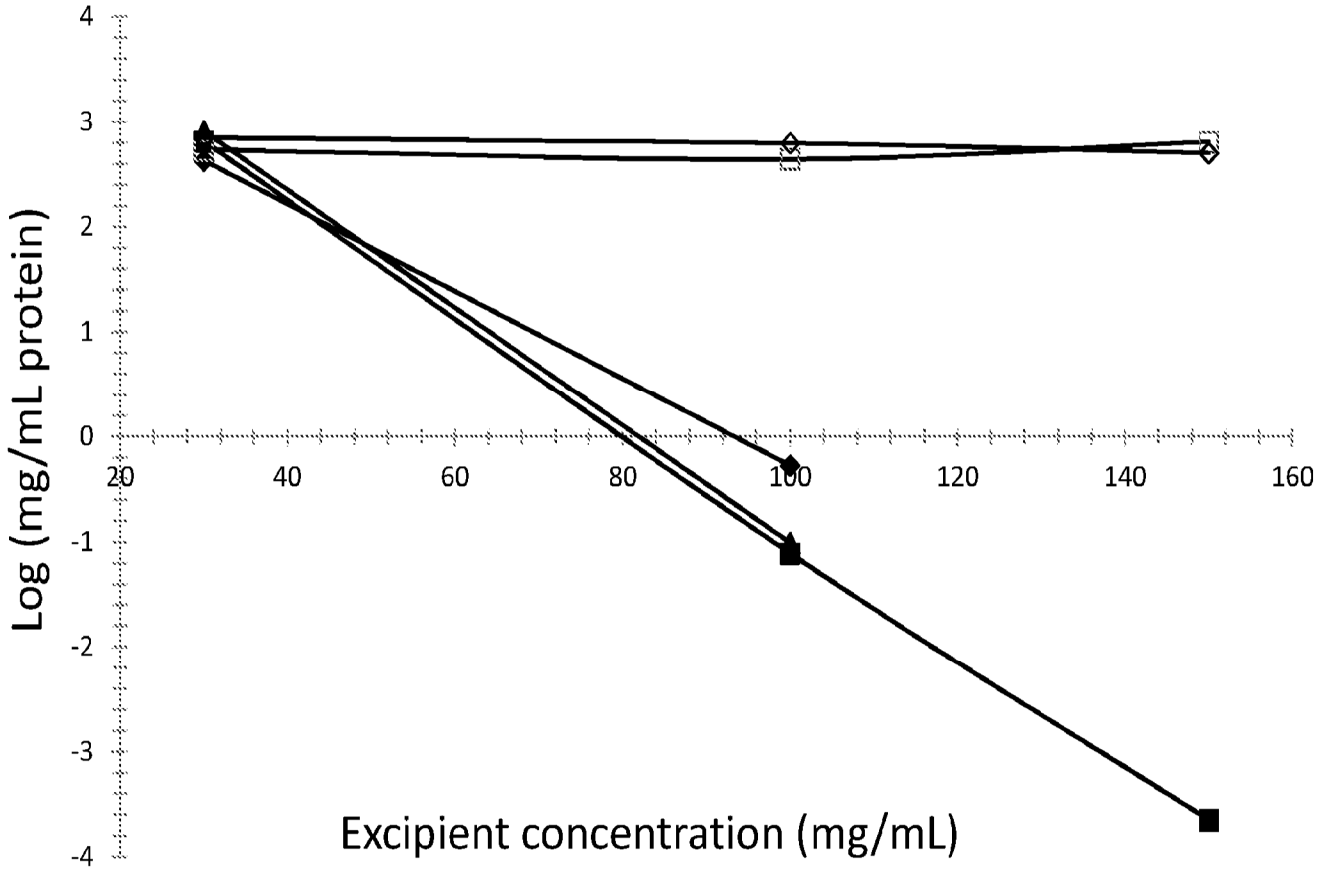

4
3
2
1
0
-1
-2
-3
-4
20
40
60
80
100
120
140
160
Log (mg/mL protein)
Excipient concentration (mg/mL)

FIGURE 1

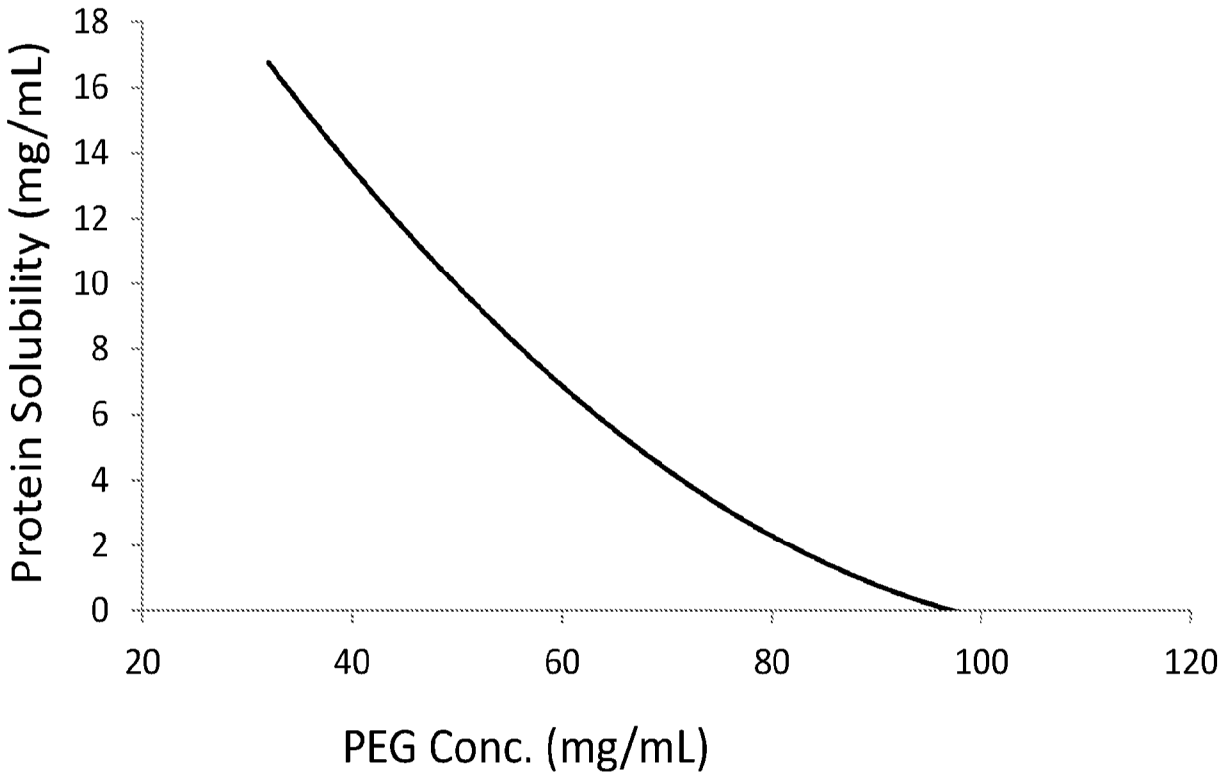

Protein Solubility (mg/mL)
18
16
14
12
10
8
6
4
2
0
20
40
60
80
100
120
PEG Conc. (mg/mL)

FIGURE 2

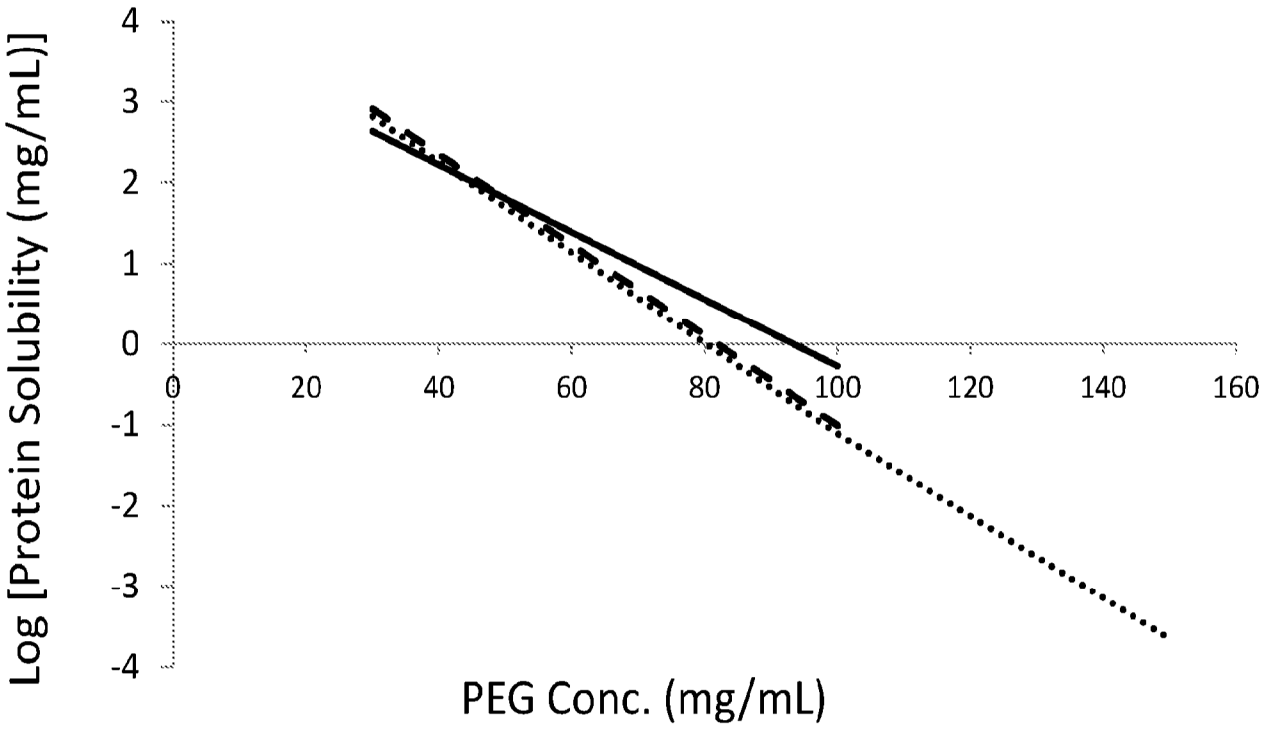

Log [Protein Solubility (mg/mL)]
4
3
2
1
0
-1
-2
-3
-4
0
20
40
60
80
100
120
140
160
PEG Conc. (mg/mL)

FIGURE 3

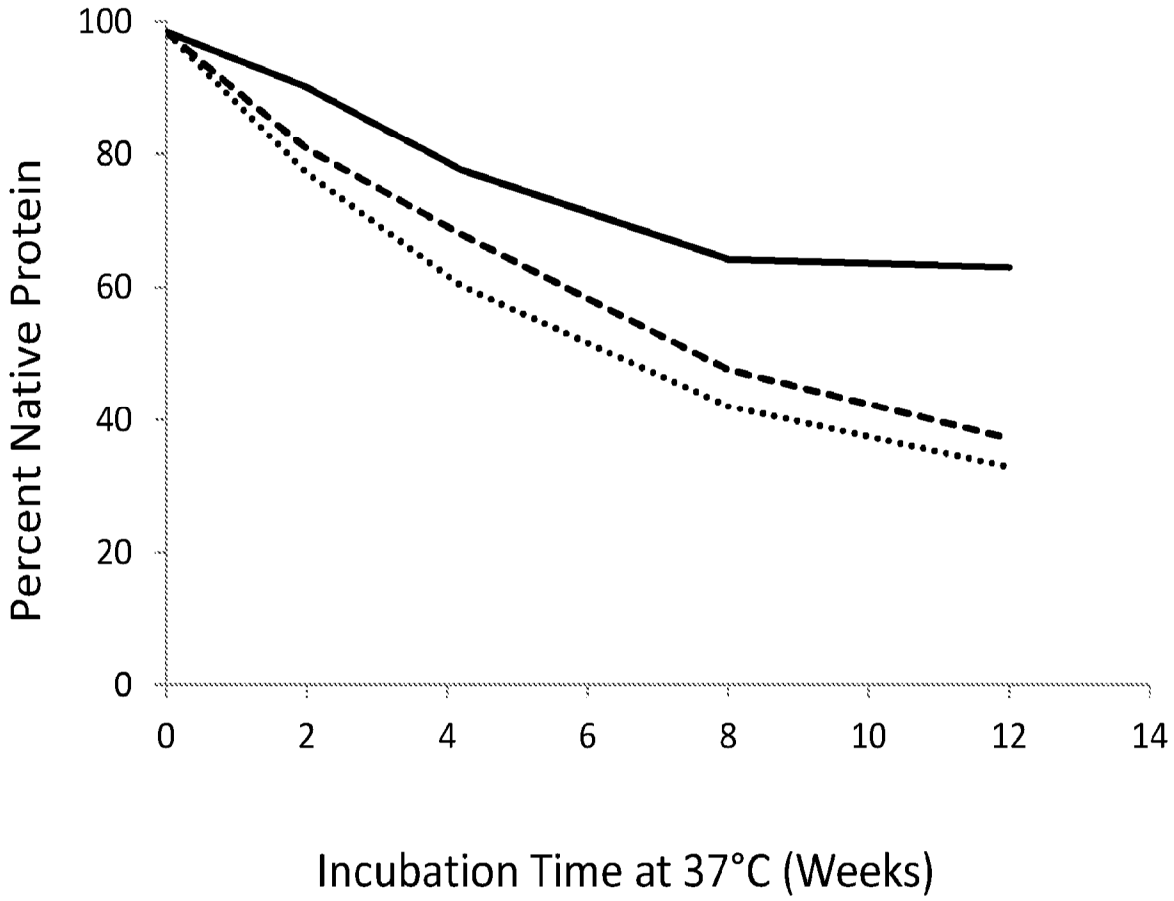

Percent Native Protein
100
80
60
40
20
0
0
2
4
6
8
10
12
14
Incubation Time at 37°C (Weeks)

FIGURE 4

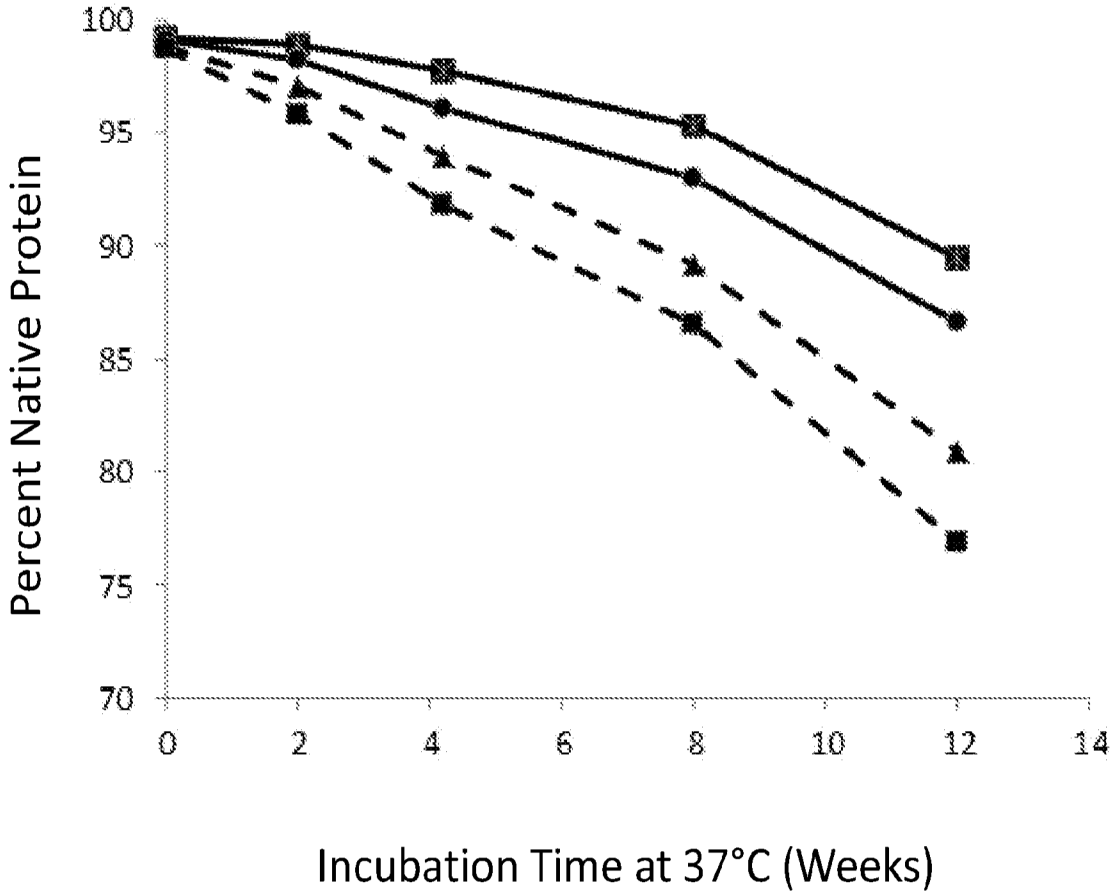

Percent Native Protein
100
95
90
85
80
75
70
0
2
4
6
8
10
12
14
Incubation Time at 37°C (Weeks)

FIGURE 5

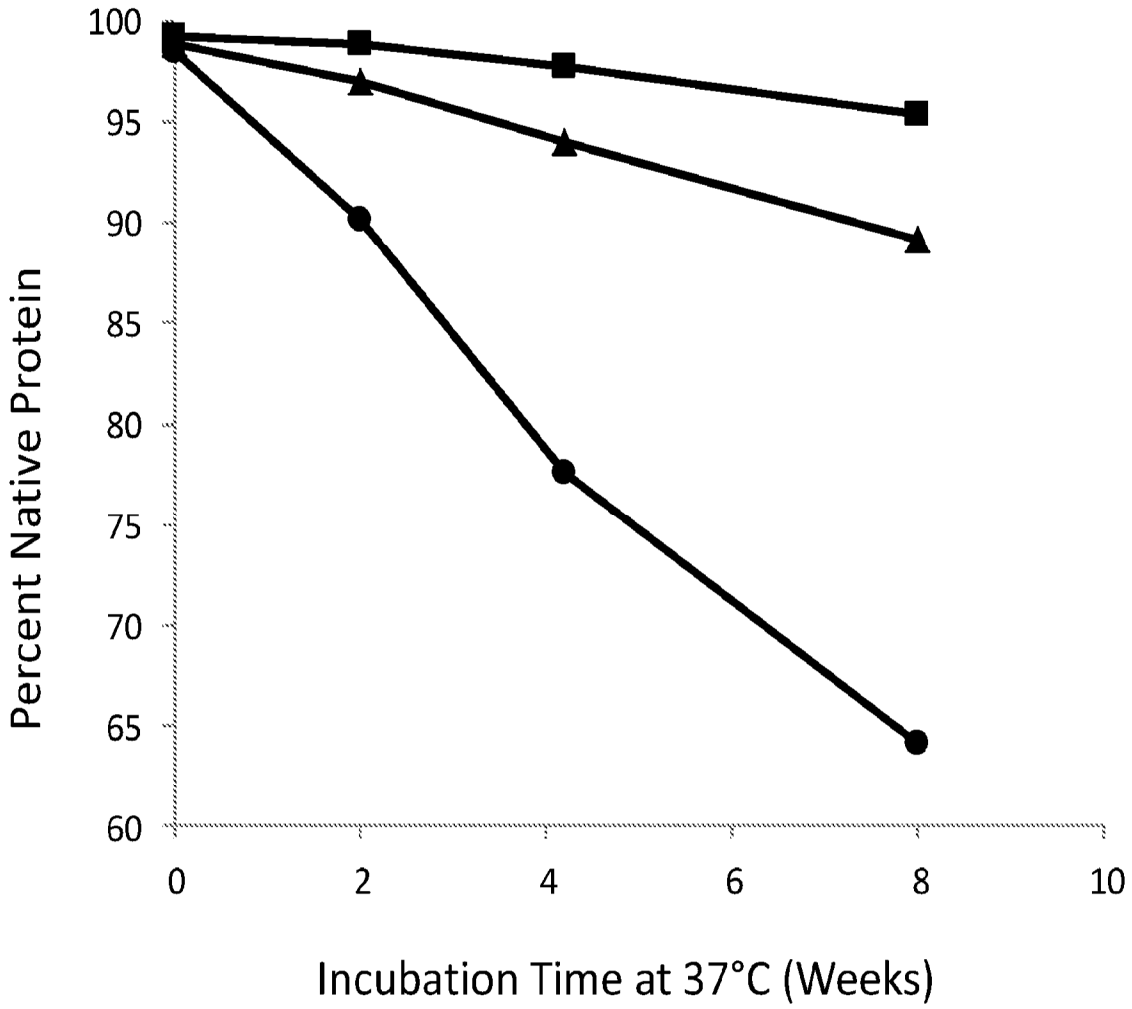

100
95
90
85
80
75
70
65
60
Percent Native Protein
0
2
4
6
8
10
Incubation Time at 37°C (Weeks)

FIGURE 6A

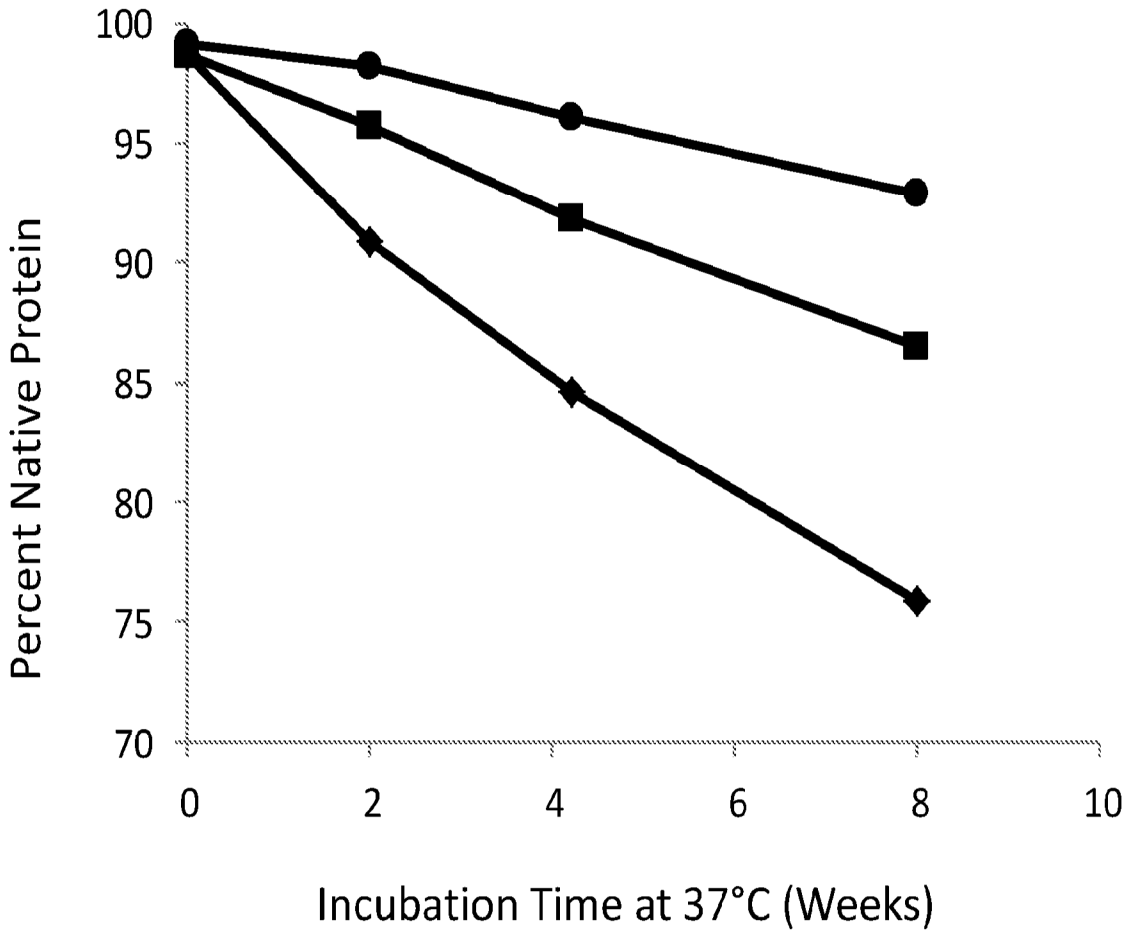

100
95
90
85
80
75
70
Percent Native Protein
0
2
4
6
8
10
Incubation Time at 37°C (Weeks)

FIGURE 6B

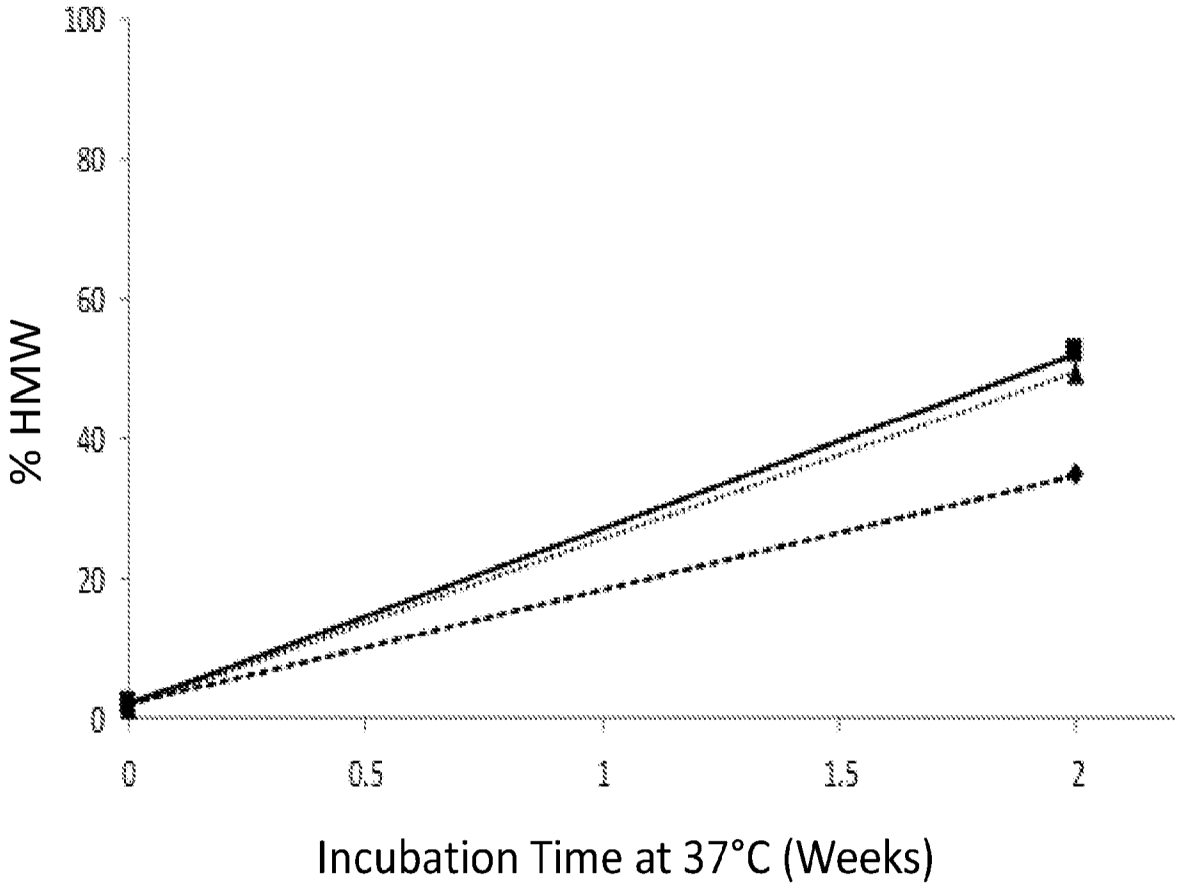


FIGURE 7

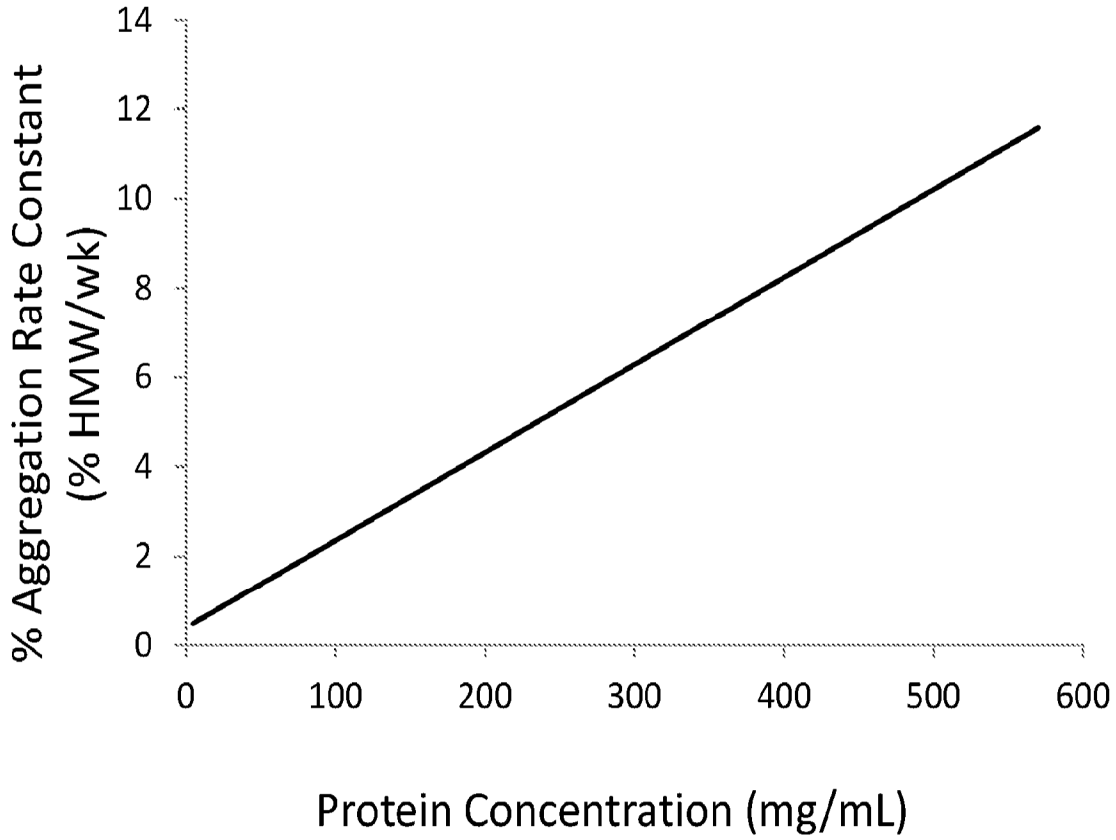

% Aggregation Rate Constant
(% HMW/wk)
14
12
10
8
6
4
2
0
0
100
200
300
400
500
600
Protein Concentration (mg/mL)

FIGURE 8

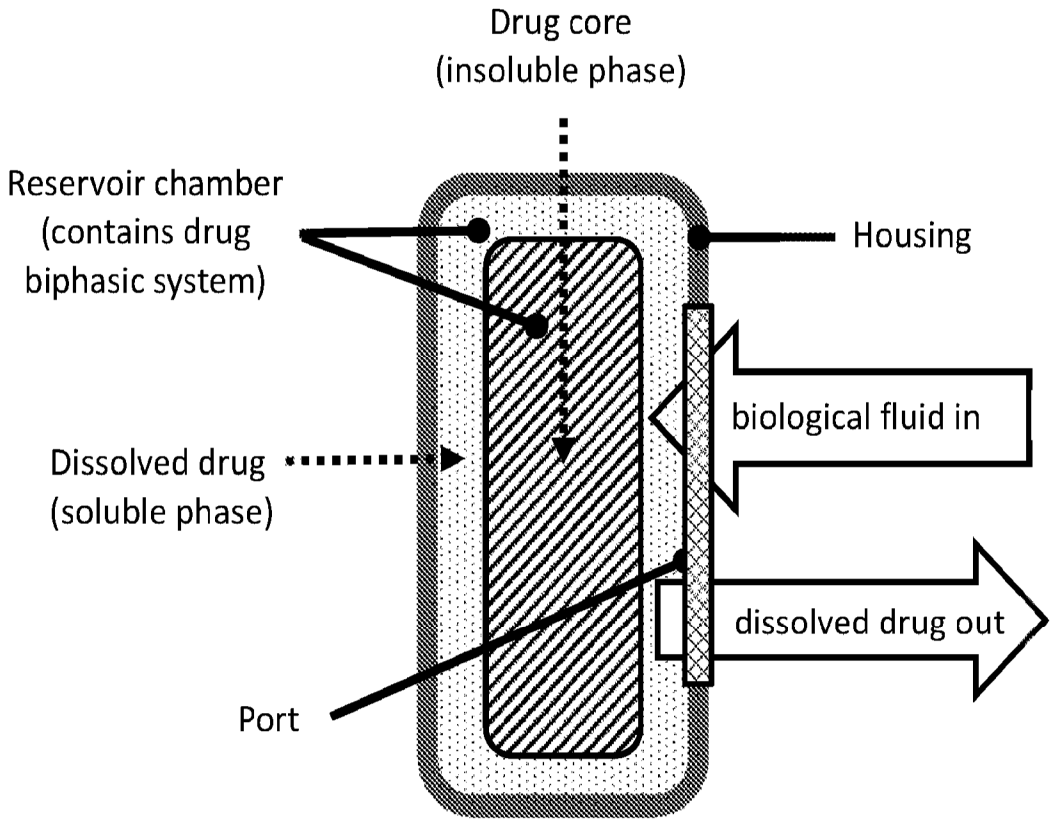

Drug core
(insoluble phase)
Reservoir chamber
(contains drug
biphasic system)
Housing
biological fluid in
Dissolved drug
(soluble phase)
dissolved drug out
Port

FIGURE 9

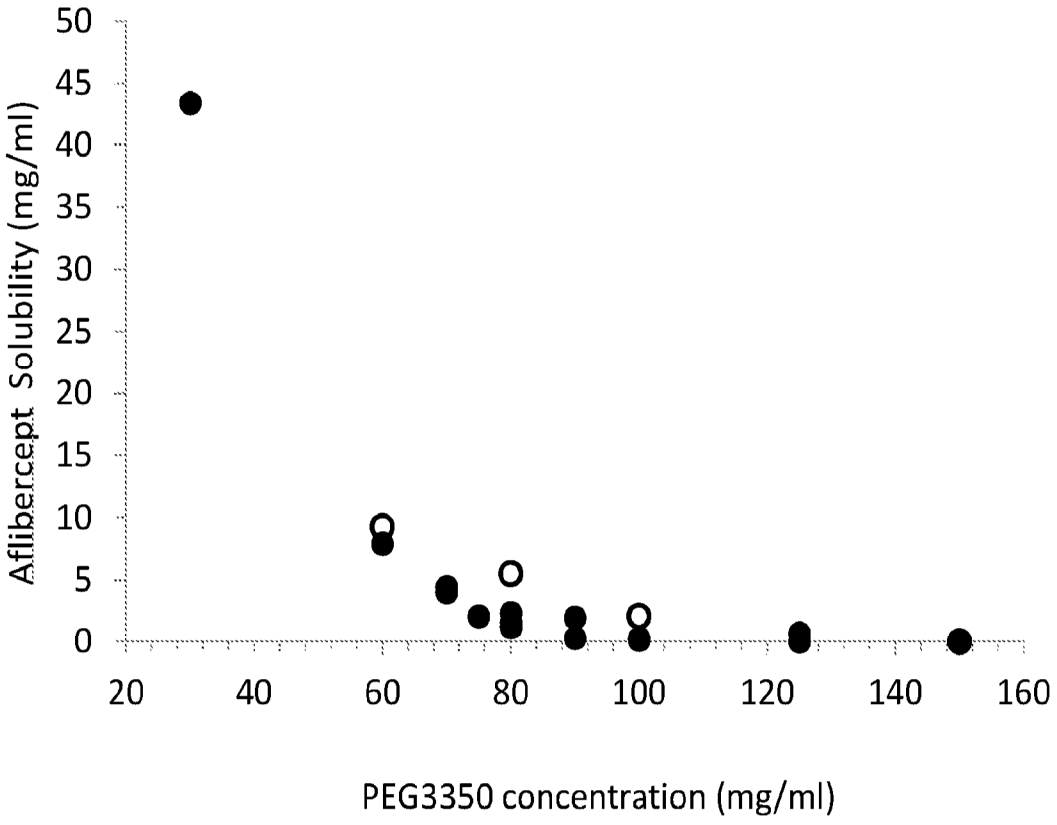

Aflibercept Solubility (mg/ml)
50
45
40
35
30
25
20
15
10
5
0
20
40
60
80
100
120
140
160
PEG3350 concentration (mg/ml)

FIGURE 10

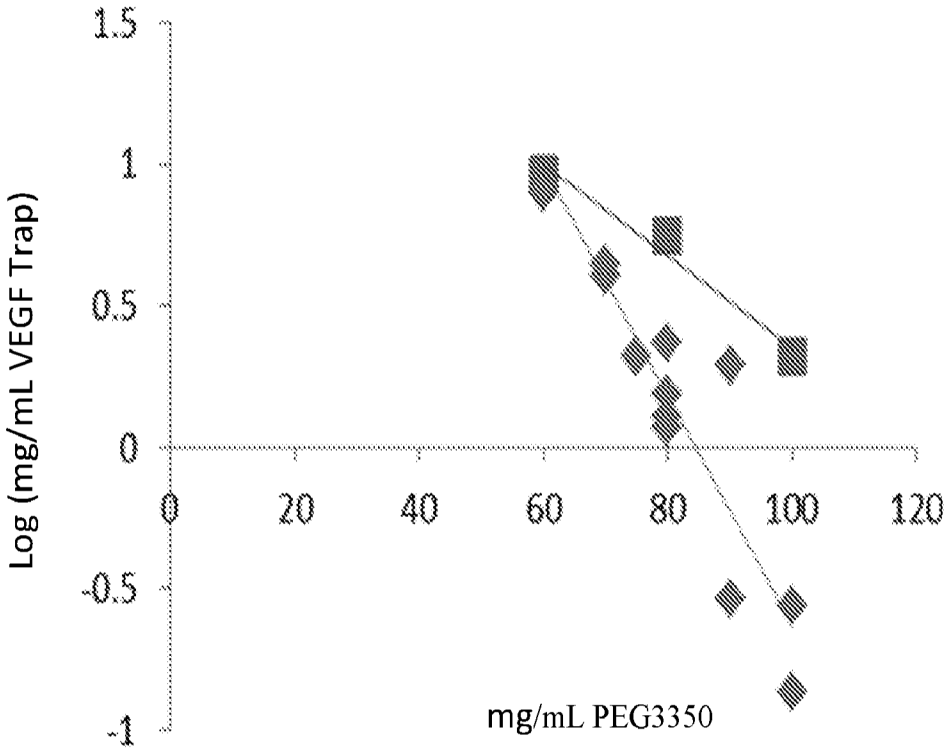

1.5
1
0.5
0
-0.5
-1
0
20
40
60
80
100
120
Log (mg/mL VEGF Trap)
mg/mL PEG3350

FIGURE 11

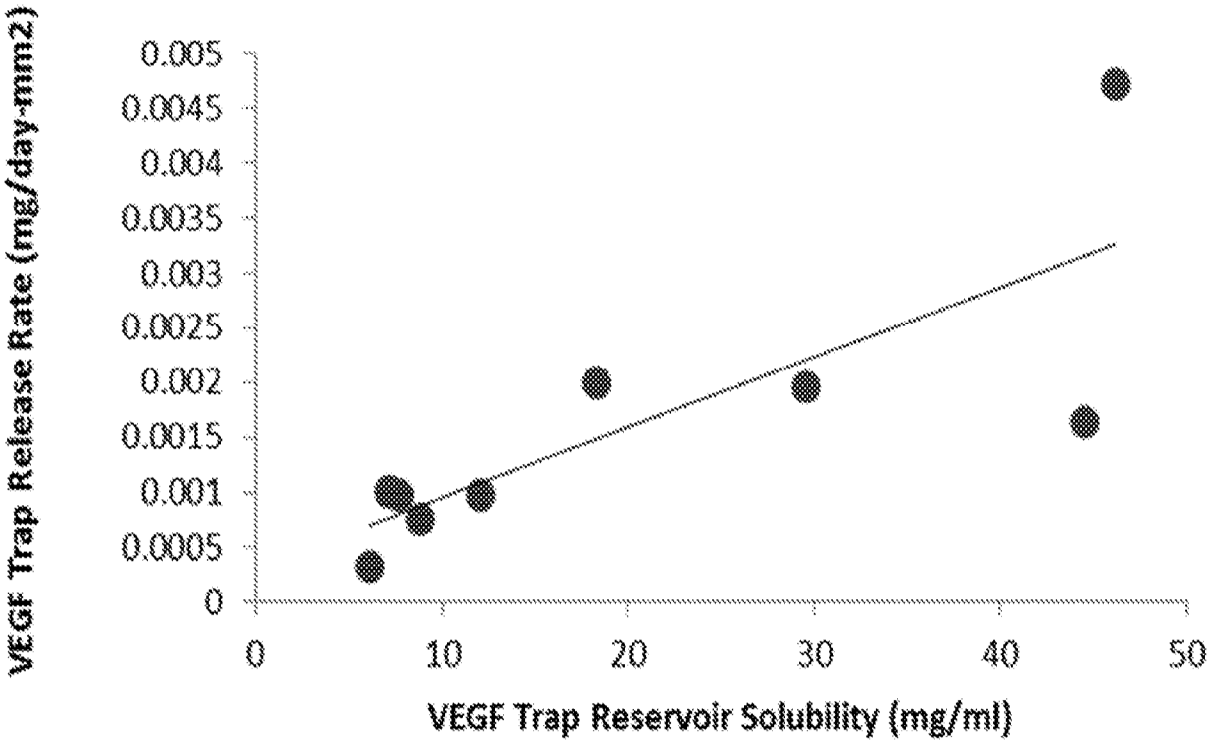

VEGF Trap Release Rate (mg/day-mm2)
0.005
0.0045
0.004
0.0035
0.003
0.0025
0.002
0.0015
0.001
0.0005
0
0
10
20
30
40
50
VEGF Trap Reservoir Solubility (mg/ml)

FIGURE 12

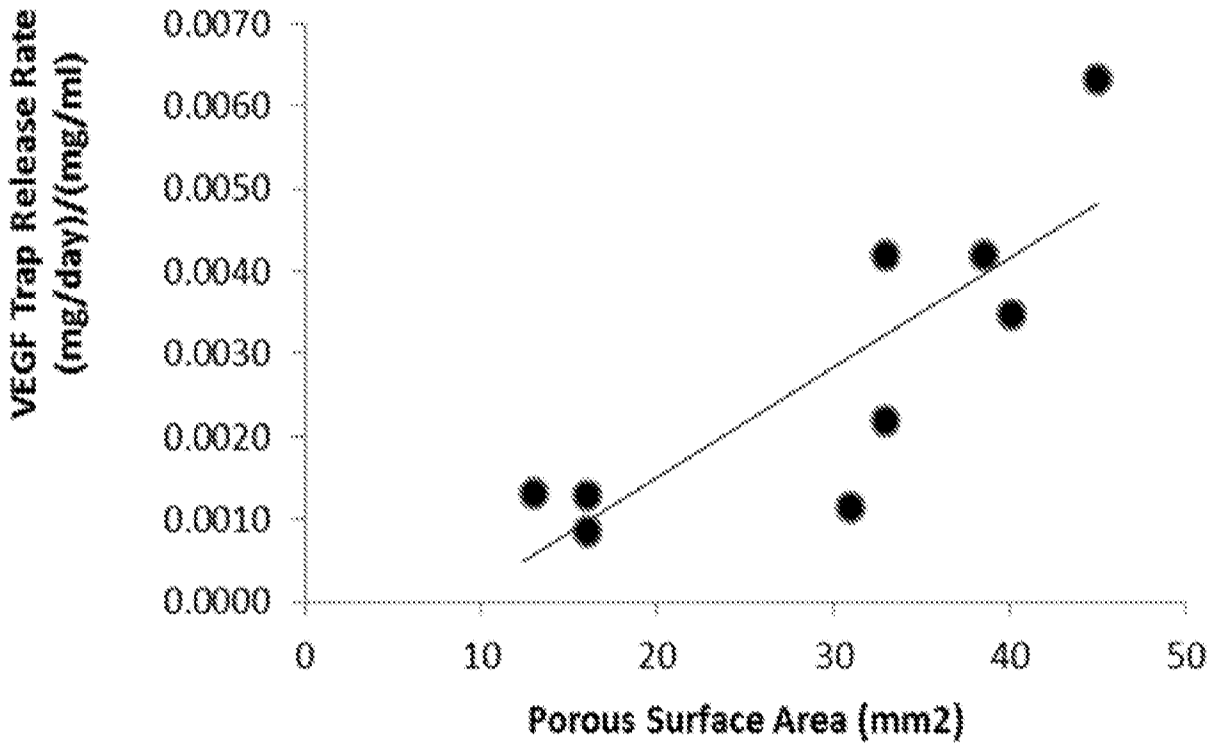

VEGF Trap Release Rate (mg/day)/(mg/ml)
0.0070
0.0060
0.0050
0.0040
0.0030
0.0020
0.0010
0.0000
0
10
20
30
40
50
Porous Surface Area (mm2)

FIGURE 13

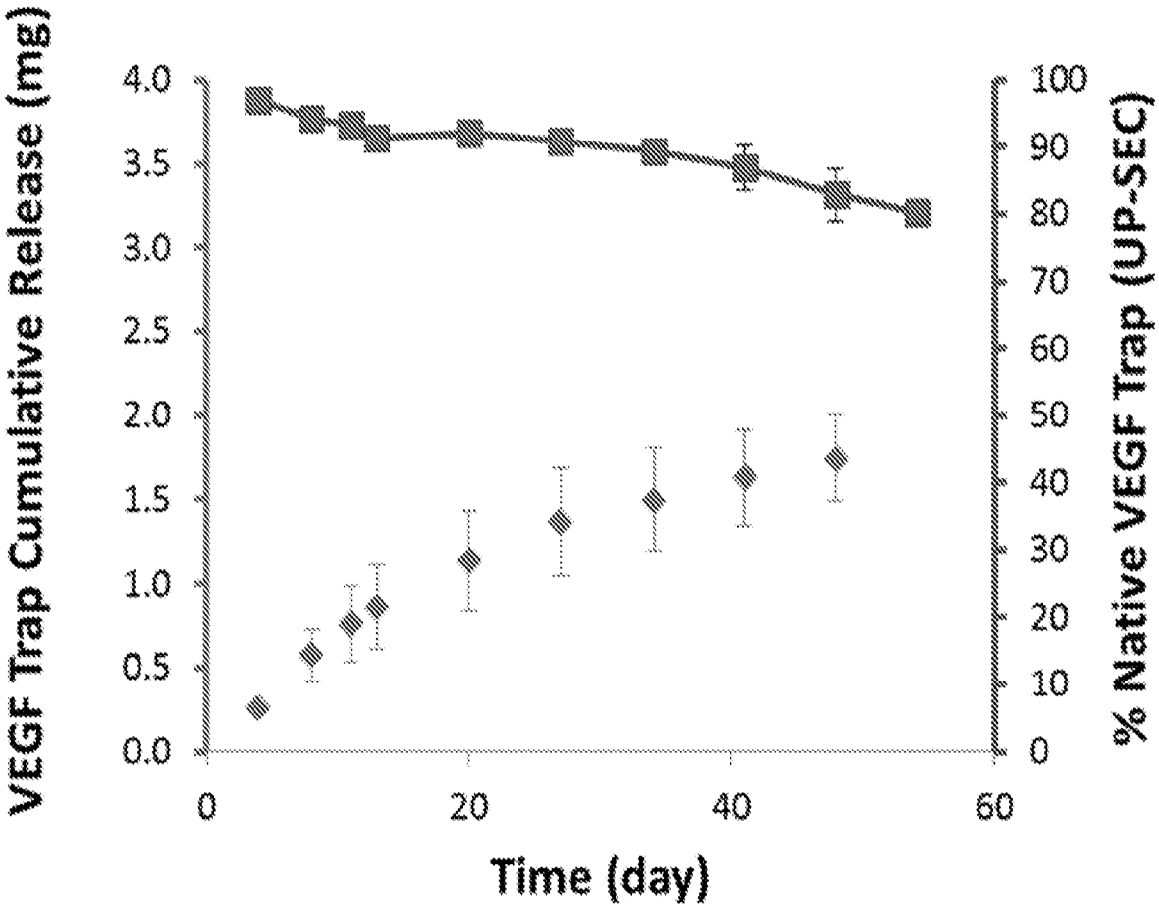

VEGF Trap Cumulative Release (mg)
4.0
3.5
3.0
2.5
2.0
1.5
1.0
0.5
0.0
0
20
40
60
Time (day)
100
90
80
70
60
50
40
30
20
10
0
% Native VEGF Trap (UP-SEC)

FIGURE 14A

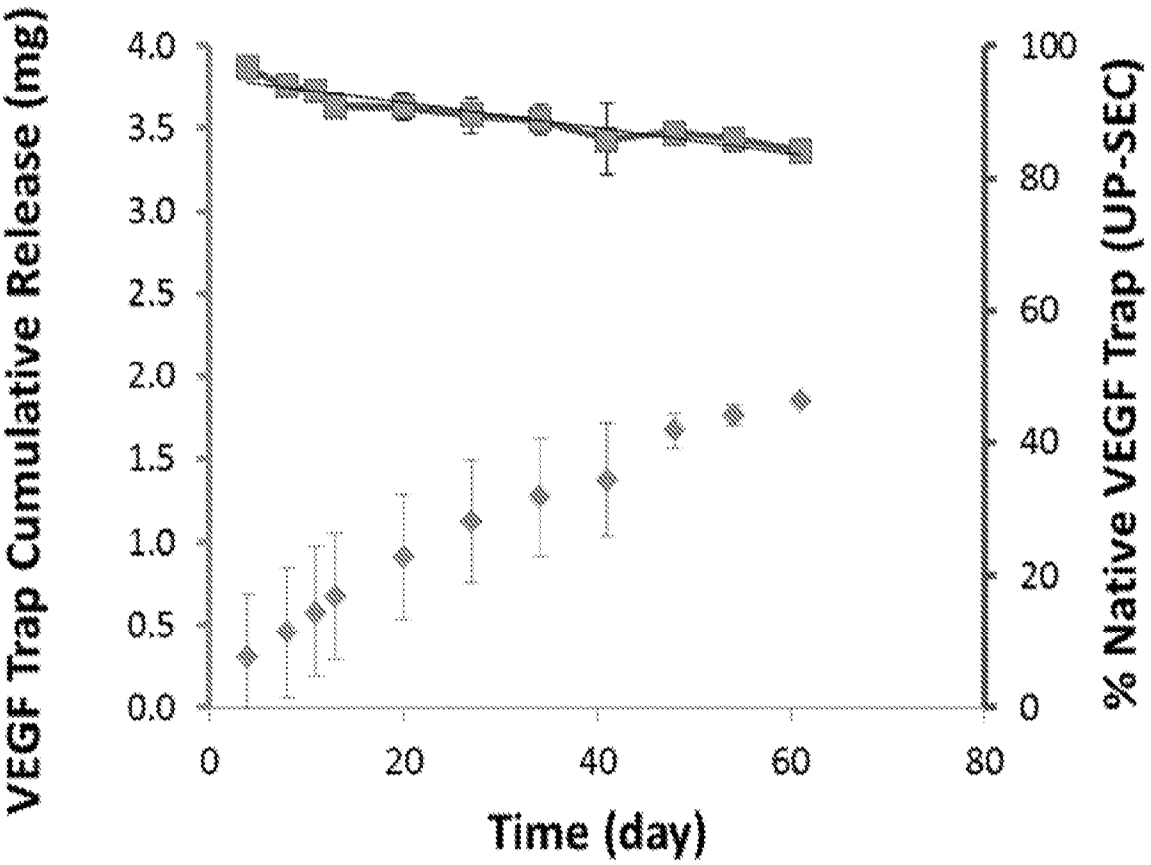

VEGF Trap Cumulative Release (mg)
4.0
3.5
3.0
2.5
2.0
1.5
1.0
0.5
0.0
0
20
40
60
80
Time (day)
100
80
60
40
20
0
% Native VEGF Trap (UP-SEC)

FIGURE 14B

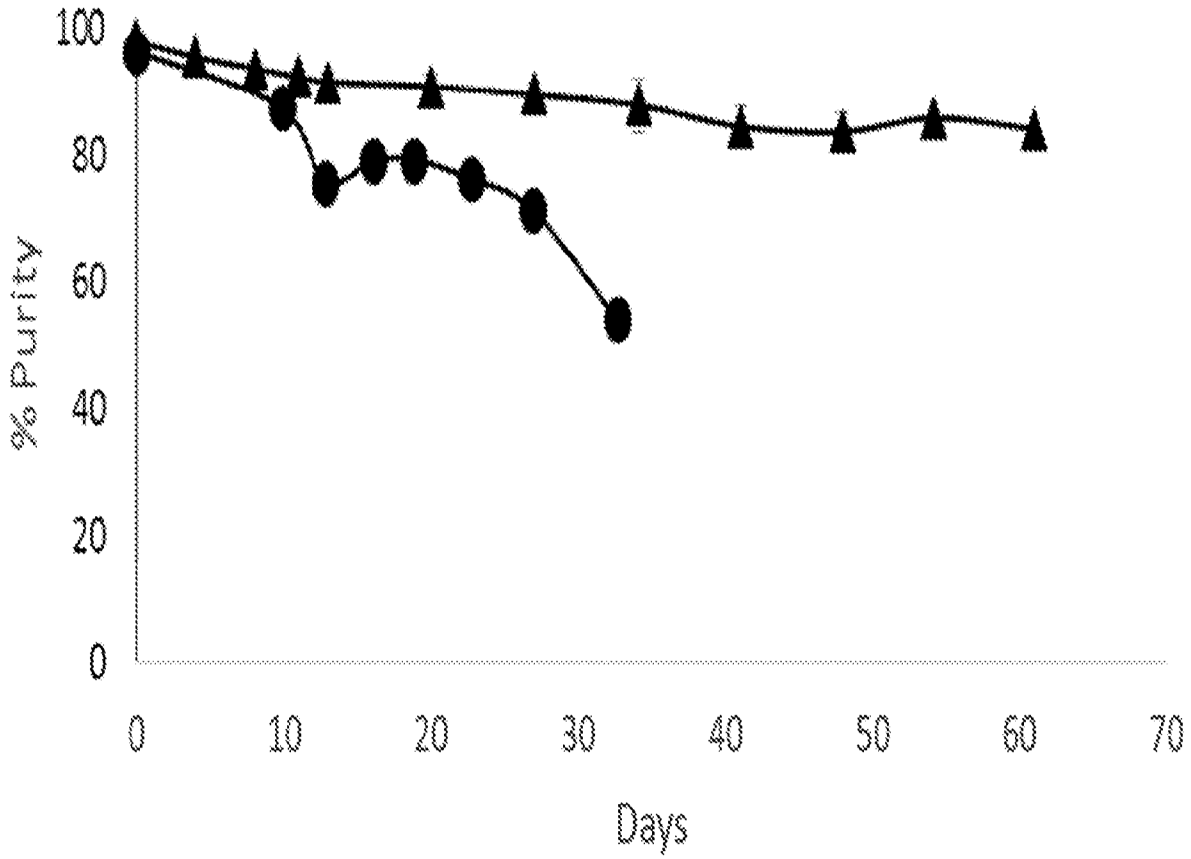


FIGURE 15