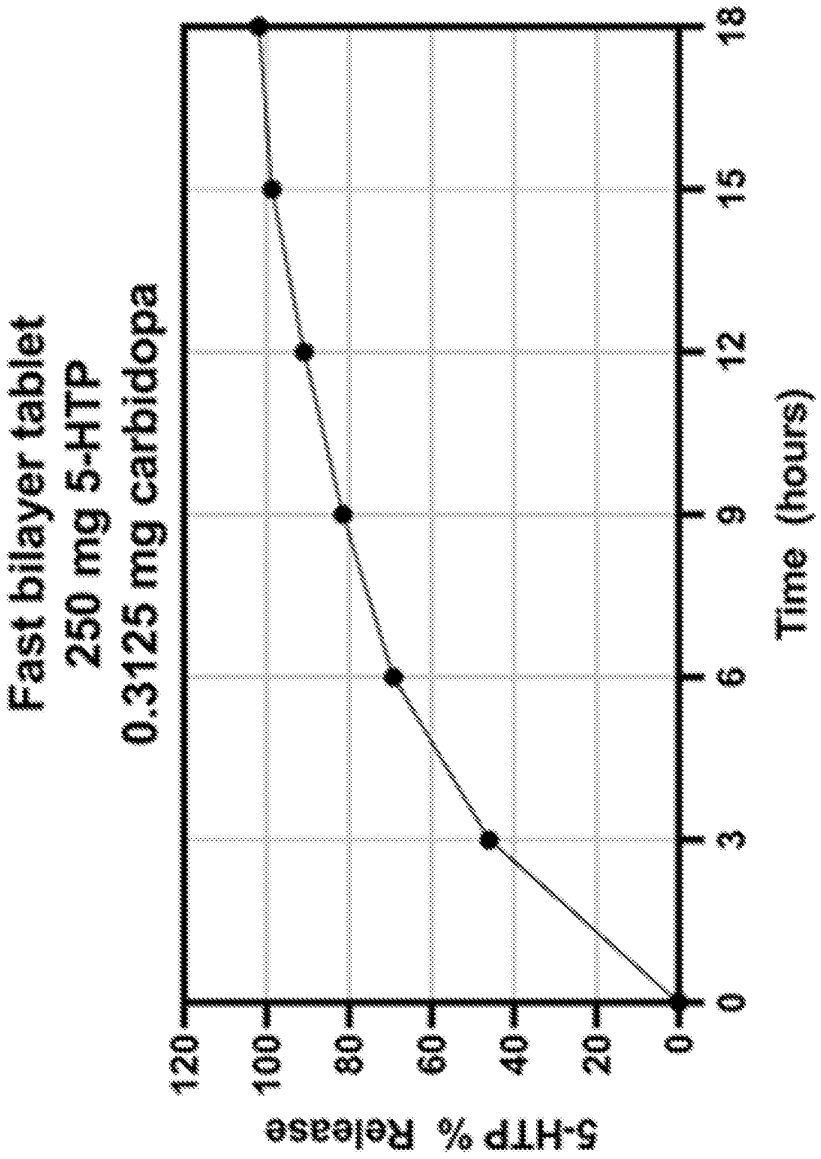

Fast bilayer tablet
250 mg 5-HTP
0.3125 mg carbidopa
5-HTP % Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 1A

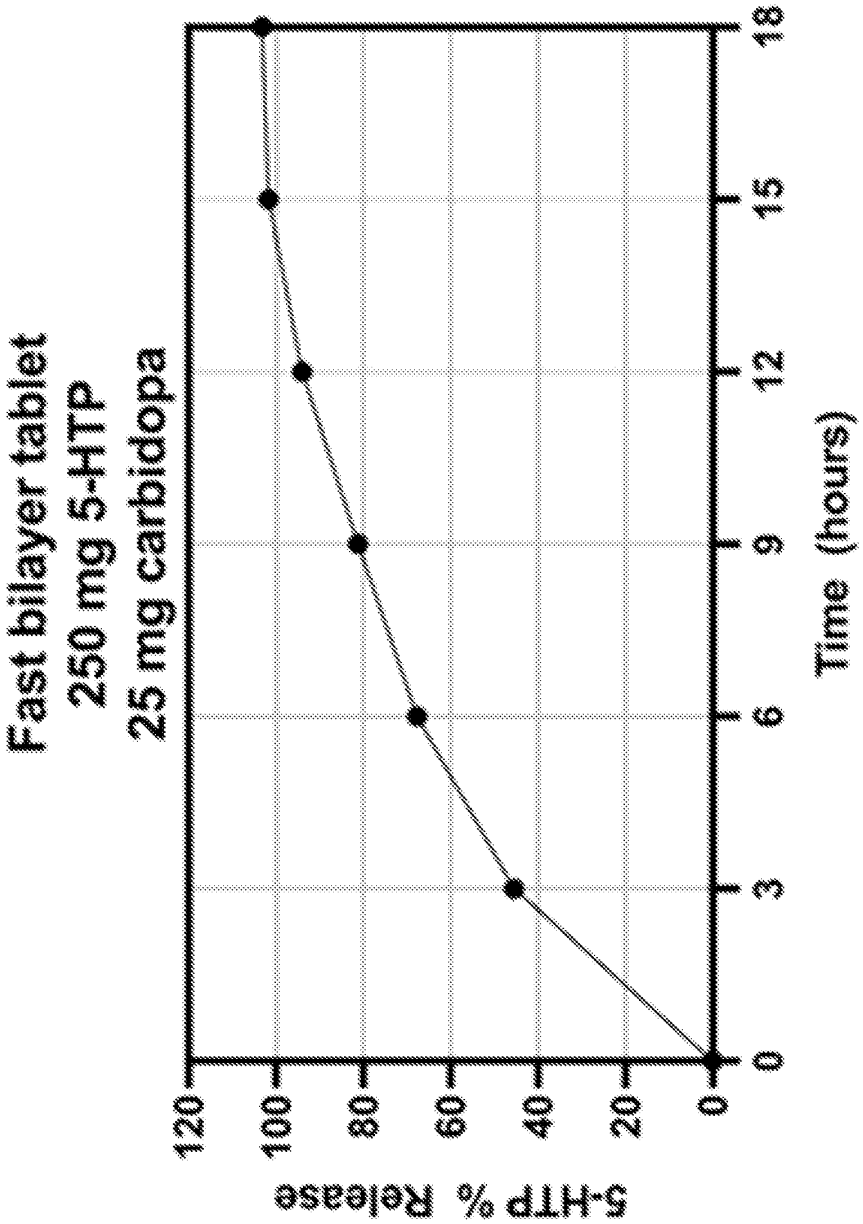

Fast bilayer tablet
250 mg 5-HTP
25 mg carbidopa
5-HTP % Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 1B

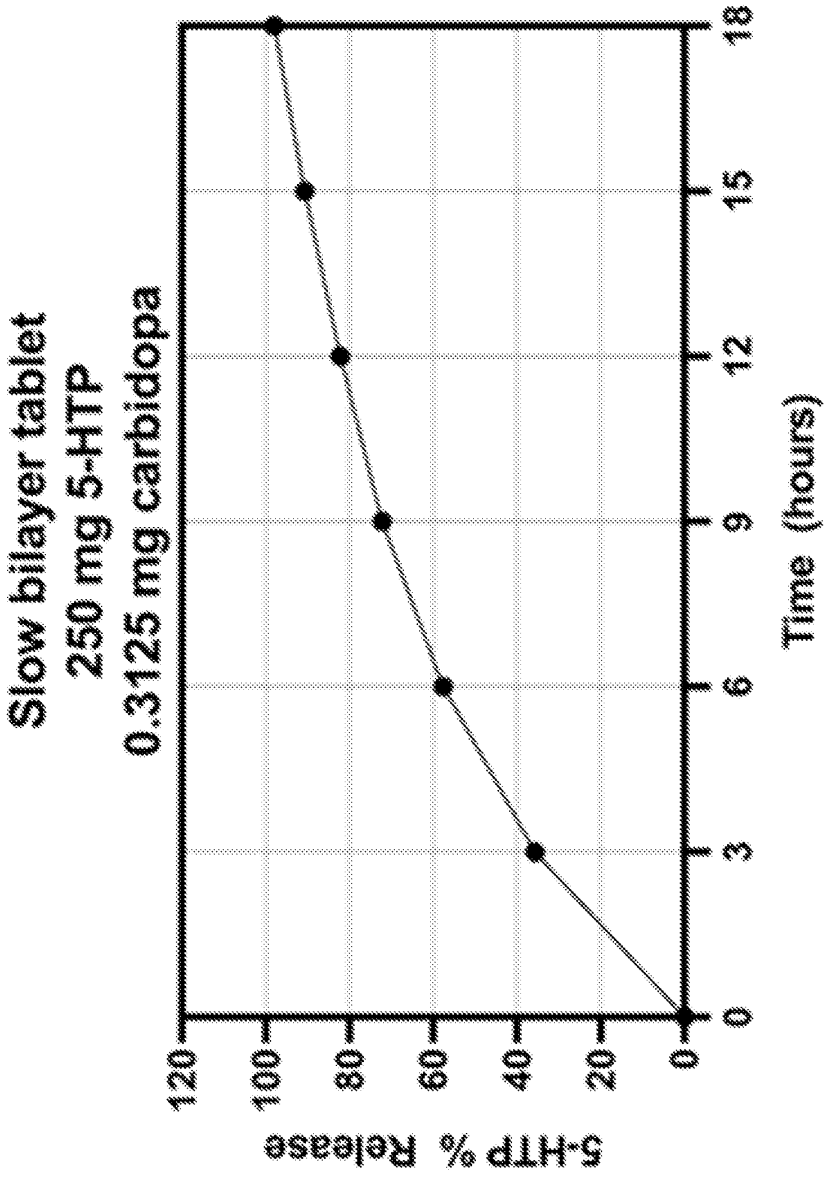

Slow bilayer tablet
250 mg 5-HTP
0.3125 mg carbidopa
5-HTP % Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 1C

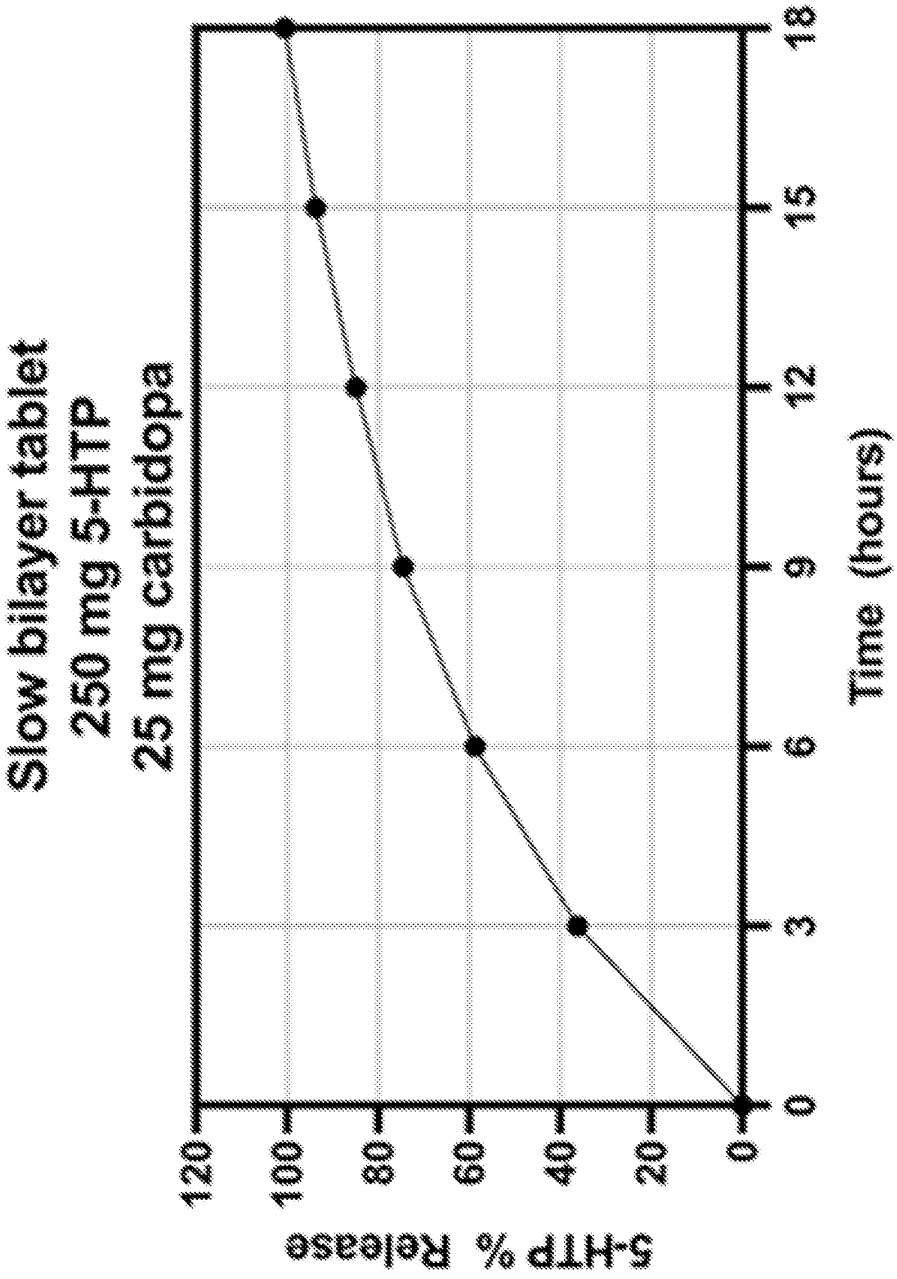

Slow bilayer tablet
250 mg 5-HTP
25 mg carbidopa
5-HTP % Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 1D

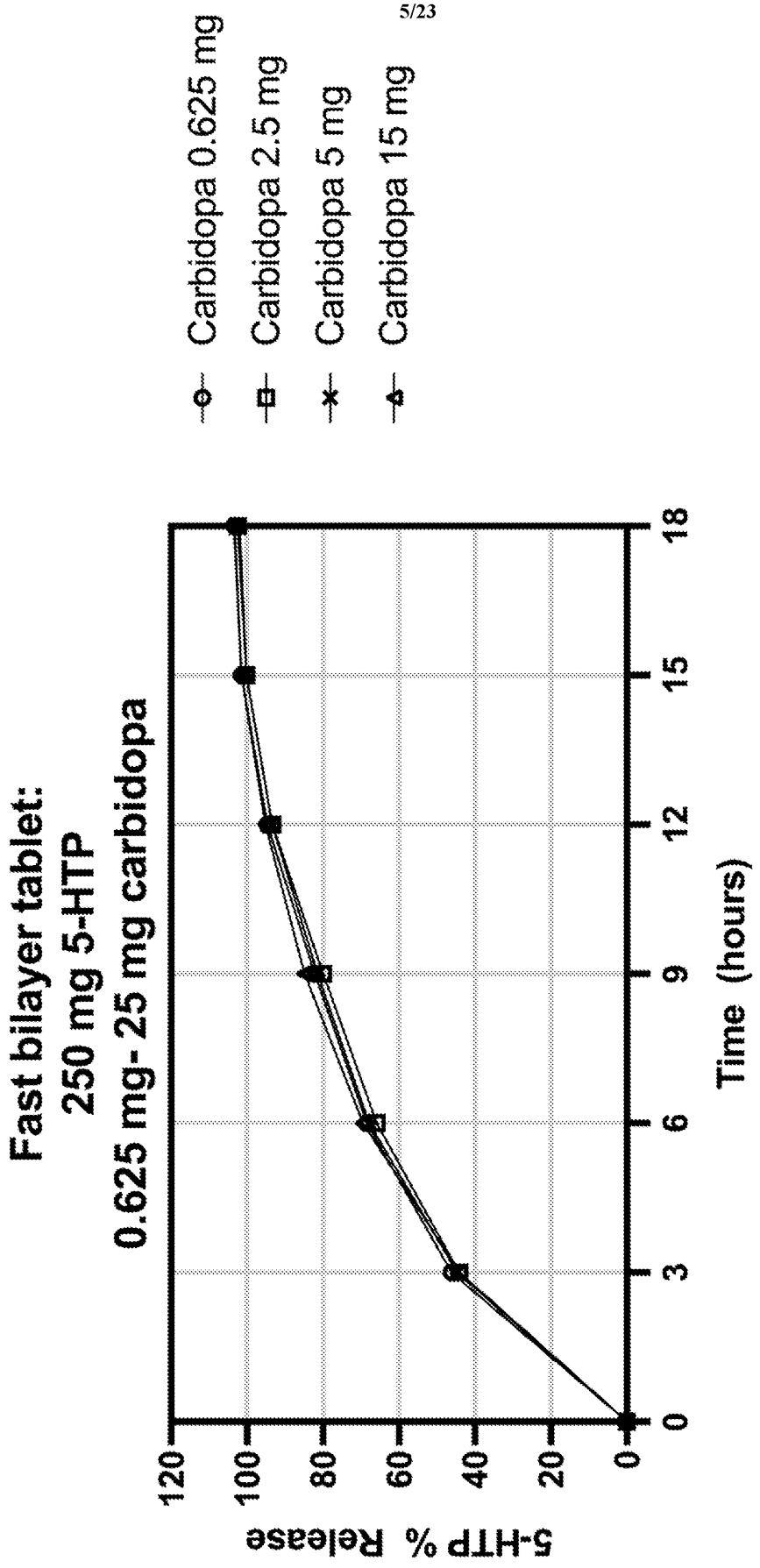

Fast bilayer tablet:
250 mg 5-HTP
0.625 mg- 25 mg carbidopa
5-HTP % Release
Time (hours)
0
20
40
60
80
100
120
0
3
6
9
12
15
18
Carbidopa 0.625 mg
Carbidopa 2.5 mg
Carbidopa 5 mg
Carbidopa 15 mg

FIG. 2

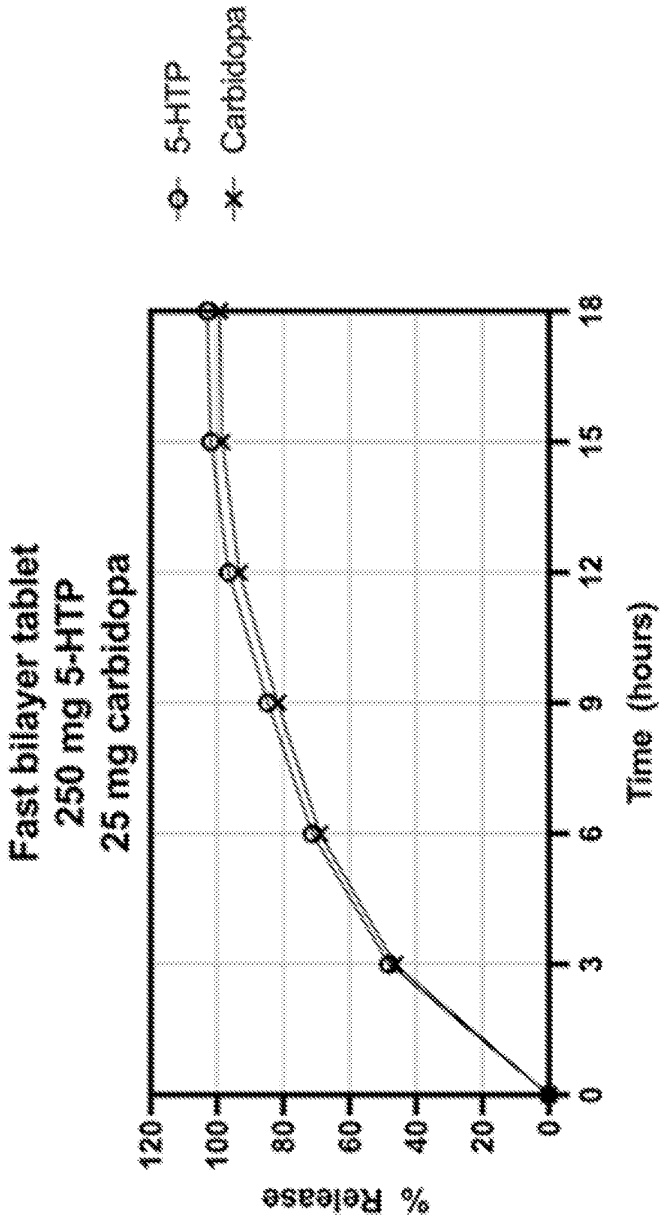

Fast bilayer tablet
250 mg 5-HTP
25 mg carbidopa
5-HTP
Carbidopa
% Release
0
20
40
60
80
100
120
Time (hours)
0
3
6
9
12
15
18

FIG. 3A

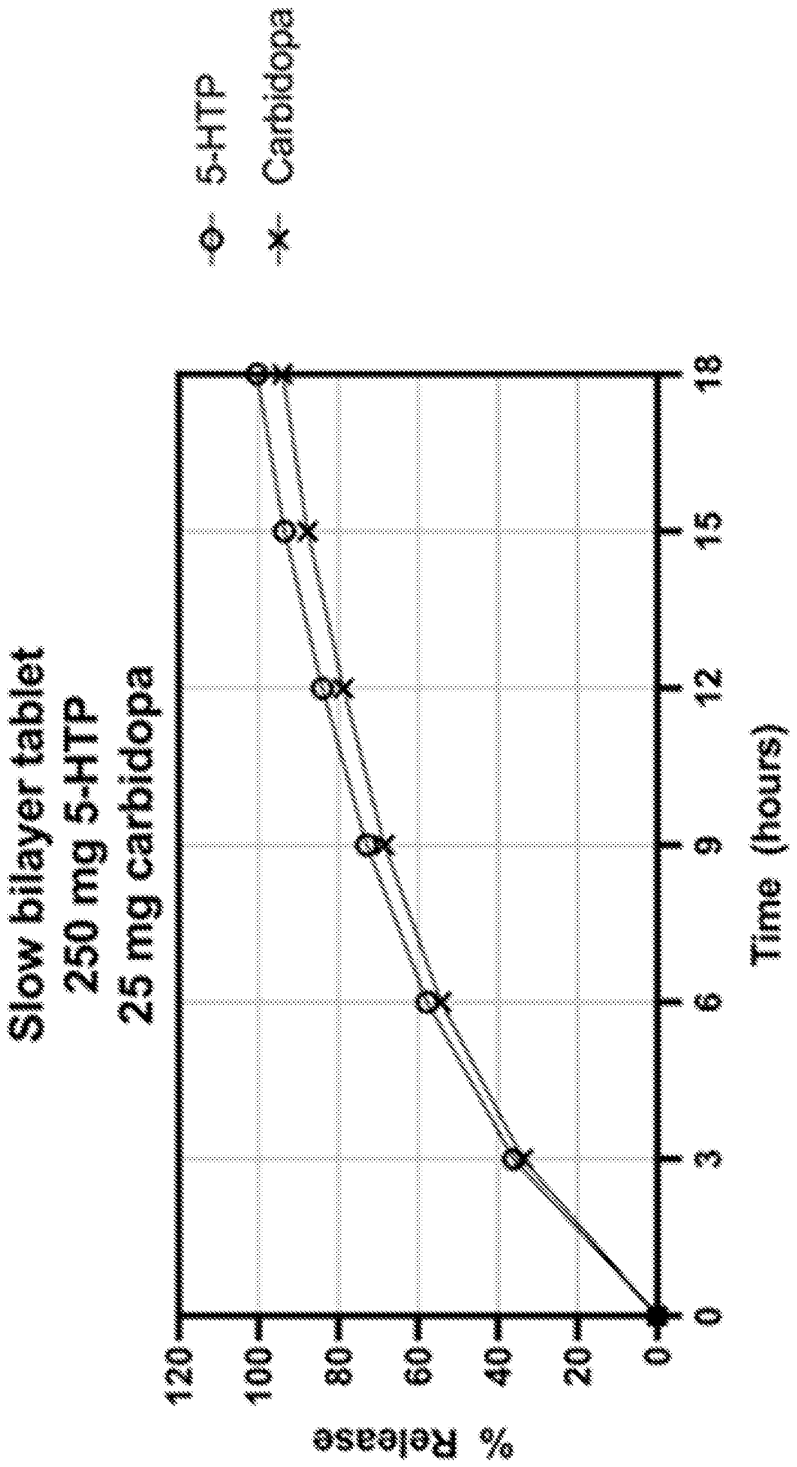

5-HTP
Carbidopa
Slow bilayer tablet
250 mg 5-HTP
25 mg carbidopa
% Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 3B

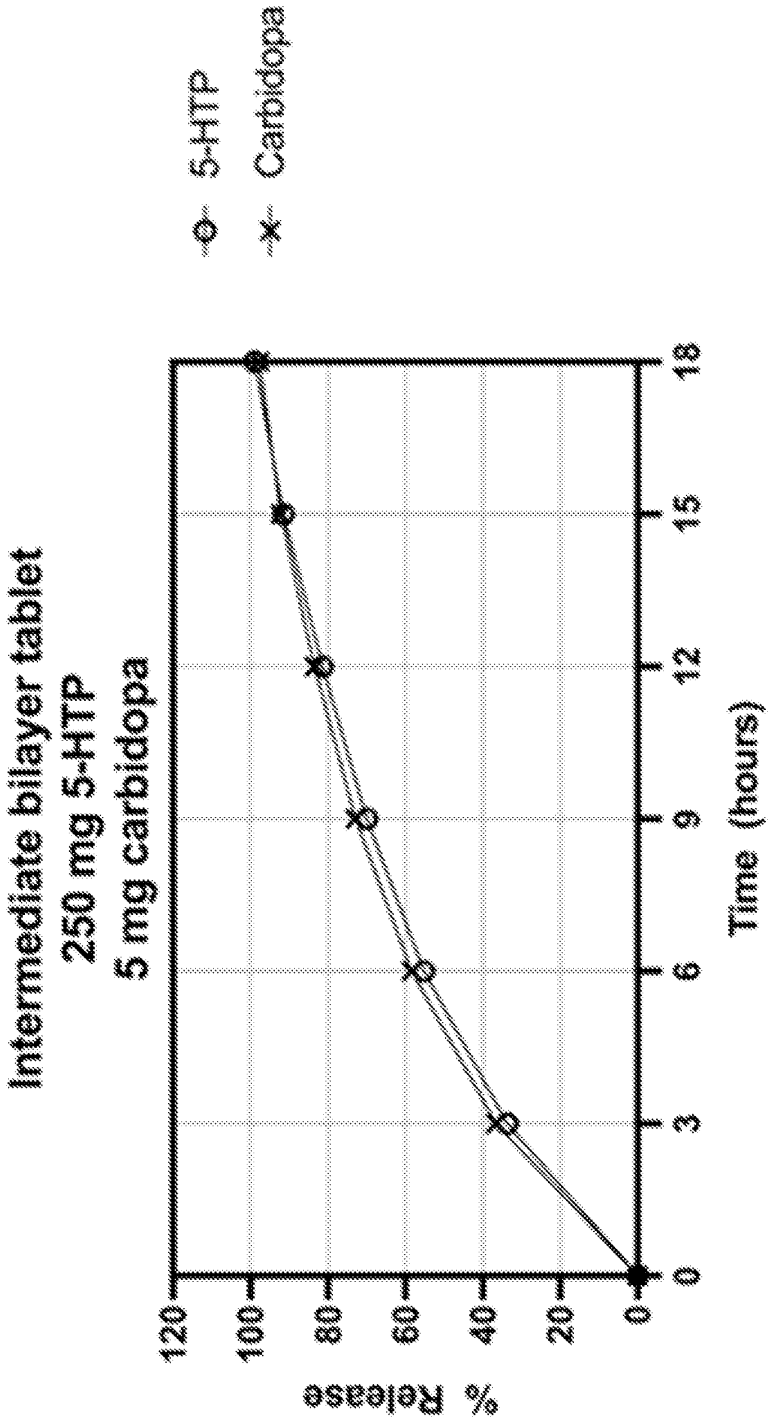

Intermediate bilayer tablet
250 mg 5-HTP
5 mg carbidopa
5-HTP
Carbidopa
% Release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 3C

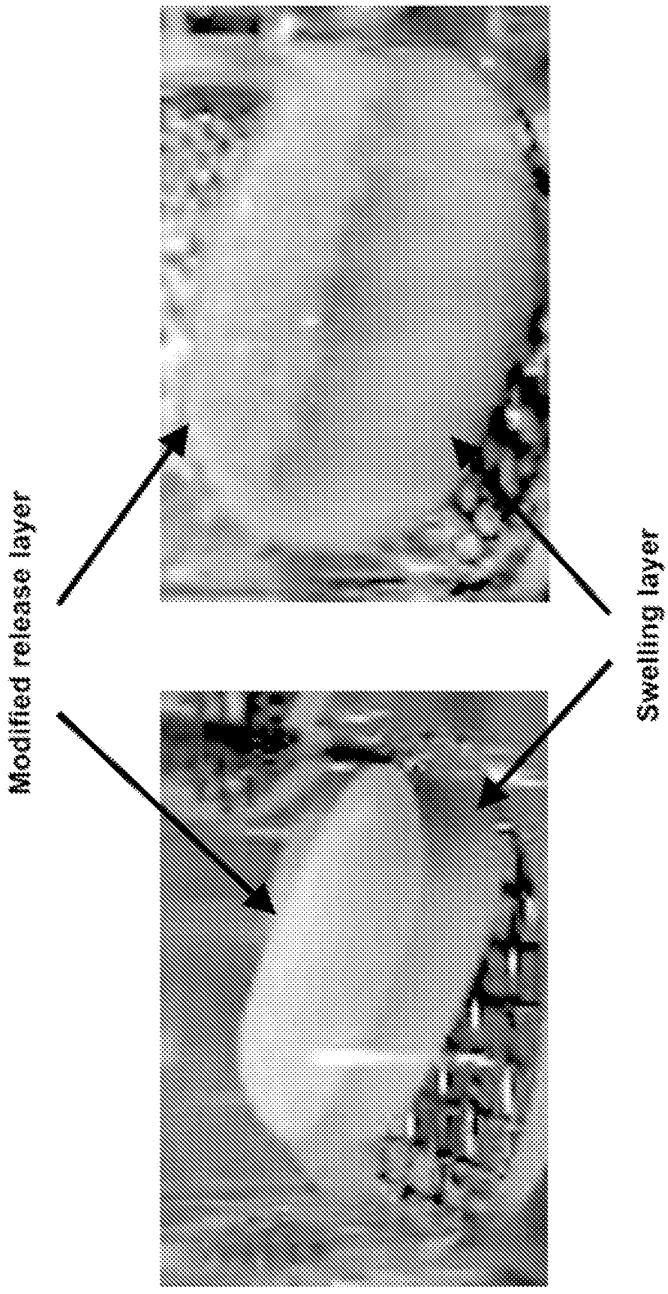


FIG. 4

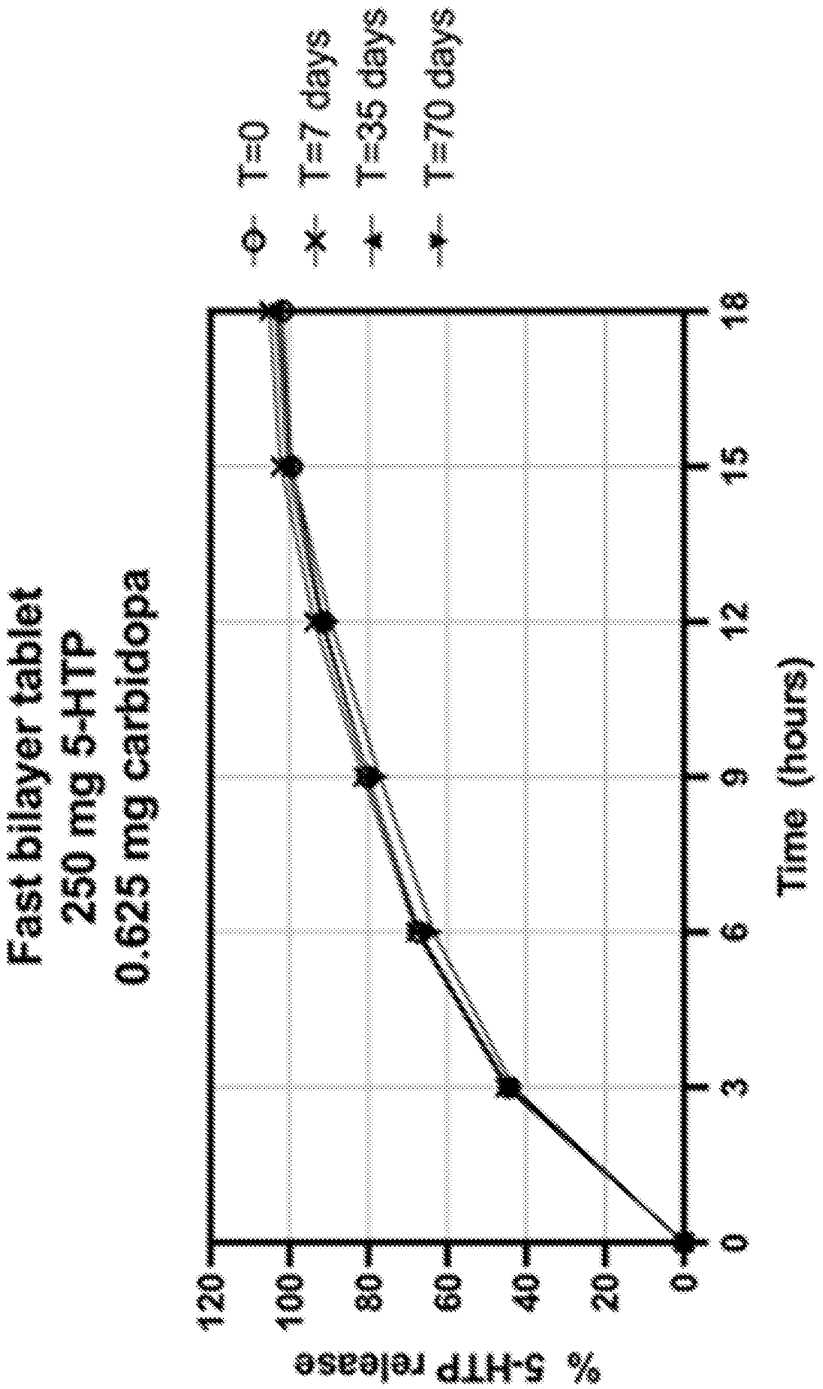

Fast bilayer tablet
250 mg 5-HTP
0.625 mg carbidopa
T=0
T=7 days
T=35 days
T=70 days
% 5-HTP release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 5A

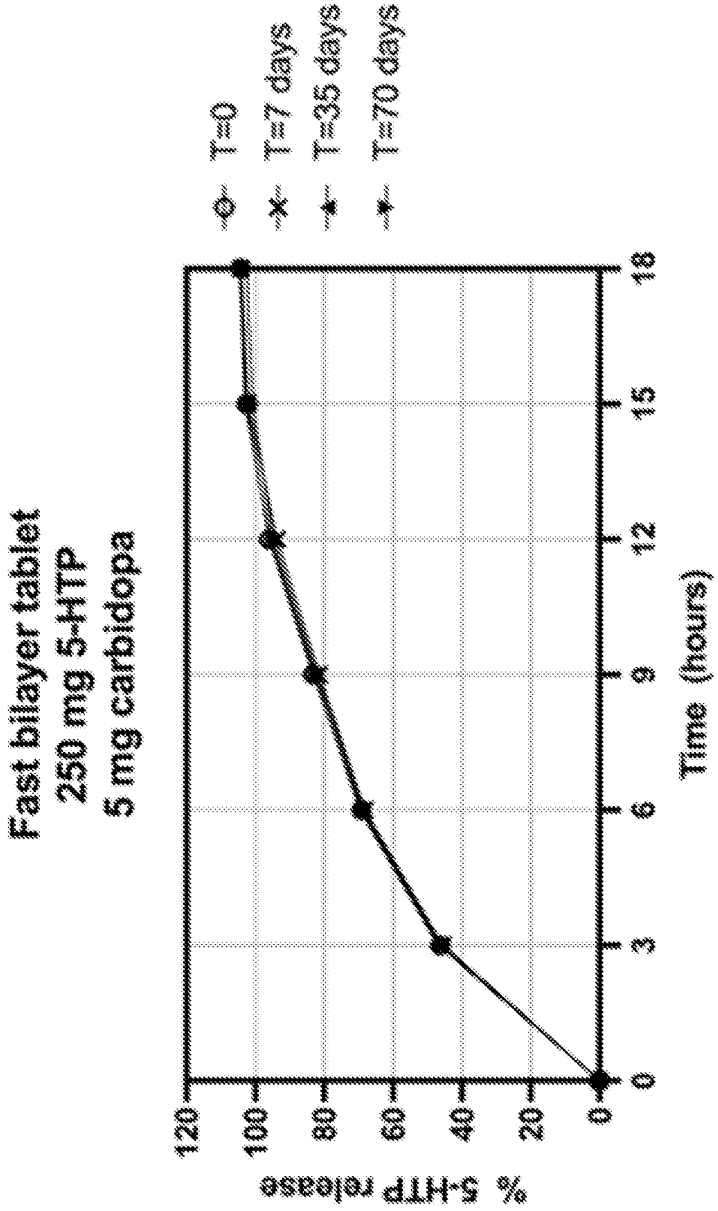

Fast bilayer tablet
250 mg 5-HTP
5 mg carbidopa
T=0
T=7 days
T=35 days
T=70 days
% 5-HTP release
0
20
40
60
80
100
120
Time (hours)
0
3
6
9
12
15
18

FIG. 5B

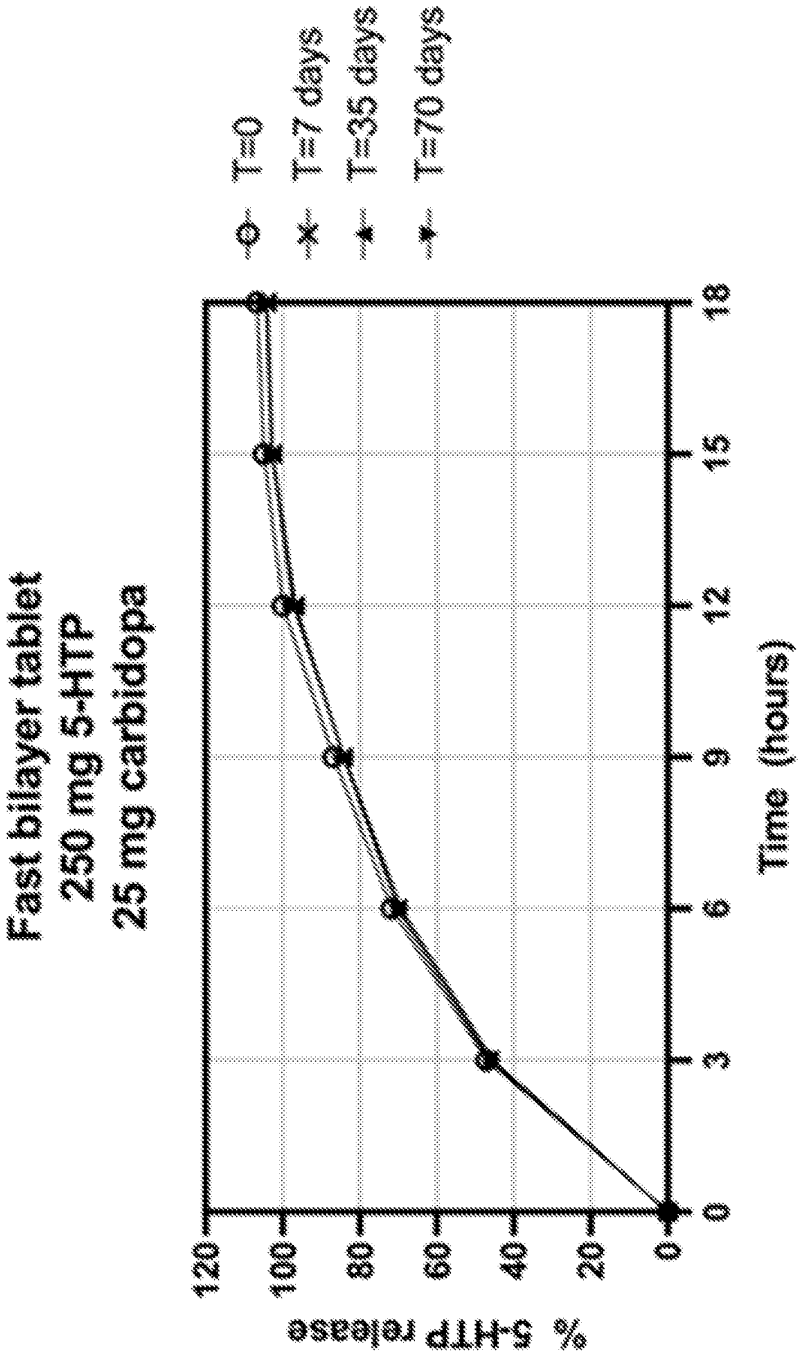

Fast bilayer tablet
250 mg 5-HTP
25 mg carbidopa
T=0
T=7 days
T=35 days
T=70 days
% 5-HTP release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 5C

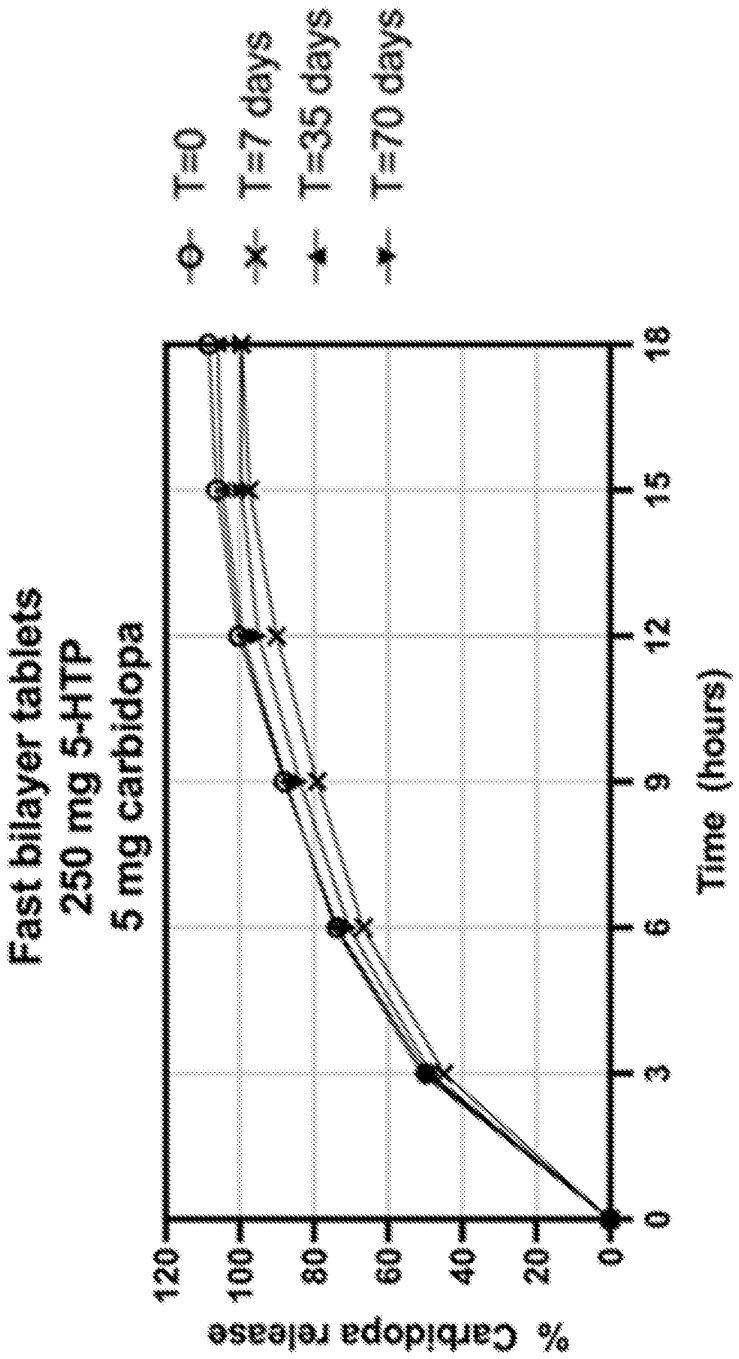

T=0
T=7 days
T=35 days
T=70 days
Fast bilayer tablets
250 mg 5-HTP
5 mg carbidopa
% Carbidopa release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 5D

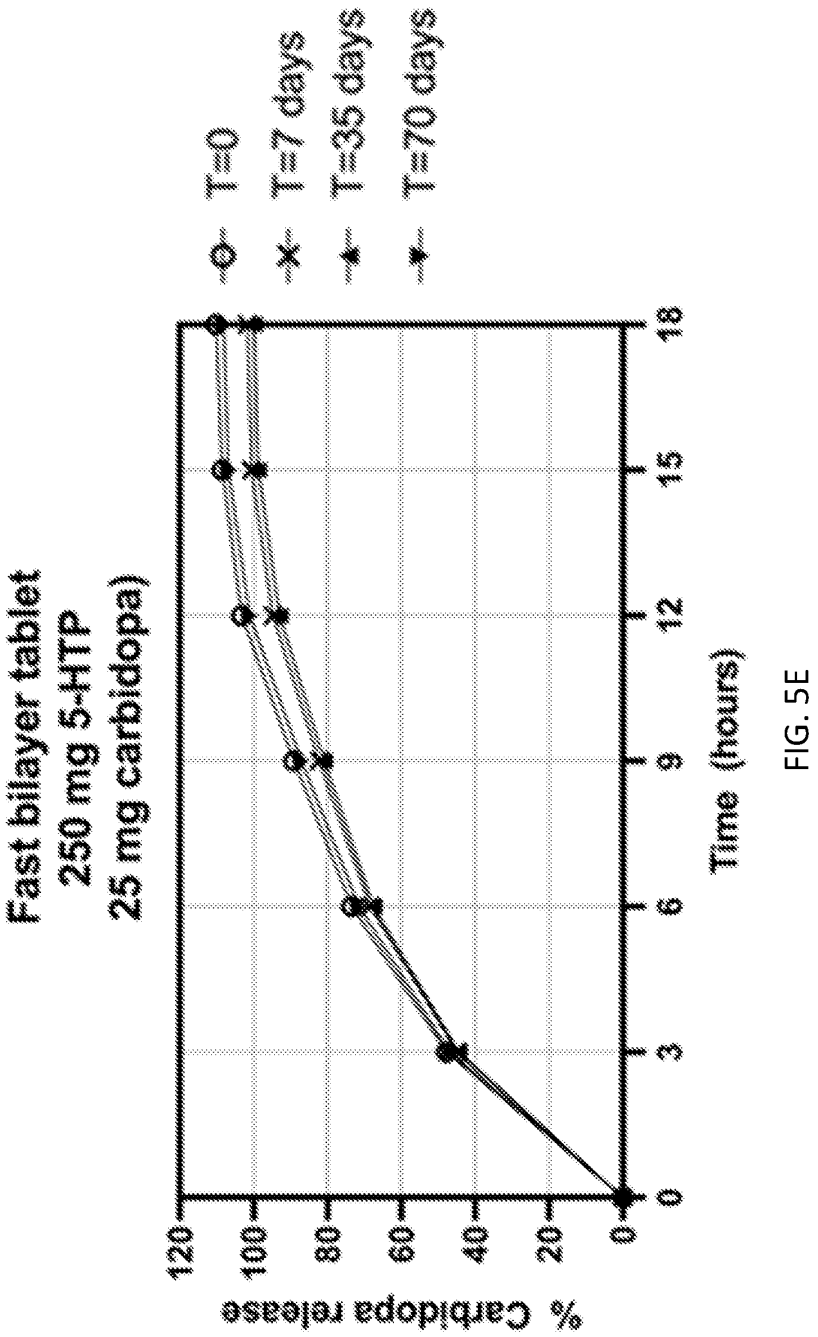

Fast bilayer tablet
250 mg 5-HTP
25 mg carbidopa)
T=0
T=7 days
T=35 days
T=70 days
% Carbidopa release
120
100
80
60
40
20
0
0
3
6
9
12
15
18
Time (hours)

FIG. 5E

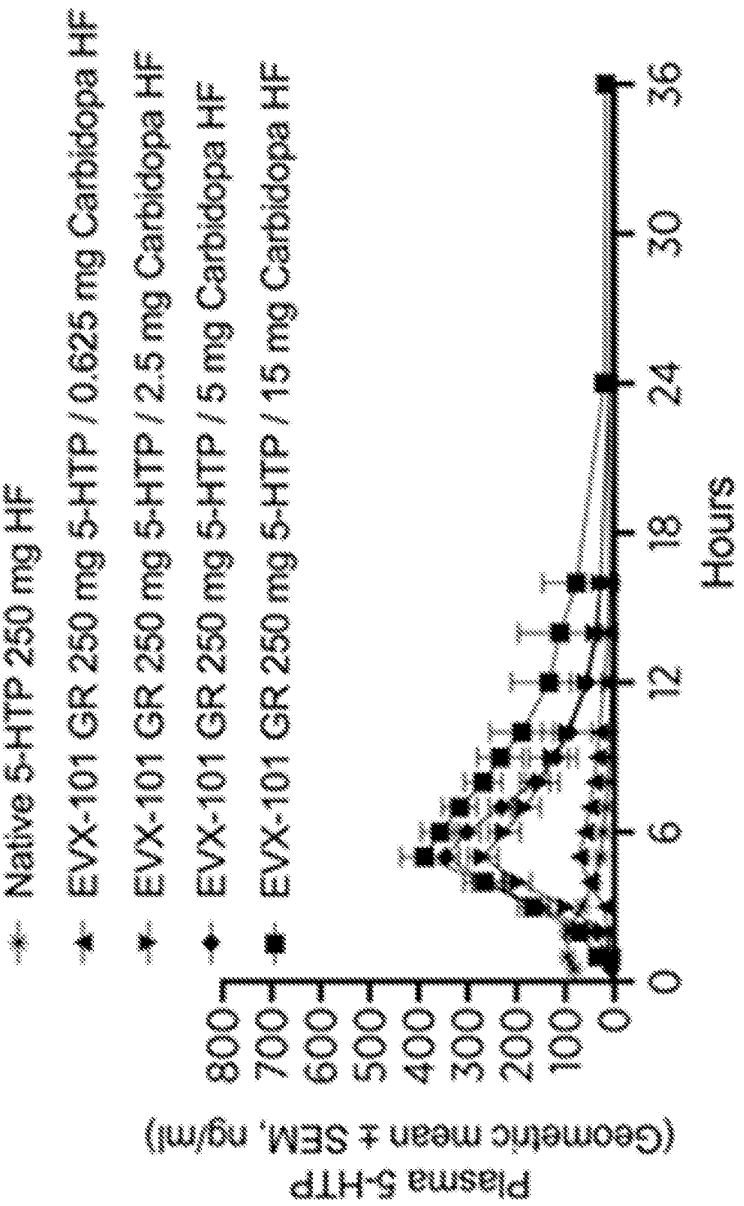

Native 5-HTP 250 mg HF
EVX-101 GR 250 mg 5-HTP / 0.625 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 2.5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
Plasma 5-HTP
(Geometric mean ± SEM, ng/ml)
800
700
600
500
400
300
200
100
0
0
6
12
18
24
30
36
Hours

FIG. 6A

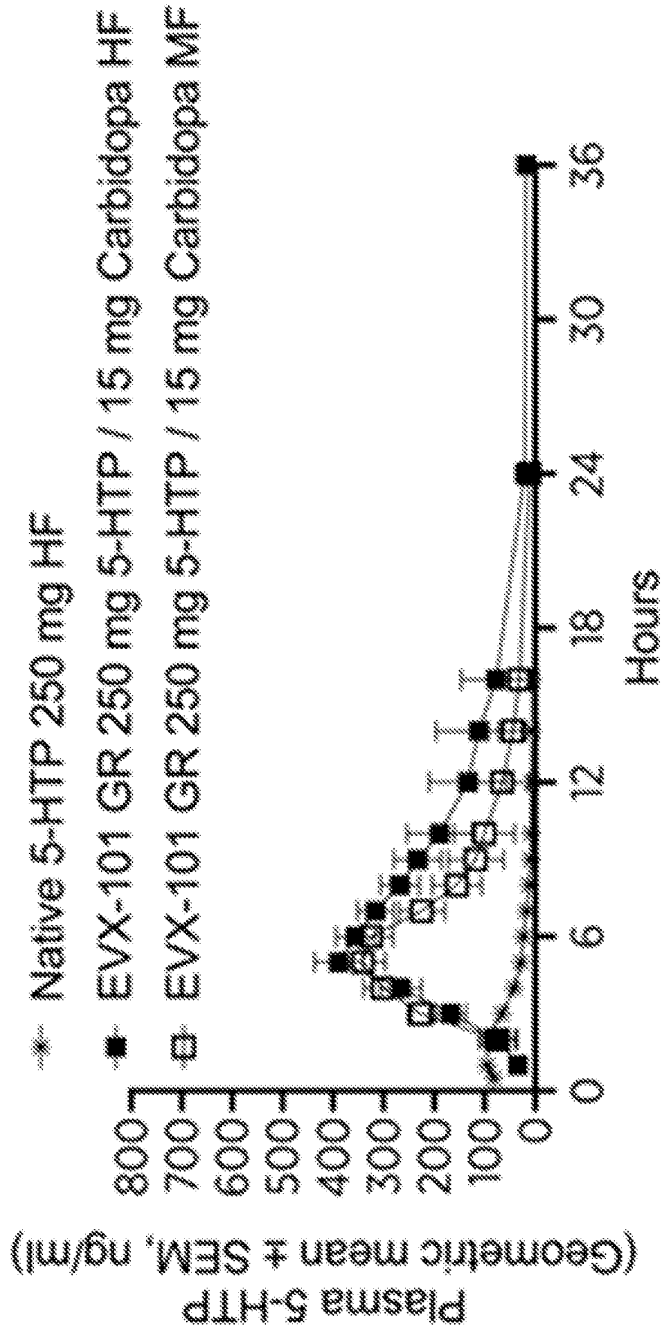

Native 5-HTP 250 mg HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa MF
Plasma 5-HTP
(Geometric mean ± SEM, ng/ml)
800
700
600
500
400
300
200
100
0
0
6
12
18
24
30
36
Hours

FIG. 6B

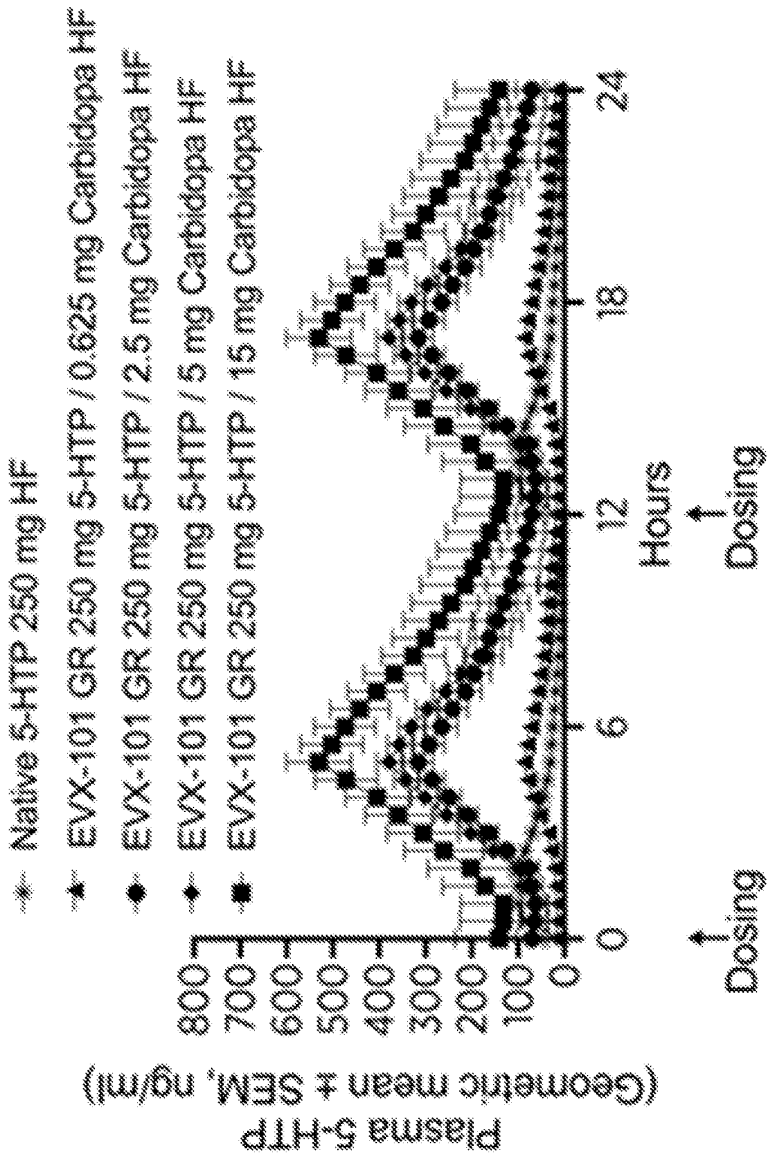

Native 5-HTP 250 mg HF
EVX-101 GR 250 mg 5-HTP / 0.625 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 2.5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
Plasma 5-HTP
(Geometric mean ± SEM, ng/ml)
800
700
600
500
400
300
200
100
0
0
6
12
18
24
Hours
Dosing
Dosing

FIG. 6C

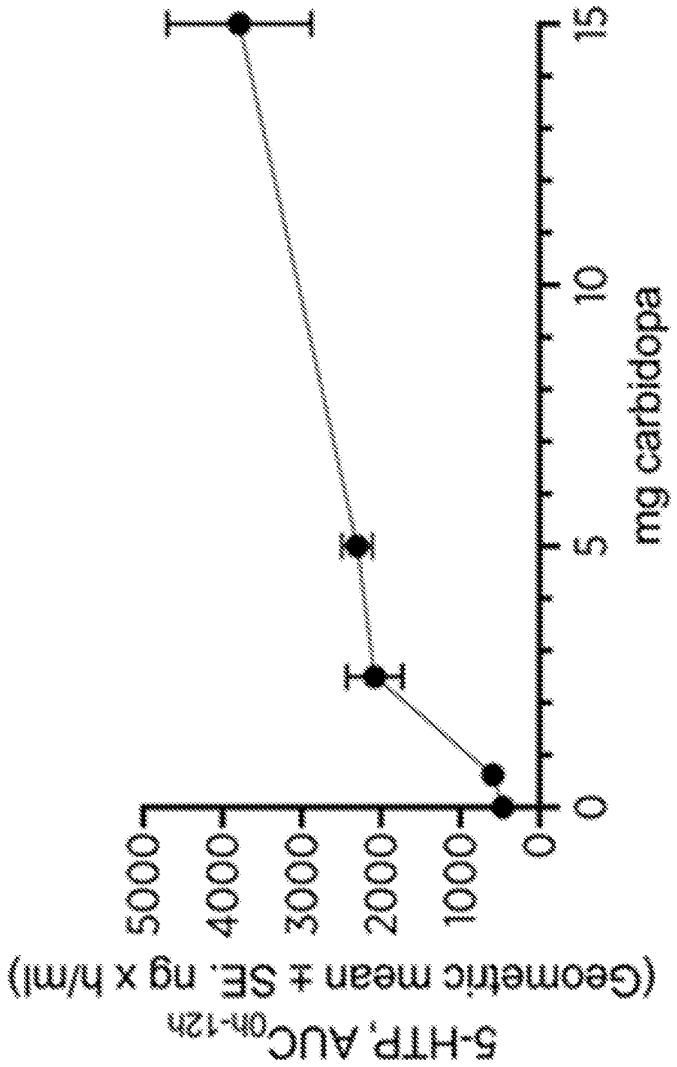

5-HTP, AUC0h-12h
(Geometric mean ± SE, ng x h/ml)
5000
4000
3000
2000
1000
0
0
5
10
15
mg carbidopa

FIG. 6D

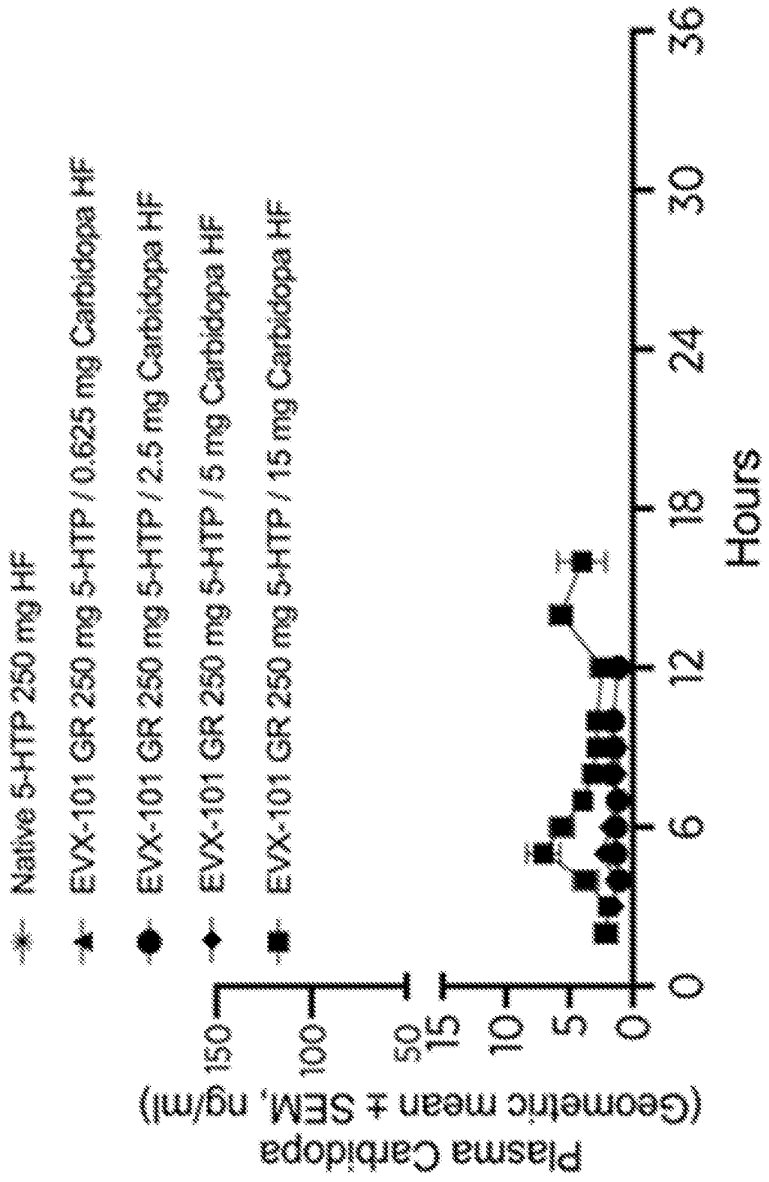

Native 5-HTP 250 mg HF
EVX-101 GR 250 mg 5-HTP / 0.625 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 2.5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
Plasma Carbidopa (Geometric mean ± SEM, ng/ml)
150
100
50
15
10
5
0
0
6
12
18
24
30
36
Hours

FIG. 7A

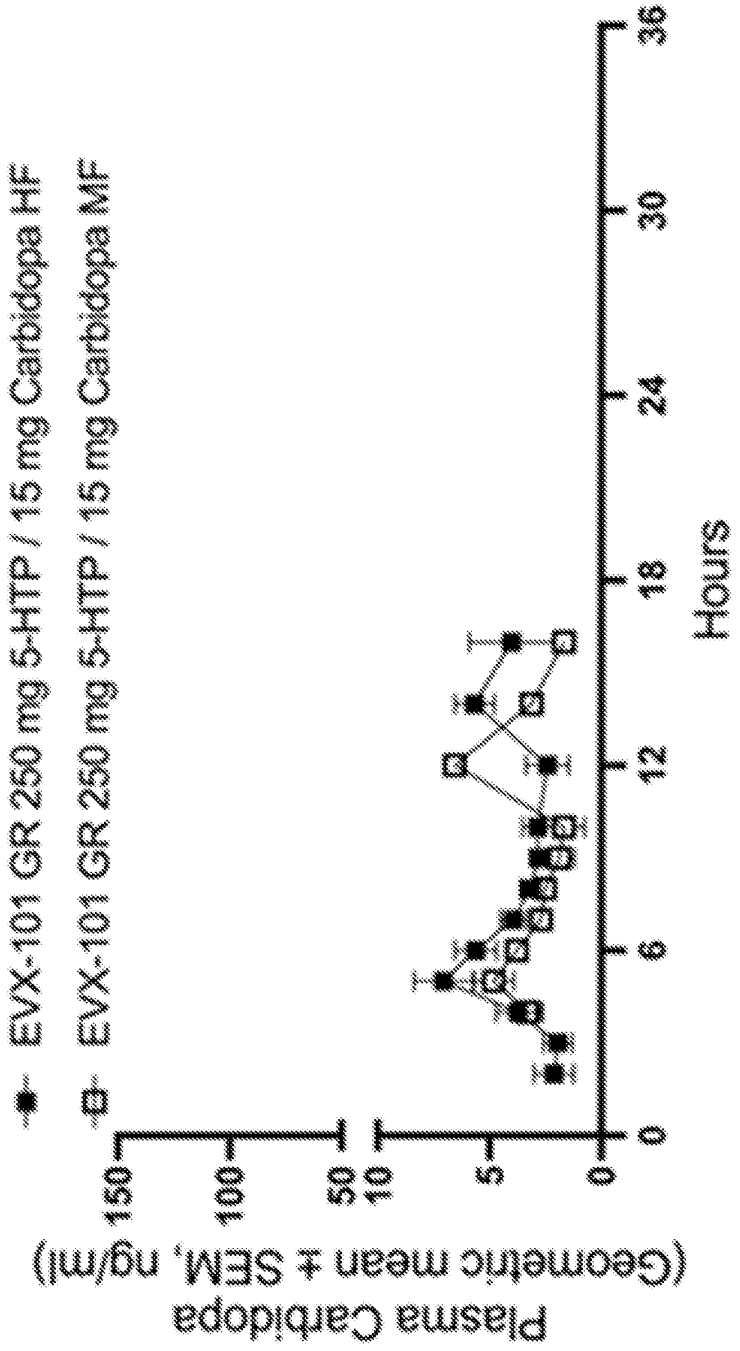

EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa MF
Plasma Carbidopa
(Geometric mean ± SEM, ng/ml)
150
100
50
10
5
0
0
6
12
18
24
30
36
Hours

FIG. 7B

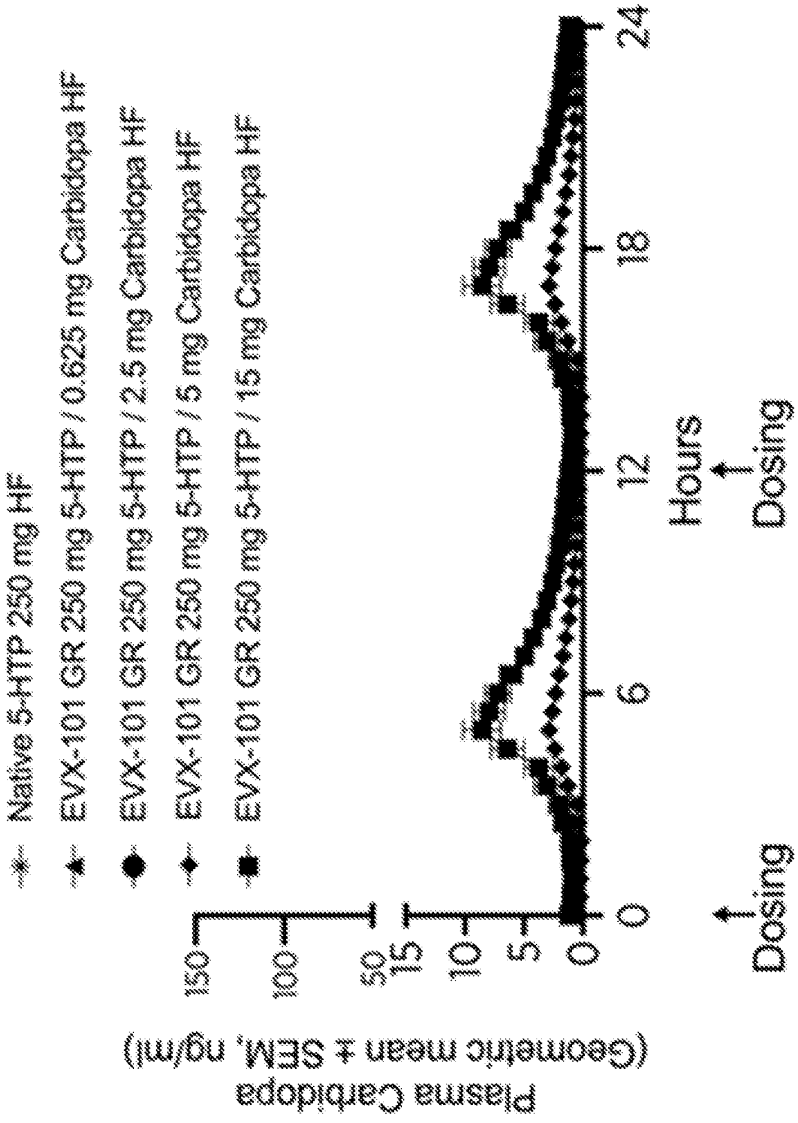

Native 5-HTP 250 mg HF
EVX-101 GR 250 mg 5-HTP / 0.625 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 2.5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 5 mg Carbidopa HF
EVX-101 GR 250 mg 5-HTP / 15 mg Carbidopa HF
Plasma Carbidopa
(Geometric mean ± SEM, ng/ml)
150
100
50
15
10
5
0
0
6
12
18
24
Hours
Dosing
Dosing

FIG. 7C

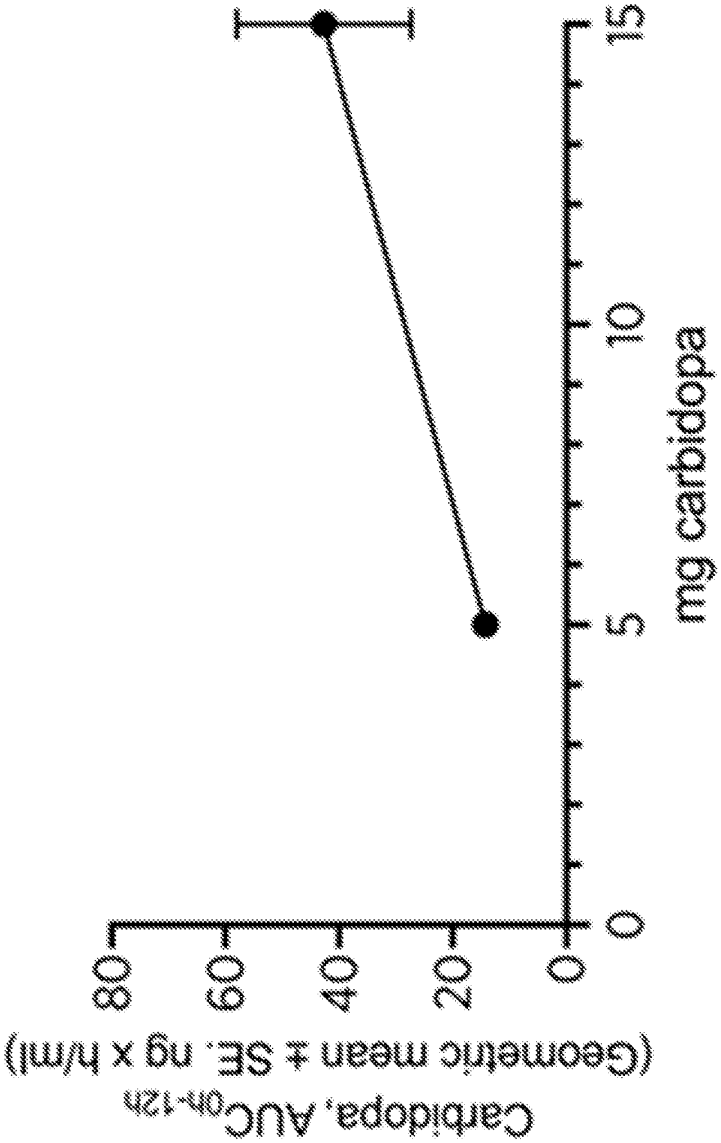

Carbidopa, AUC0h-12h
(Geometric mean ± SE, ng x h/ml)
0
20
40
60
80
0
5
10
15
mg carbidopa

FIG. 7D

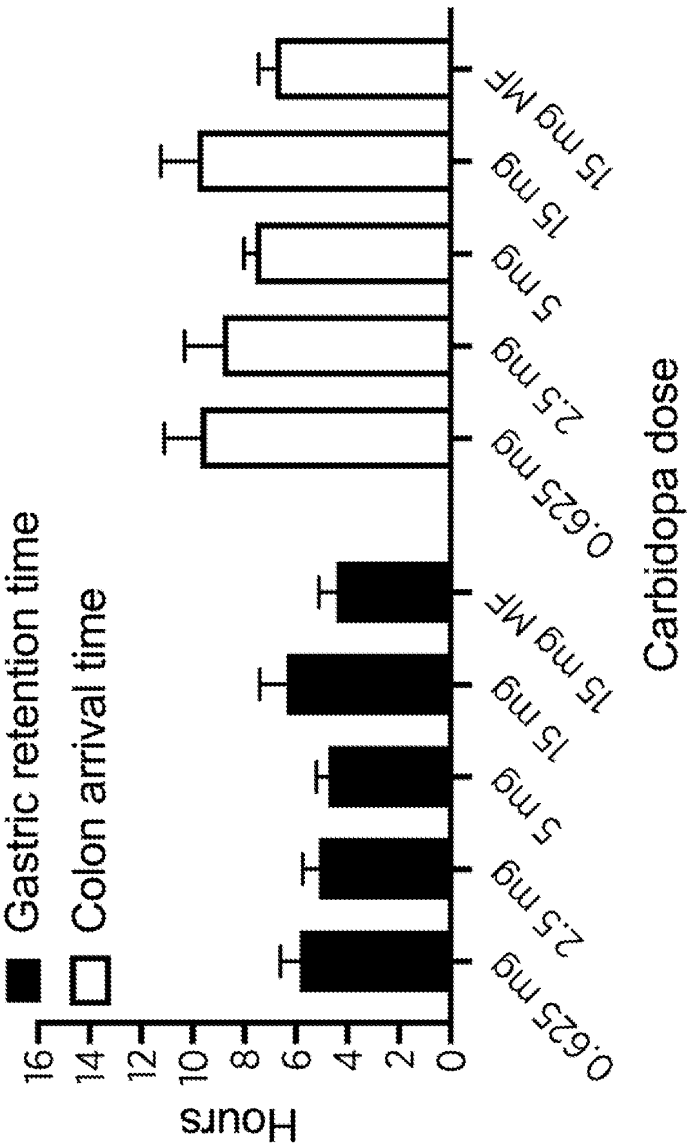

Gastric retention time
Colon arrival time
Hours
16
14
12
10
8
6
4
2
0
0.625 mg
2.5 mg
5 mg
15 mg
15 mg ME
0.625 mg
2.5 mg
5 mg
15 mg
15 mg ME
Carbidopa dose

FIG. 8