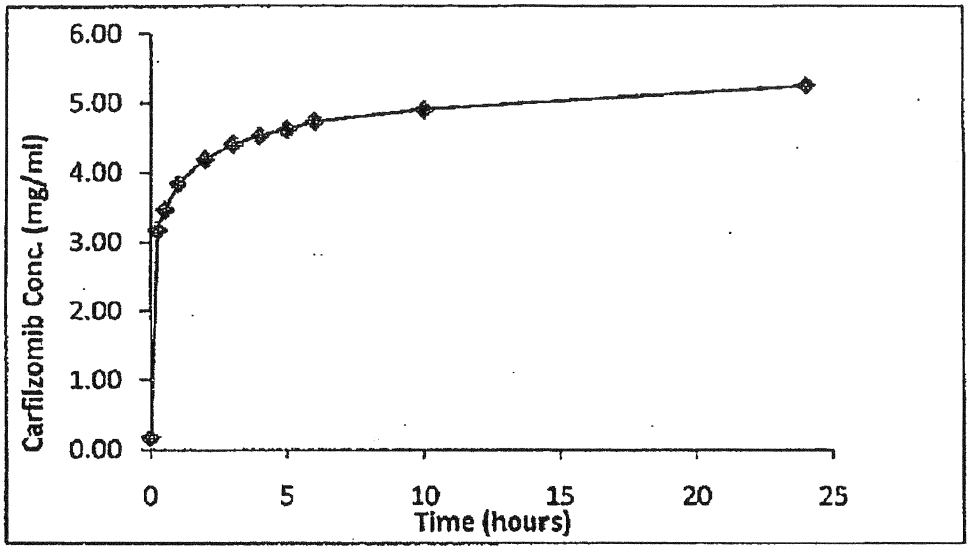

6.00
5.00
4.00
3.00
2.00
1.00
0.00
Carfilzomib Conc. (mg/ml)
0
5
10
15
20
25
Time (hours)

FIG. 1

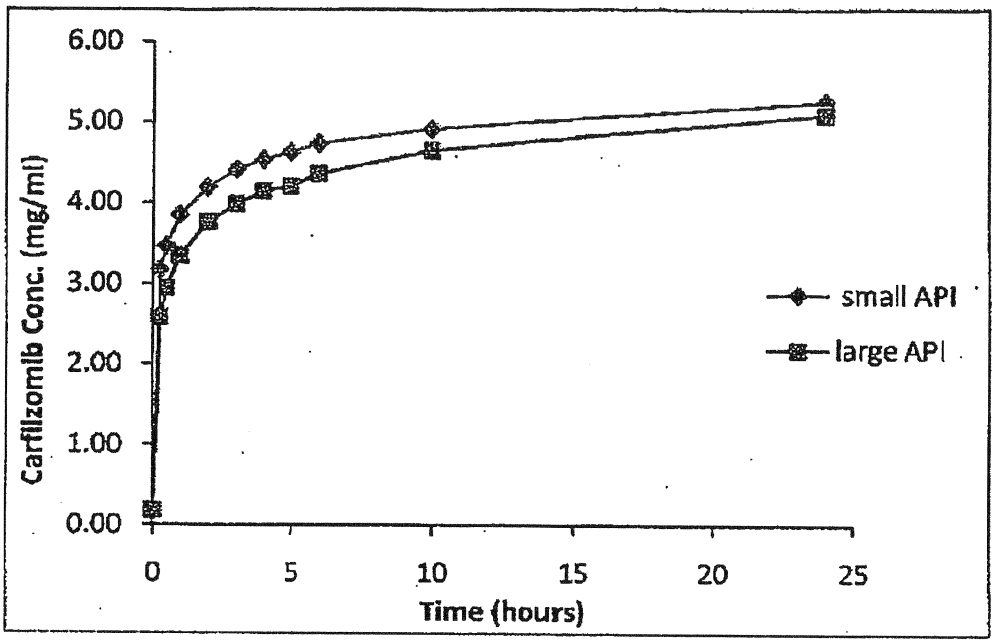

6.00
5.00
4.00
3.00
2.00
1.00
0.00
Carfilzomib Conc. (mg/ml)
0
5
10
15
20
25
Time (hours)
small API
large API

FIG. 2

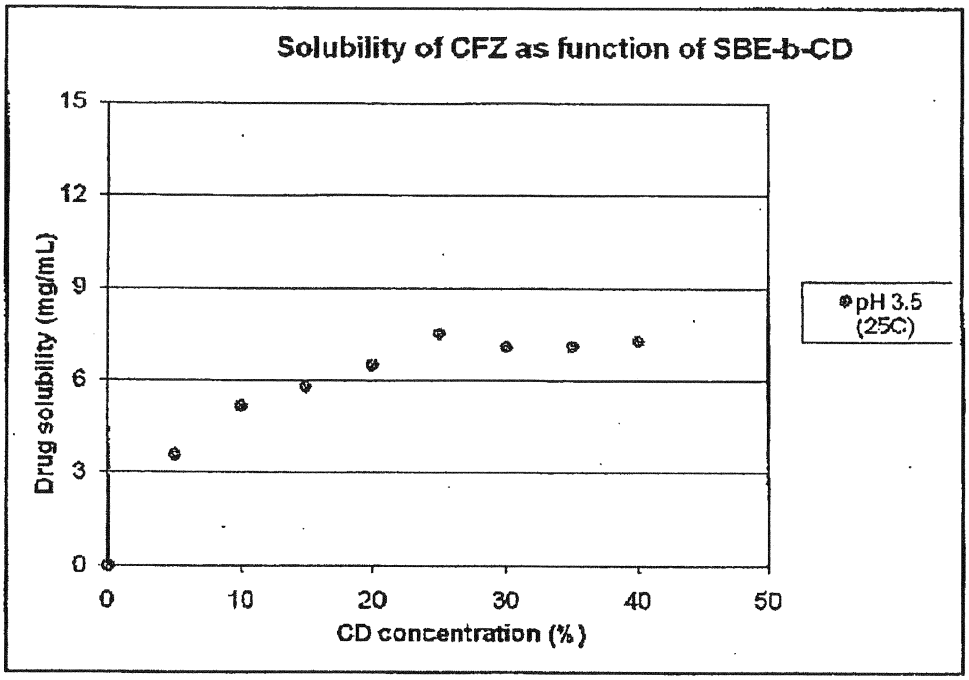

Solubility of CFZ as function of SBE-b-CD
15
12
9
6
3
0
Drug solubility (mg/mL)
0
10
20
30
40
50
CD concentration (%)
pH 3.5 (25C)

FIG. 3

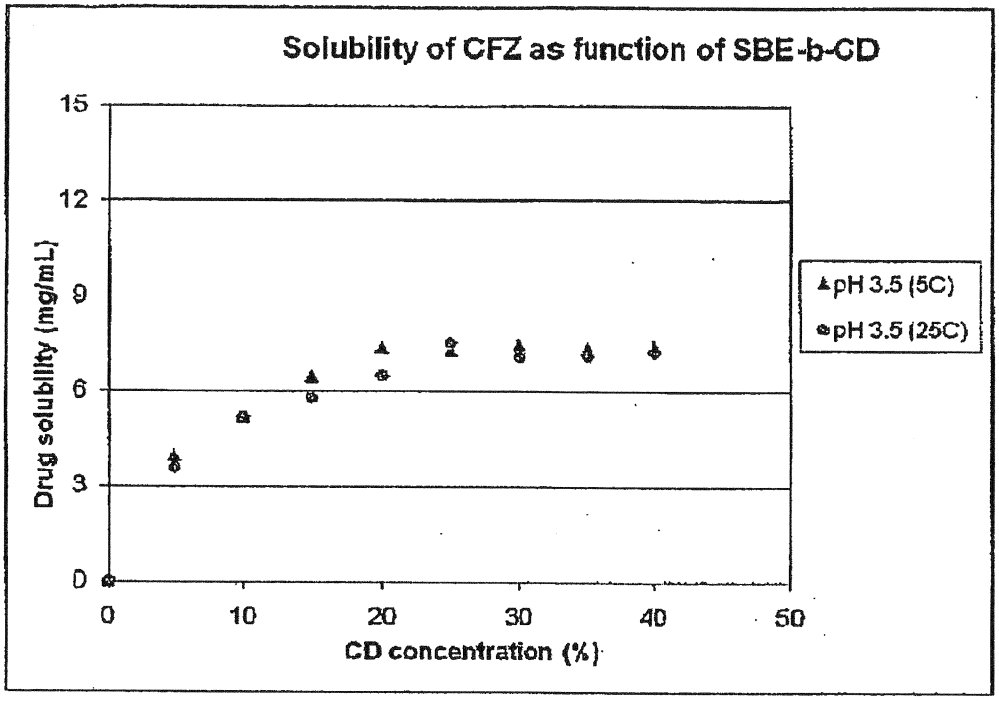

Solubility of CFZ as function of SBE-b-CD
15
12
9
6
3
0
Drug solubility (mg/mL)
0
10
20
30
40
50
CD concentration (%)
pH 3.5 (5C)
pH 3.5 (25C)

FIG. 4

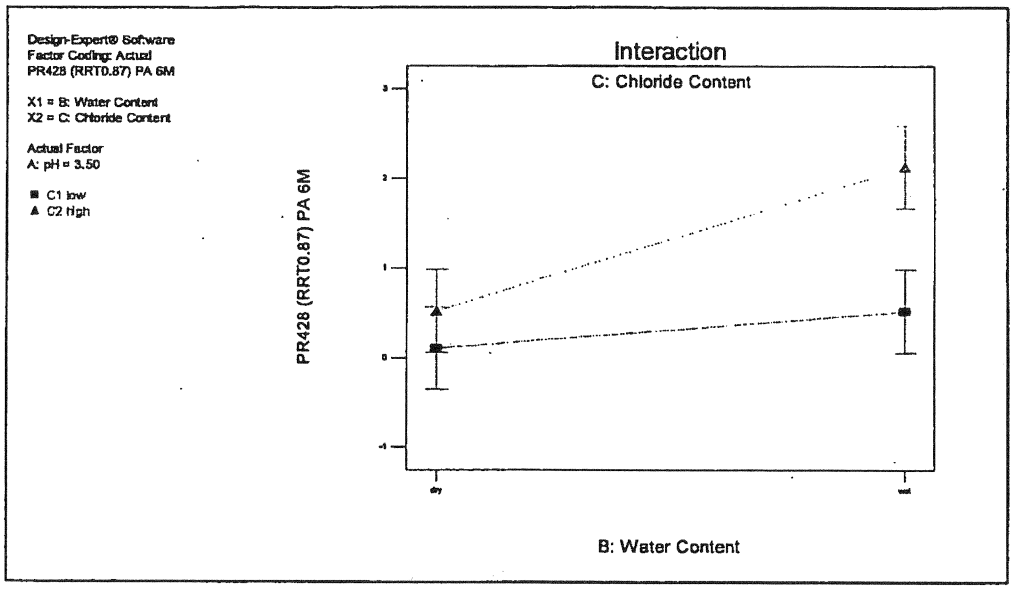

Design-Expert® Software
Factor Coding: Actual
PR428 (RRT0.87) PA 6M
X1 = B: Water Content
X2 = C: Chloride Content
Actual Factor
A: pH = 3.50
C1 low
C2 high
Interaction
C: Chloride Content
PR428 (RRT0.87) PA 6M
B: Water Content

FIG. 5

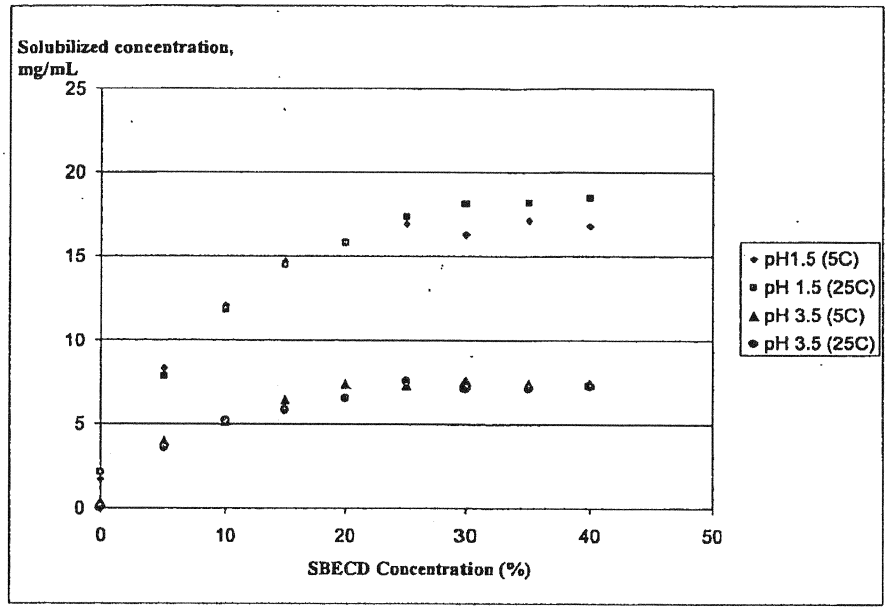

Solubilized concentration, mg/mL
25
20
15
10
5
0
0
10
20
30
40
50
SBECD Concentration (%)
pH1.5 (5C)
pH 1.5 (25C)
pH 3.5 (5C)
pH 3.5 (25C)

FIG. 6

FIG. 7

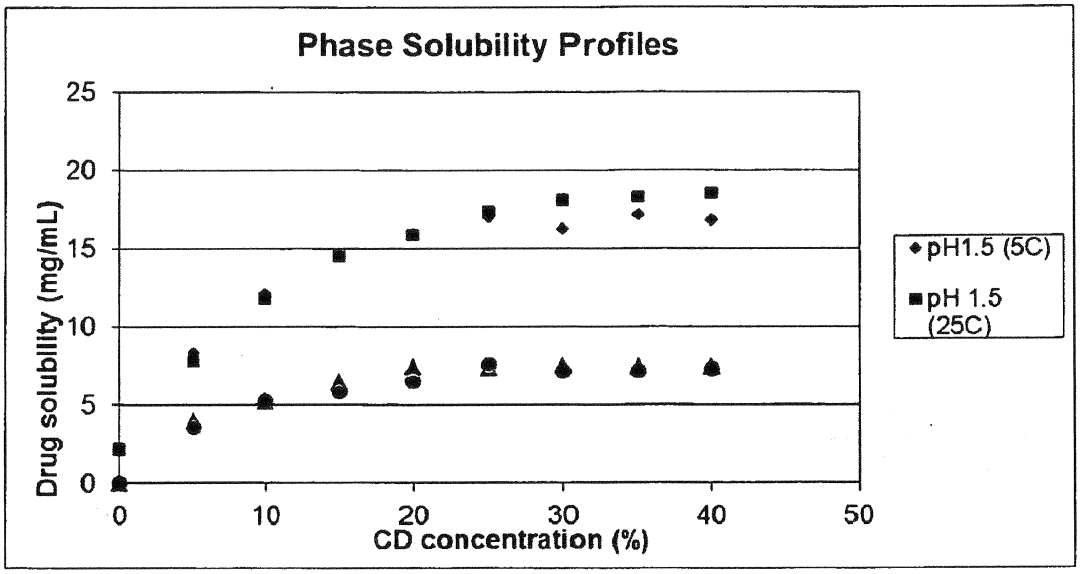

Phase Solubility Profiles
Drug solubility (mg/mL)
25
20
15
10
5
0
0
10
20
30
40
50
CD concentration (%)
pH1.5 (5C)
pH 1.5 (25C)

FIG. 8

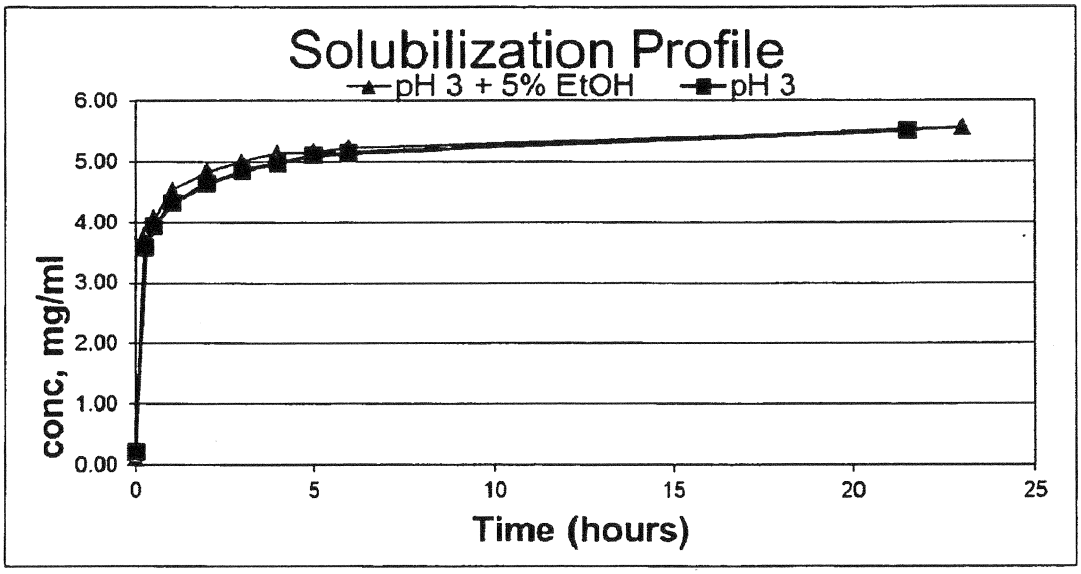

Solubilization Profile
pH 3 + 5% EtOH
pH 3
conc, mg/ml
6.00
5.00
4.00
3.00
2.00
1.00
0.00
0
5
10
15
20
25
Time (hours)

FIG. 9

Molar solubilized carfilzomib versus complexation indexed free cyclodextrin

