Figure 1

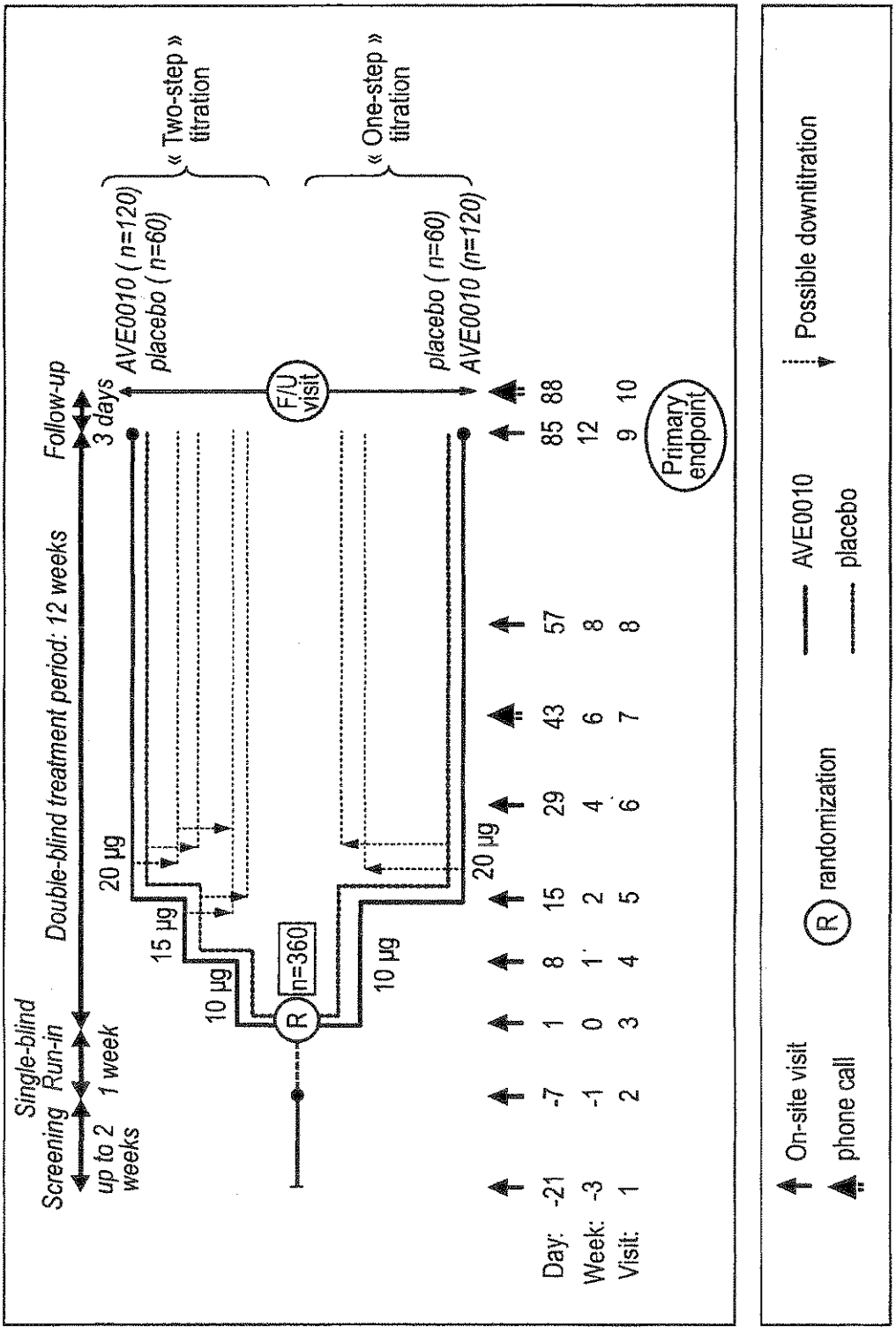

Single-blind
Screening Run-in
Double-blind treatment period: 12 weeks
Follow-up
up to 2 weeks
1 week
3 days
20 µg
15 µg
10 µg
R
n=360
10 µg
20 µg
F/U visit
AVE0010 (n=120)
placebo (n=60)
« Two-step » titration
placebo (n=60)
AVE0010 (n=120)
« One-step » titration
Primary endpoint
Day: -21 -7 1 8 15 29 43 57 85 88
Week: -3 -1 0 1 2 4 6 8 12
Visit: 1 2 3 4 5 6 7 8 9 10
On-site visit
phone call
R randomization
AVE0010
placebo
Possible downtitration

Figure 2

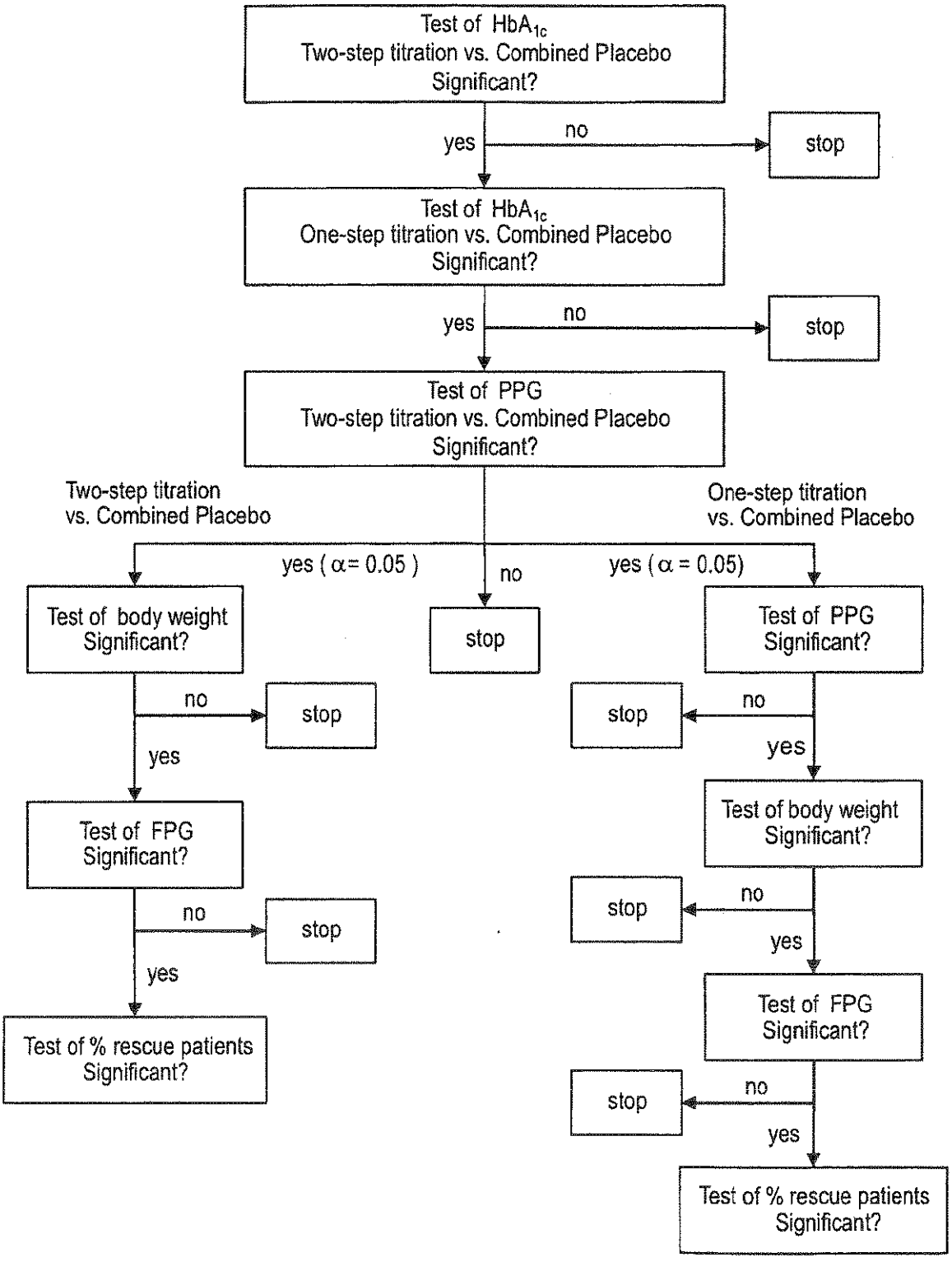

Test of HbA_{1c}
Two-step titration vs. Combined Placebo
Significant?
yes
no
stop
Test of HbA_{1c}
One-step titration vs. Combined Placebo
Significant?
yes
no
stop
Test of PPG
Two-step titration vs. Combined Placebo
Significant?
Two-step titration
vs. Combined Placebo
One-step titration
vs. Combined Placebo
yes (α= 0.05)
no
yes (α = 0.05)
Test of body weight
Significant?
stop
Test of PPG
Significant?
no
stop
yes
stop
no
yes
Test of FPG
Significant?
Test of body weight
Significant?
no
stop
stop
no
yes
yes
Test of FPG
Significant?
Test of % rescue patients
Significant?
stop
no
yes
Test of % rescue patients
Significant?

Figure 3

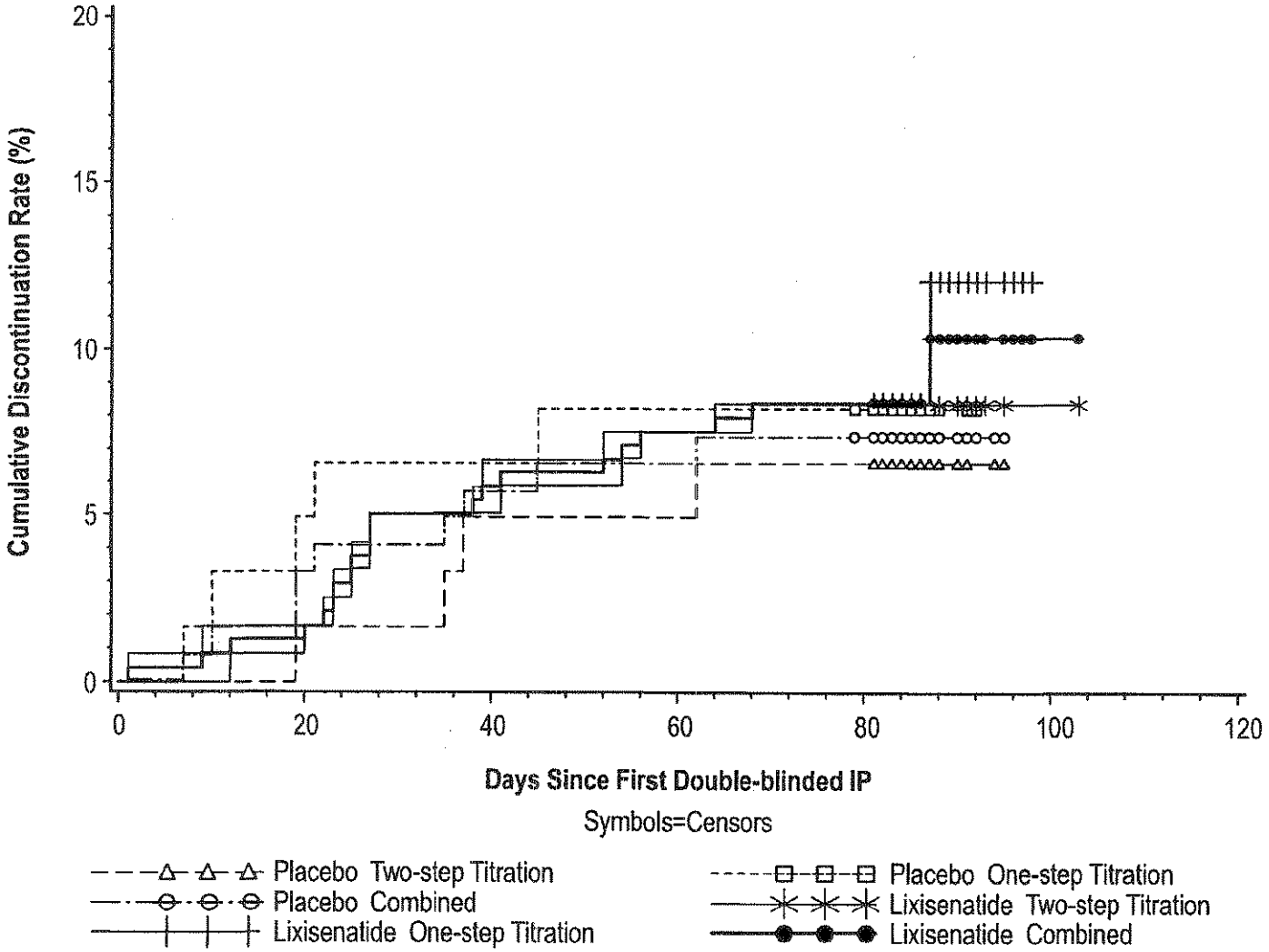

Cumulative Discontinuation Rate (%)
20
15
10
5
0
0
20
40
60
80
100
120
Days Since First Double-blinded IP
Symbols=Censors
Placebo Two-step Titration
Placebo Combined
Lixisenatide One-step Titration
Placebo One-step Titration
Lixisenatide Two-step Titration
Lixisenatide Combined

Figure 4

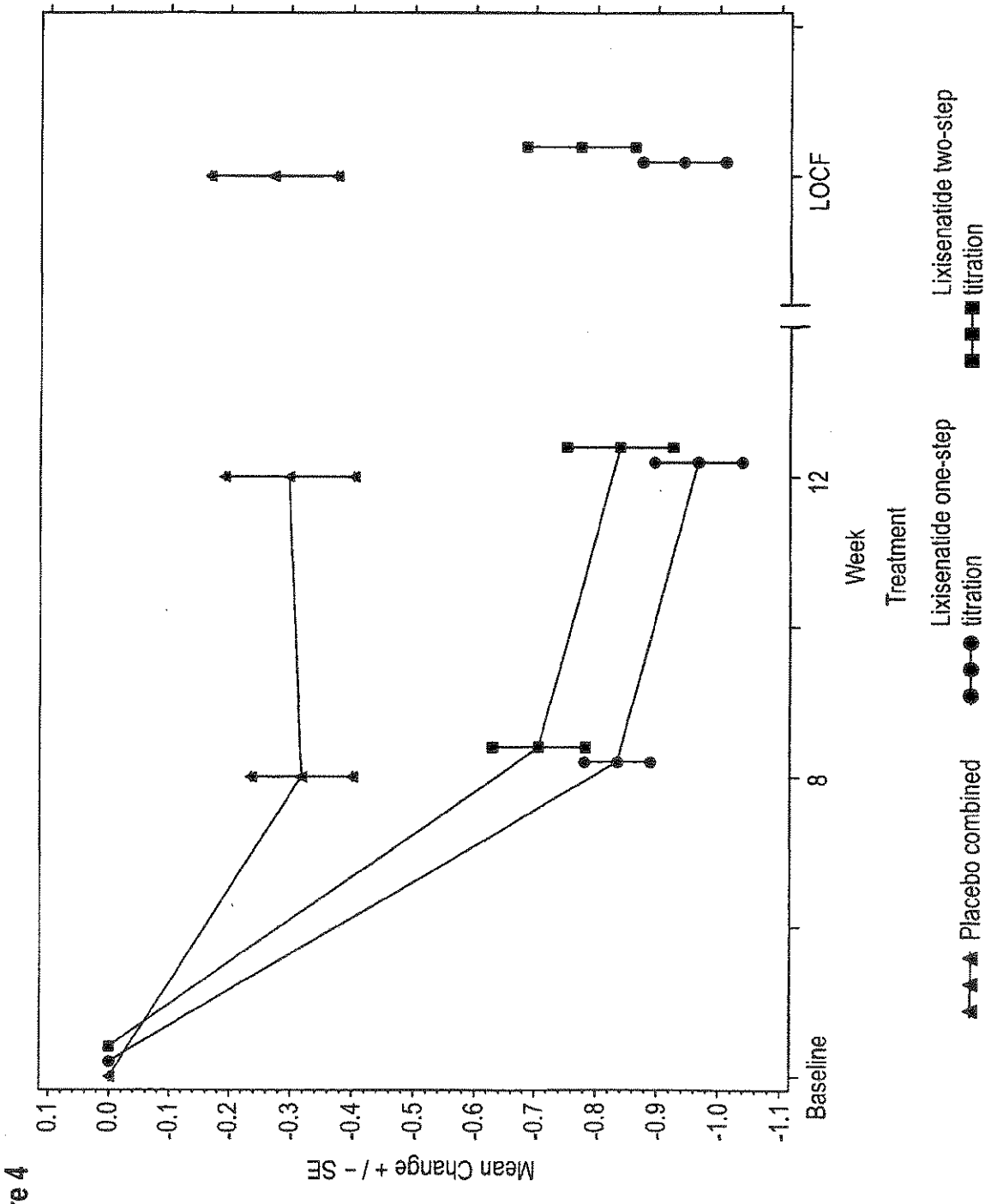

Mean Change + / - SE
0.1
0.0
-0.1
-0.2
-0.3
-0.4
-0.5
-0.6
-0.7
-0.8
-0.9
-1.0
-1.1
Baseline
8
12
LOCF
Week
Treatment
Placebo combined
Lixisenatide one-step titration
Lixisenatide two-step titration

Figure 5

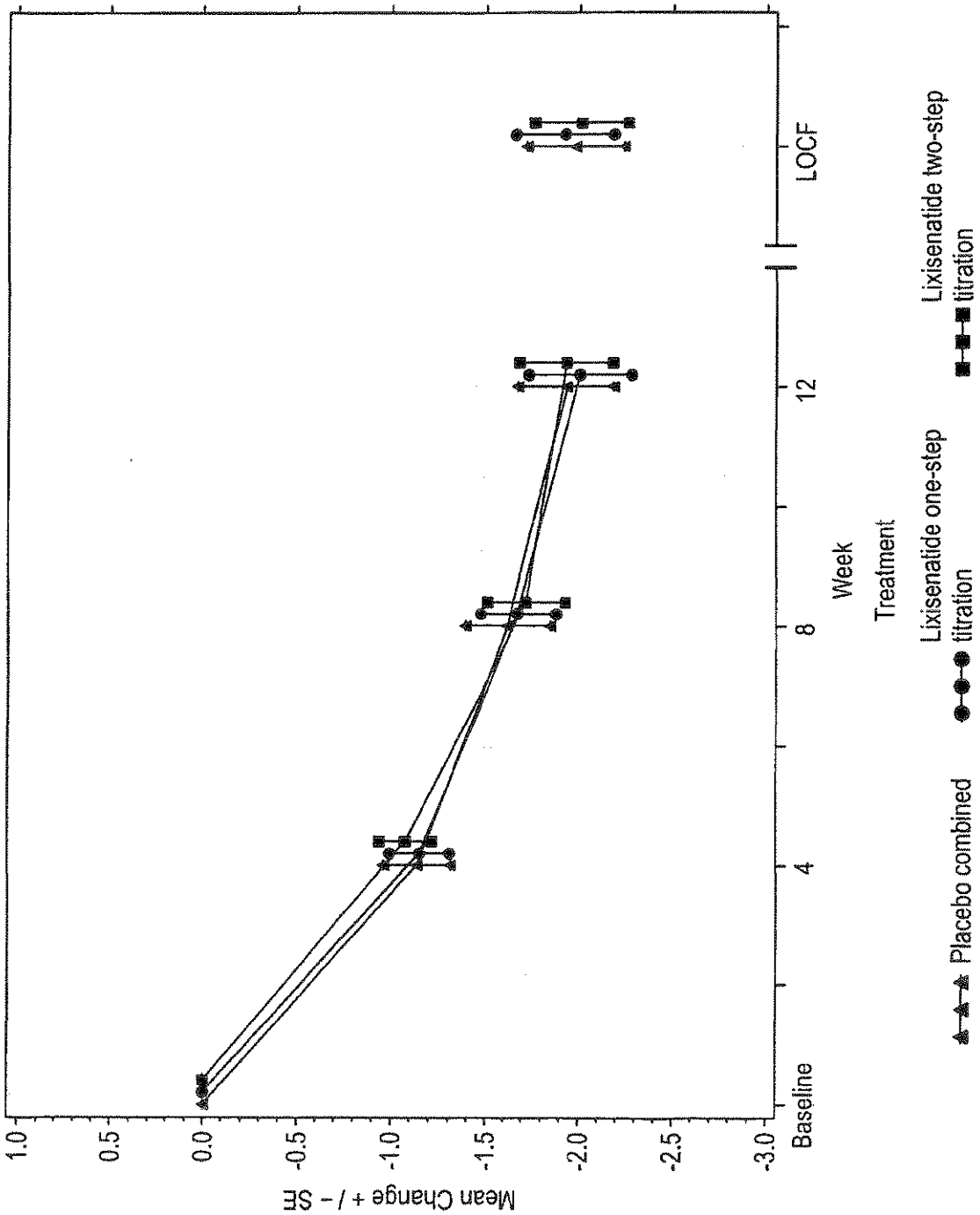
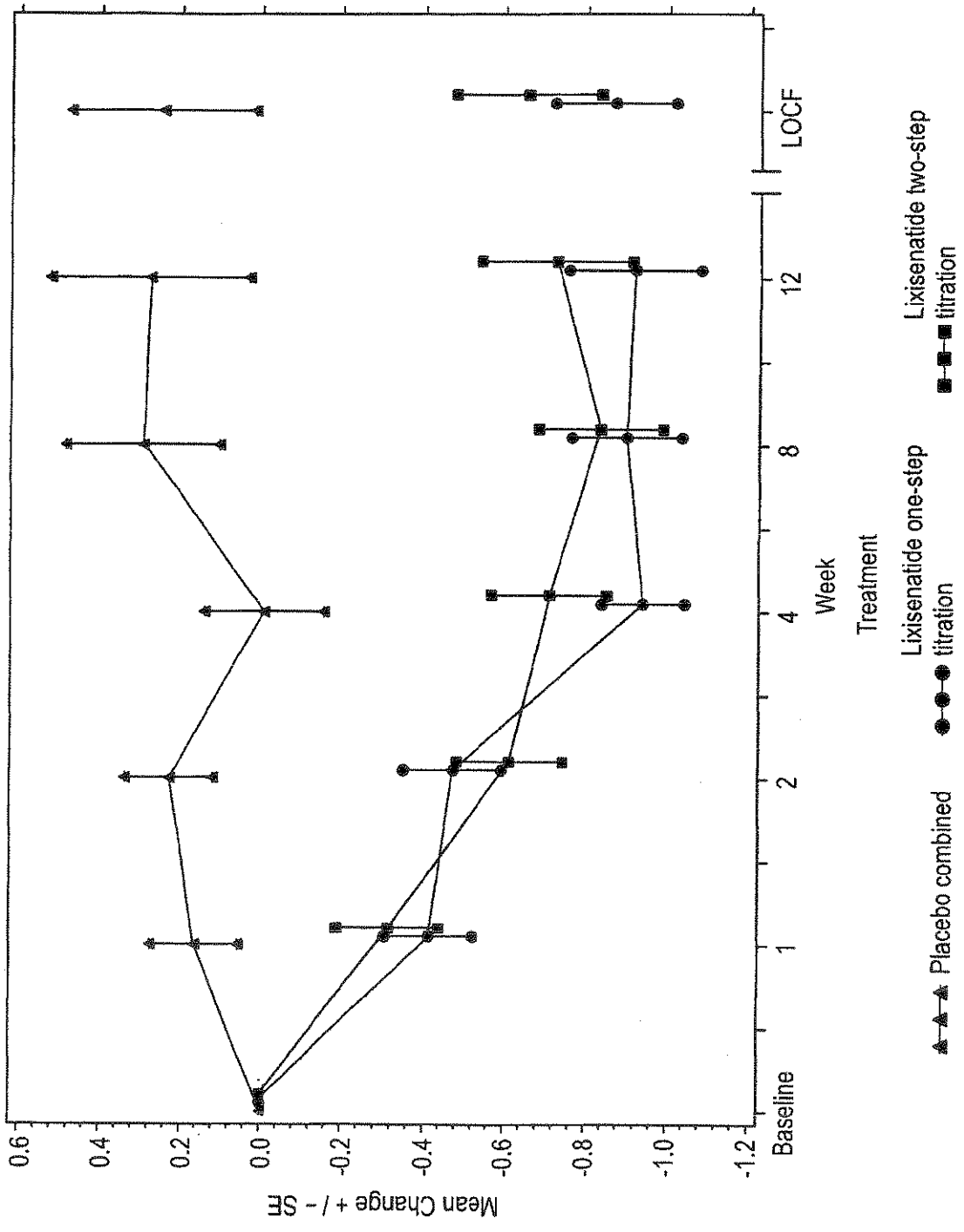

Mean Change + / - SE
1.0
0.5
0.0
-0.5
-1.0
-1.5
-2.0
-2.5
-3.0
Baseline
4
8
12
LOCF
Week
Treatment
Placebo combined
Lixisenatide one-step titration
Lixisenatide two-step titration

Figure 6


Mean Change + / - SE
0.6
0.4
0.2
0.0
-0.2
-0.4
-0.6
-0.8
-1.0
-1.2
Baseline
1
2
4
8
12
LOCF
Week
Treatment
Placebo combined
Lixisenatide one-step
titration
Lixisenatide two-step
titration