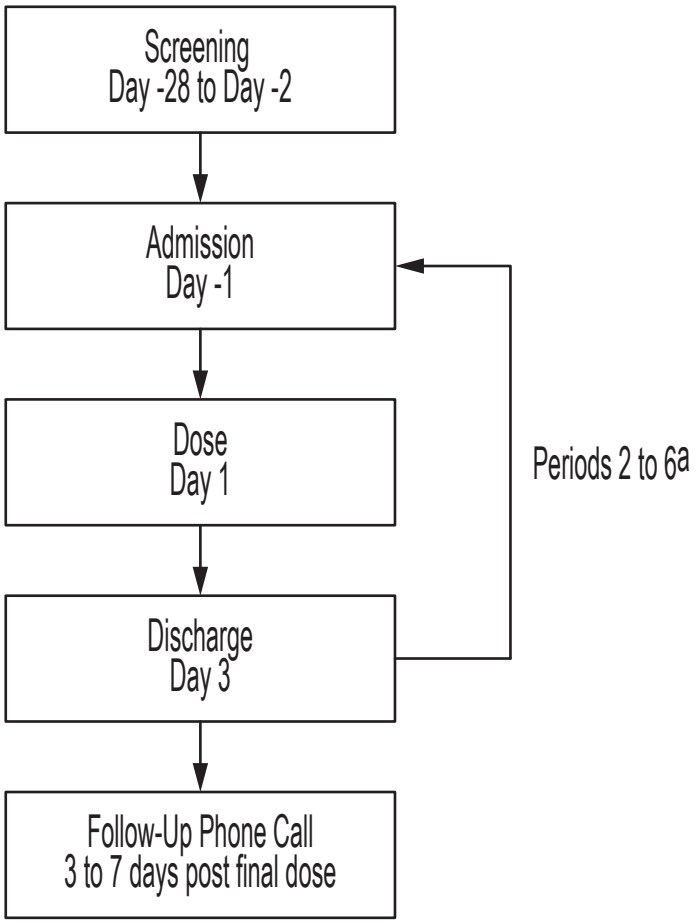

Screening
Day -28 to Day -2
Admission
Day -1
Dose
Day 1
Discharge
Day 3
Follow-Up Phone Call
3 to 7 days post final dose
Periods 2 to 6a

FIG. 1

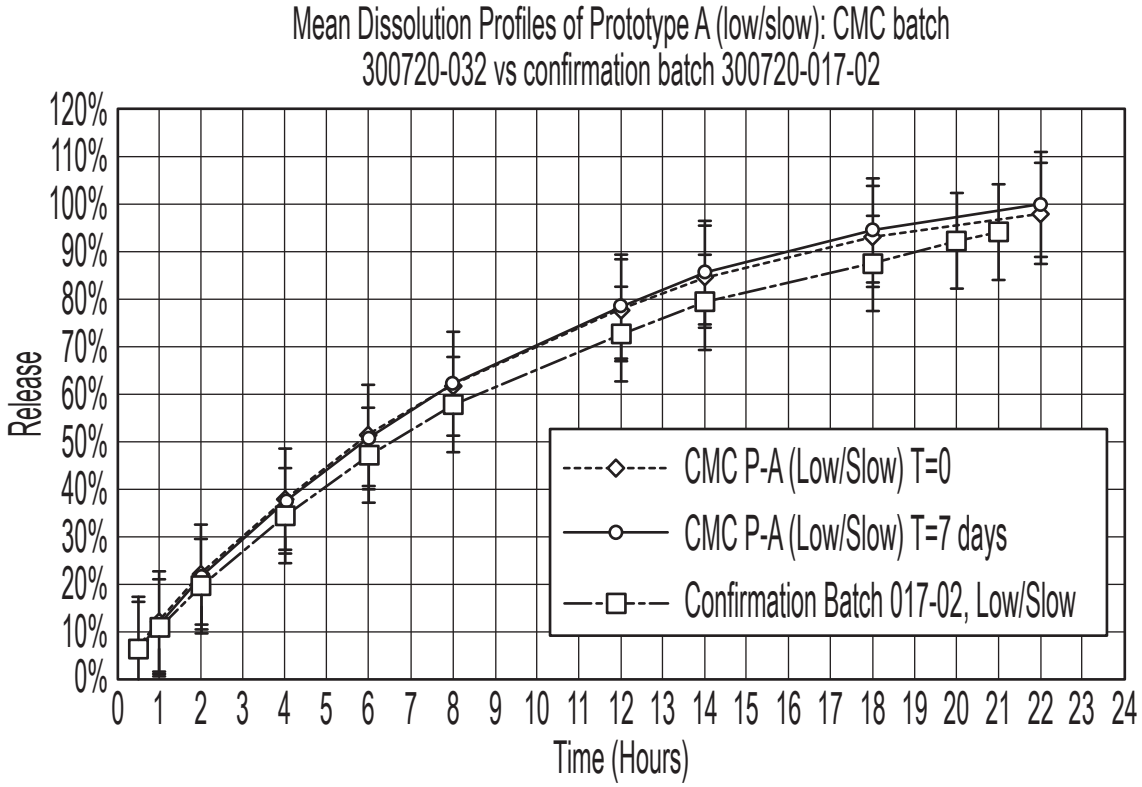

Mean Dissolution Profiles of Prototype A (low/slow): CMC batch 300720-032 vs confirmation batch 300720-017-02
Release
120%
110%
100%
90%
80%
70%
60%
50%
40%
30%
20%
10%
0%
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24
Time (Hours)
CMC P-A (Low/Slow) T=0
CMC P-A (Low/Slow) T=7 days
Confirmation Batch 017-02, Low/Slow

FIG. 2

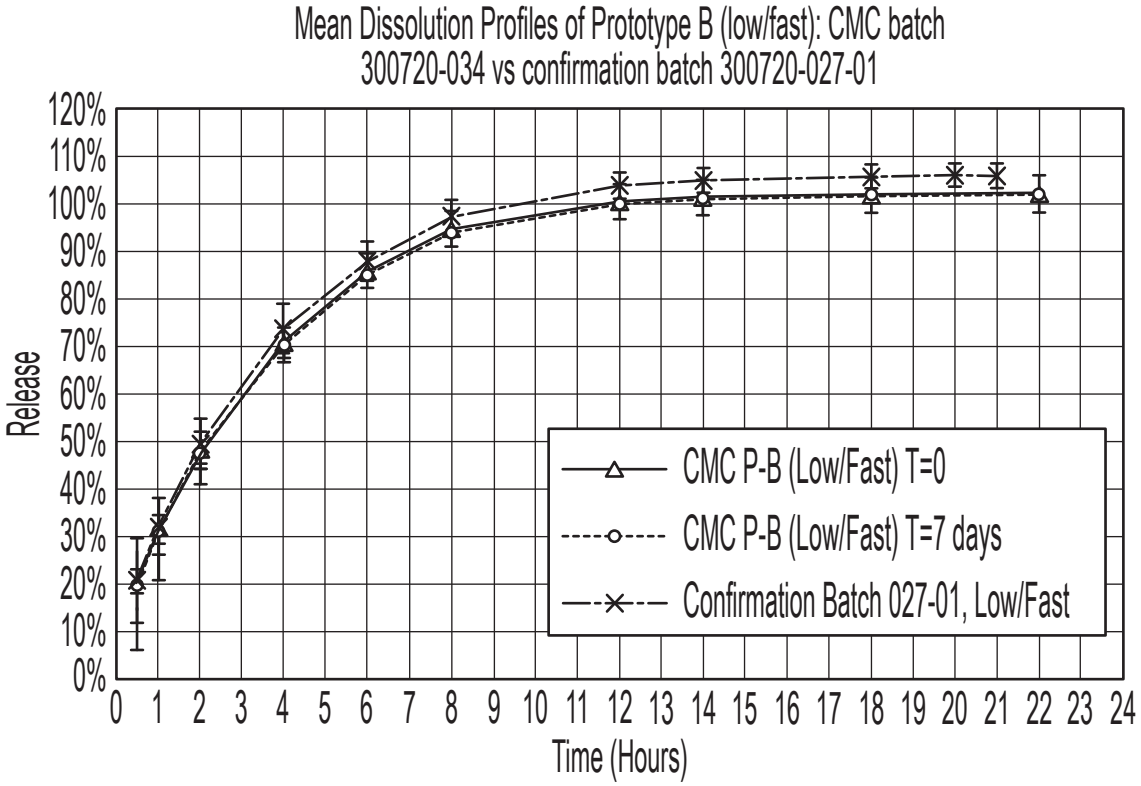

Mean Dissolution Profiles of Prototype B (low/fast): CMC batch 300720-034 vs confirmation batch 300720-027-01
Release
120%
110%
100%
90%
80%
70%
60%
50%
40%
30%
20%
10%
0%
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24
Time (Hours)
CMC P-B (Low/Fast) T=0
CMC P-B (Low/Fast) T=7 days
Confirmation Batch 027-01, Low/Fast

FIG. 3

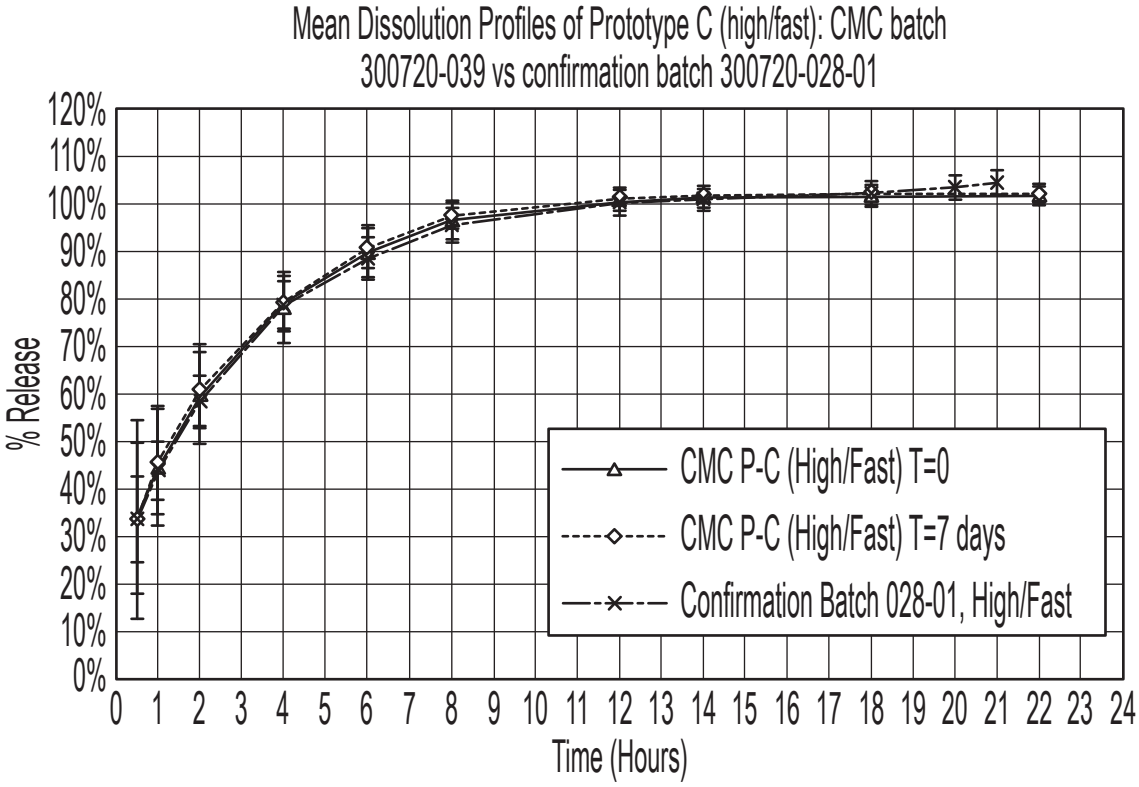

Mean Dissolution Profiles of Prototype C (high/fast): CMC batch 300720-039 vs confirmation batch 300720-028-01
% Release
120%
110%
100%
90%
80%
70%
60%
50%
40%
30%
20%
10%
0%
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24
Time (Hours)
CMC P-C (High/Fast) T=0
CMC P-C (High/Fast) T=7 days
Confirmation Batch 028-01, High/Fast

FIG. 4

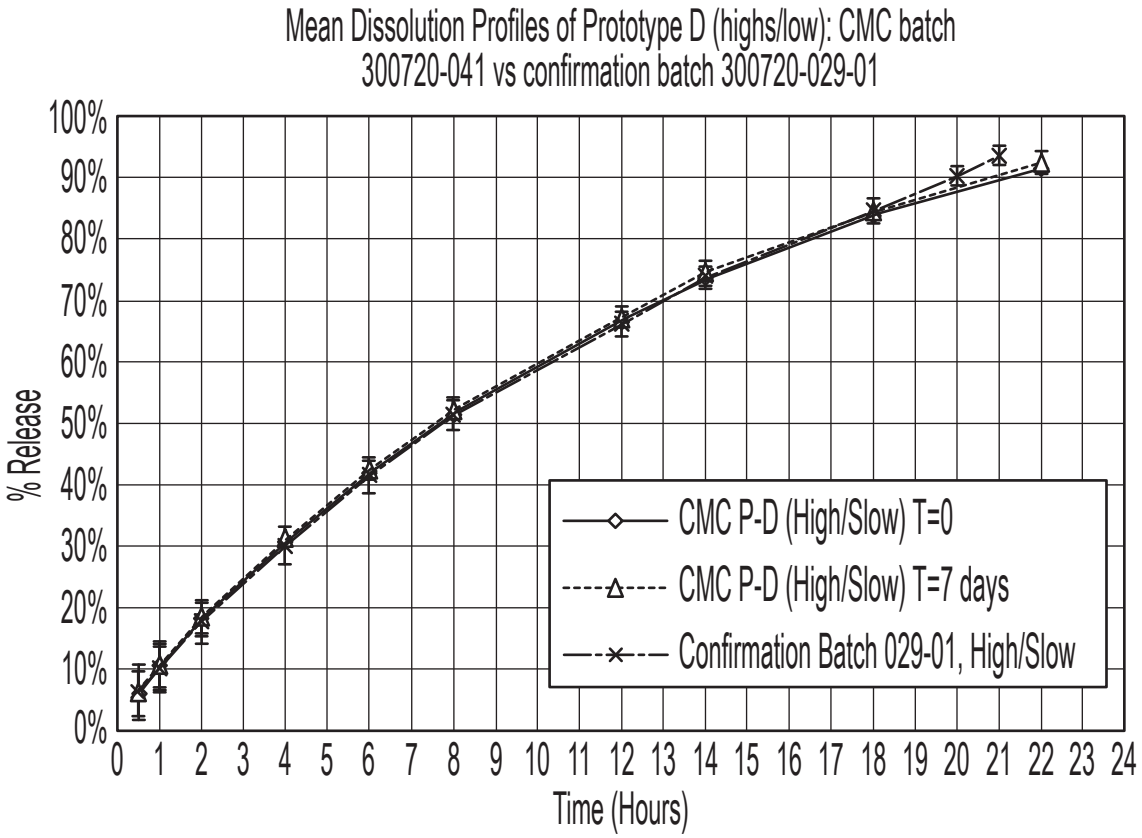

Mean Dissolution Profiles of Prototype D (highs/low): CMC batch 300720-041 vs confirmation batch 300720-029-01
% Release
100%
90%
80%
70%
60%
50%
40%
30%
20%
10%
0%
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24
Time (Hours)
CMC P-D (High/Slow) T=0
CMC P-D (High/Slow) T=7 days
Confirmation Batch 029-01, High/Slow

FIG. 5

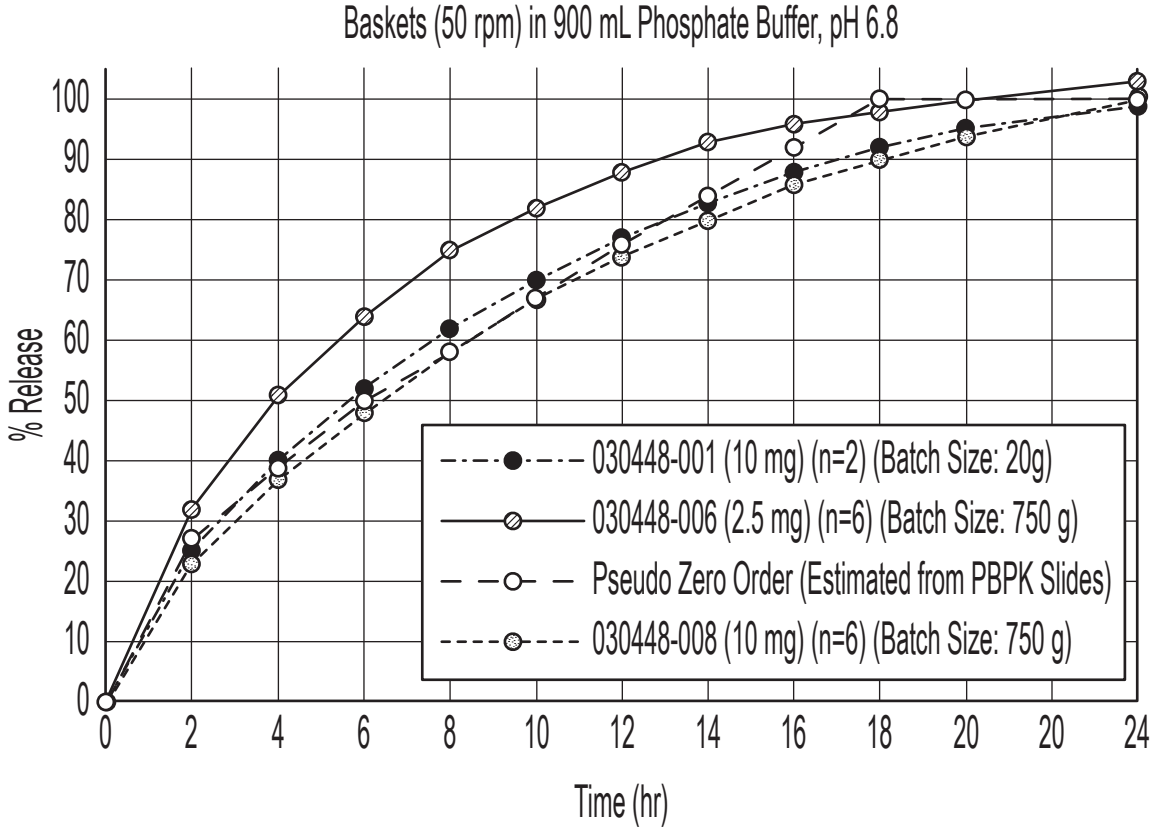

Baskets (50 rpm) in 900 mL Phosphate Buffer, pH 6.8
% Release
Time (hr)
0
10
20
30
40
50
60
70
80
90
100
0
2
4
6
8
10
12
14
16
18
20
20
24
030448-001 (10 mg) (n=2) (Batch Size: 20g)
030448-006 (2.5 mg) (n=6) (Batch Size: 750 g)
Pseudo Zero Order (Estimated from PBPK Slides)
030448-008 (10 mg) (n=6) (Batch Size: 750 g)

FIG. 6

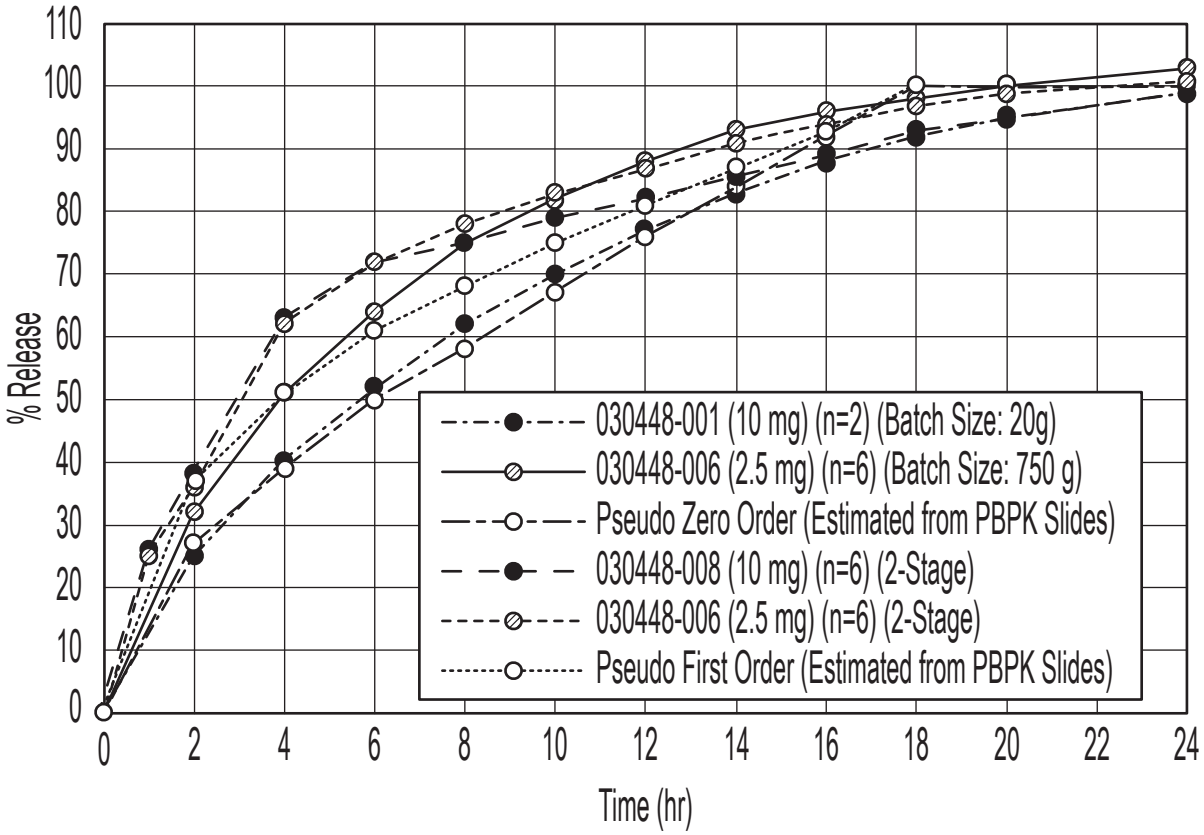

110
100
90
80
70
60
50
40
30
20
10
0
% Release
0
2
4
6
8
10
12
14
16
18
20
22
24
Time (hr)
030448-001 (10 mg) (n=2) (Batch Size: 20g)
030448-006 (2.5 mg) (n=6) (Batch Size: 750 g)
Pseudo Zero Order (Estimated from PBPK Slides)
030448-008 (10 mg) (n=6) (2-Stage)
030448-006 (2.5 mg) (n=6) (2-Stage)
Pseudo First Order (Estimated from PBPK Slides)

FIG. 7