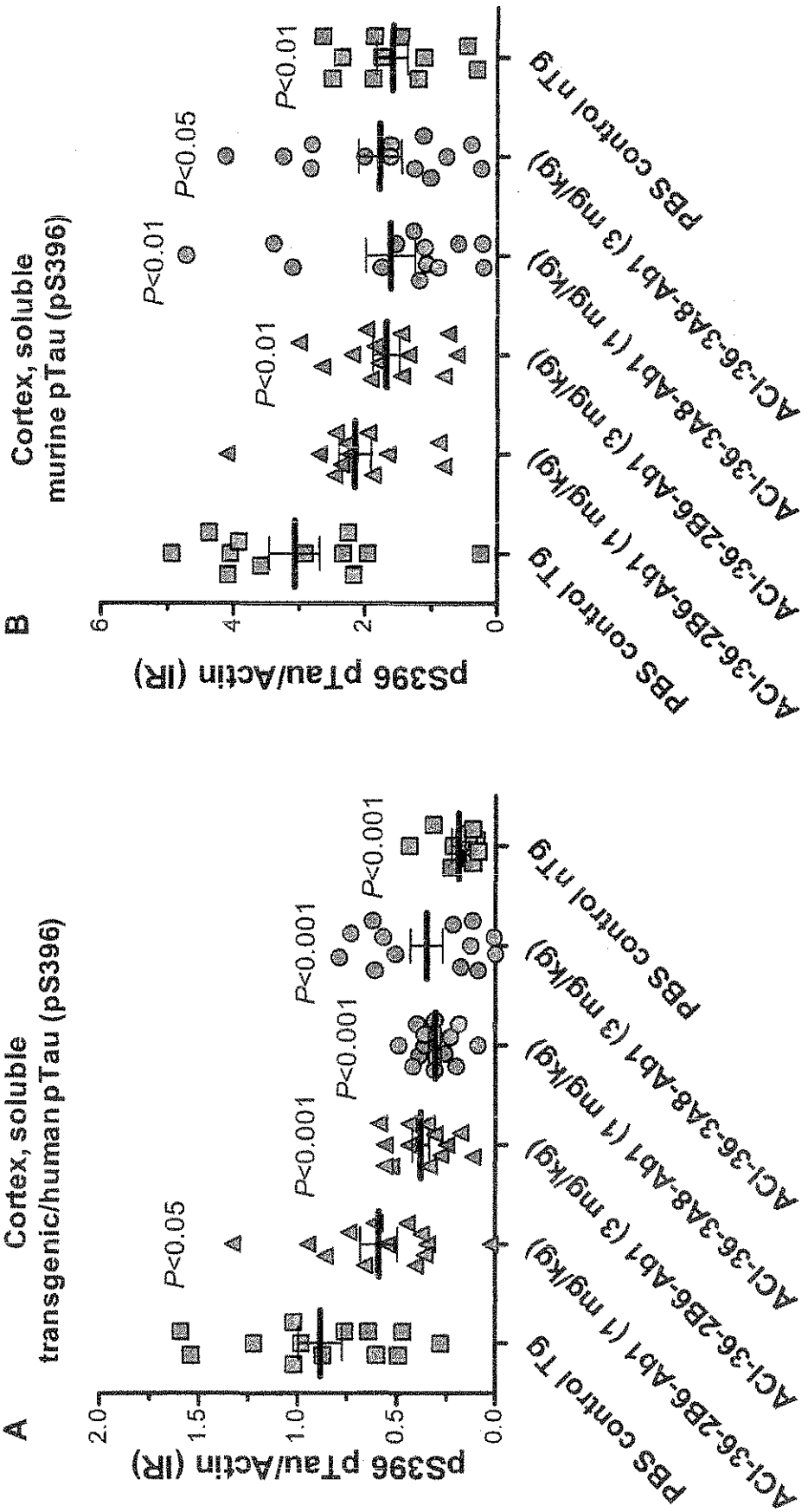
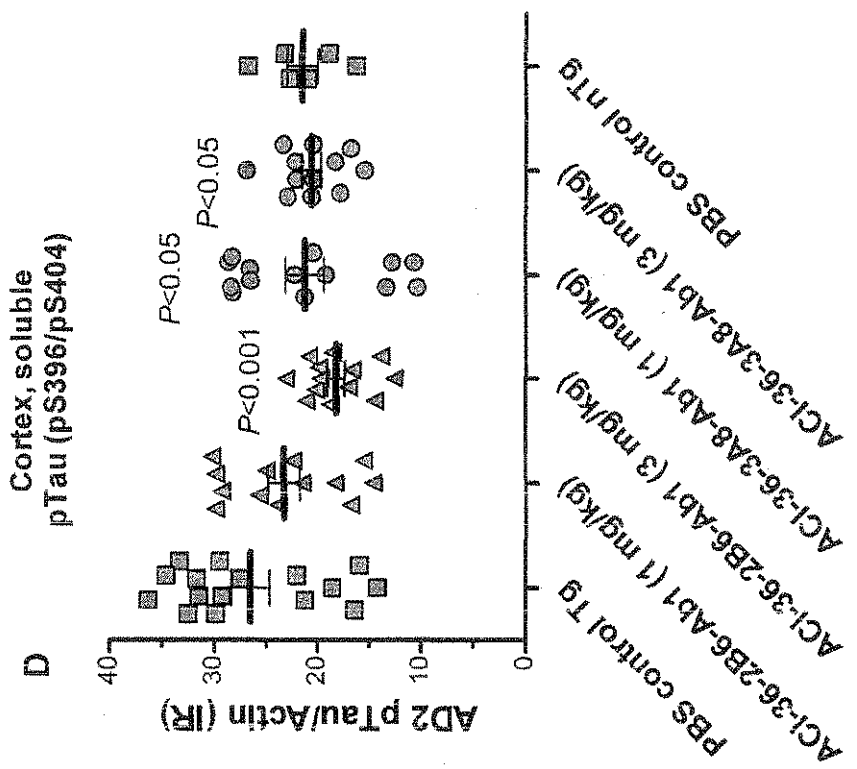

A
Cortex, soluble
transgenic/human pTau (pS396)
pS396 pTau/Actin (IR)
2.0
1.5
1.0
0.5
0.0
P<0.05
P<0.001
P<0.001
P<0.001
P<0.001
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTG
B
Cortex, soluble
murine pTau (pS396)
pS396 pTau/Actin (IR)
6
4
2
0
P<0.01
P<0.05
P<0.01
P<0.01
P<0.01
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTG

FIGURE 5-1


D
Cortex, soluble
pTau (pS396/pS404)
AD2 pTau/Actin (IR)
P<0.05
P<0.001
P<0.05
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTG

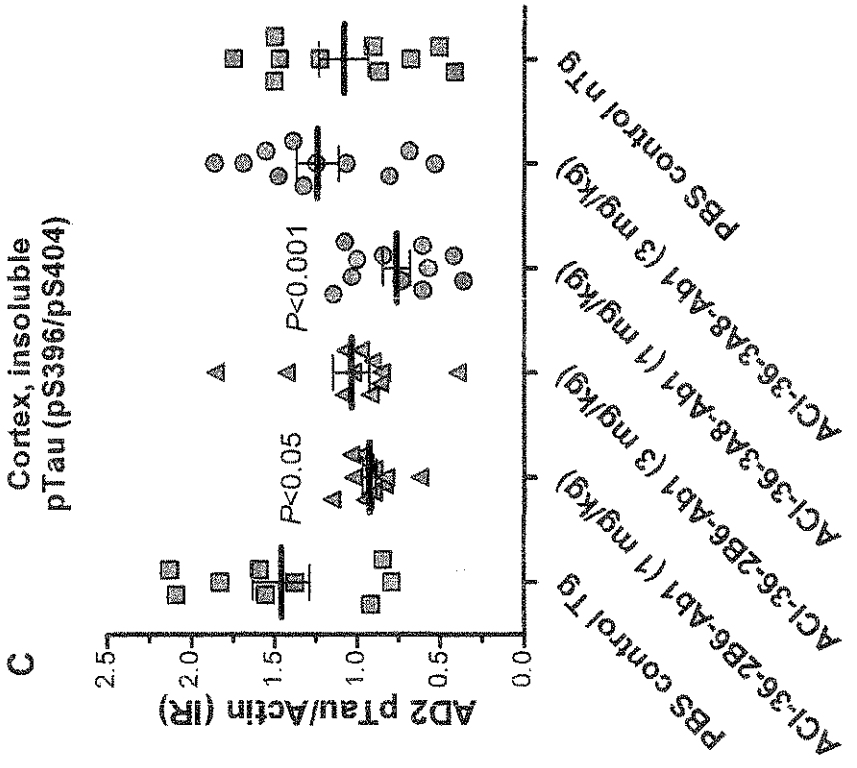

C
Cortex, insoluble
pTau (pS396/pS404)
AD2 pTau/Actin (IR)
P<0.05
P<0.001
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTG

FIGURE 5-2

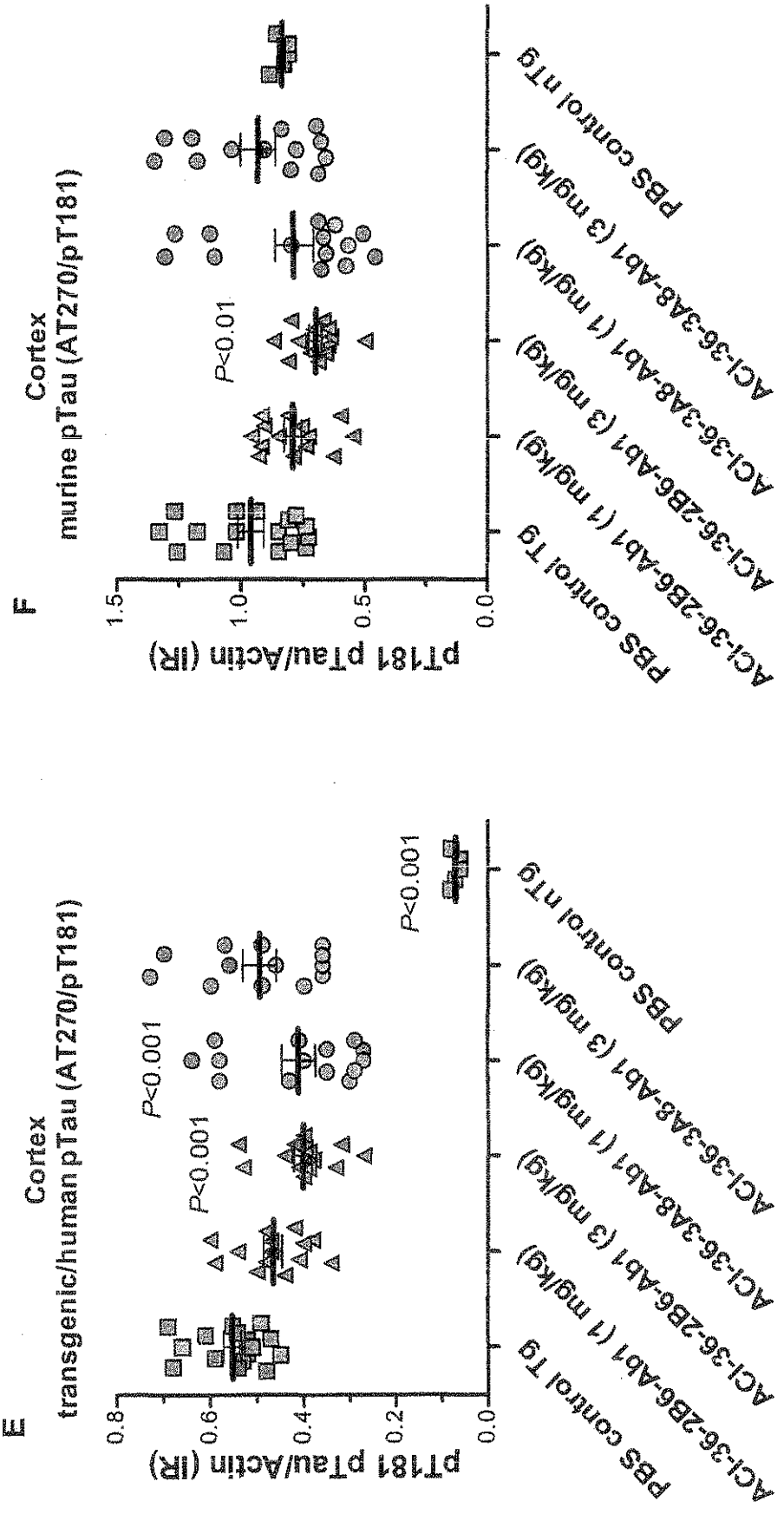

E
Cortex
transgenic/human pTau (AT270/pT181)
pT181 pTau/Actin (IR)
0.0
0.2
0.4
0.6
0.8
P<0.001
P<0.001
P<0.001
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTg
F
Cortex
murine pTau (AT270/pT181)
pT181 pTau/Actin (IR)
0.0
0.5
1.0
1.5
P<0.01
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTg

FIGURE 5-3

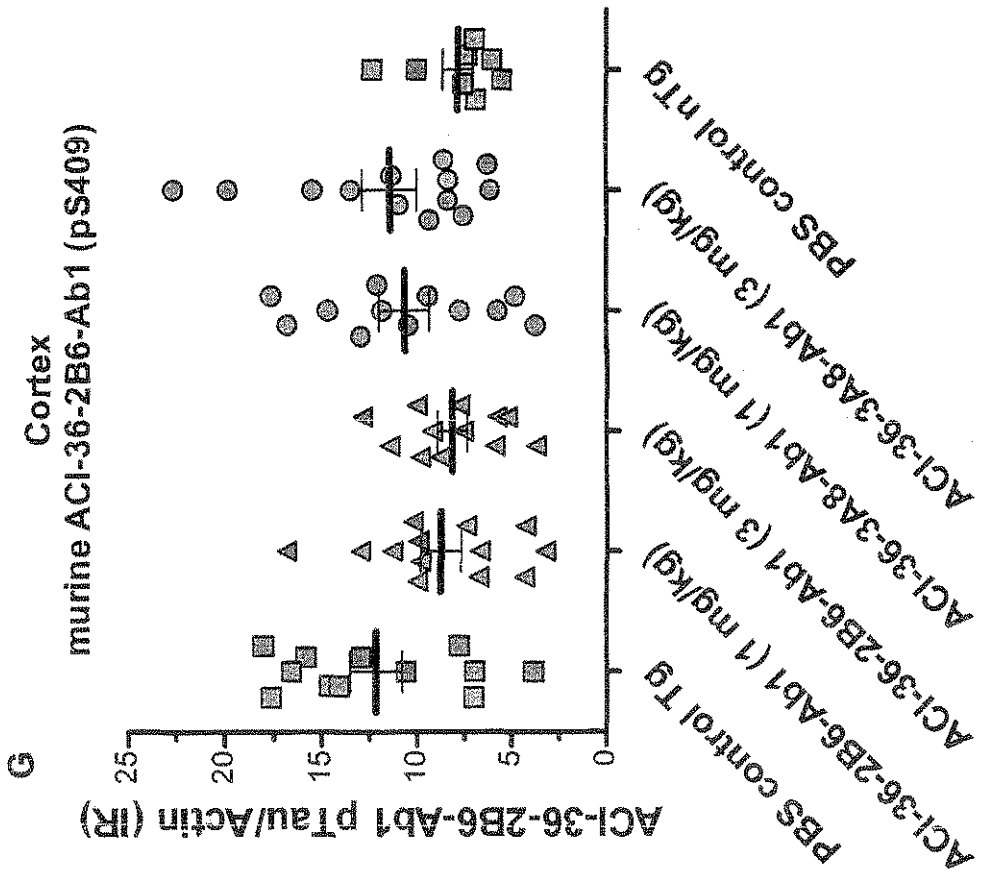

G
Cortex
murine ACI-36-2B6-Ab1 (pS409)
ACI-36-2B6-Ab1 pTau/Actin (IR)
25
20
15
10
5
0
PBS control TG
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
PBS control nTG

FIGURE 5-4

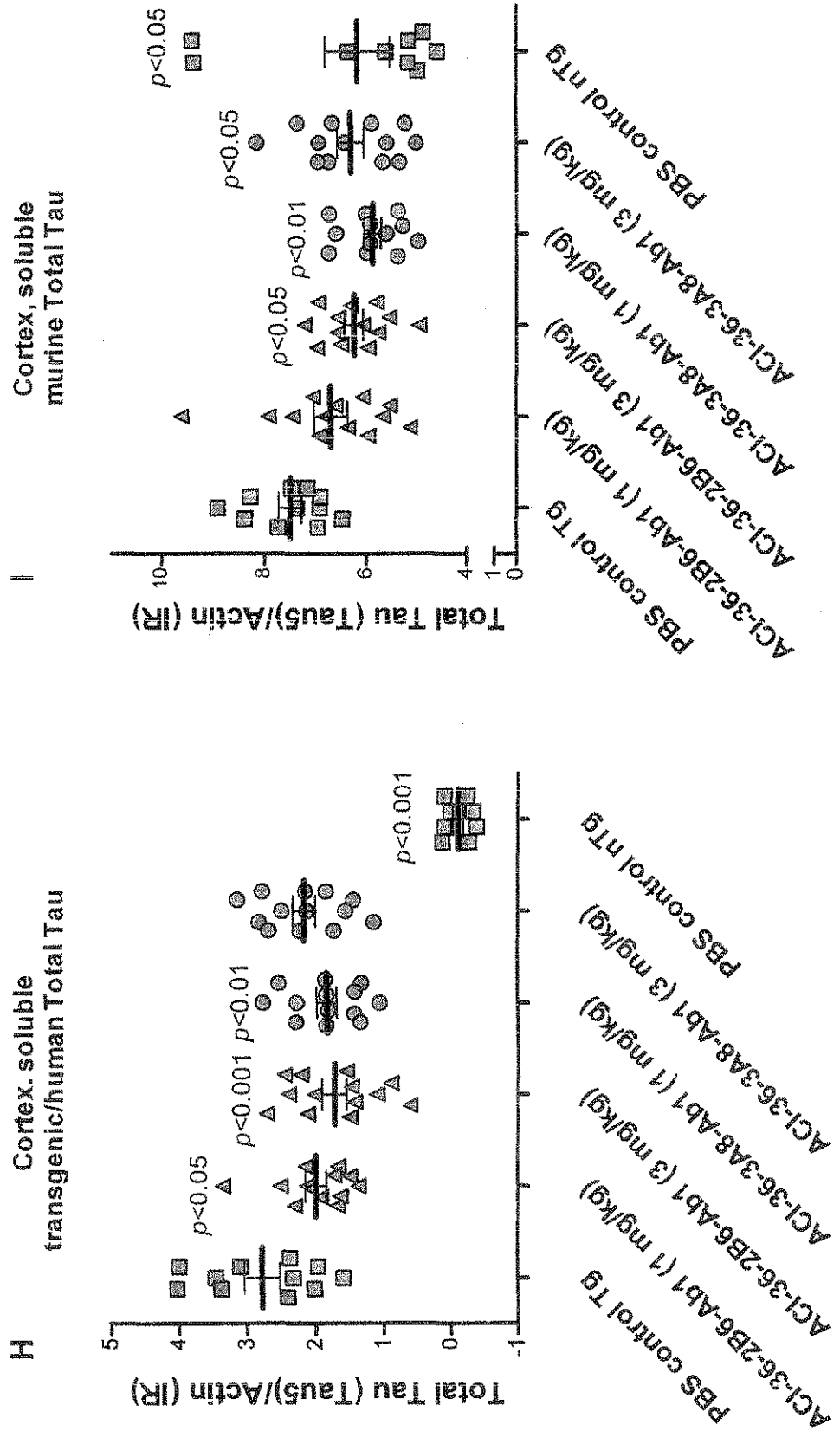

I
Cortex, soluble
murine Total Tau
Total Tau (Tau5)/Actin (IR)
p<0.05
p<0.05
p<0.01
p<0.05
p<0.05
PBS control nTg
ACI-36-3A8-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-2B6-Ab1 (1 mg/kg)
PBS control Tg
H
Cortex. soluble
transgenic/human Total Tau
Total Tau (Tau5)/Actin (IR)
p<0.05
p<0.001
p<0.01
p<0.001
PBS control nTg
ACI-36-3A8-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-2B6-Ab1 (1 mg/kg)
PBS control Tg

FIGURE 5-5

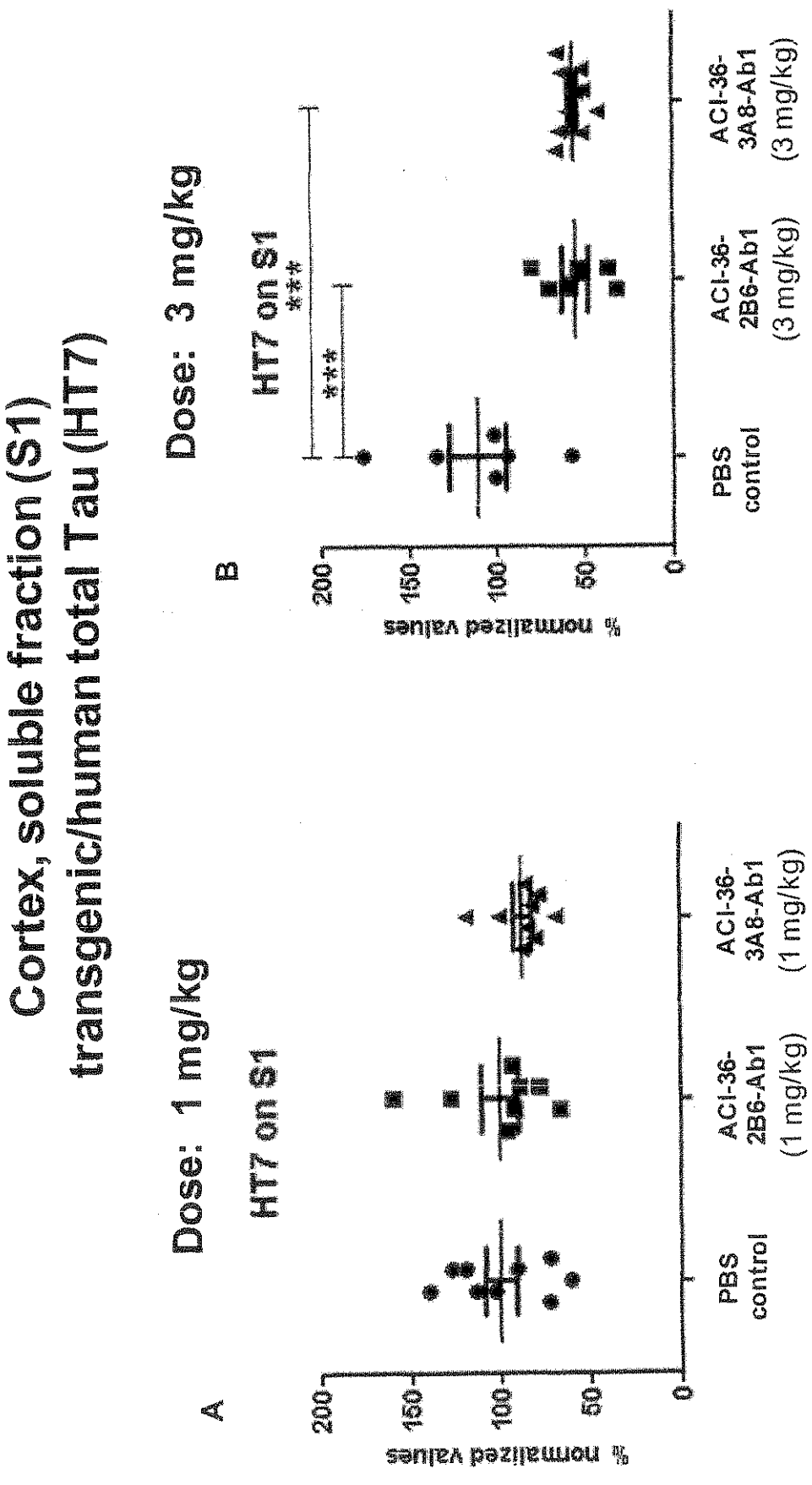

Cortex, soluble fraction (S1)
transgenic/human total Tau (HT7)
A
Dose: 1 mg/kg
HT7 on S1
% normalized values
200
150
100
50
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
B
Dose: 3 mg/kg
HT7 on S1
***
***
% normalized values
200
150
100
50
0
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-1

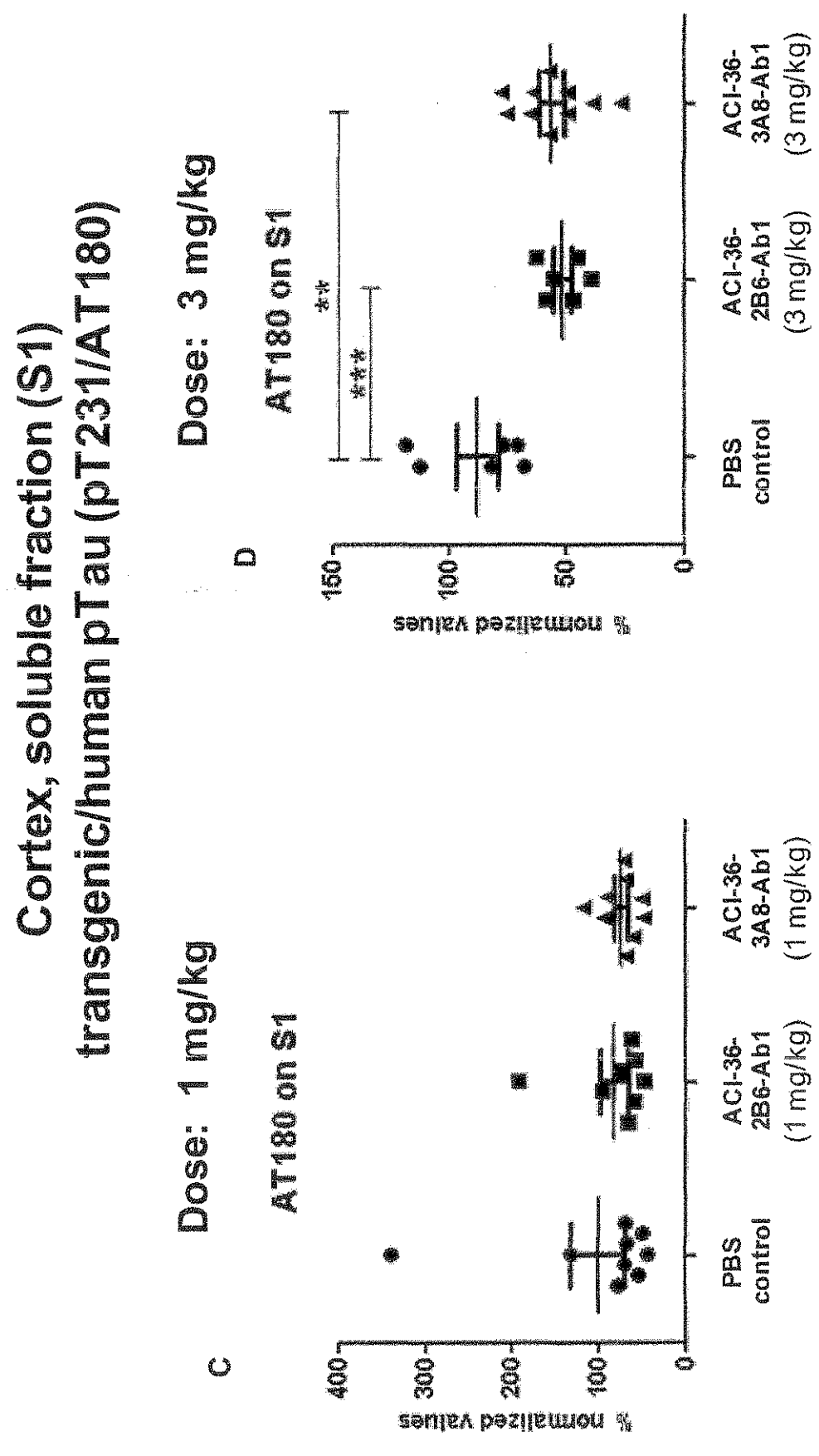

Cortex, soluble fraction (S1)
transgenic/human pTau (pT231/AT180)
C
Dose: 1 mg/kg
AT180 on S1
% normalized values
400
300
200
100
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
D
Dose: 3 mg/kg
AT180 on S1
% normalized values
150
100
50
0
**
***
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-2

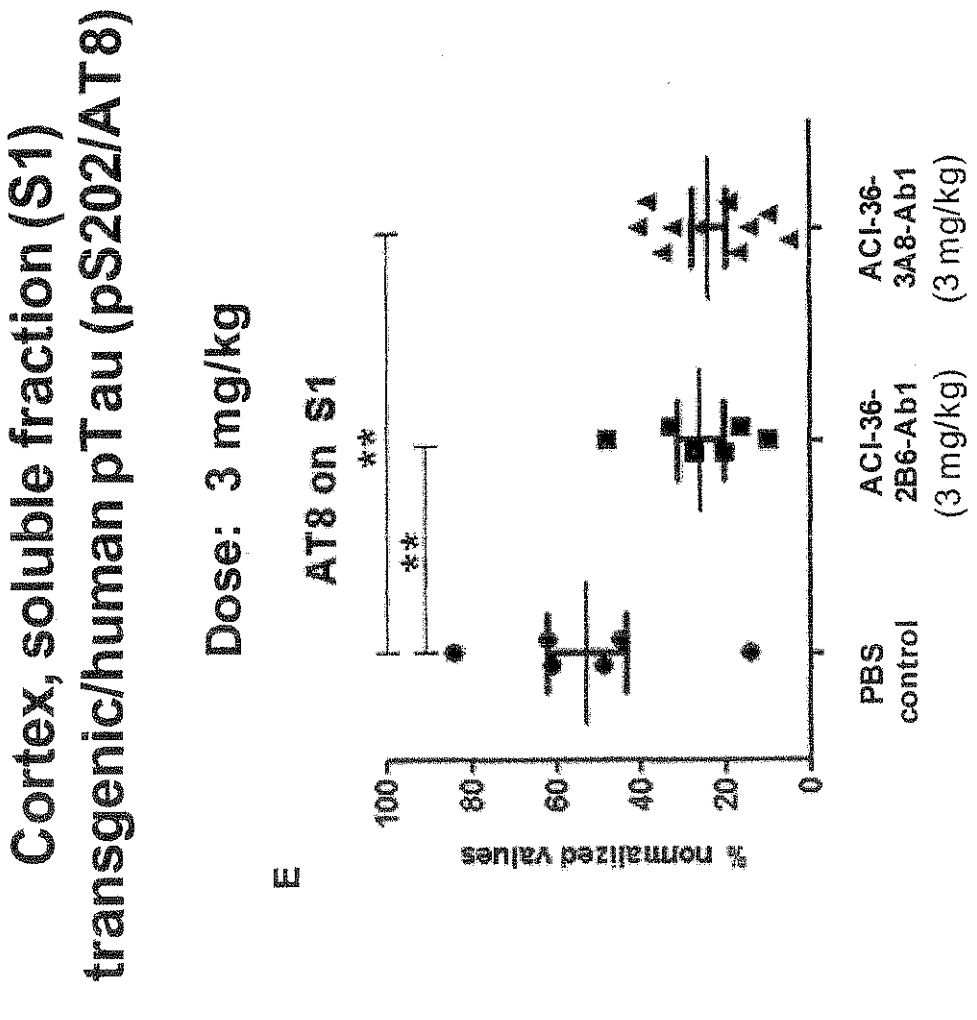

Cortex, soluble fraction (S1)
transgenic/human pTau (pS202/AT8)
Dose: 3 mg/kg
E
AT8 on S1
**
**
% normalized values
100
80
60
40
20
0
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-3

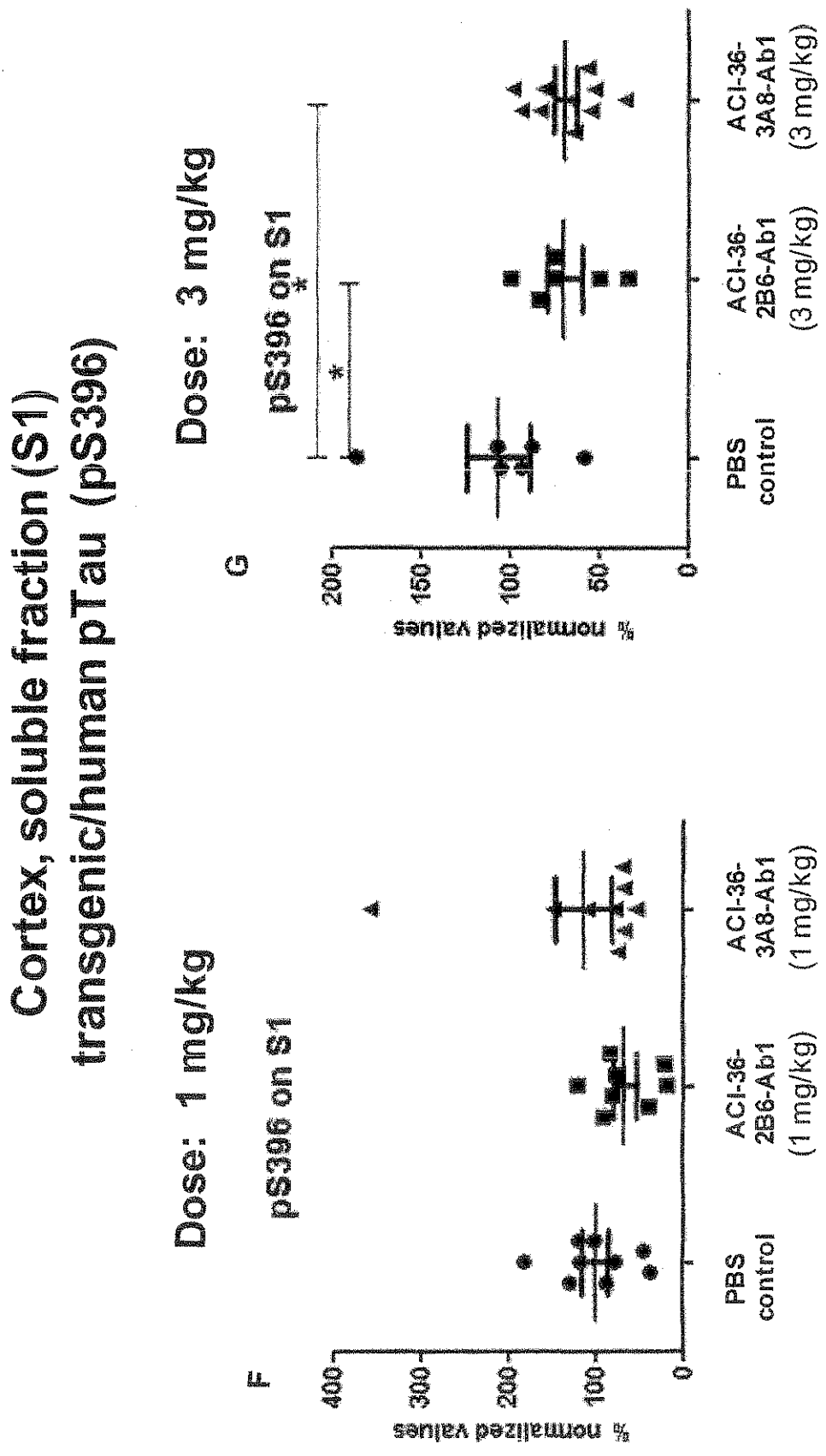

Cortex, soluble fraction (S1)
transgenic/human pTau (pS396)
F
Dose: 1 mg/kg
pS396 on S1
% normalized values
400
300
200
100
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
G
Dose: 3 mg/kg
pS396 on S1
*
*
% normalized values
200
150
100
50
0
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-4

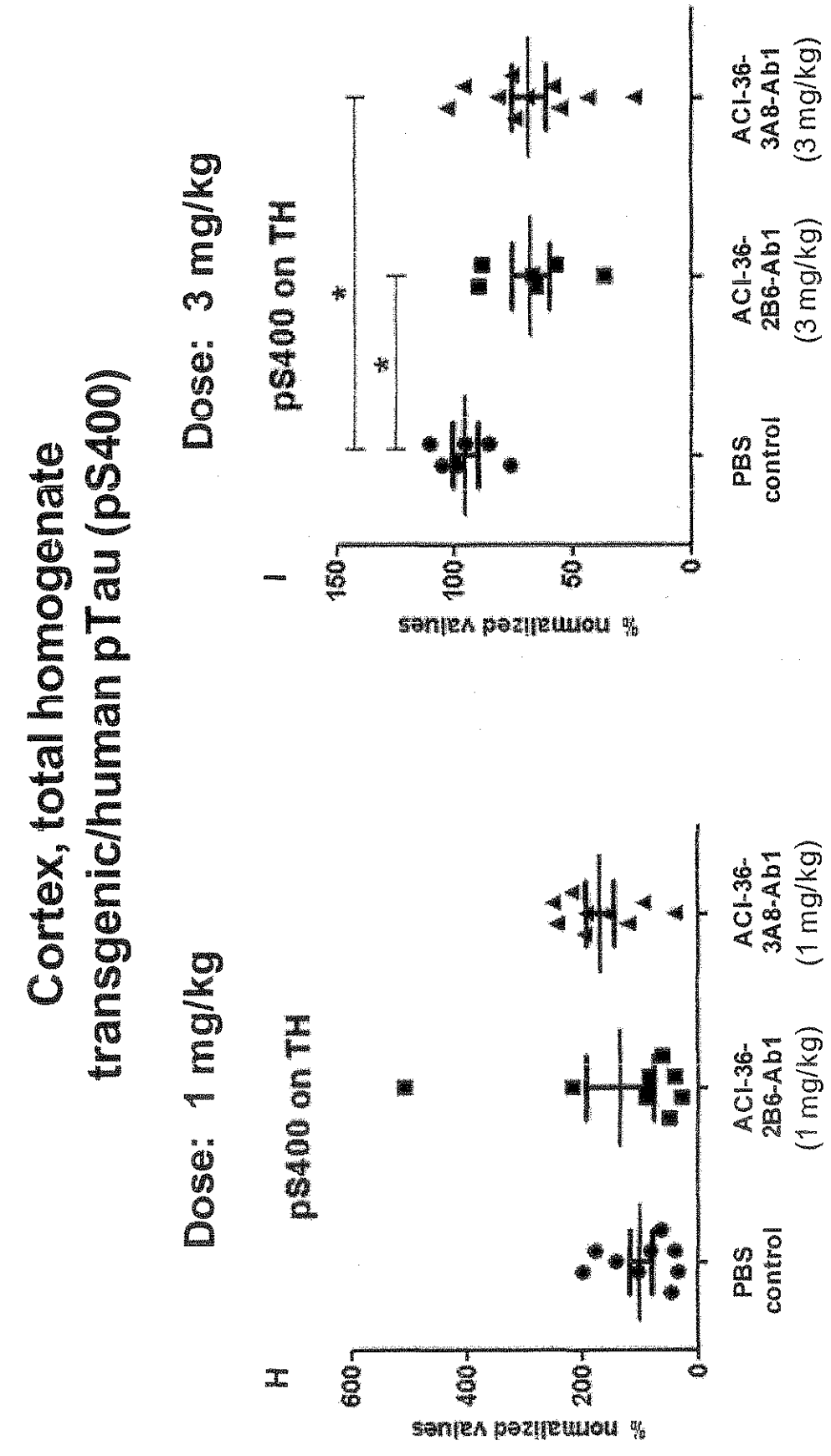

Cortex, total homogenate
transgenic/human pTau (pS400)
Dose: 1 mg/kg
Dose: 3 mg/kg
H
I
pS400 on TH
% normalized values
600
400
200
0
150
100
50
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)
*
*

FIGURE 6-5

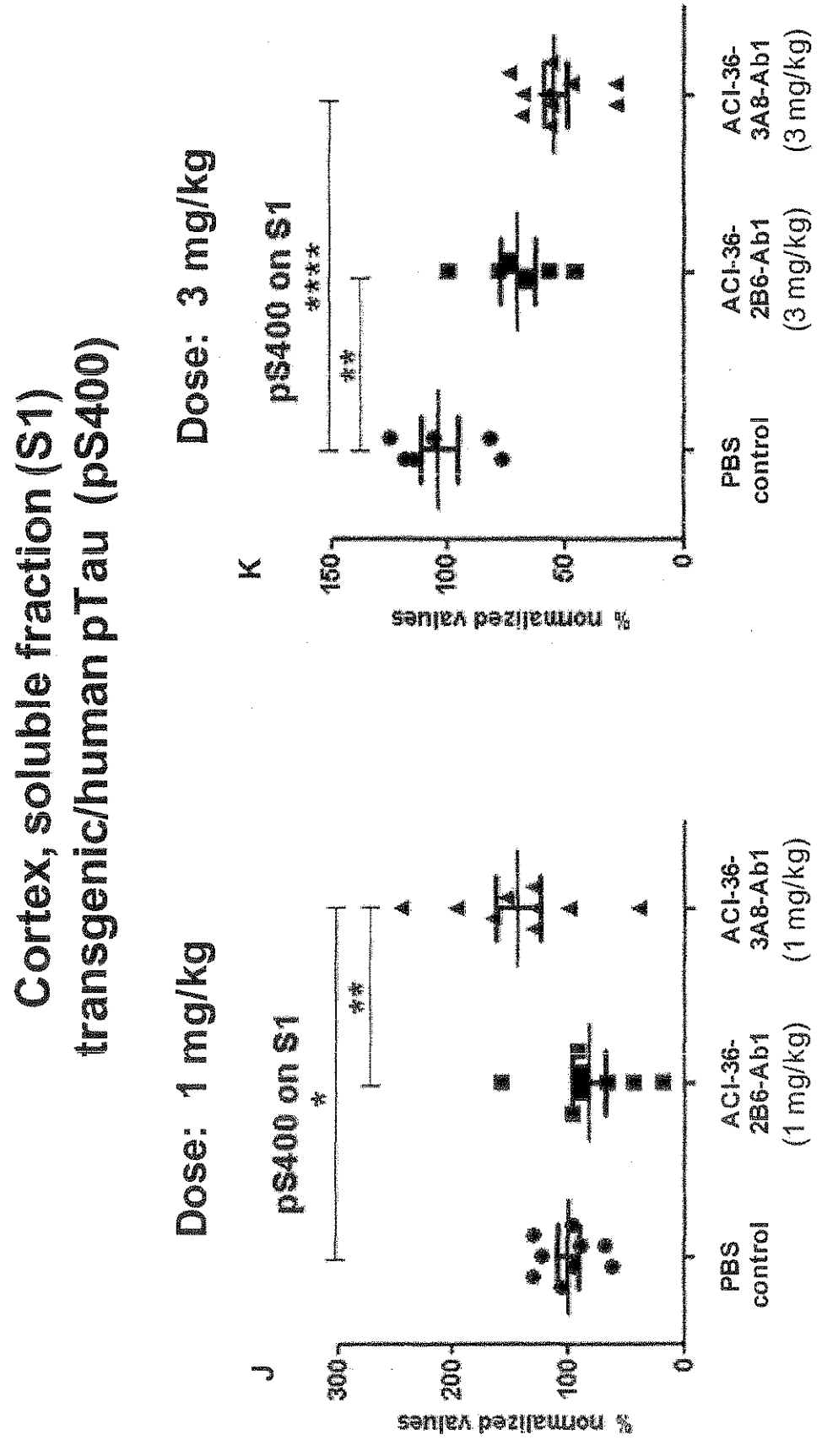

Cortex, soluble fraction (S1)
transgenic/human pTau (pS400)
J
Dose: 1 mg/kg
pS400 on S1
*
**
% normalized values
300
200
100
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
K
Dose: 3 mg/kg
pS400 on S1
****
**
% normalized values
150
100
60
0
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-6

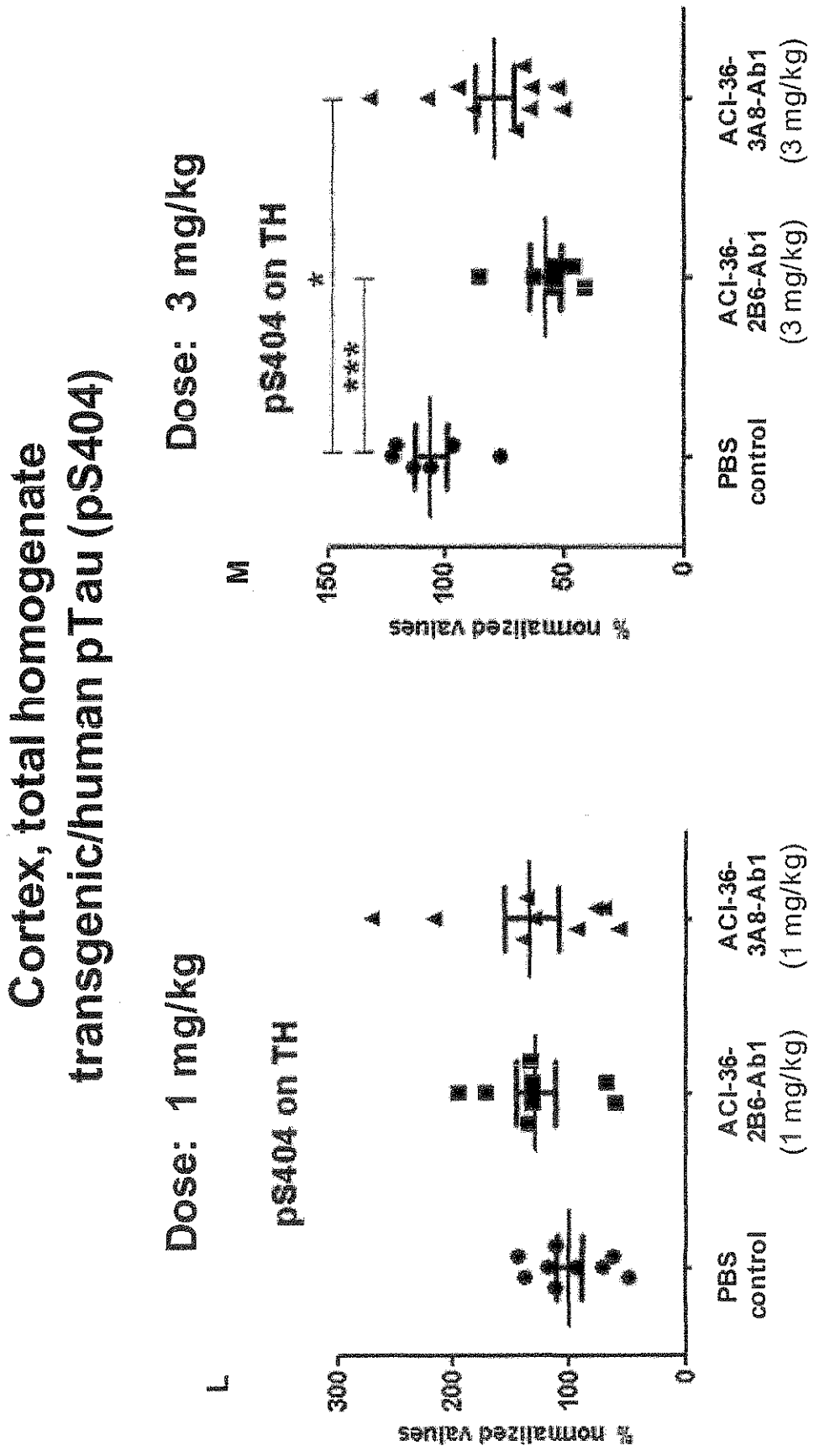

Cortex, total homogenate
transgenic/human pTau (pS404)
L
Dose: 1 mg/kg
pS404 on TH
% normalized values
300
200
100
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
M
Dose: 3 mg/kg
pS404 on TH
% normalized values
150
100
50
0
***
*
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-7

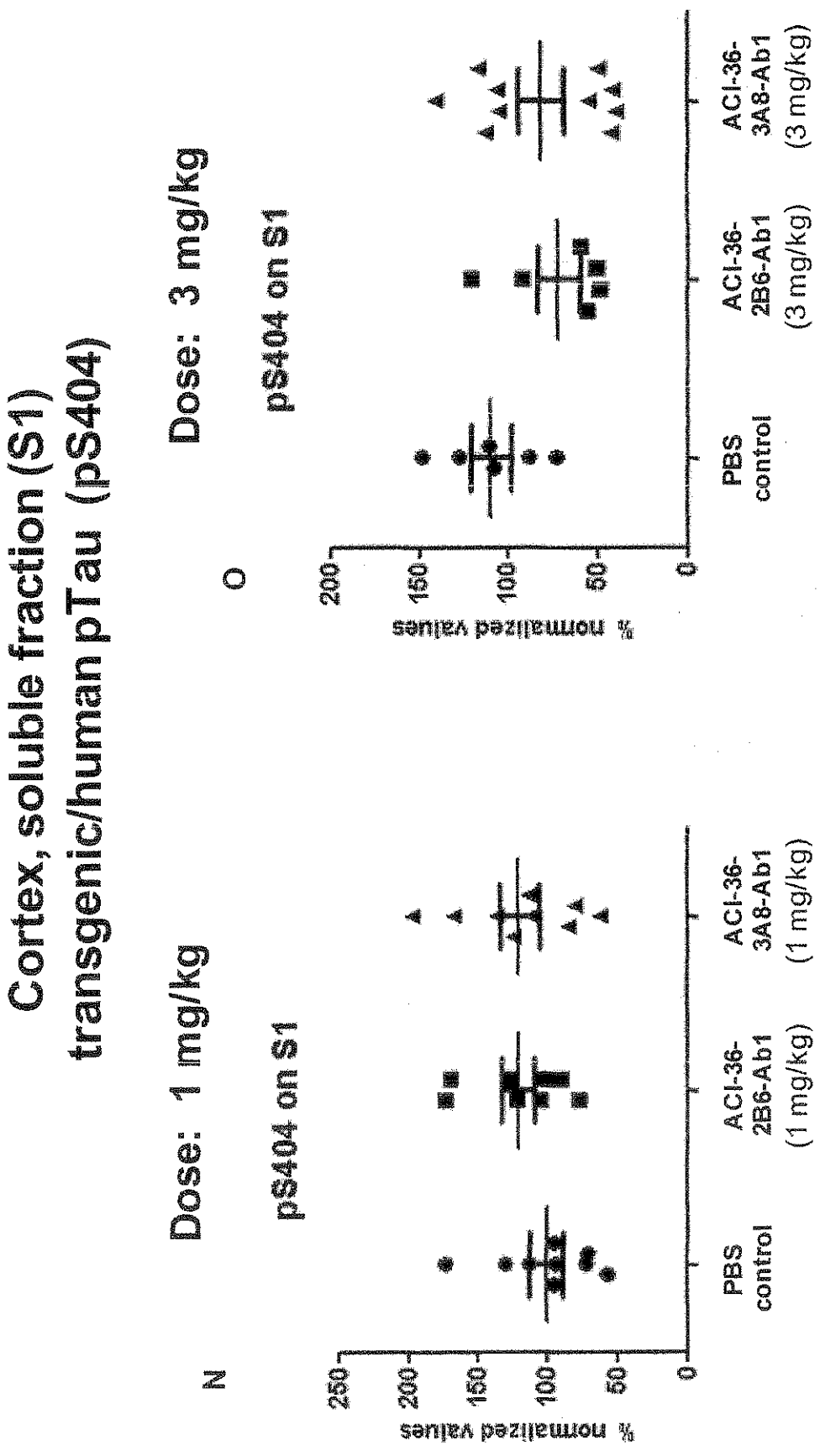

Cortex, soluble fraction (S1)
transgenic/human pTau (pS404)
N
Dose: 1 mg/kg
pS404 on S1
% normalized values
250
200
150
100
50
0
PBS control
ACI-36-2B6-Ab1 (1 mg/kg)
ACI-36-3A8-Ab1 (1 mg/kg)
O
Dose: 3 mg/kg
pS404 on S1
% normalized values
200
150
100
50
0
PBS control
ACI-36-2B6-Ab1 (3 mg/kg)
ACI-36-3A8-Ab1 (3 mg/kg)

FIGURE 6-8

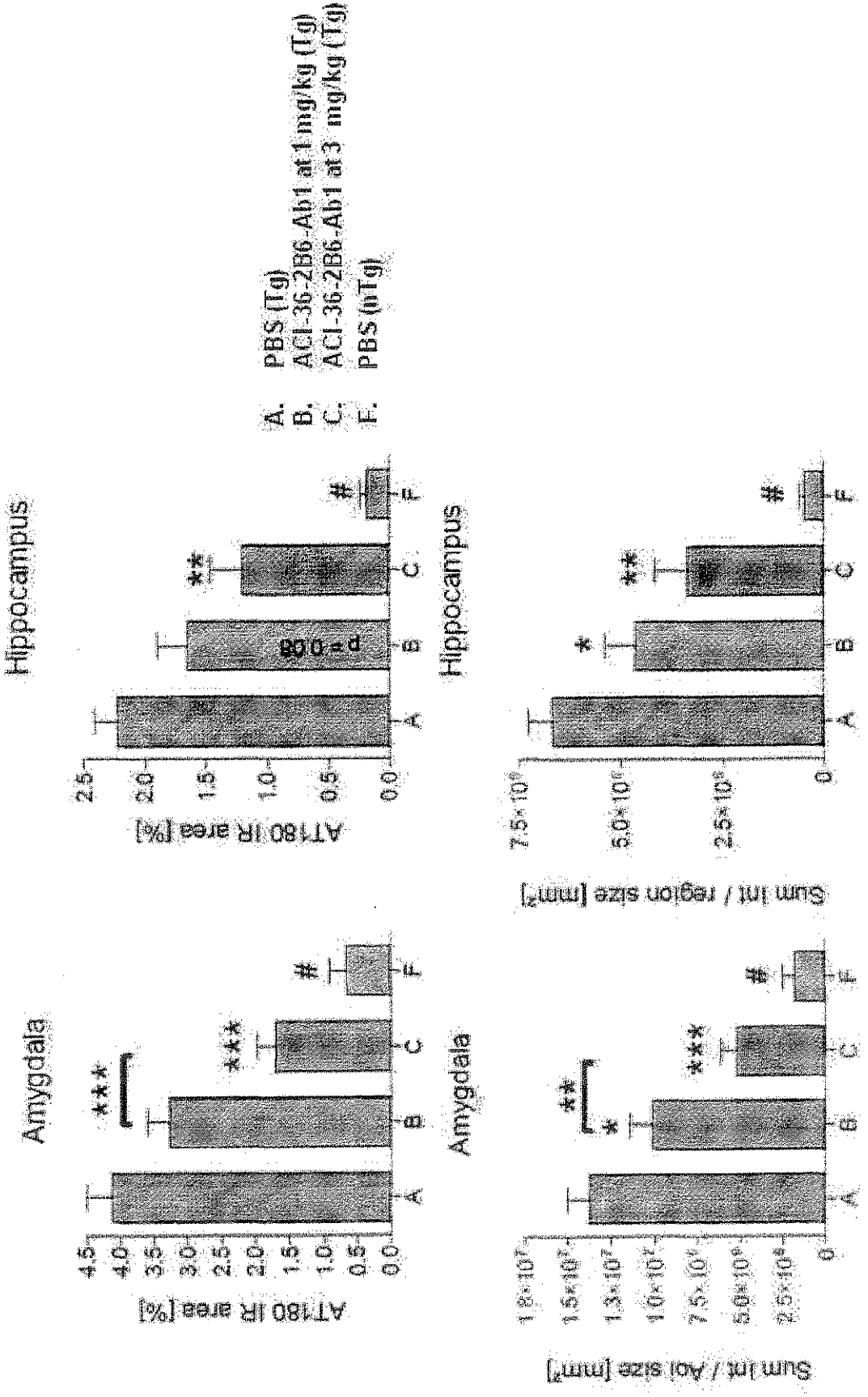

A. PBS (Tg)
B. ACI-36-2B6-Ab1 at 1 mg/kg (Tg)
C. ACI-36-2B6-Ab1 at 3 mg/kg (Tg)
F. PBS (nTg)
Hippocampus
AT180 IR area [%]
p = 0.08
Hippocampus
Sum Int / region size [mm²]
Amygdala
AT180 IR area [%]
Amygdala
Sum Int / Aoi size [mm²]

FIGURE 7

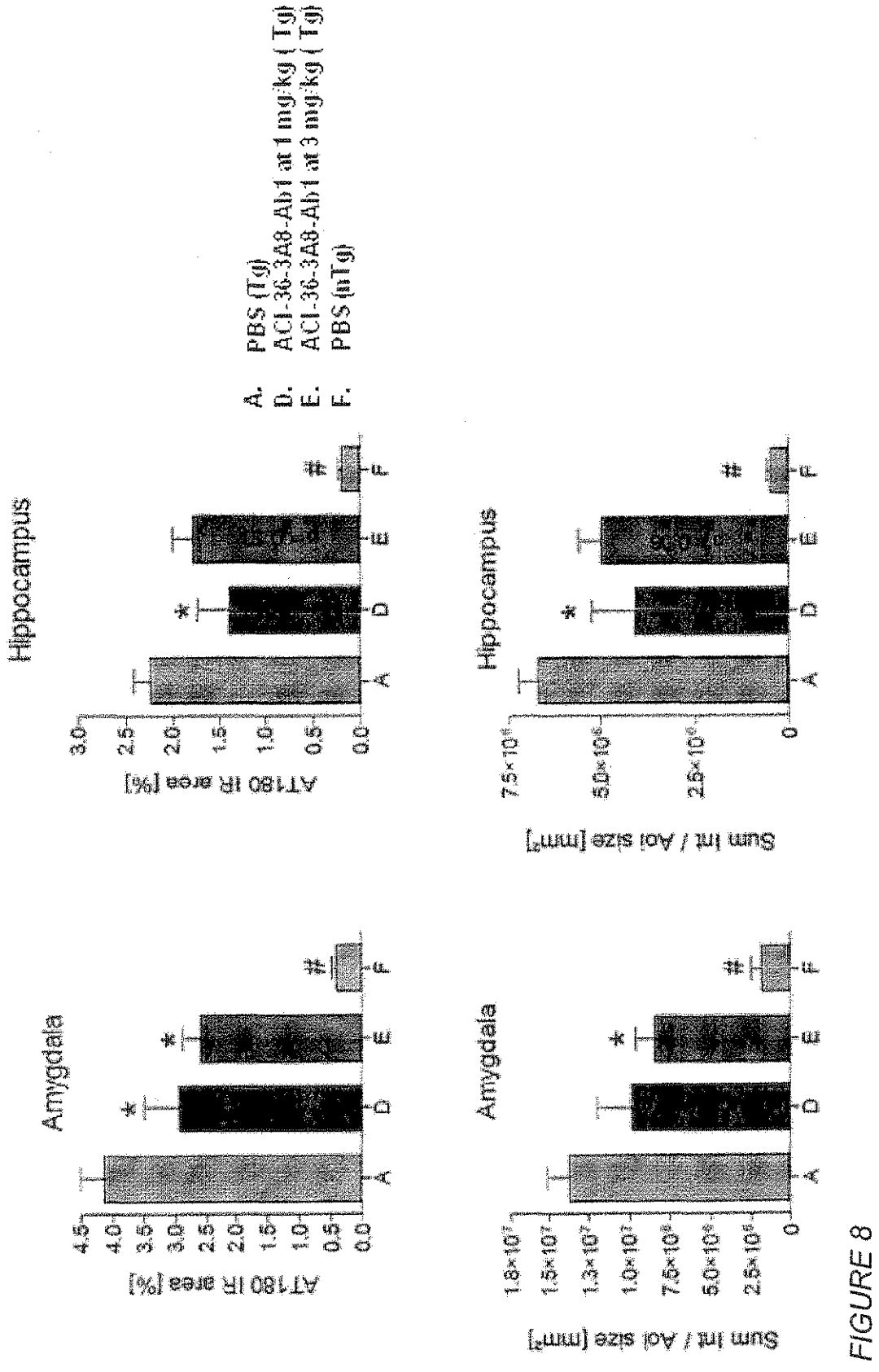
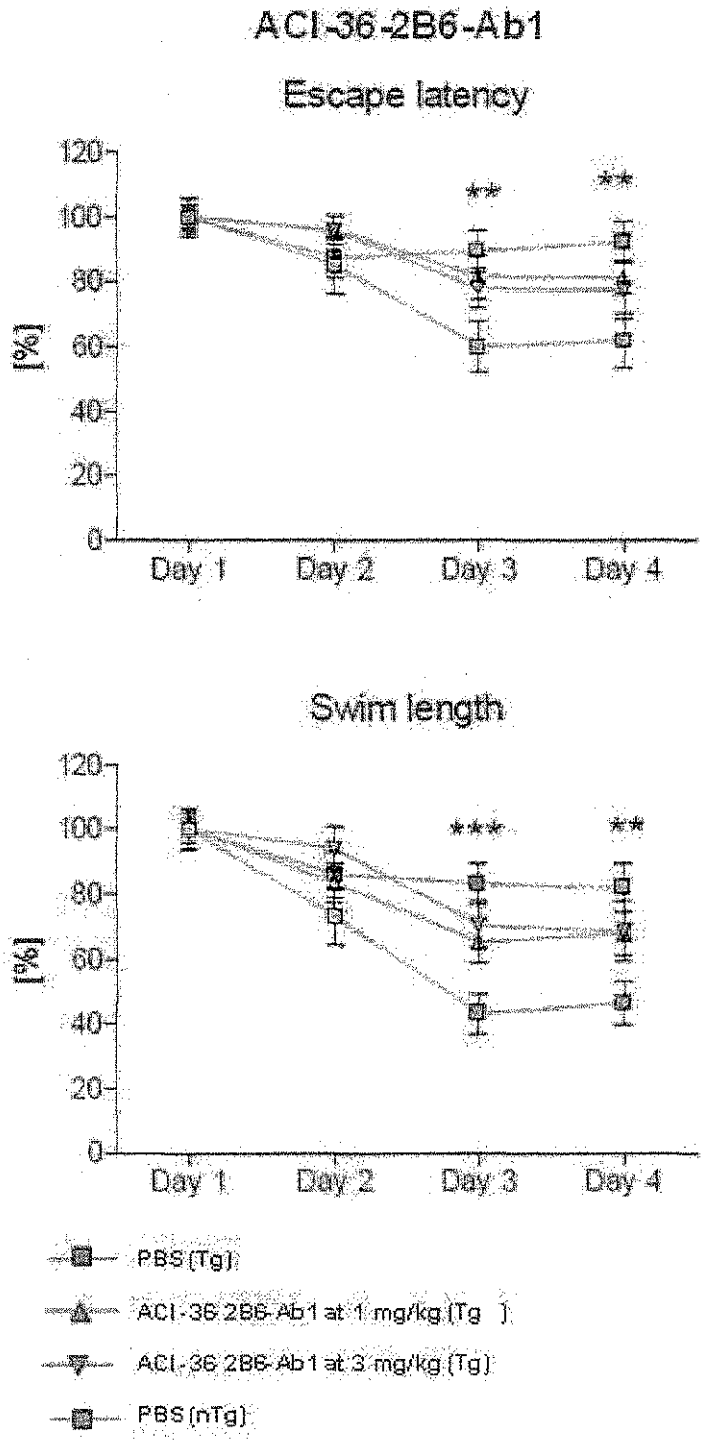
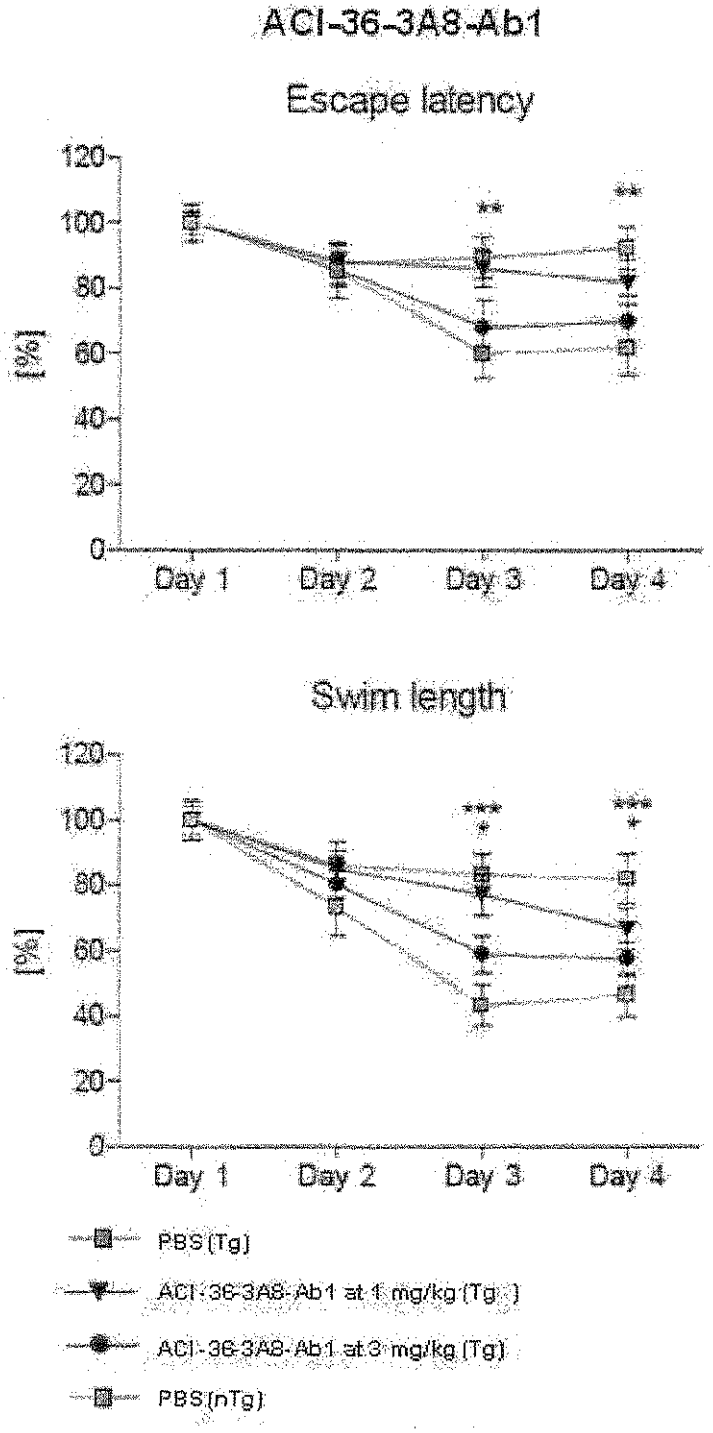

A. PBS (Tg)
D. ACI-36-3A8-Ab1 at 1 mg/kg (Tg)
E. ACI-36-3A8-Ab1 at 3 mg/kg (Tg)
F. PBS (nTg)
Hippocampus
AT180 IR area [%]
Sum int / Aol size [mm²]
Amygdala
AT180 IR area [%]
Sum int / Aol size [mm²]

FIGURE 8



	Escape latency			Swim length		
	Day 2	Day 3	Day 4	Day 2	Day 3	Day 4
A vs B	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05
A vs C	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05
A vs F	P > 0.05	P<0.01	P<0.01	P > 0.05	P<0.001	P<0.01

FIGURE 9



	Escape latency			Swim length		
	Day 2	Day 3	Day 4	Day 2	Day 3	Day 4
A vs D	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P > 0.05
A vs E	P > 0.05	P > 0.05	P > 0.05	P > 0.05	P < 0.05	P < 0.05
A vs F	P > 0.05	P<0.01	P<0.01	P > 0.05	P<0.001	P<0.001

FIGURE 10