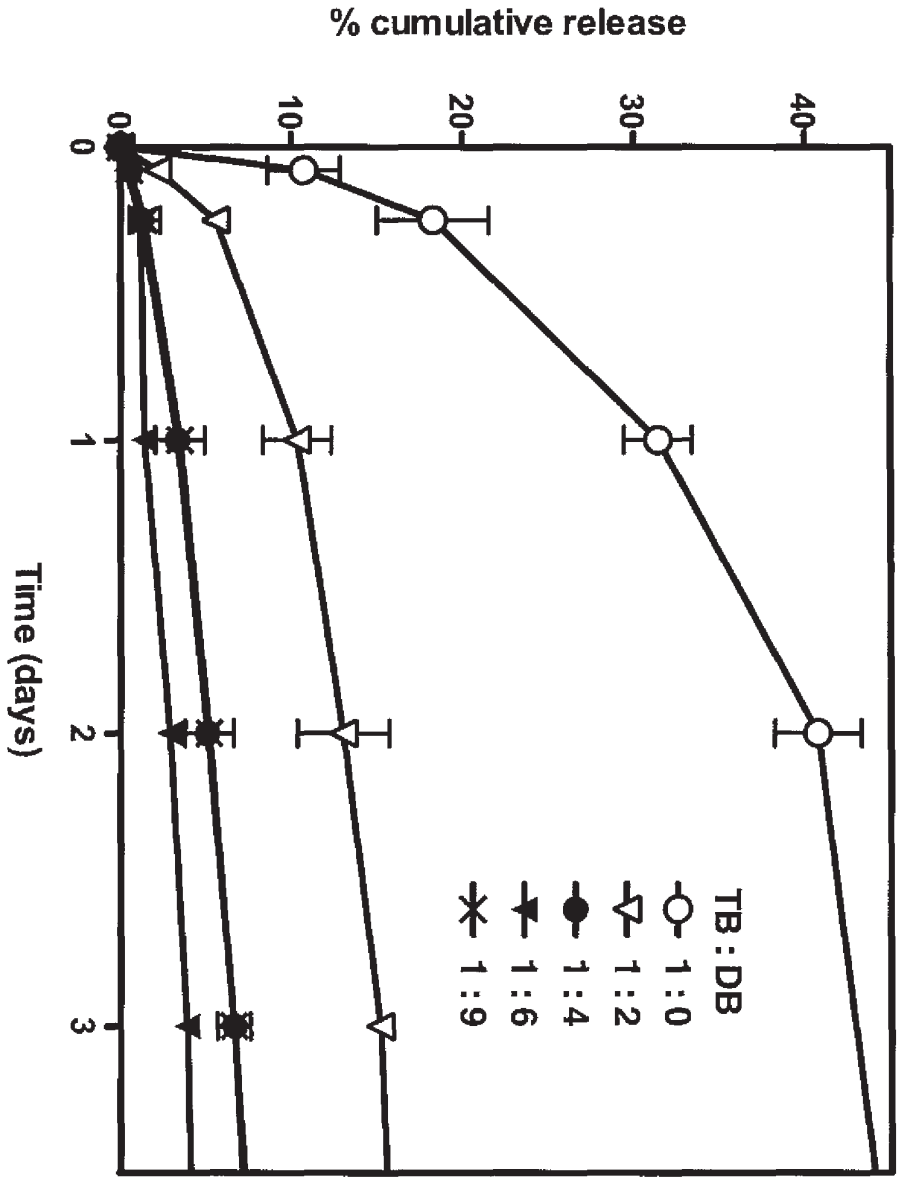

% cumulative release
0
10
20
30
40
Time (days)
0
1
2
3
TB : DB
1 : 0
1 : 2
1 : 4
1 : 6
1 : 9

FIGURE 1

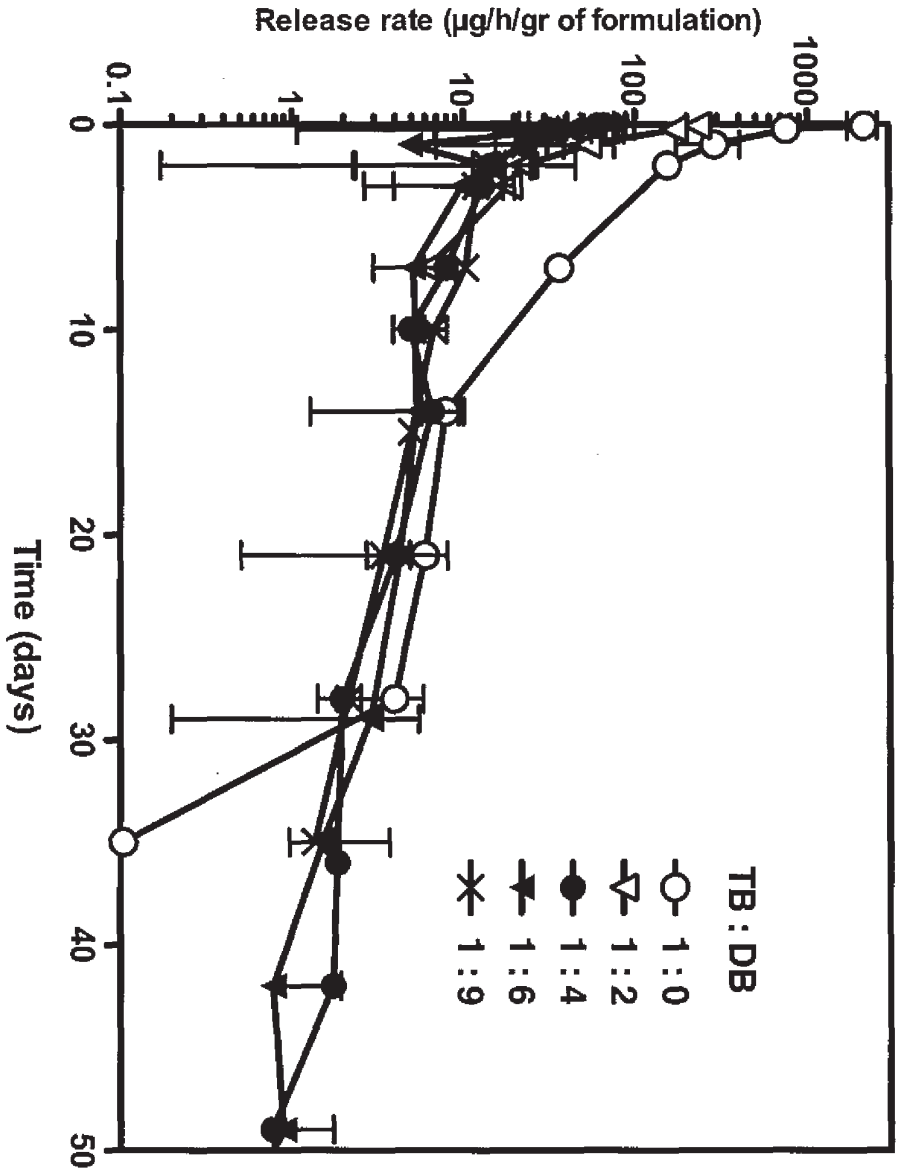

Release rate (µg/h/gr of formulation)
0.1
1
10
100
1000
0
10
20
30
40
50
Time (days)
TB : DB
1 : 0
1 : 2
1 : 4
1 : 6
1 : 9

FIGURE 2

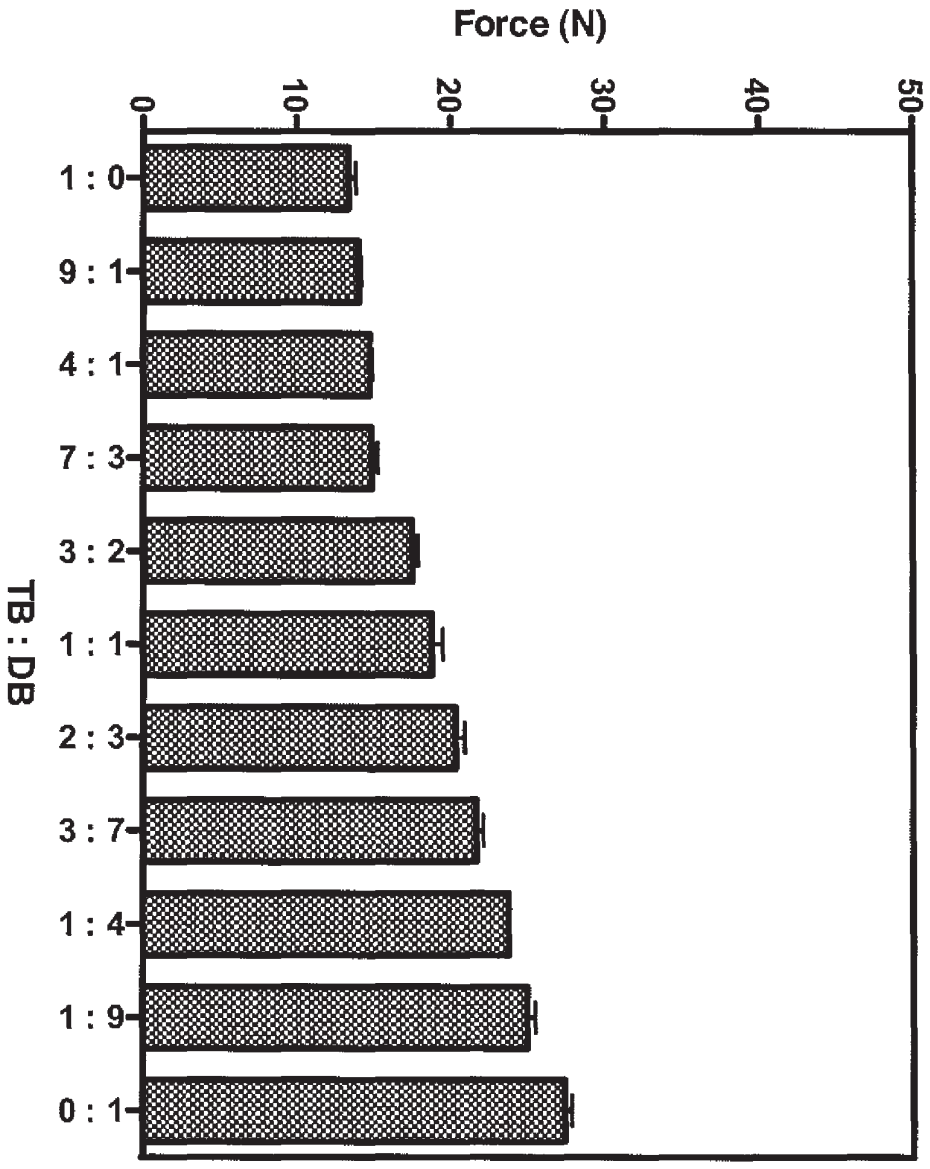


FIGURE 3

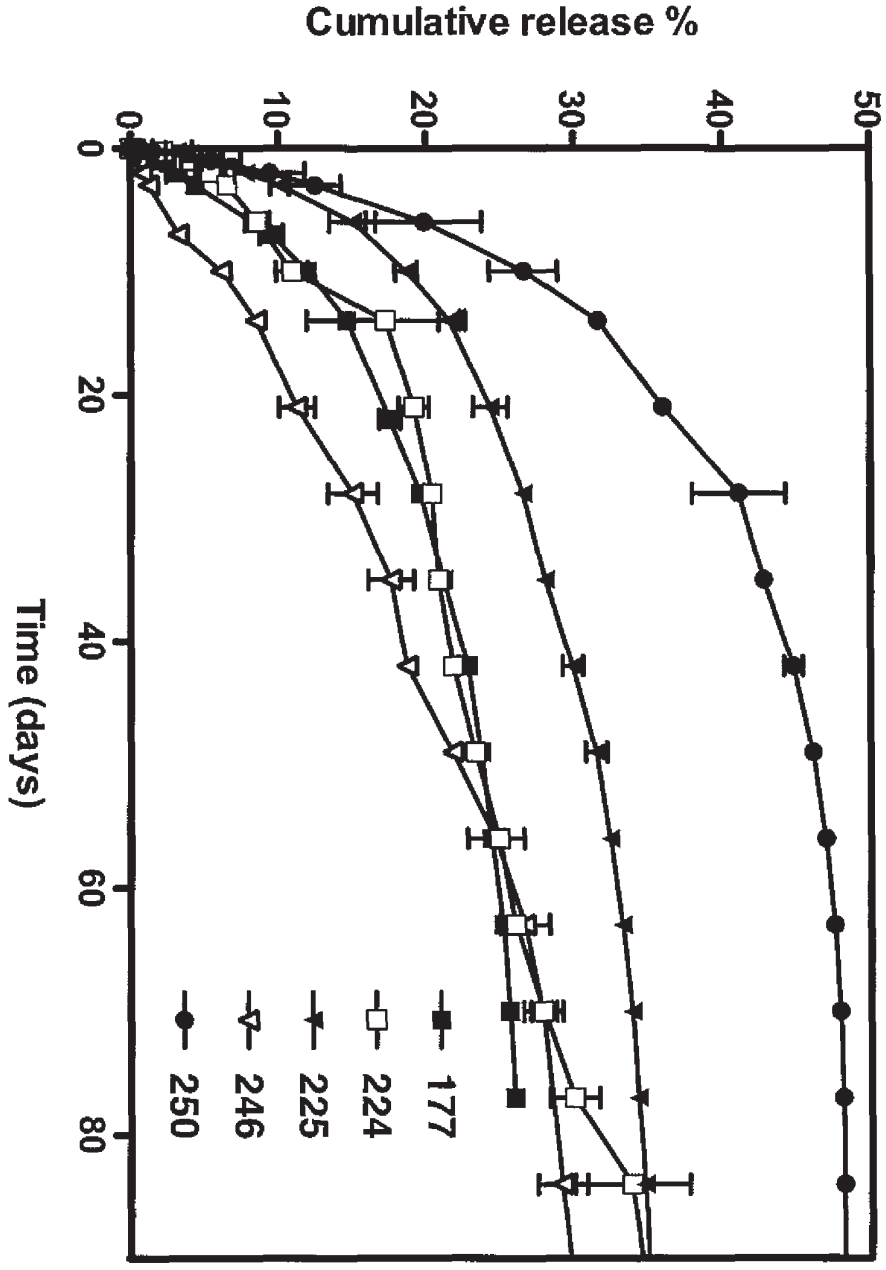

Cumulative release %
0
10
20
30
40
50
Time (days)
0
20
40
60
80
250
246
225
224
177

FIGURE 4

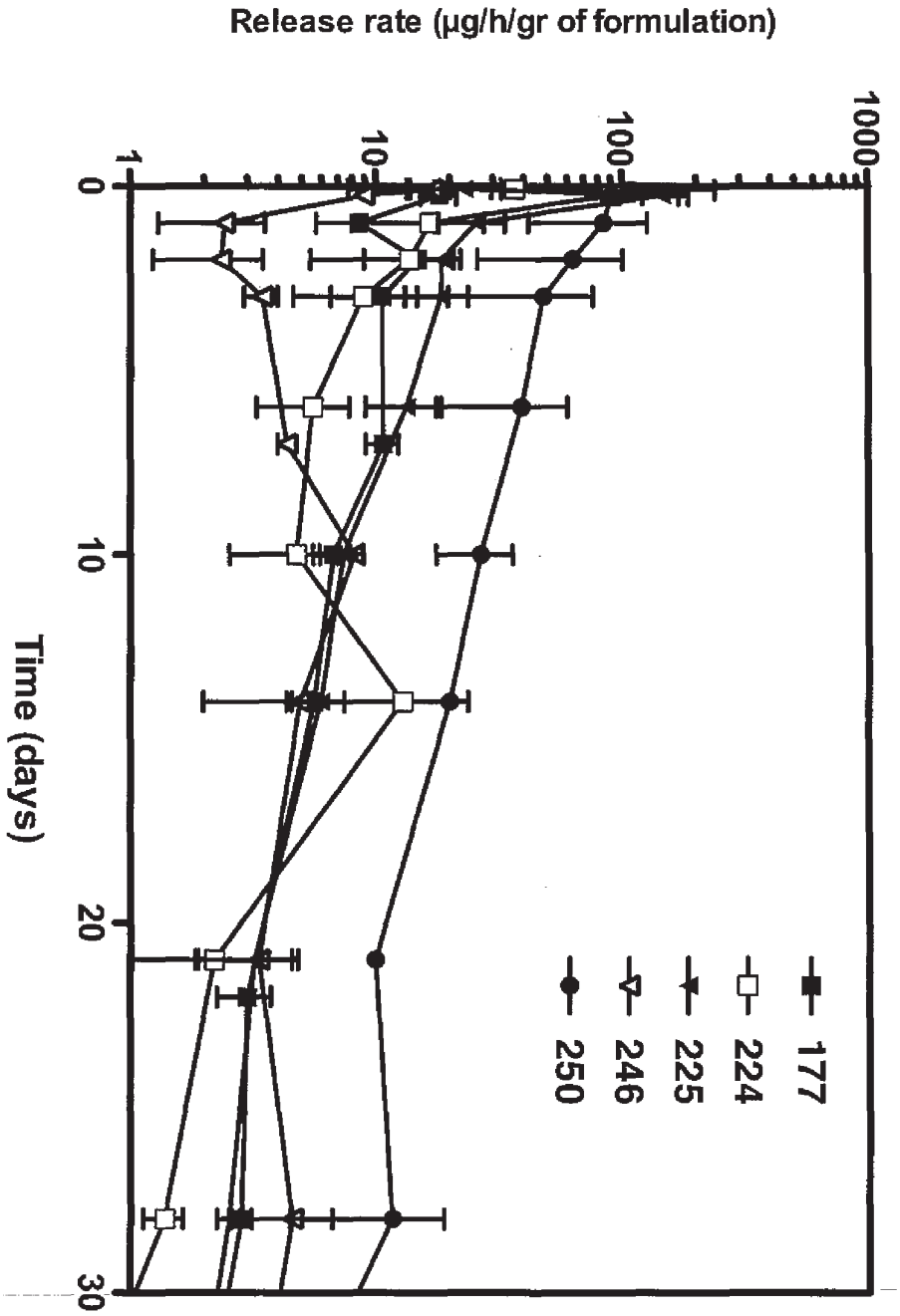

Release rate (µg/h/gr of formulation)
1
10
100
1000
0
10
20
30
Time (days)
177
224
225
246
250

FIGURE 5

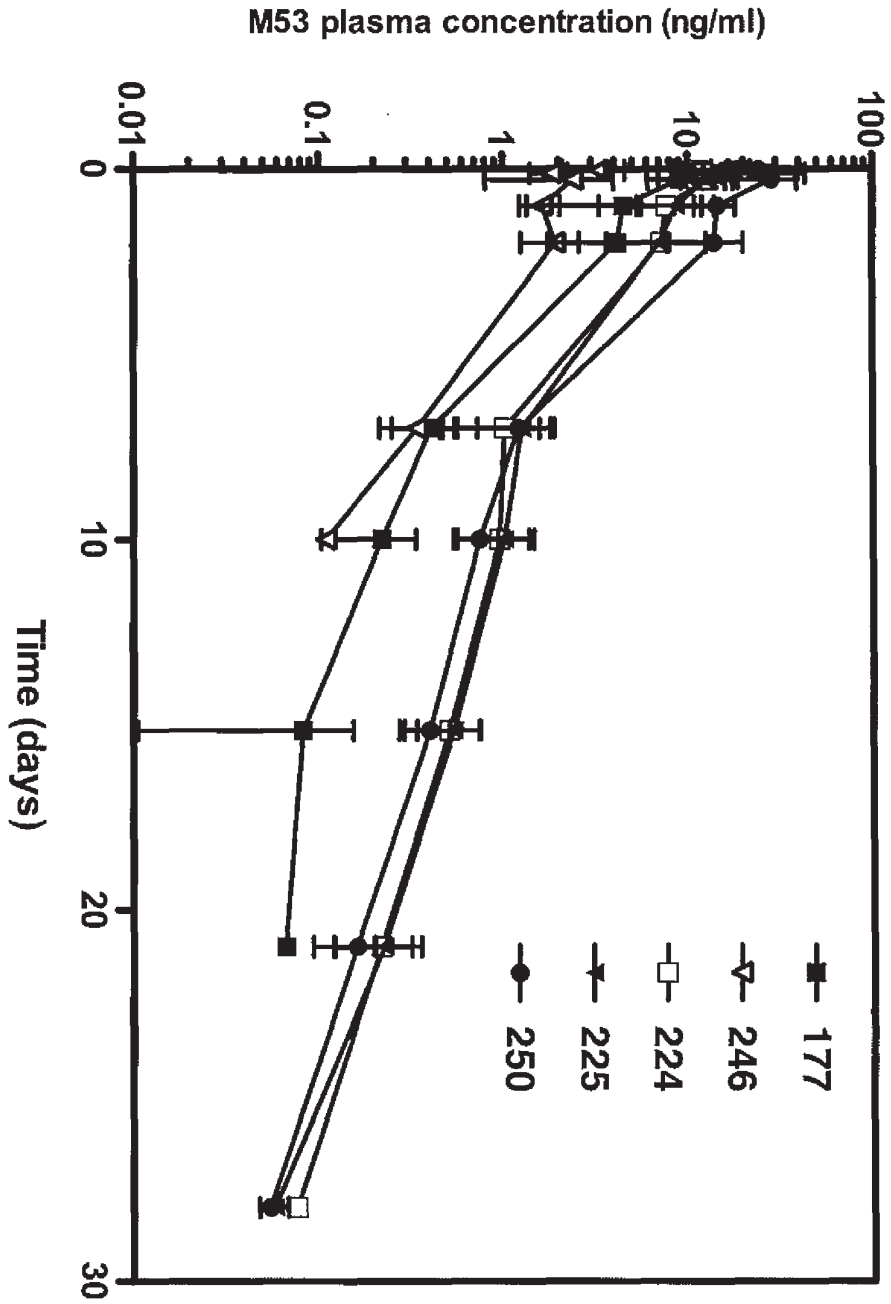

M53 plasma concentration (ng/ml)
0.01
0.1
1
10
100
0
10
20
30
Time (days)
250
225
224
246
177

FIGURE 6

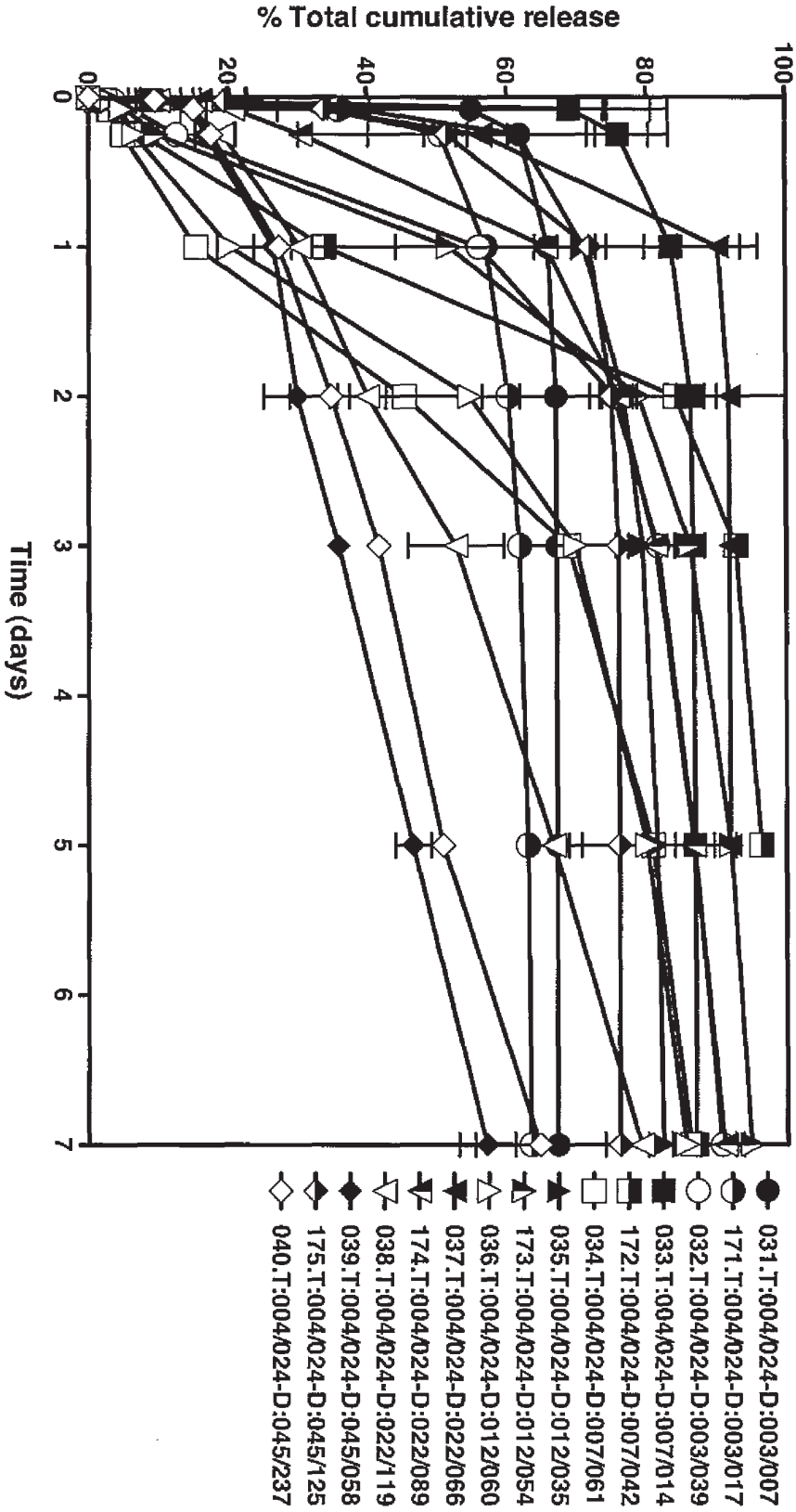

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
031.T:004/024-D:003/007
171.T:004/024-D:003/017
032.T:004/024-D:003/039
033.T:004/024-D:007/014
172.T:004/024-D:007/042
034.T:004/024-D:007/061
035.T:004/024-D:012/035
173.T:004/024-D:012/054
036.T:004/024-D:012/060
037.T:004/024-D:022/066
174.T:004/024-D:022/089
038.T:004/024-D:022/119
039.T:004/024-D:045/058
175.T:004/024-D:045/125
040.T:004/024-D:045/237

FIGURE 7

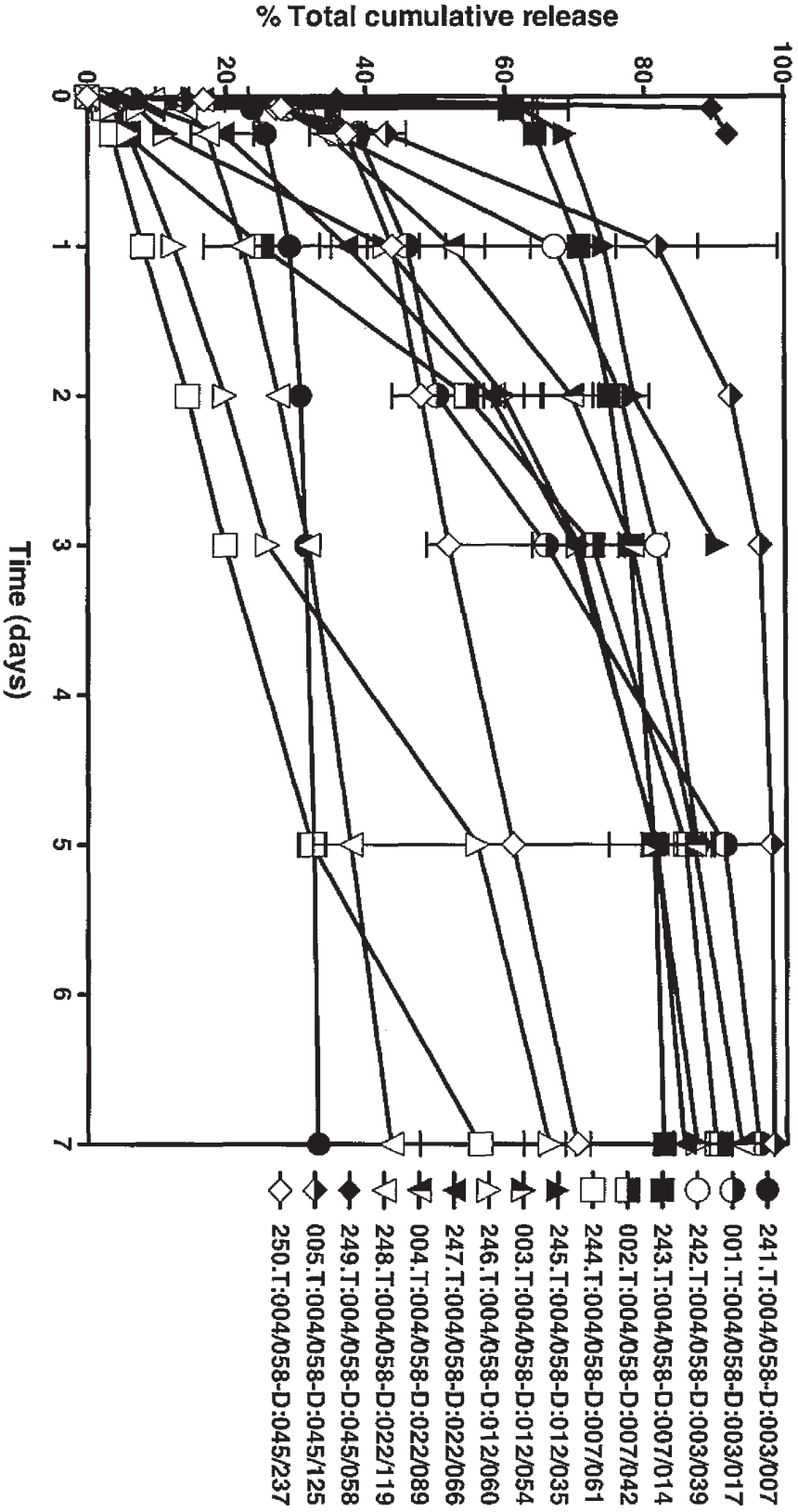

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
241.T:004/058-D:003/007
001.T:004/058-D:003/017
242.T:004/058-D:003/039
243.T:004/058-D:007/014
002.T:004/058-D:007/042
244.T:004/058-D:007/061
245.T:004/058-D:012/035
003.T:004/058-D:012/054
246.T:004/058-D:012/060
247.T:004/058-D:022/066
004.T:004/058-D:022/089
248.T:004/058-D:022/119
249.T:004/058-D:045/058
005.T:004/058-D:045/125
250.T:004/058-D:045/237

FIGURE 8

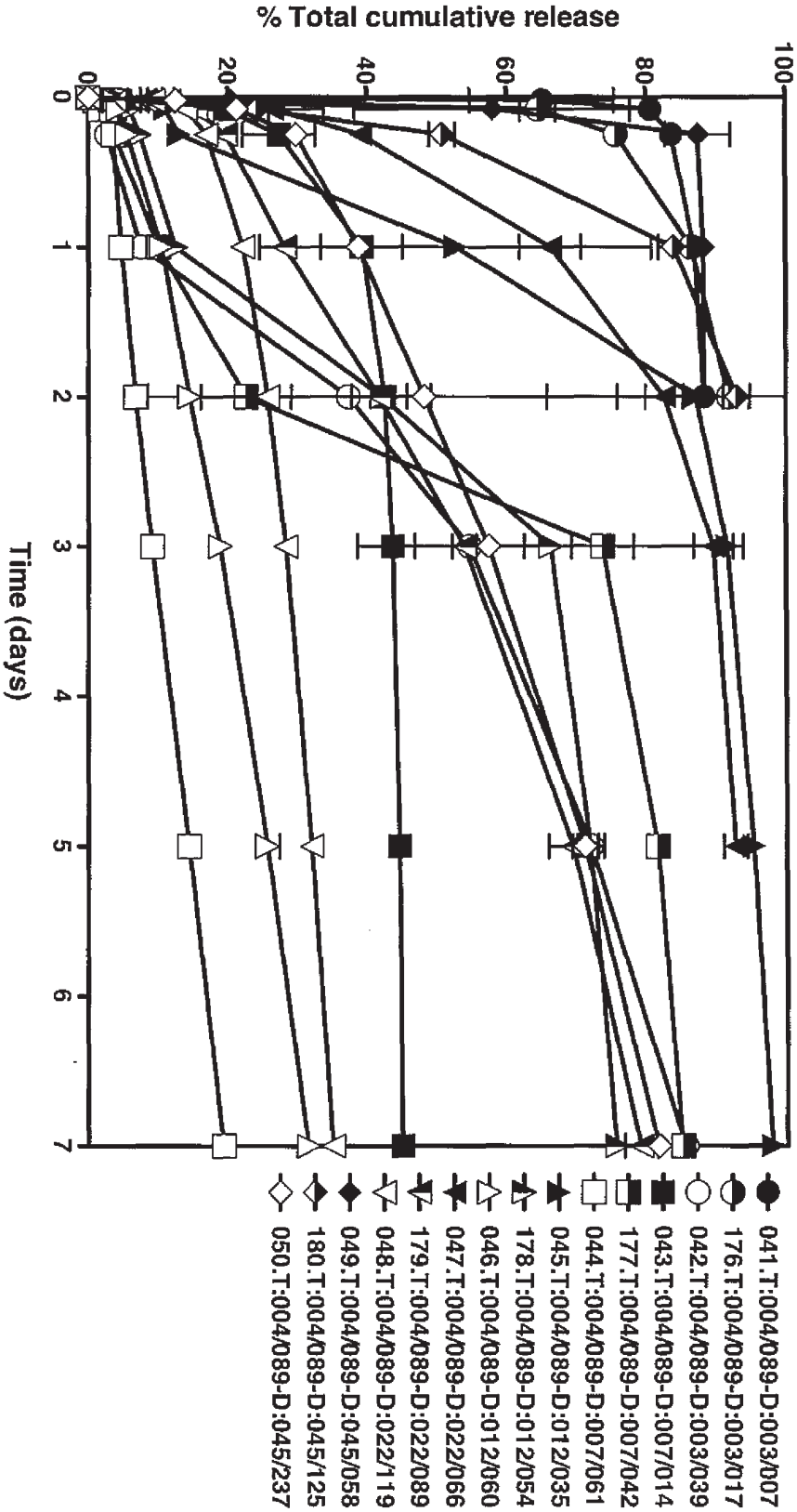

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
041.T:004/089-D:003/007
176.T:004/089-D:003/017
042.T:004/089-D:003/039
043.T:004/089-D:007/014
177.T:004/089-D:007/042
044.T:004/089-D:007/061
045.T:004/089-D:012/035
178.T:004/089-D:012/054
046.T:004/089-D:012/060
047.T:004/089-D:022/066
179.T:004/089-D:022/089
048.T:004/089-D:022/119
049.T:004/089-D:045/058
180.T:004/089-D:045/125
050.T:004/089-D:045/237

FIGURE 9

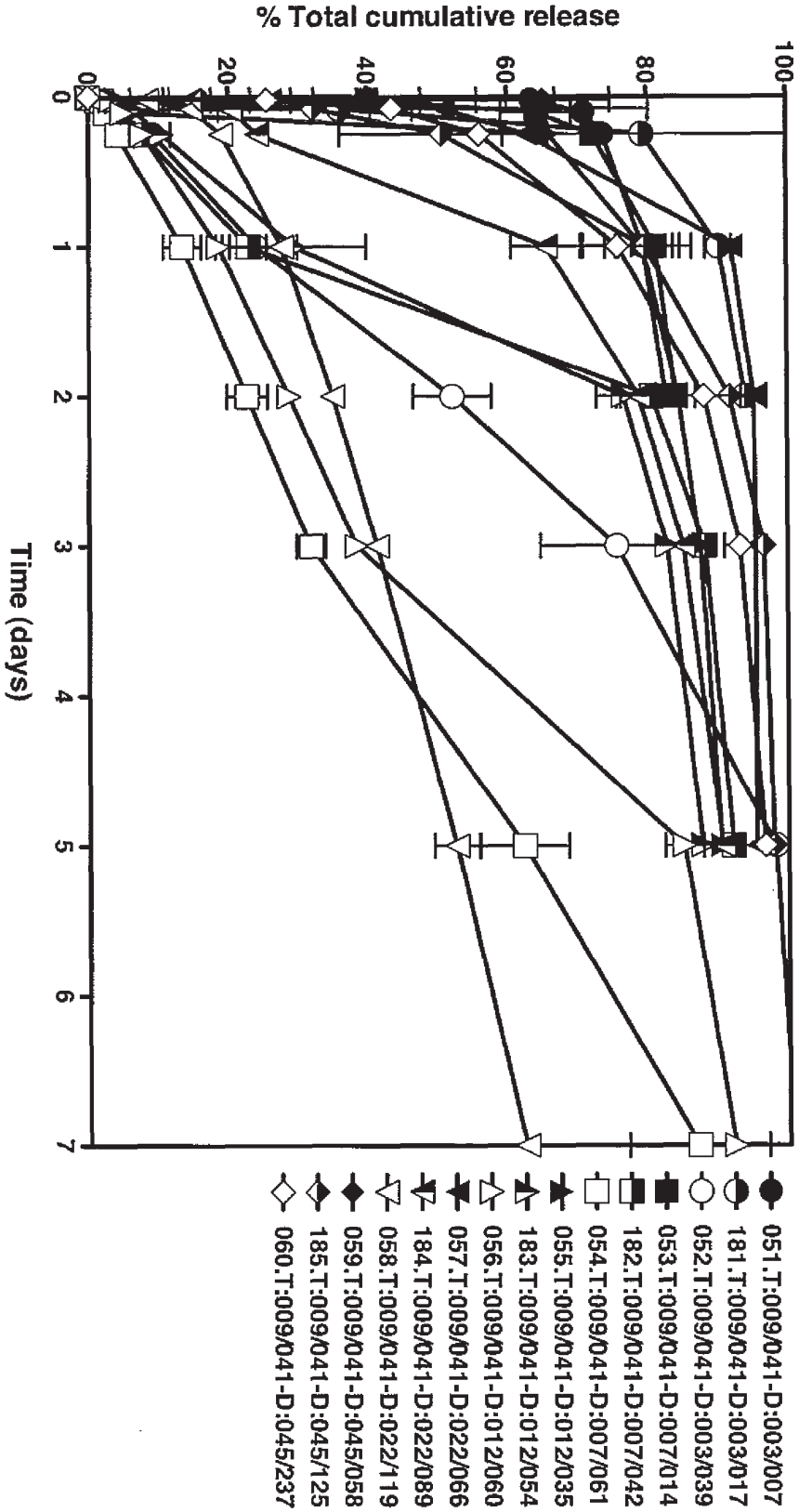

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
051.T:009/041-D:003/007
181.T:009/041-D:003/017
052.T:009/041-D:003/039
053.T:009/041-D:007/014
182.T:009/041-D:007/042
054.T:009/041-D:007/061
055.T:009/041-D:012/035
183.T:009/041-D:012/054
056.T:009/041-D:012/060
057.T:009/041-D:022/066
184.T:009/041-D:022/089
058.T:009/041-D:022/119
059.T:009/041-D:045/058
185.T:009/041-D:045/125
060.T:009/041-D:045/237

FIGURE 10

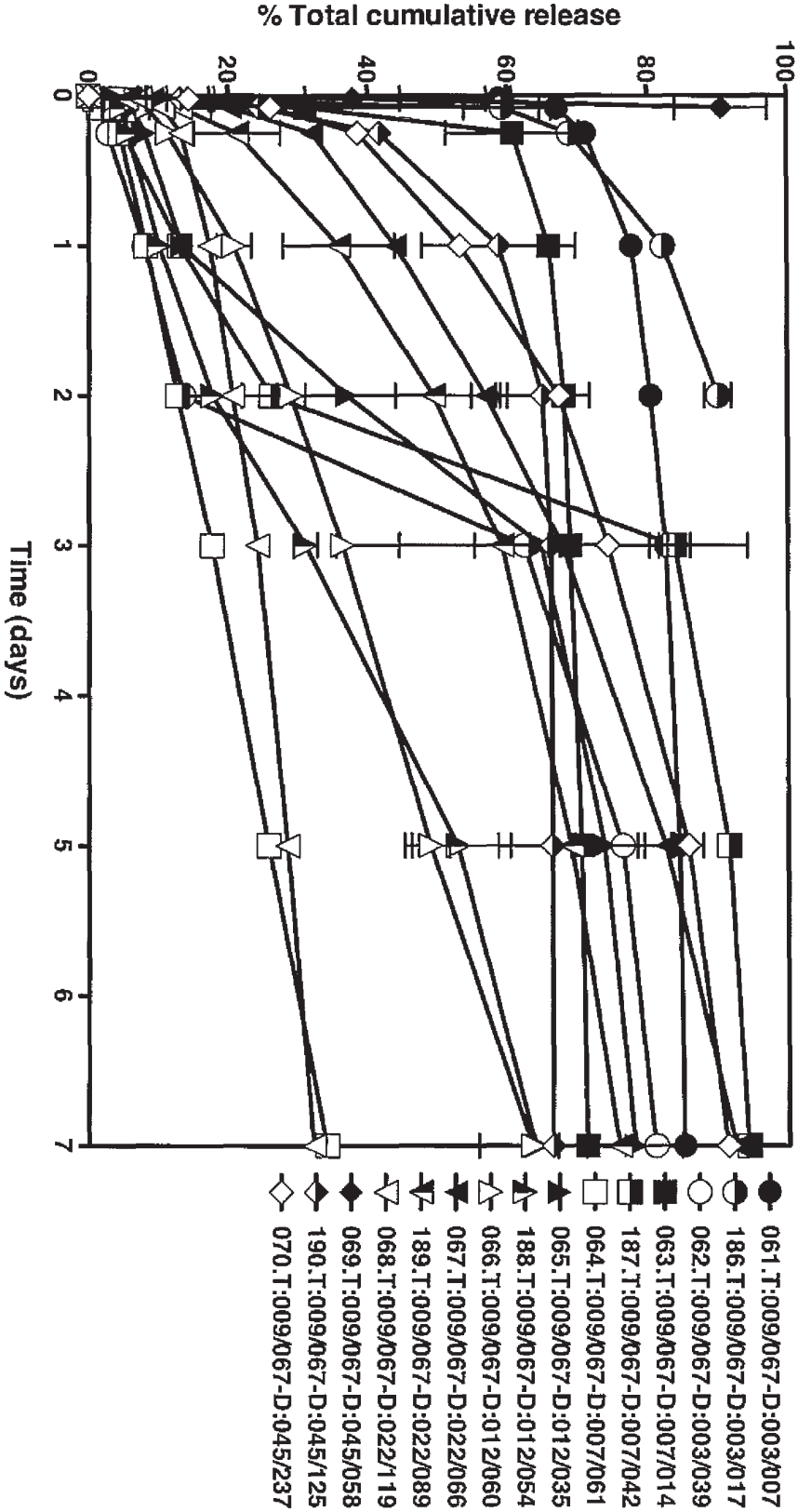

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
061.T:009/067-D:003/007
186.T:009/067-D:003/017
062.T:009/067-D:003/039
063.T:009/067-D:007/014
187.T:009/067-D:007/042
064.T:009/067-D:007/061
065.T:009/067-D:012/035
188.T:009/067-D:012/054
066.T:009/067-D:012/060
067.T:009/067-D:022/066
189.T:009/067-D:022/089
068.T:009/067-D:022/119
069.T:009/067-D:045/058
190.T:009/067-D:045/125
070.T:009/067-D:045/237

FIGURE 11

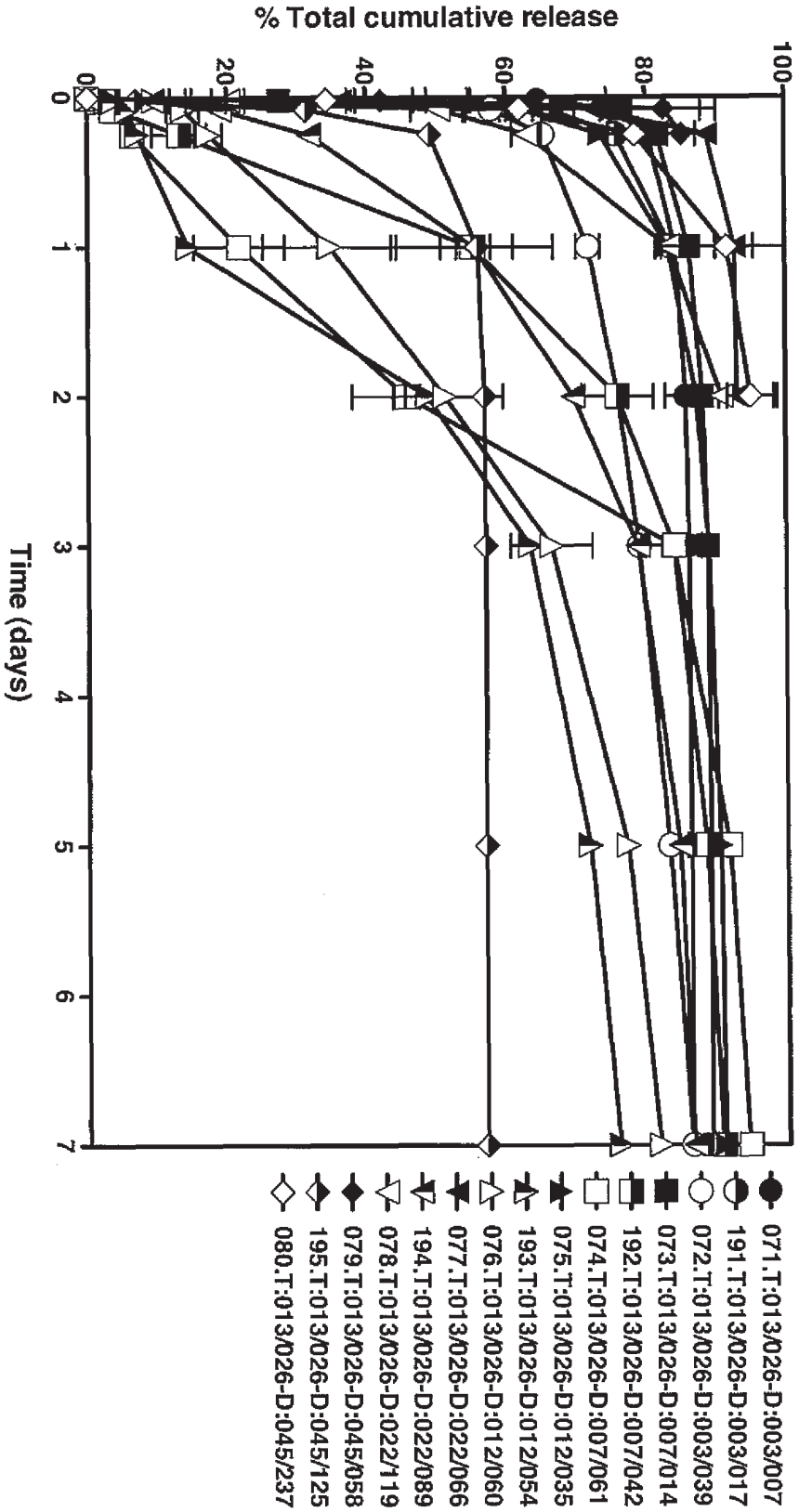

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
071.T:013/026-D:003/007
191.T:013/026-D:003/017
072.T:013/026-D:003/039
073.T:013/026-D:007/014
192.T:013/026-D:007/042
074.T:013/026-D:007/061
075.T:013/026-D:012/035
193.T:013/026-D:012/054
076.T:013/026-D:012/060
077.T:013/026-D:022/066
194.T:013/026-D:022/089
078.T:013/026-D:022/119
079.T:013/026-D:045/058
195.T:013/026-D:045/125
080.T:013/026-D:045/237

FIGURE 12

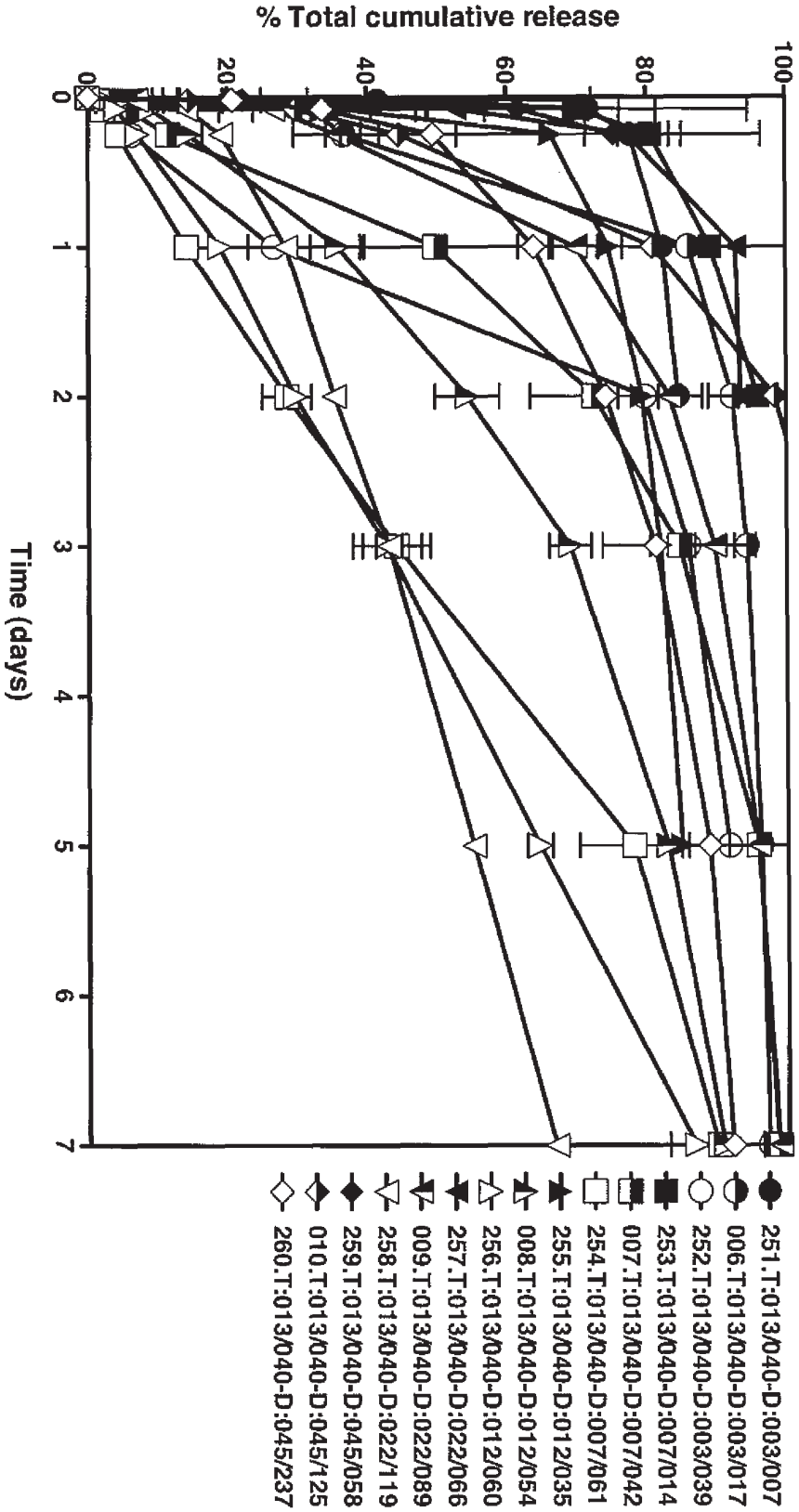

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
251.T:013/040-D:003/007
006.T:013/040-D:003/017
252.T:013/040-D:003/039
253.T:013/040-D:007/014
007.T:013/040-D:007/042
254.T:013/040-D:007/061
255.T:013/040-D:012/035
008.T:013/040-D:012/054
256.T:013/040-D:012/060
257.T:013/040-D:022/066
009.T:013/040-D:022/089
258.T:013/040-D:022/119
259.T:013/040-D:045/058
010.T:013/040-D:045/125
260.T:013/040-D:045/237

FIGURE 13

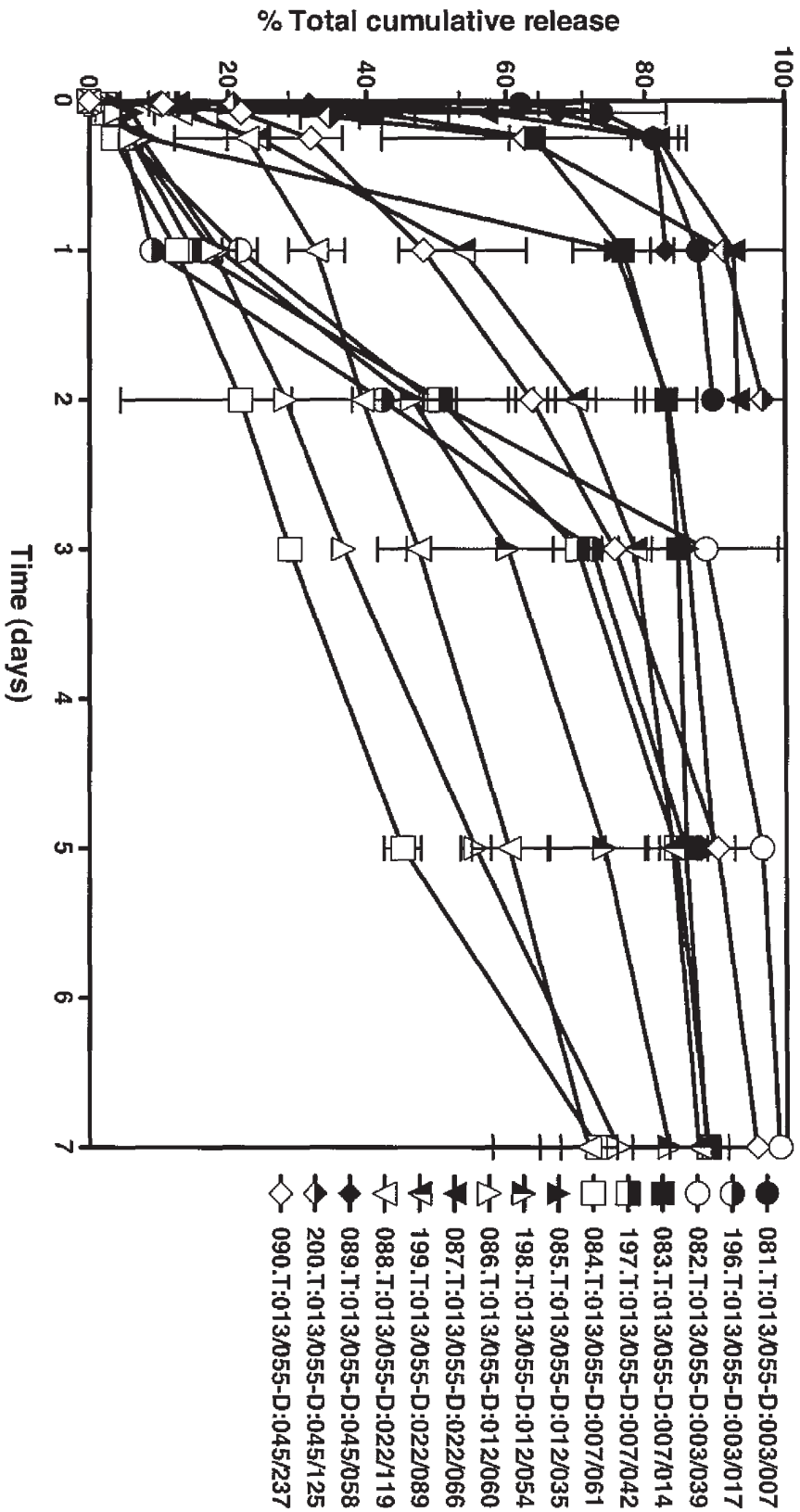

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
081.T:013/055-D:003/007
196.T:013/055-D:003/017
082.T:013/055-D:003/039
083.T:013/055-D:007/014
197.T:013/055-D:007/042
084.T:013/055-D:007/061
085.T:013/055-D:012/035
198.T:013/055-D:012/054
086.T:013/055-D:012/060
087.T:013/055-D:022/066
199.T:013/055-D:022/089
088.T:013/055-D:022/119
089.T:013/055-D:045/058
200.T:013/055-D:045/125
090.T:013/055-D:045/237

FIGURE 14

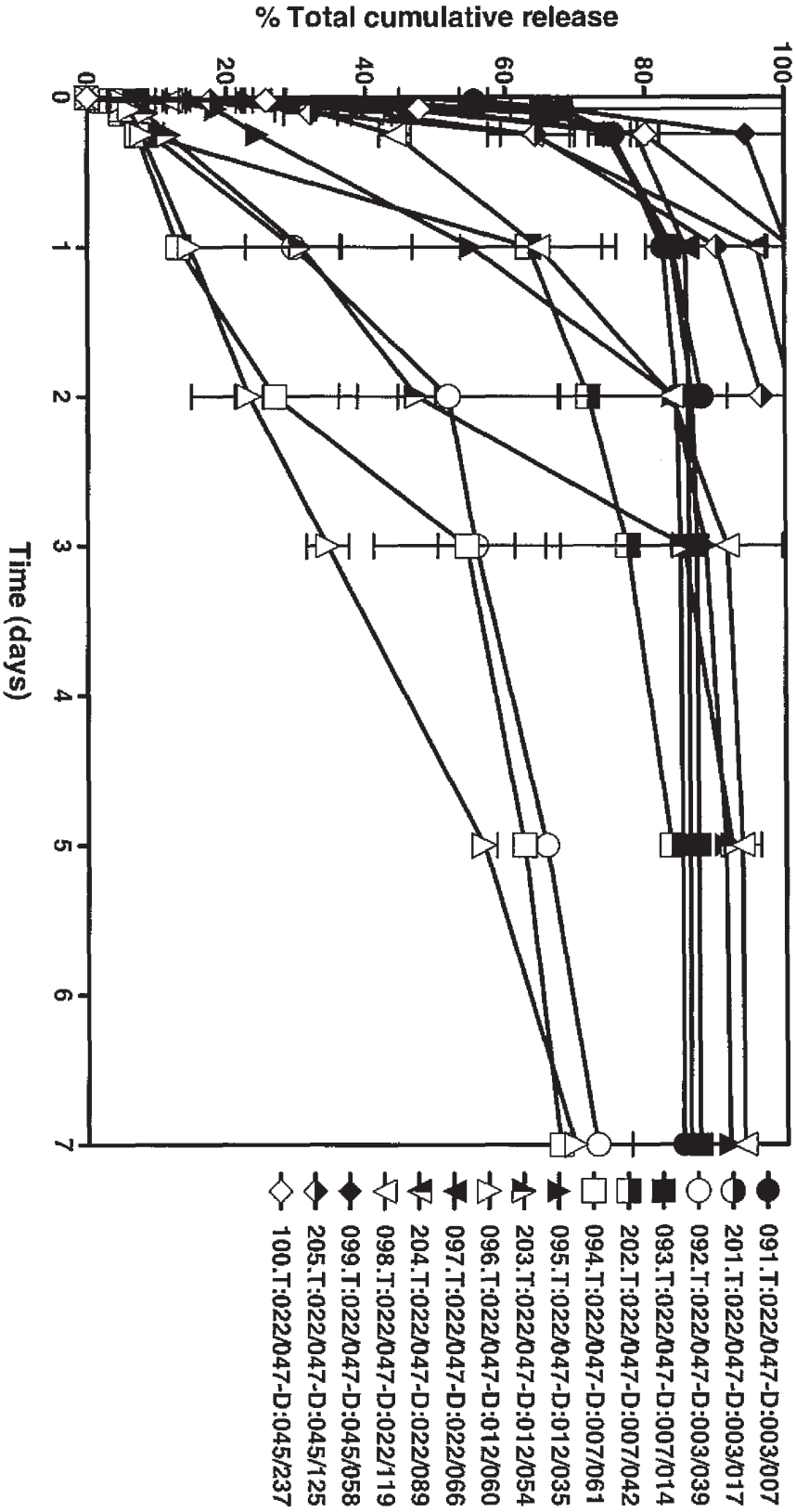

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
091.T:022/047-D:003/007
201.T:022/047-D:003/017
092.T:022/047-D:003/039
093.T:022/047-D:007/014
202.T:022/047-D:007/042
094.T:022/047-D:007/061
095.T:022/047-D:012/035
203.T:022/047-D:012/054
096.T:022/047-D:012/060
097.T:022/047-D:022/066
204.T:022/047-D:022/089
098.T:022/047-D:022/119
099.T:022/047-D:045/058
205.T:022/047-D:045/125
100.T:022/047-D:045/237

FIGURE 15

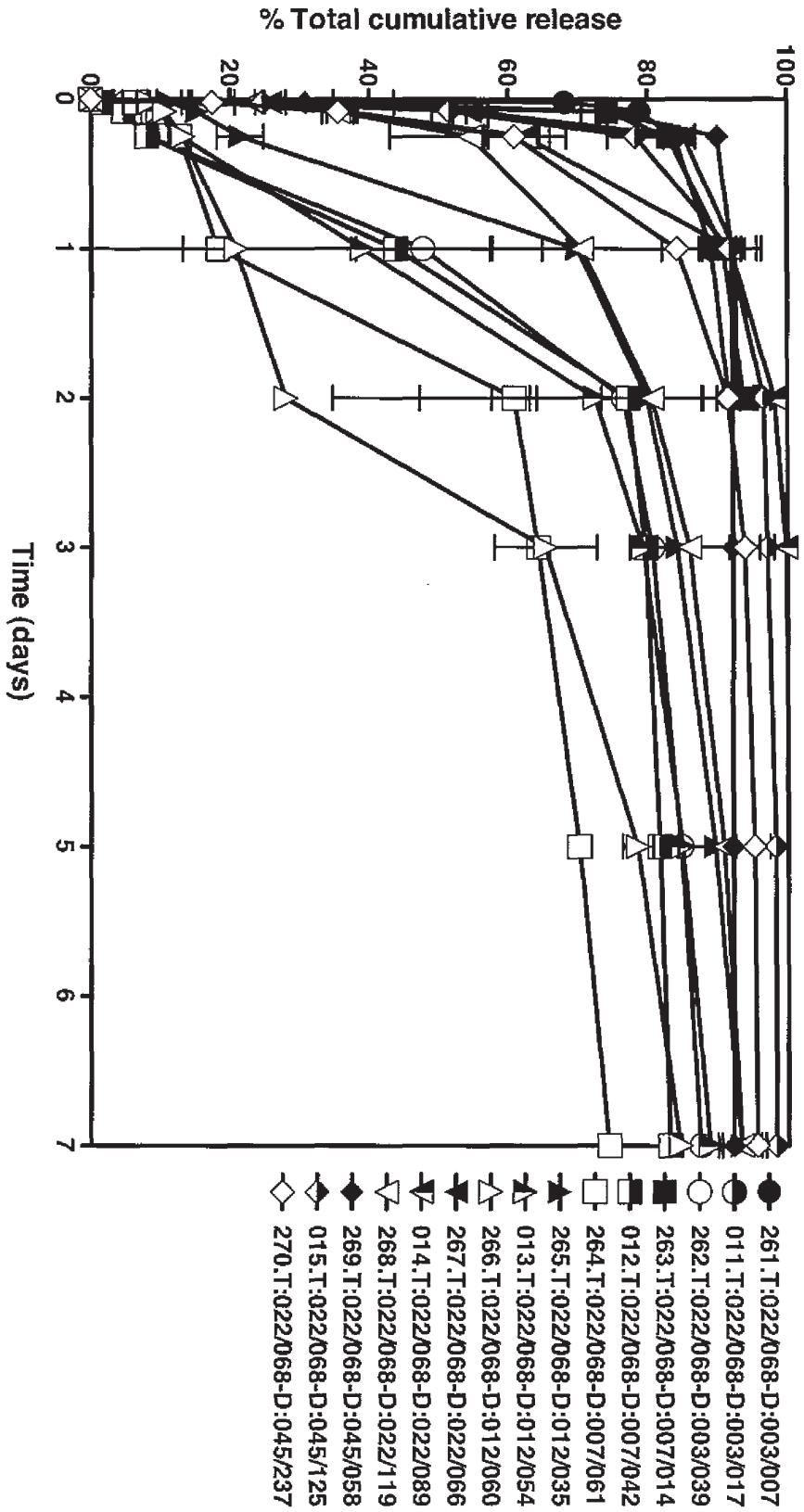

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
261.T:022/068-D:003/007
011.T:022/068-D:003/017
262.T:022/068-D:003/039
263.T:022/068-D:007/014
012.T:022/068-D:007/042
264.T:022/068-D:007/061
265.T:022/068-D:012/035
013.T:022/068-D:012/054
266.T:022/068-D:012/060
267.T:022/068-D:022/066
014.T:022/068-D:022/089
268.T:022/068-D:022/119
269.T:022/068-D:045/058
015.T:022/068-D:045/125
270.T:022/068-D:045/237

FIGURE 16

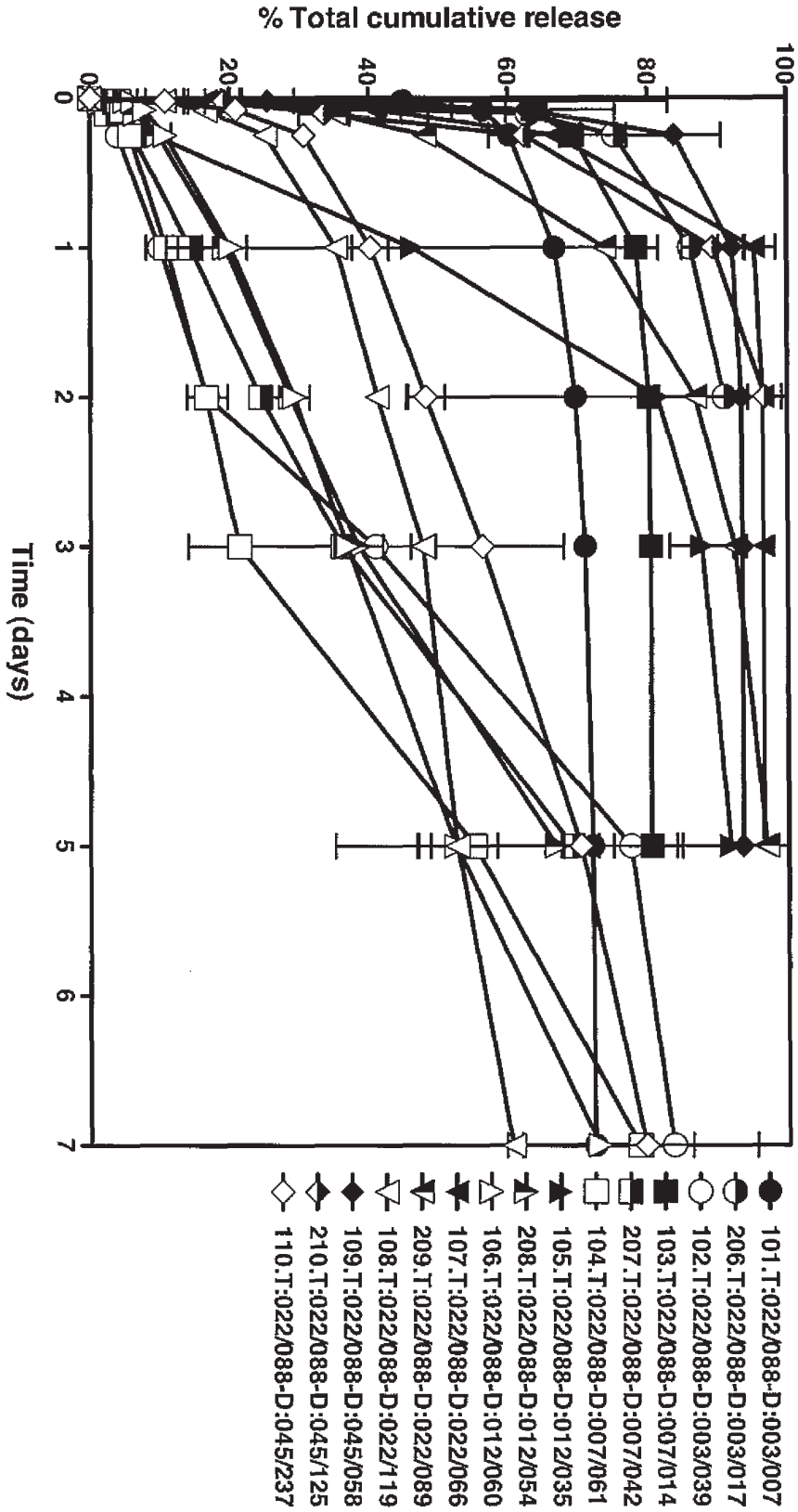

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
101.T:022/088-D:003/007
206.T:022/088-D:003/017
102.T:022/088-D:003/039
103.T:022/088-D:007/014
207.T:022/088-D:007/042
104.T:022/088-D:007/061
105.T:022/088-D:012/035
208.T:022/088-D:012/054
106.T:022/088-D:012/060
107.T:022/088-D:022/066
209.T:022/088-D:022/089
108.T:022/088-D:022/119
109.T:022/088-D:045/058
210.T:022/088-D:045/125
110.T:022/088-D:045/237

FIGURE 17

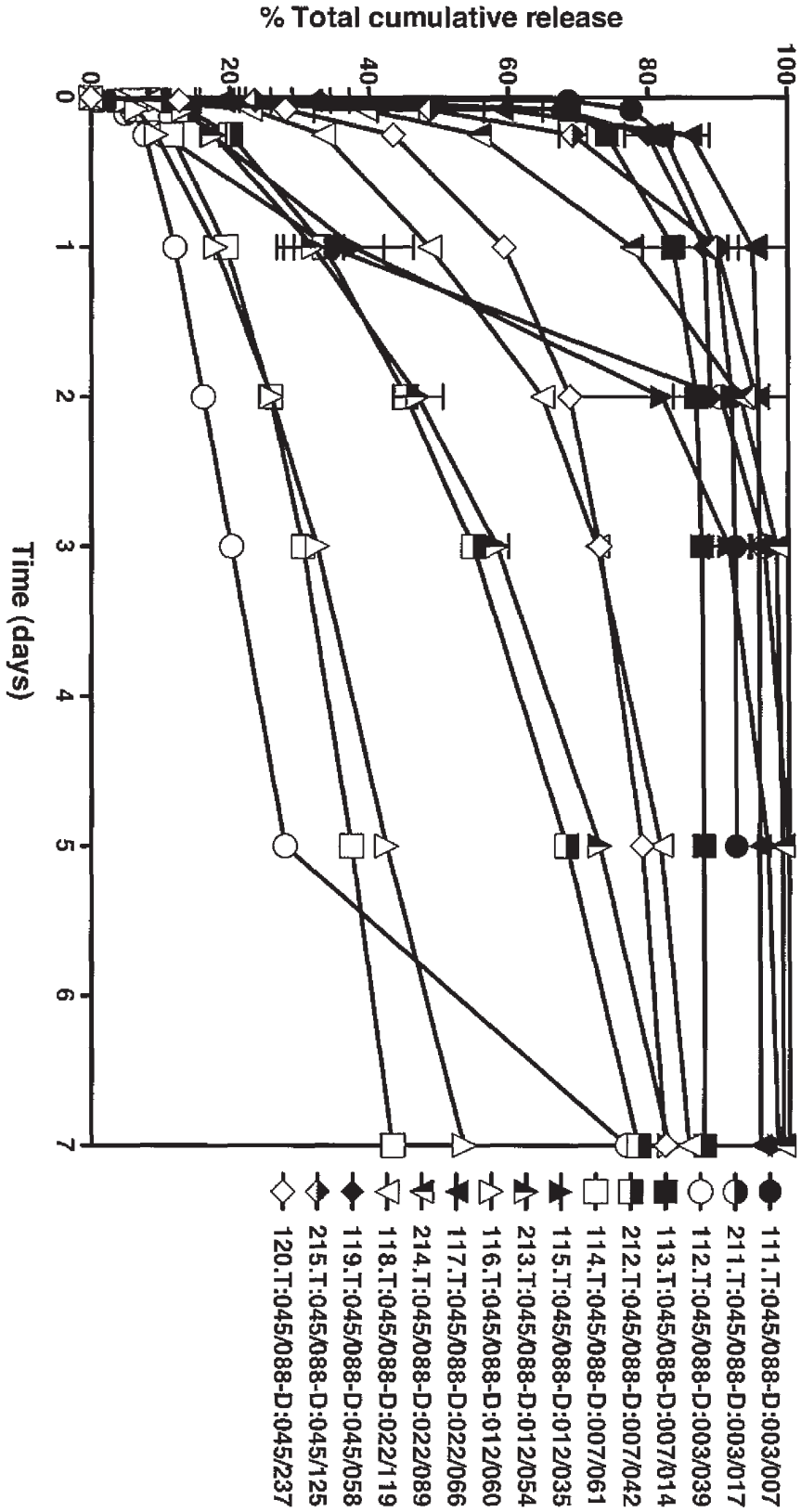

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
111.T:045/088-D:003/007
211.T:045/088-D:003/017
112.T:045/088-D:003/039
113.T:045/088-D:007/014
212.T:045/088-D:007/042
114.T:045/088-D:007/061
115.T:045/088-D:012/035
213.T:045/088-D:012/054
116.T:045/088-D:012/060
117.T:045/088-D:022/066
214.T:045/088-D:022/089
118.T:045/088-D:022/119
119.T:045/088-D:045/058
215.T:045/088-D:045/125
120.T:045/088-D:045/237

FIGURE 18

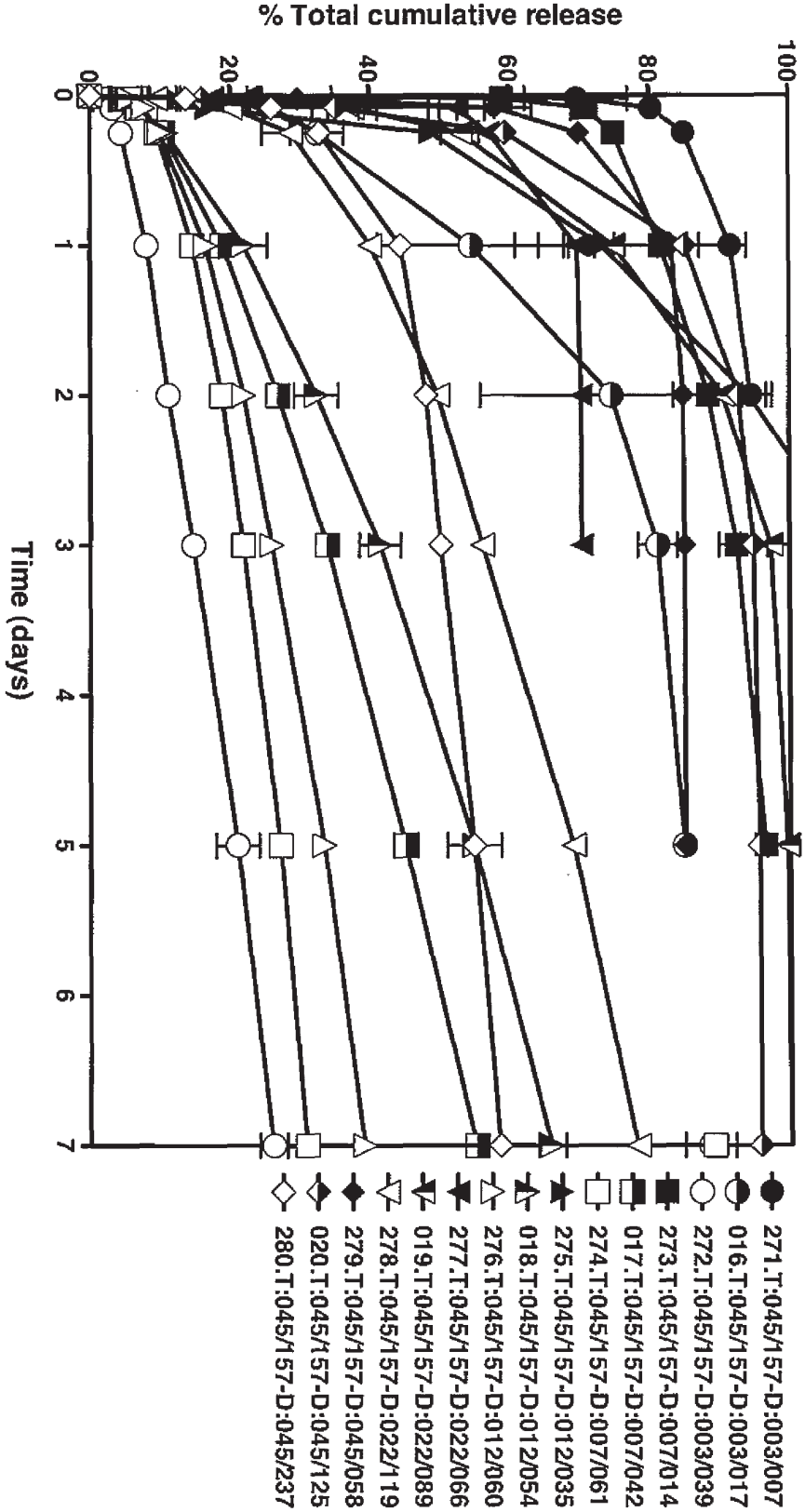

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
271.T:045/157-D:003/007
016.T:045/157-D:003/017
272.T:045/157-D:003/039
273.T:045/157-D:007/014
017.T:045/157-D:007/042
274.T:045/157-D:007/061
275.T:045/157-D:012/035
018.T:045/157-D:012/054
276.T:045/157-D:012/060
277.T:045/157-D:022/066
019.T:045/157-D:022/089
278.T:045/157-D:022/119
279.T:045/157-D:045/058
020.T:045/157-D:045/125
280.T:045/157-D:045/237

FIGURE 19

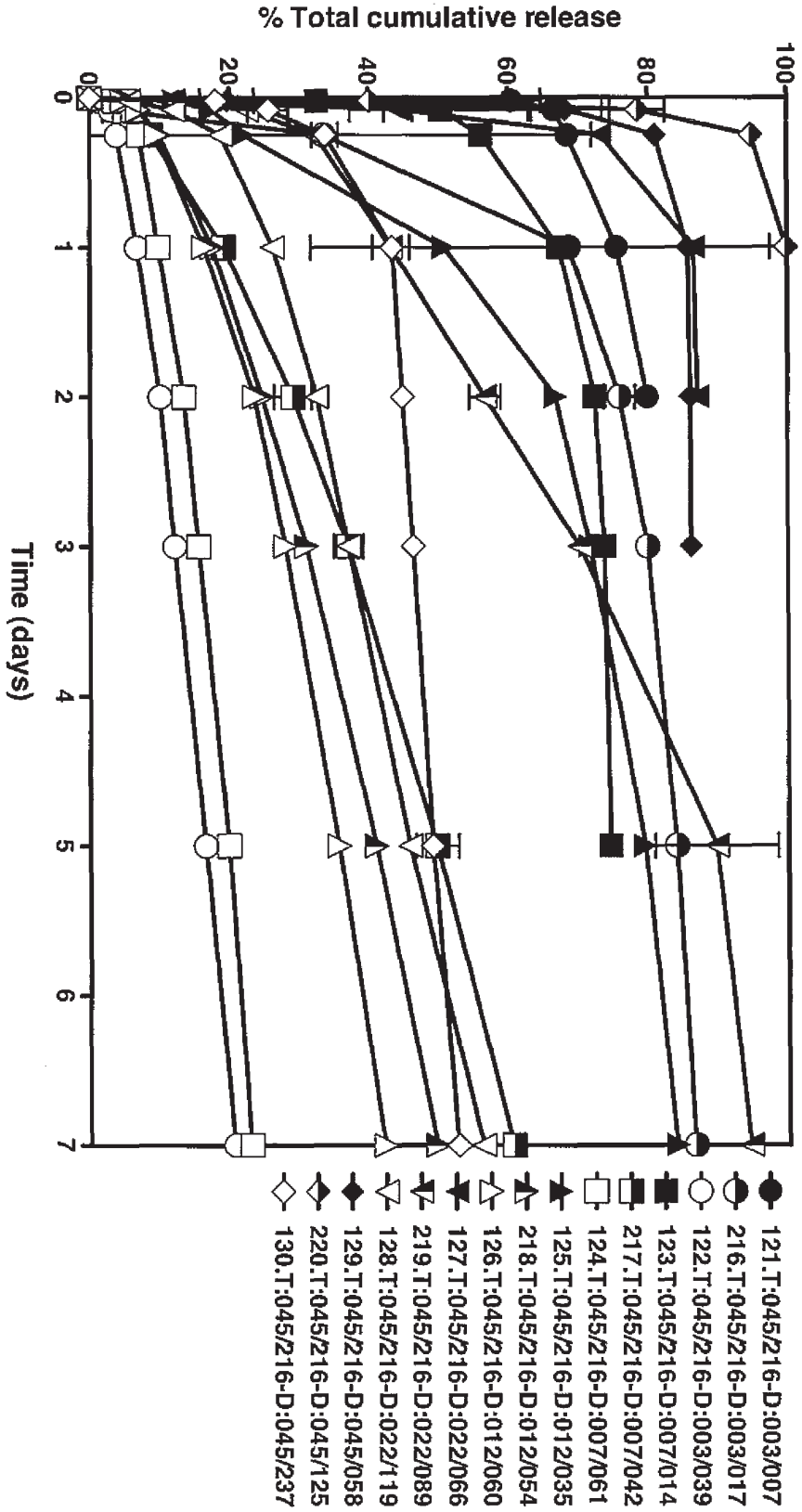

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
121.T:045/216-D:003/007
216.T:045/216-D:003/017
122.T:045/216-D:003/039
123.T:045/216-D:007/014
217.T:045/216-D:007/042
124.T:045/216-D:007/061
125.T:045/216-D:012/035
218.T:045/216-D:012/054
126.T:045/216-D:012/060
127.T:045/216-D:022/066
219.T:045/216-D:022/089
128.T:045/216-D:022/119
129.T:045/216-D:045/058
220.T:045/216-D:045/125
130.T:045/216-D:045/237

FIGURE 20

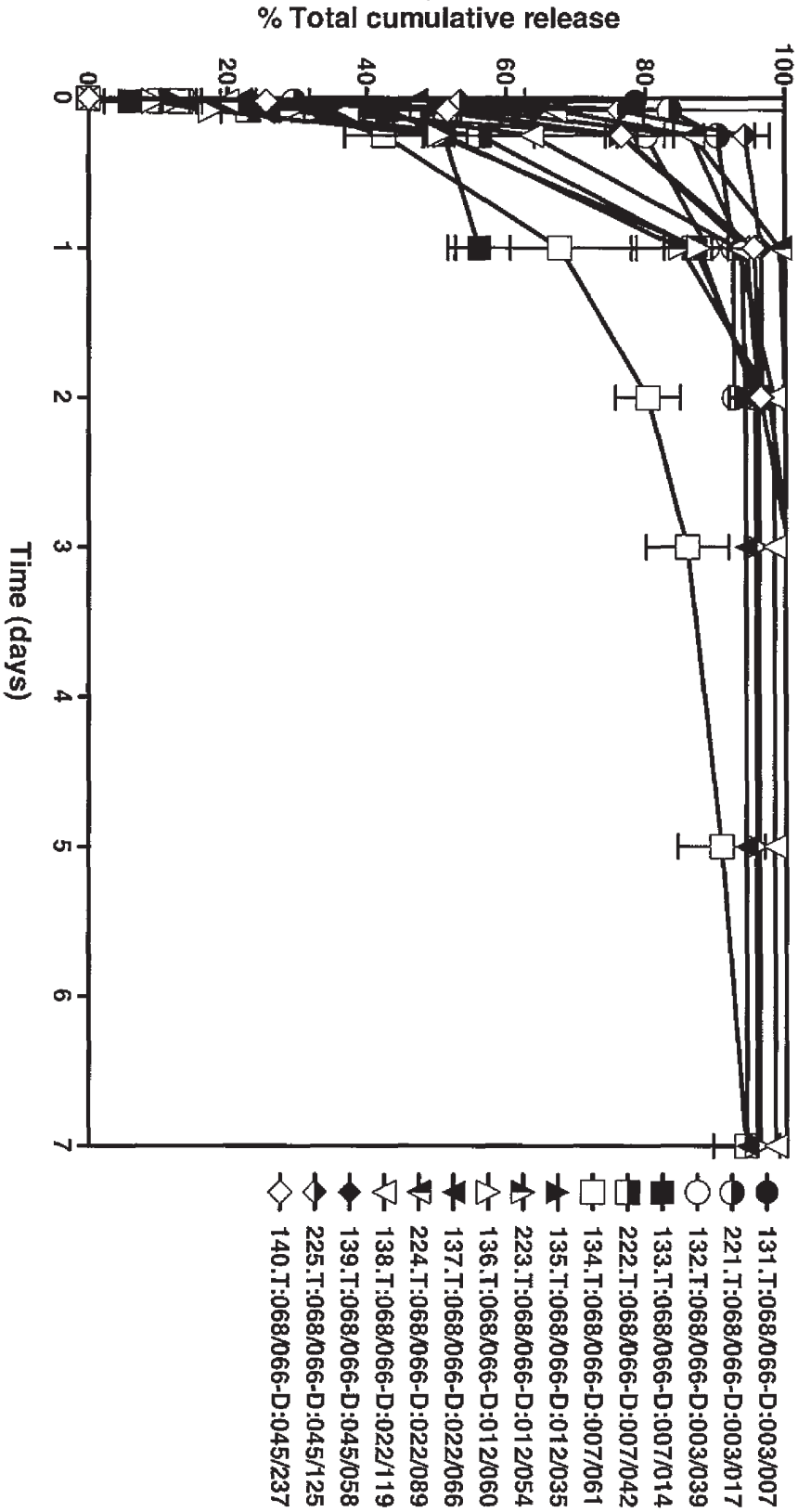

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
131.T:068/066-D:003/007
221.T:068/066-D:003/017
132.T:068/066-D:003/039
133.T:068/066-D:007/014
222.T:068/066-D:007/042
134.T:068/066-D:007/061
135.T:068/066-D:012/035
223.T:068/066-D:012/054
136.T:068/066-D:012/060
137.T:068/066-D:022/066
224.T:068/066-D:022/089
138.T:068/066-D:022/119
139.T:068/066-D:045/058
225.T:068/066-D:045/125
140.T:068/066-D:045/237

FIGURE 21

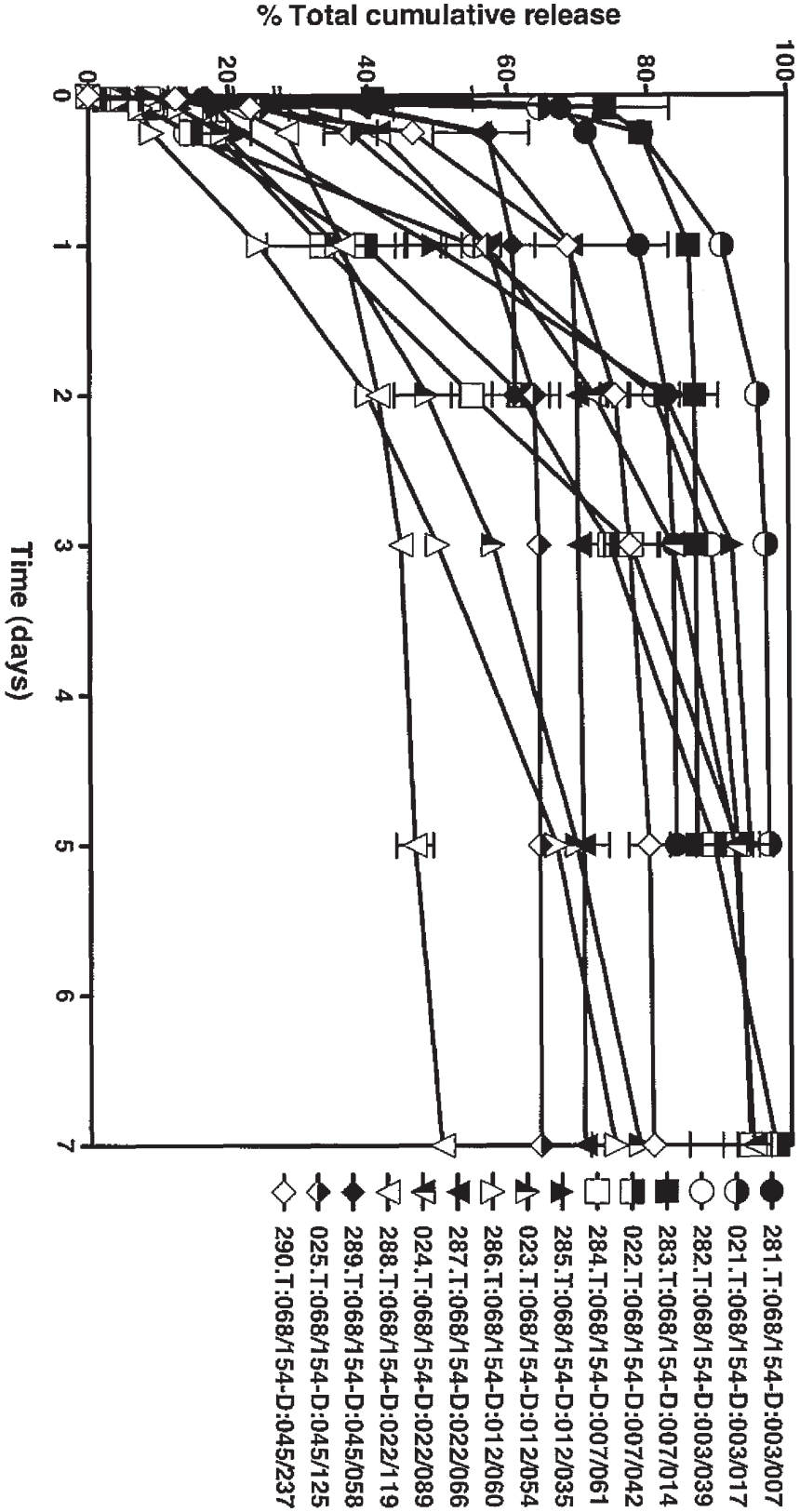

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
281.T:068/154-D:003/007
021.T:068/154-D:003/017
282.T:068/154-D:003/039
283.T:068/154-D:007/014
022.T:068/154-D:007/042
284.T:068/154-D:007/061
285.T:068/154-D:012/035
023.T:068/154-D:012/054
286.T:068/154-D:012/060
287.T:068/154-D:022/066
024.T:068/154-D:022/089
288.T:068/154-D:022/119
289.T:068/154-D:045/058
025.T:068/154-D:045/125
290.T:068/154-D:045/237

FIGURE 22

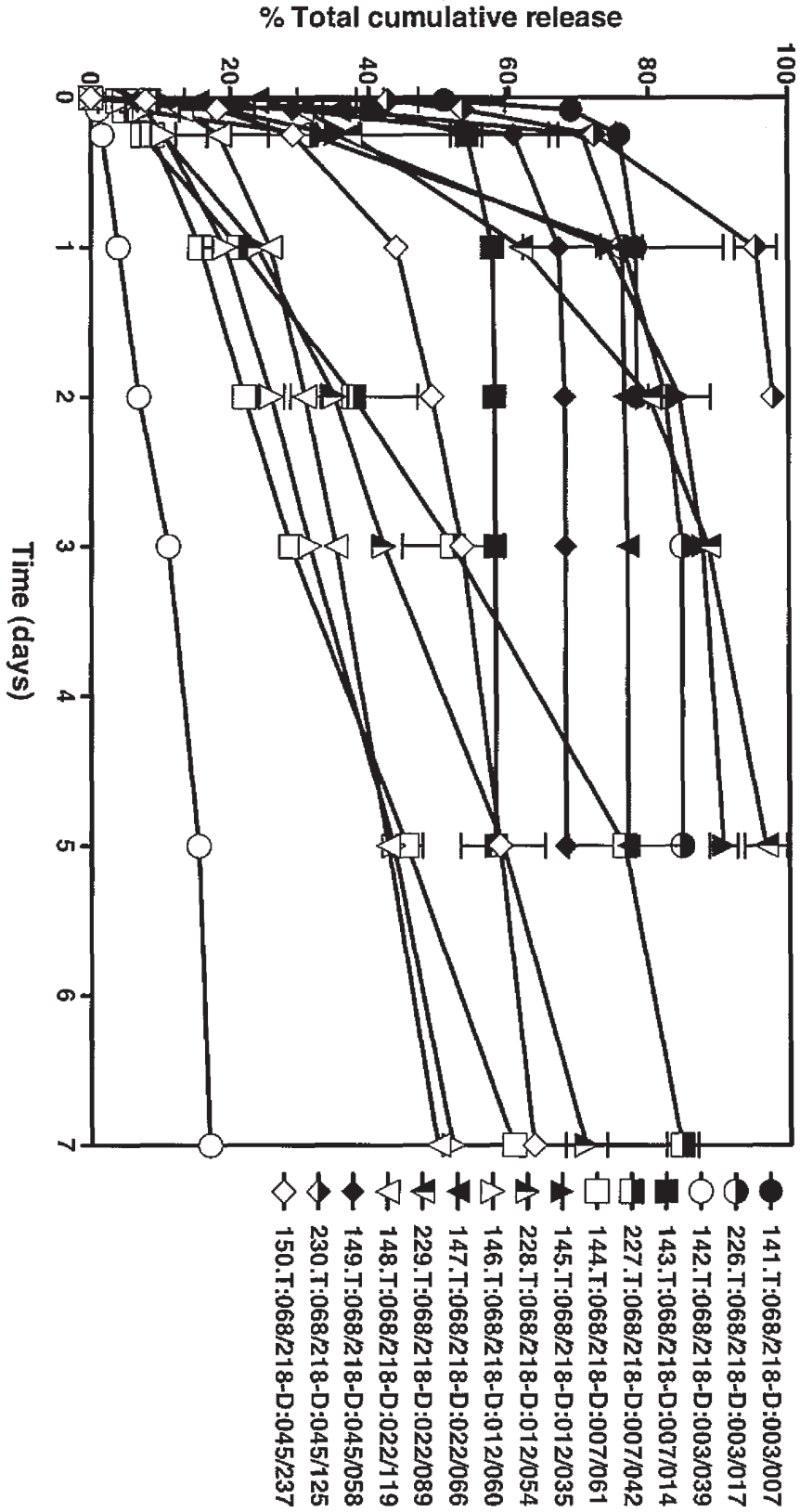

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
141.T:068/218-D:003/007
226.T:068/218-D:003/017
142.T:068/218-D:003/039
143.T:068/218-D:007/014
227.T:068/218-D:007/042
144.T:068/218-D:007/061
145.T:068/218-D:012/035
228.T:068/218-D:012/054
146.T:068/218-D:012/060
147.T:068/218-D:022/066
229.T:068/218-D:022/089
148.T:068/218-D:022/119
149.T:068/218-D:045/058
230.T:068/218-D:045/125
150.T:068/218-D:045/237

FIGURE 23

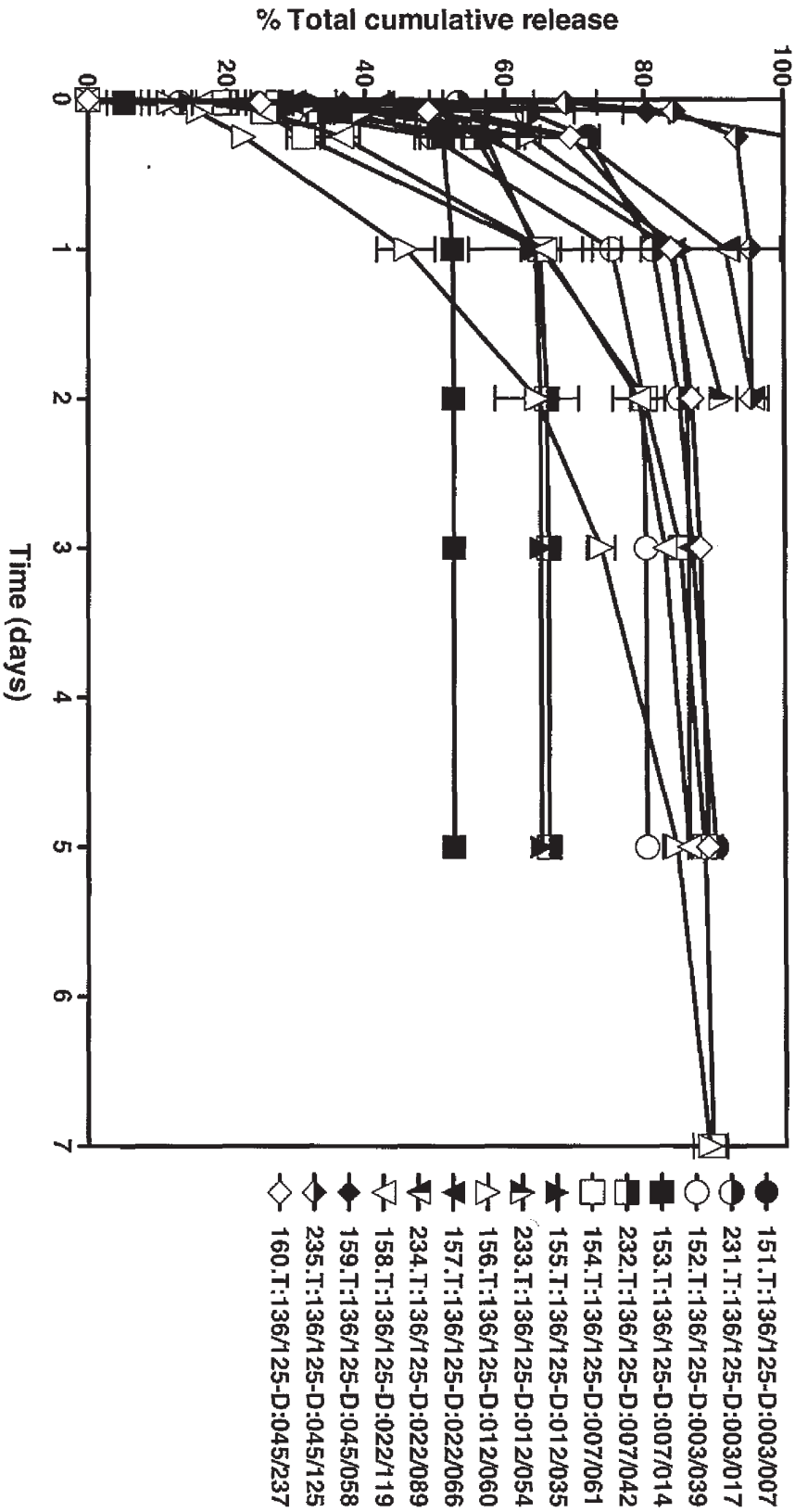

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
151.T:136/125-D:003/007
231.T:136/125-D:003/017
152.T:136/125-D:003/039
153.T:136/125-D:007/014
232.T:136/125-D:007/042
154.T:136/125-D:007/061
155.T:136/125-D:012/035
233.T:136/125-D:012/054
156.T:136/125-D:012/060
157.T:136/125-D:022/066
234.T:136/125-D:022/089
158.T:136/125-D:022/119
159.T:136/125-D:045/058
235.T:136/125-D:045/125
160.T:136/125-D:045/237

FIGURE 24

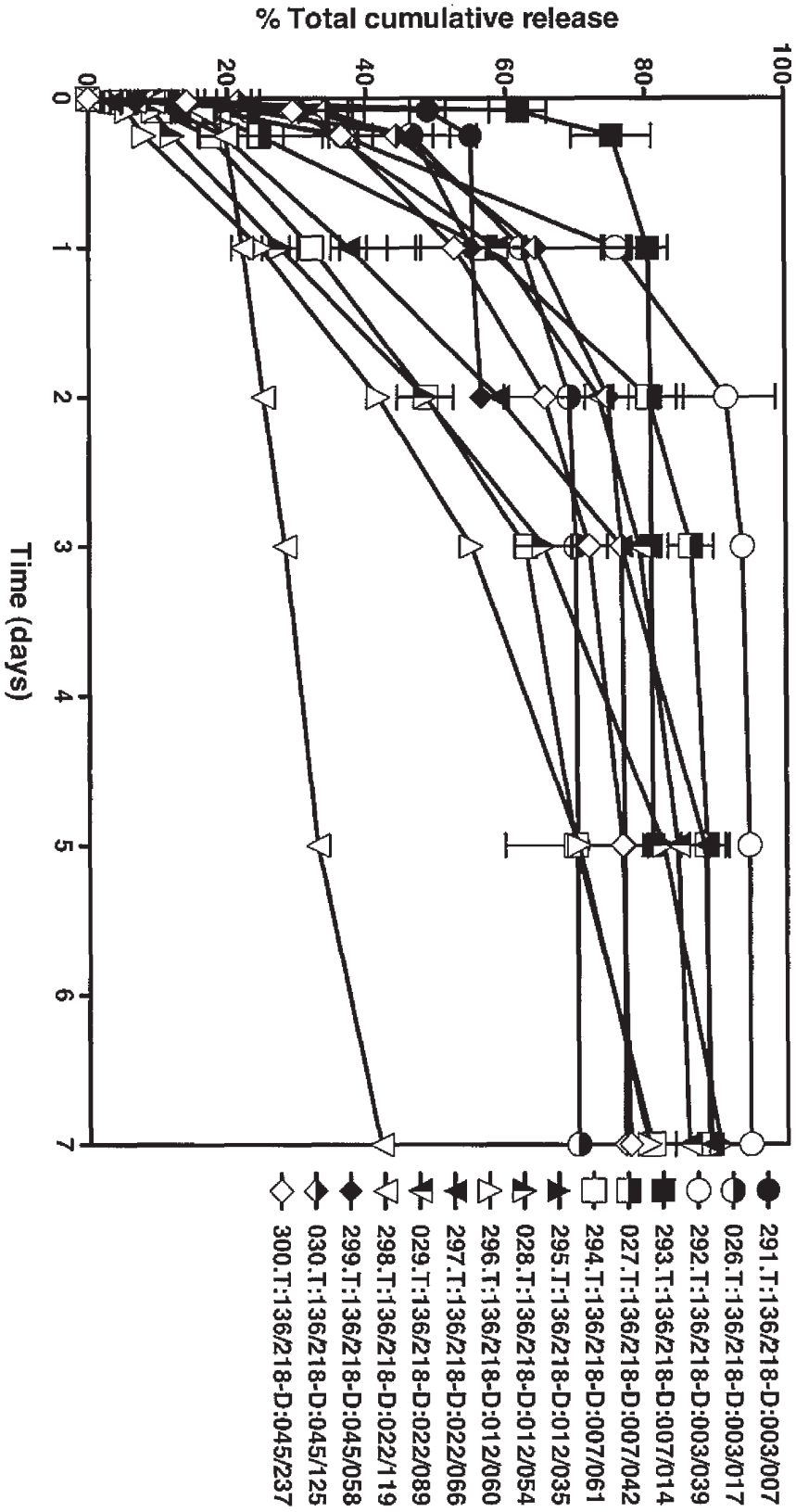

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
291.T:136/218-D:003/007
026.T:136/218-D:003/017
292.T:136/218-D:003/039
293.T:136/218-D:007/014
027.T:136/218-D:007/042
294.T:136/218-D:007/061
295.T:136/218-D:012/035
028.T:136/218-D:012/054
296.T:136/218-D:012/060
297.T:136/218-D:022/066
029.T:136/218-D:022/089
298.T:136/218-D:022/119
299.T:136/218-D:045/058
030.T:136/218-D:045/125
300.T:136/218-D:045/237

FIGURE 25

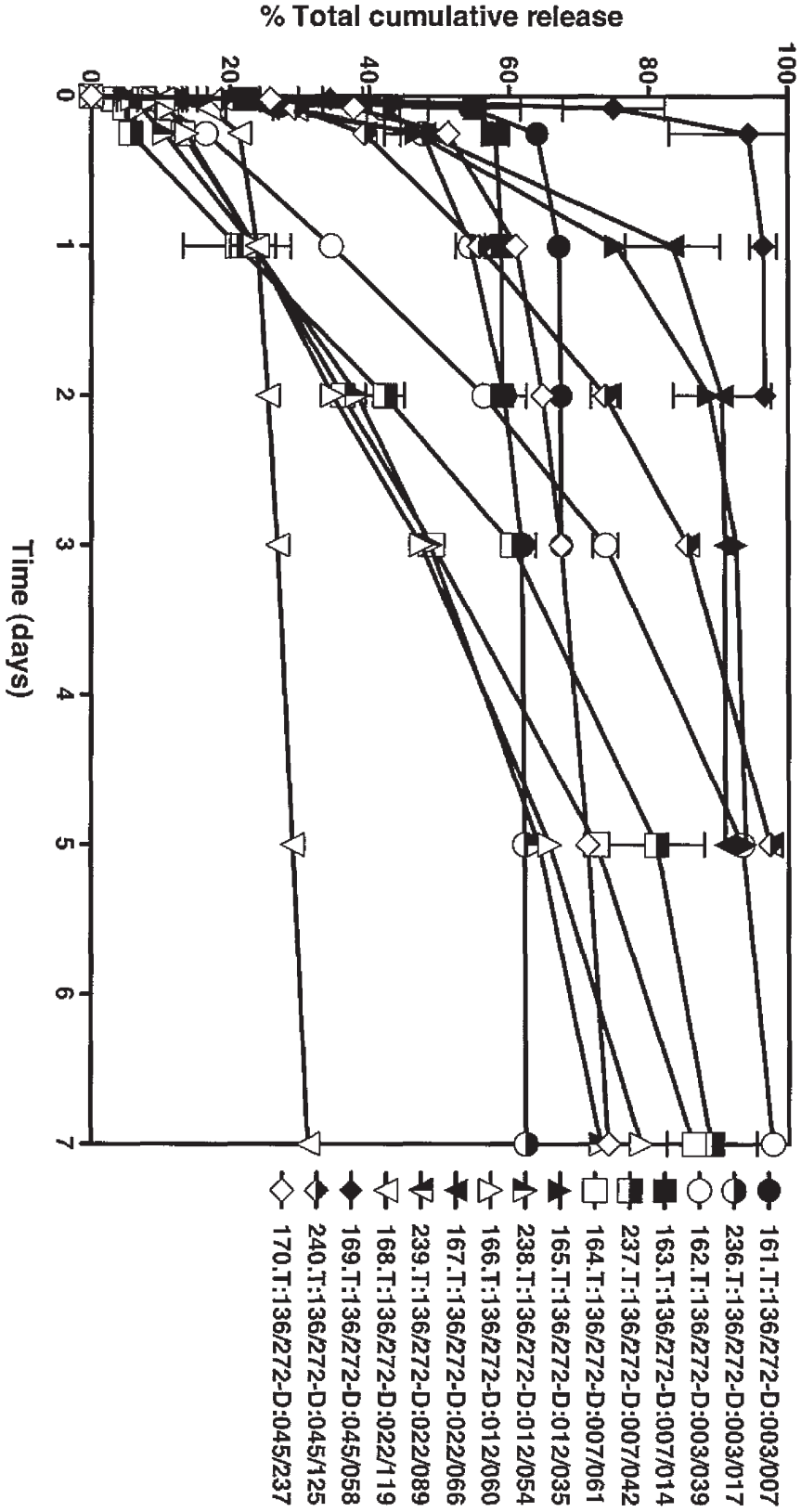

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
161.T:136/272-D:003/007
236.T:136/272-D:003/017
162.T:136/272-D:003/039
163.T:136/272-D:007/014
237.T:136/272-D:007/042
164.T:136/272-D:007/061
165.T:136/272-D:012/035
238.T:136/272-D:012/054
166.T:136/272-D:012/060
167.T:136/272-D:022/066
239.T:136/272-D:022/089
168.T:136/272-D:022/119
169.T:136/272-D:045/058
240.T:136/272-D:045/125
170.T:136/272-D:045/237

FIGURE 26

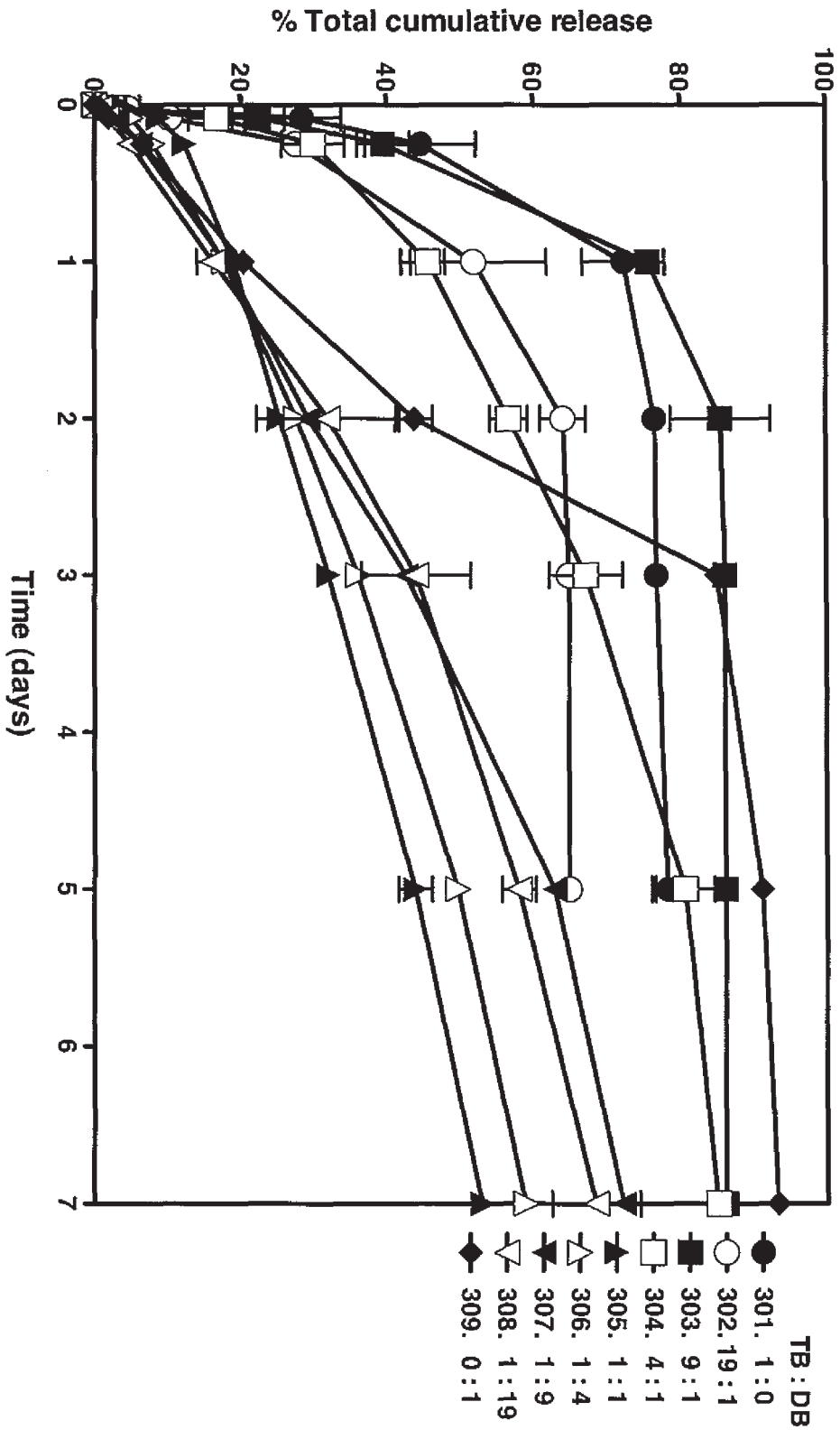

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
TB : DB
301. 1 : 0
302. 19 : 1
303. 9 : 1
304. 4 : 1
305. 1 : 1
306. 1 : 4
307. 1 : 9
308. 1 : 19
309. 0 : 1

FIGURE 27

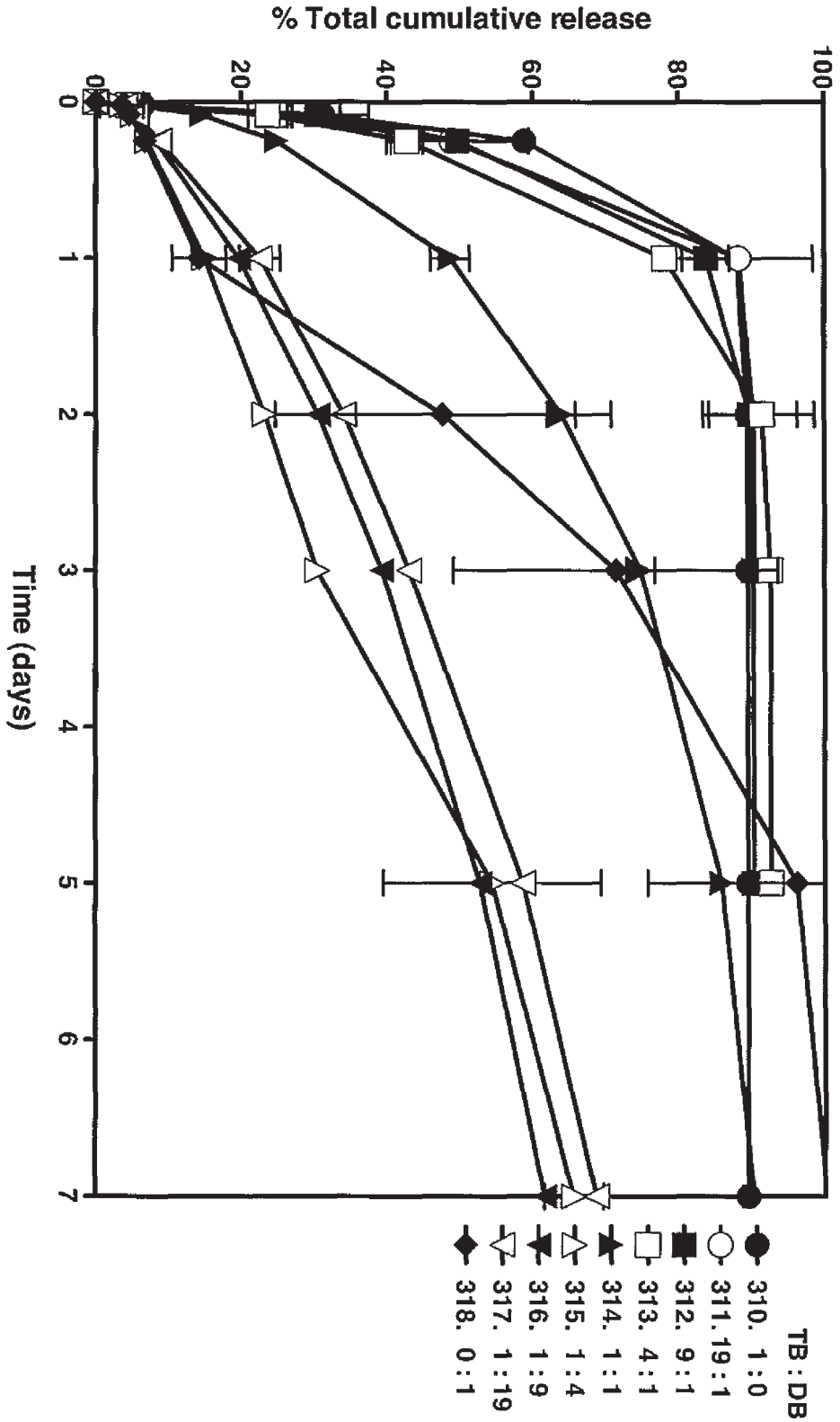

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
TB : DB
310. 1 : 0
311. 19 : 1
312. 9 : 1
313. 4 : 1
314. 1 : 1
315. 1 : 4
316. 1 : 9
317. 1 : 19
318. 0 : 1

FIGURE 28

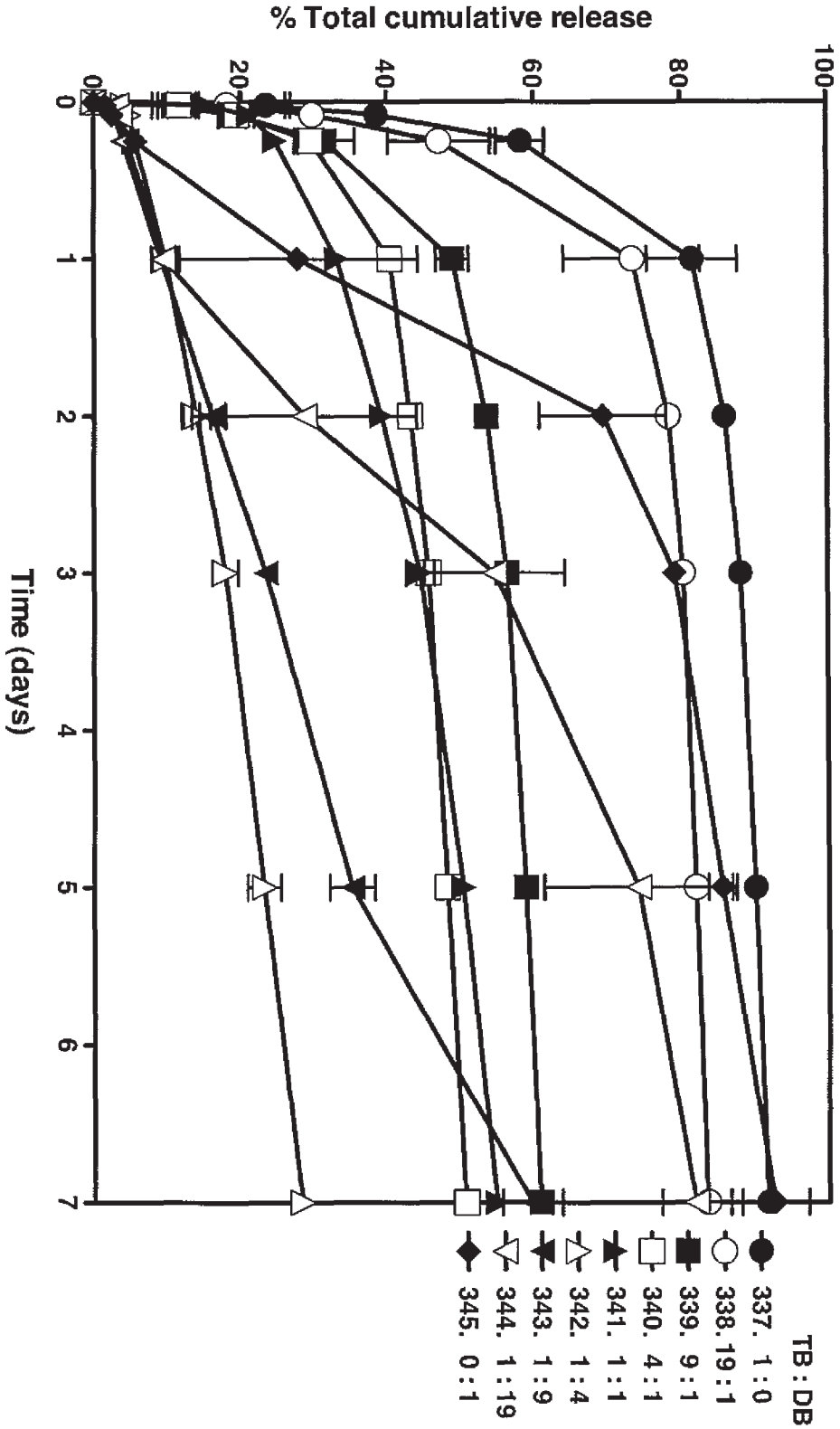

% Total cumulative release
0
20
40
60
80
100
Time (days)
0
1
2
3
4
5
6
7
TB : DB
337. 1 : 0
338. 19 : 1
339. 9 : 1
340. 4 : 1
341. 1 : 1
342. 1 : 4
343. 1 : 9
344. 1 : 19
345. 0 : 1

FIGURE 29

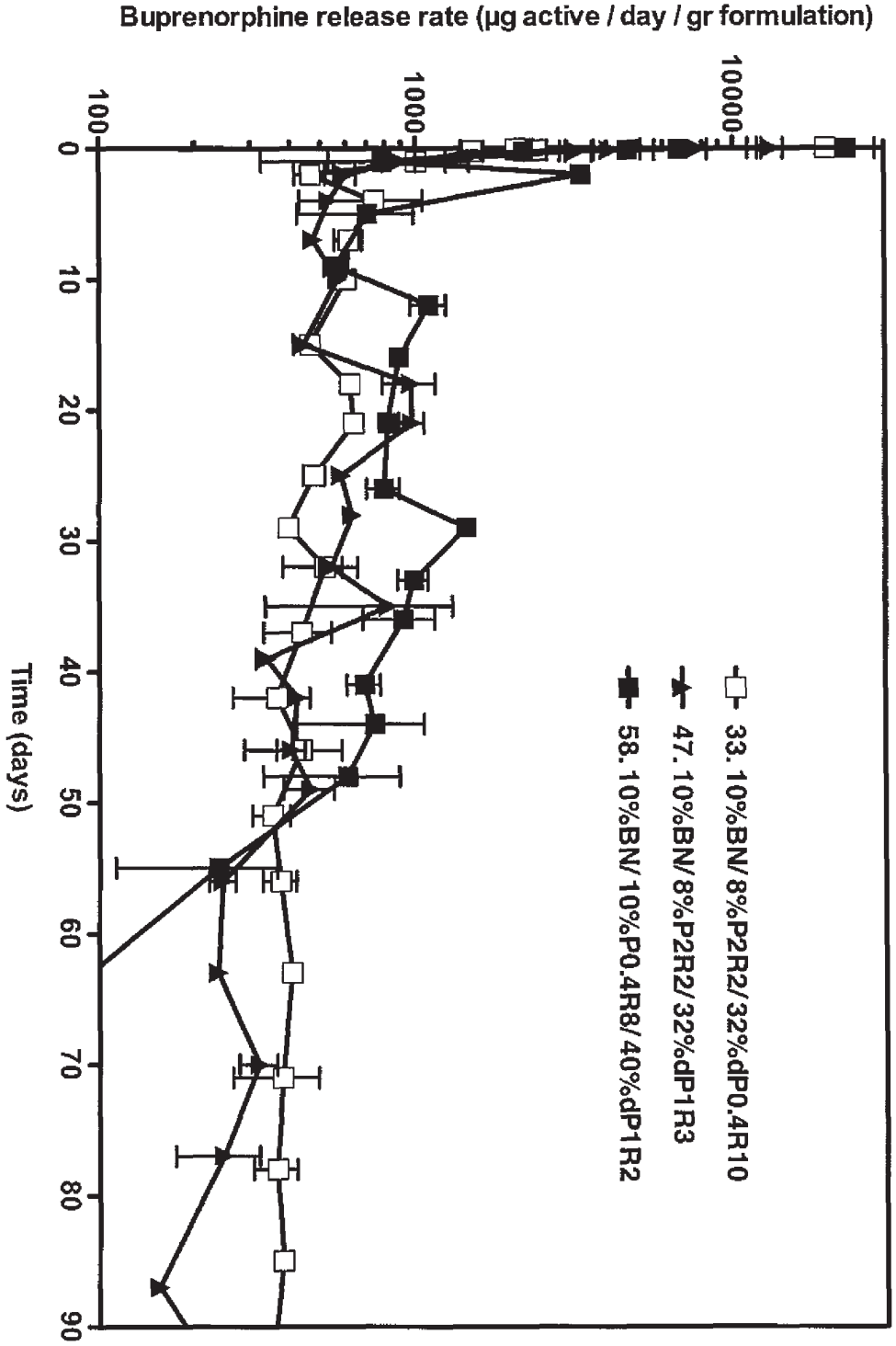

Buprenorphine release rate (µg active / day / gr formulation)
100
1000
10000
0
10
20
30
40
50
60
70
80
90
Time (days)
33.10%BN/8%P2R2/32%dP0.4R10
47.10%BN/8%P2R2/32%dP1R3
58.10%BN/10%P0.4R8/40%dP1R2

FIGURE 30

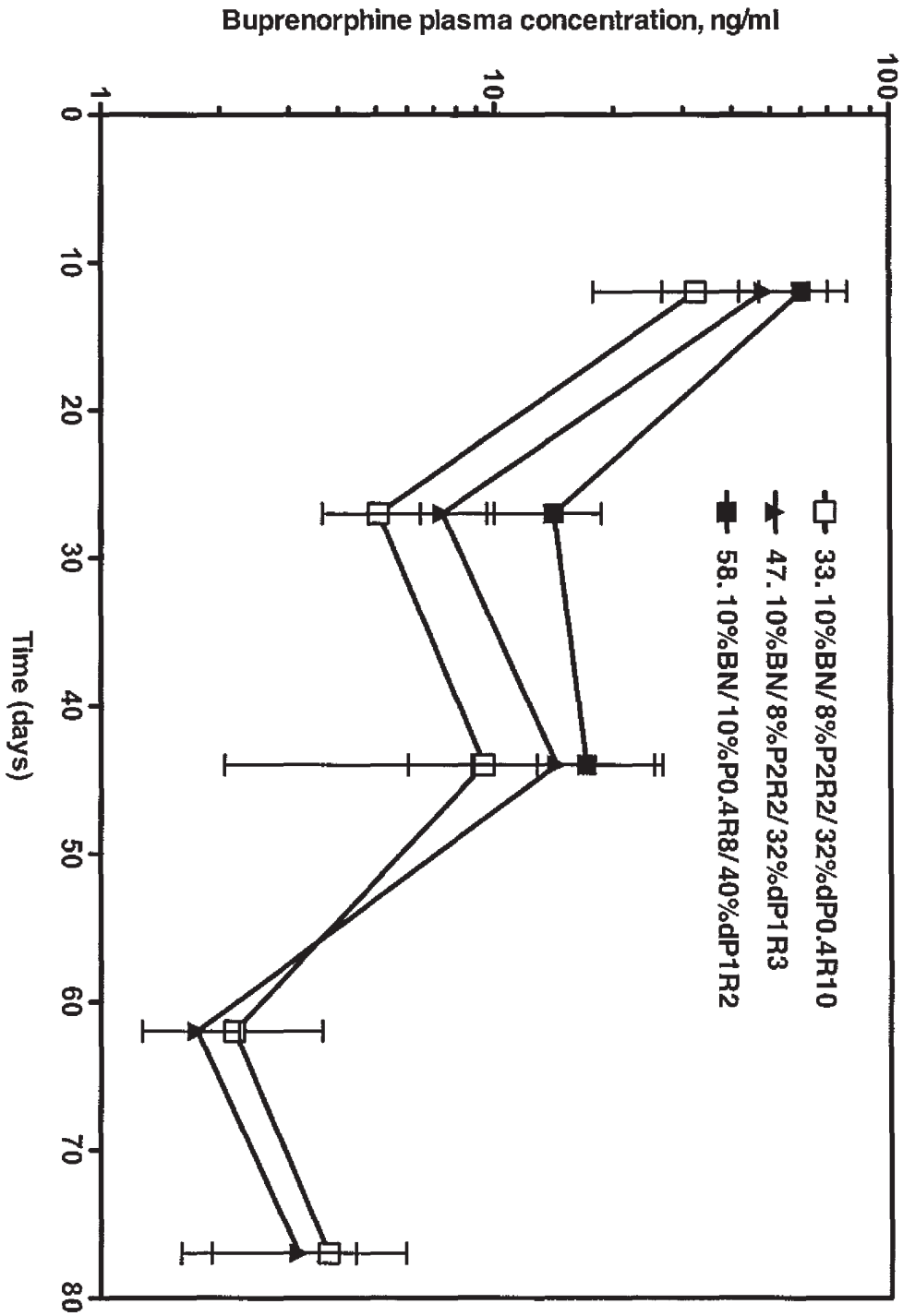

Buprenorphine plasma concentration, ng/ml
1
10
100
0
10
20
30
40
50
60
70
80
Time (days)
33. 10%BN/8%P2R2/32%dP0.4R10
47. 10%BN/8%P2R2/32%dP1R3
58. 10%BN/10%P0.4R8/40%dP1R2

FIGURE 31

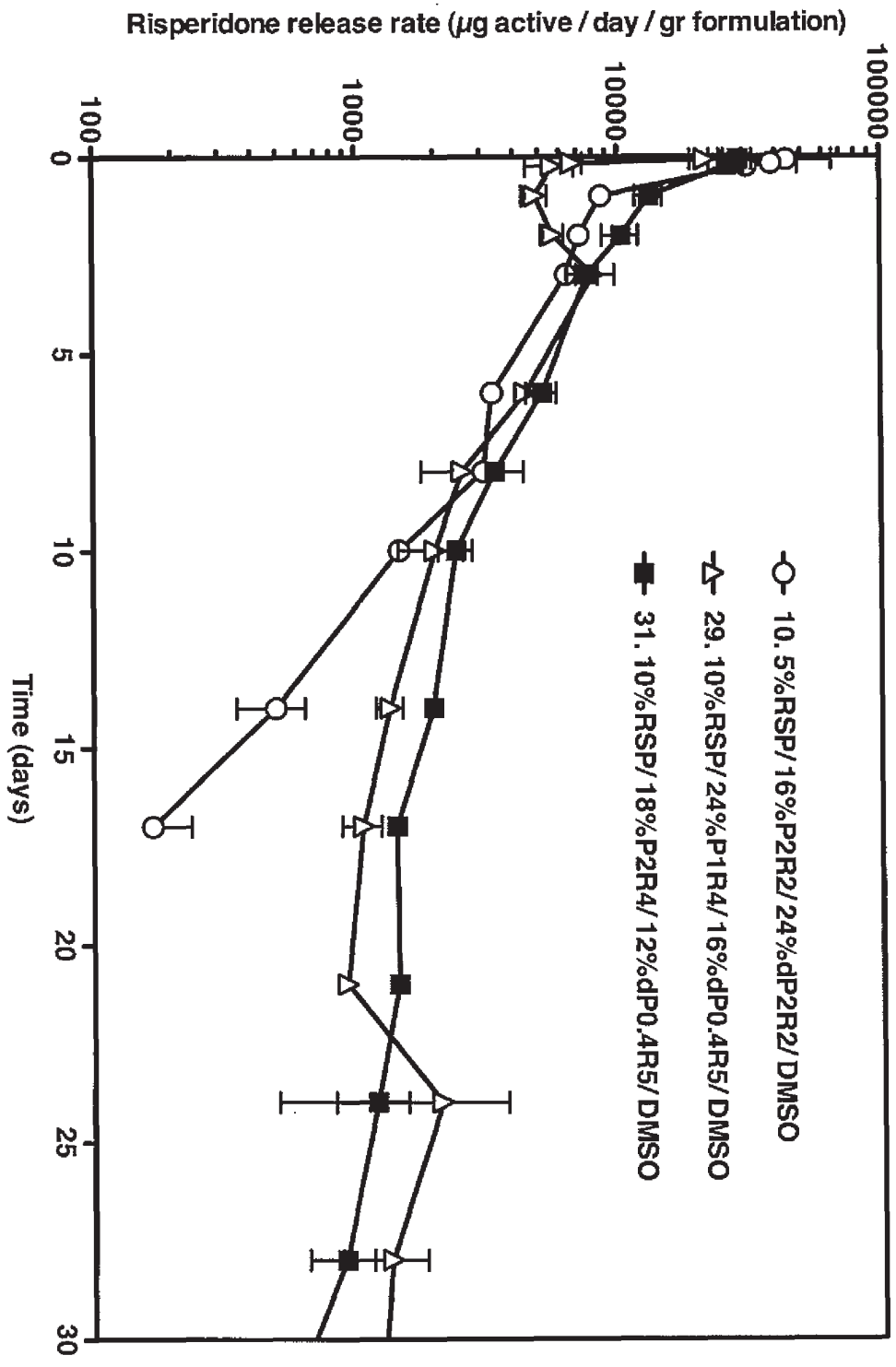

Risperidone release rate (µg active / day / gr formulation)
100
1000
10000
100000
0
5
10
15
20
25
30
Time (days)
10.5%RSP/16%P2R2/24%dP2R2/DMSO
29.10%RSP/24%P1R4/16%dP0.4R5/DMSO
31.10%RSP/18%P2R4/12%dP0.4R5/DMSO

FIGURE 32

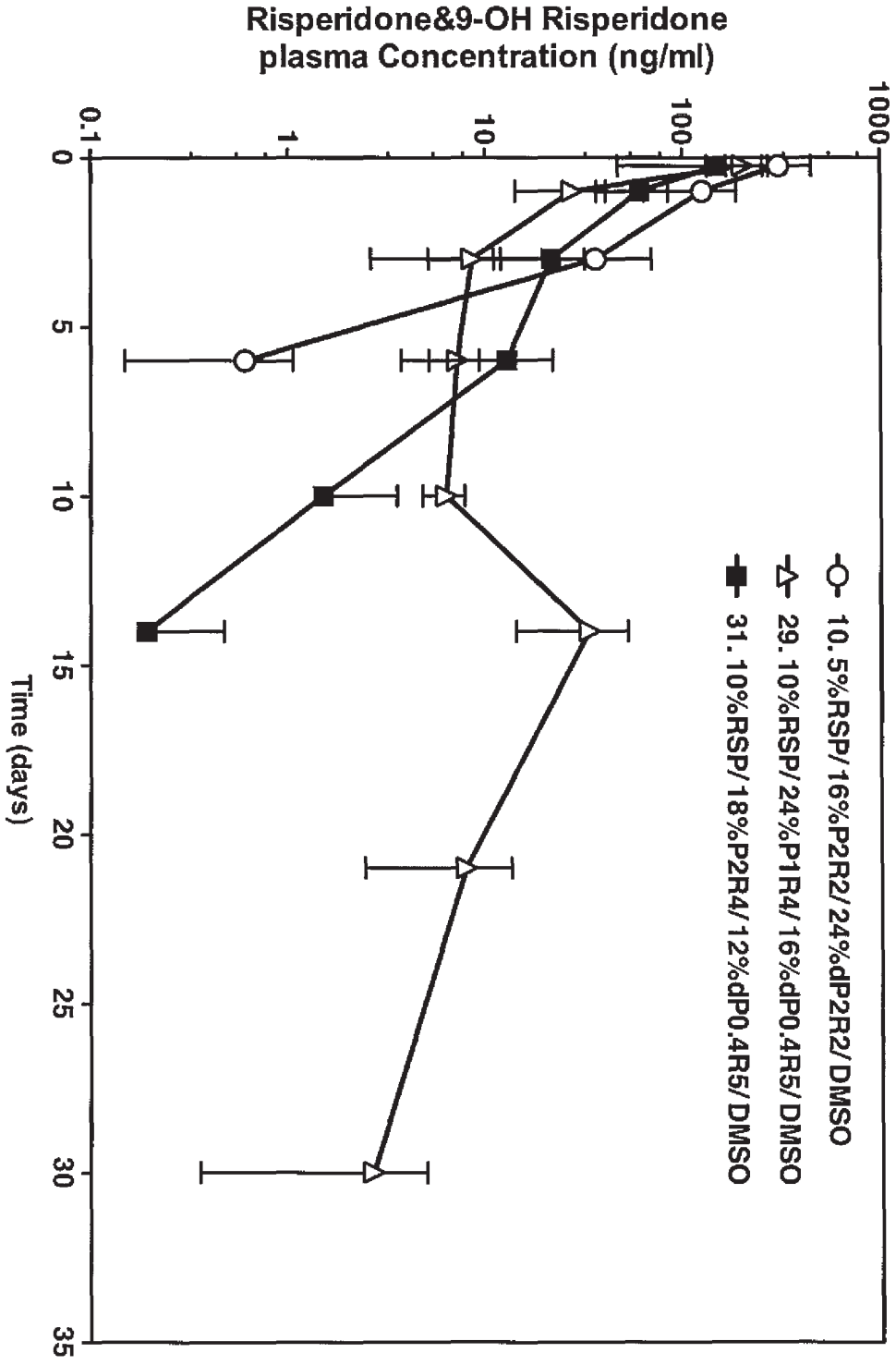

Risperidone&9-OH Risperidone
plasma Concentration (ng/ml)
0.1
1
10
100
1000
0
5
10
15
20
25
30
35
Time (days)
10. 5%RSP/16%P2R2/24%dP2R2/DMSO
29. 10%RSP/24%P1R4/16%dP0.4R5/DMSO
31. 10%RSP/18%P2R4/12%dP0.4R5/DMSO

FIGURE 33

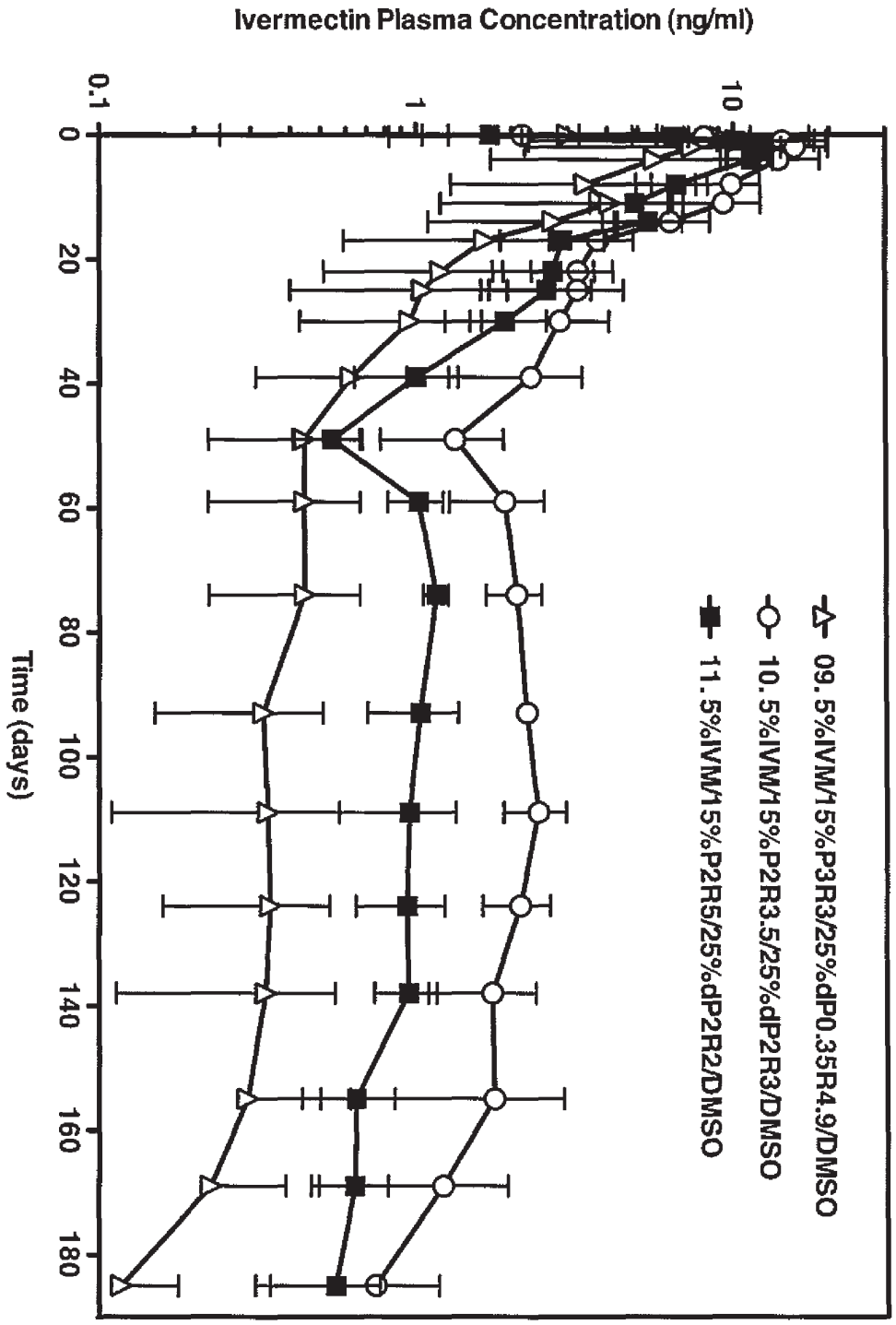

Ivermectin Plasma Concentration (ng/ml)
0.1
1
10
0
20
40
60
80
100
120
140
160
180
Time (days)
09.5%IVM/15%P3R3/25%dP0.35R4.9/DMSO
10.5%IVM/15%P2R3.5/25%dP2R3/DMSO
11.5%IVM/15%P2R5/25%dP2R2/DMSO

FIGURE 34

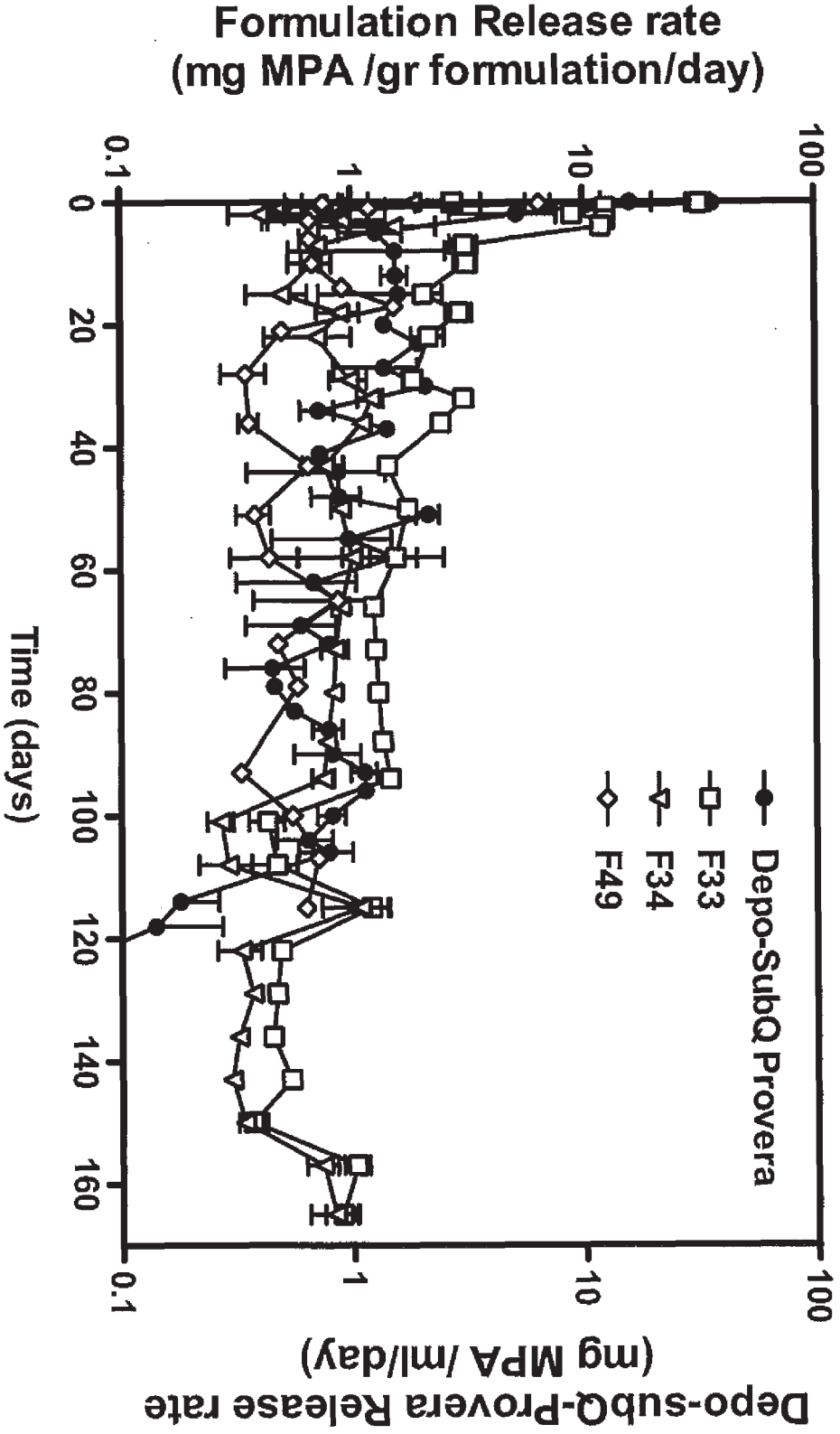

Formulation Release rate
(mg MPA /gr formulation/day)
Depo-subQ-Provera Release rate
(mg MPA /ml/day)
Time (days)
0.1
1
10
100
0
20
40
60
80
100
120
140
160
Depo-SubQ Provera
F33
F34
F49

FIGURE 35

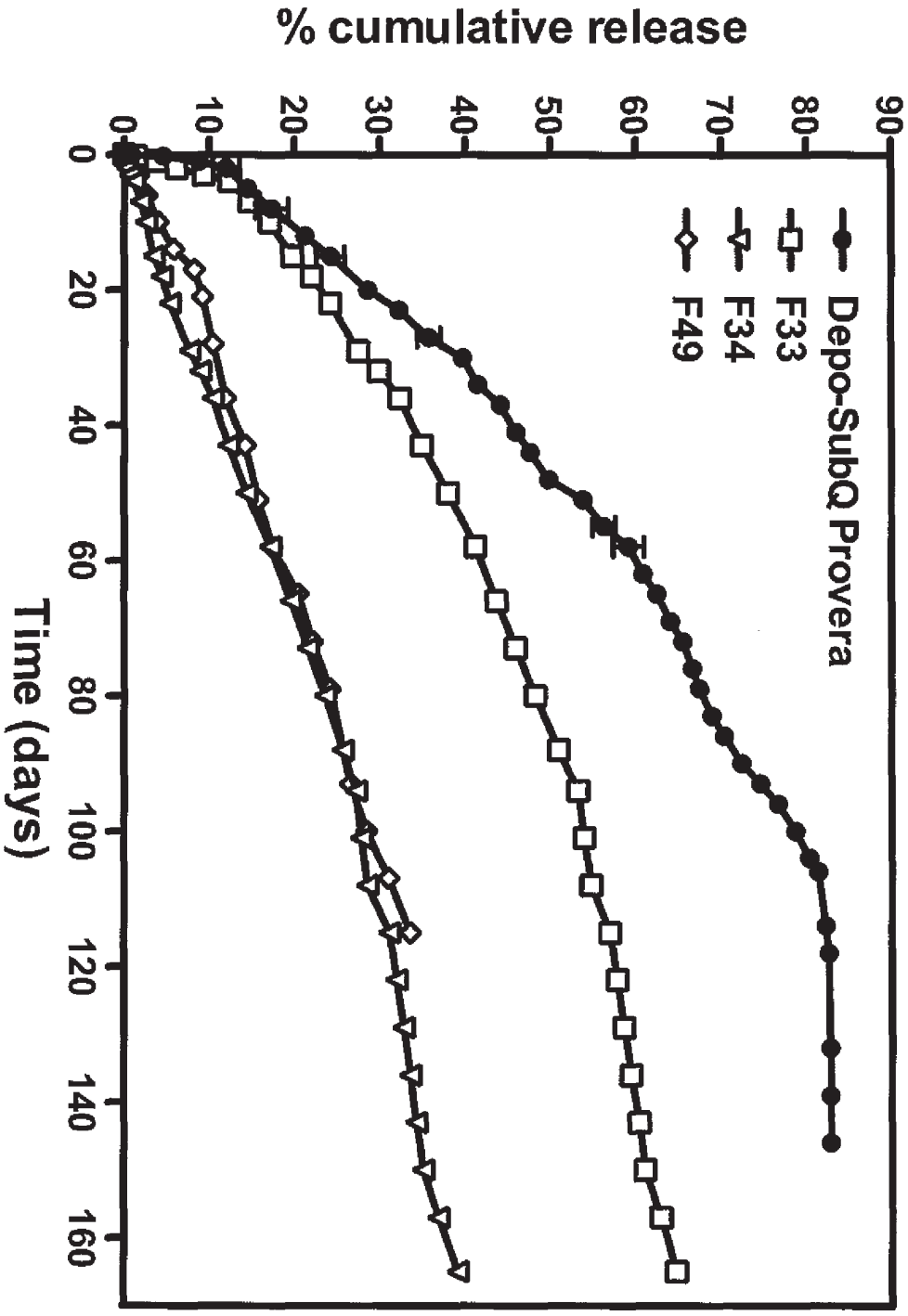

% cumulative release
0
10
20
30
40
50
60
70
80
90
Time (days)
0
20
40
60
80
100
120
140
160
Depo-SubQ Provera
F33
F34
F49

FIGURE 36

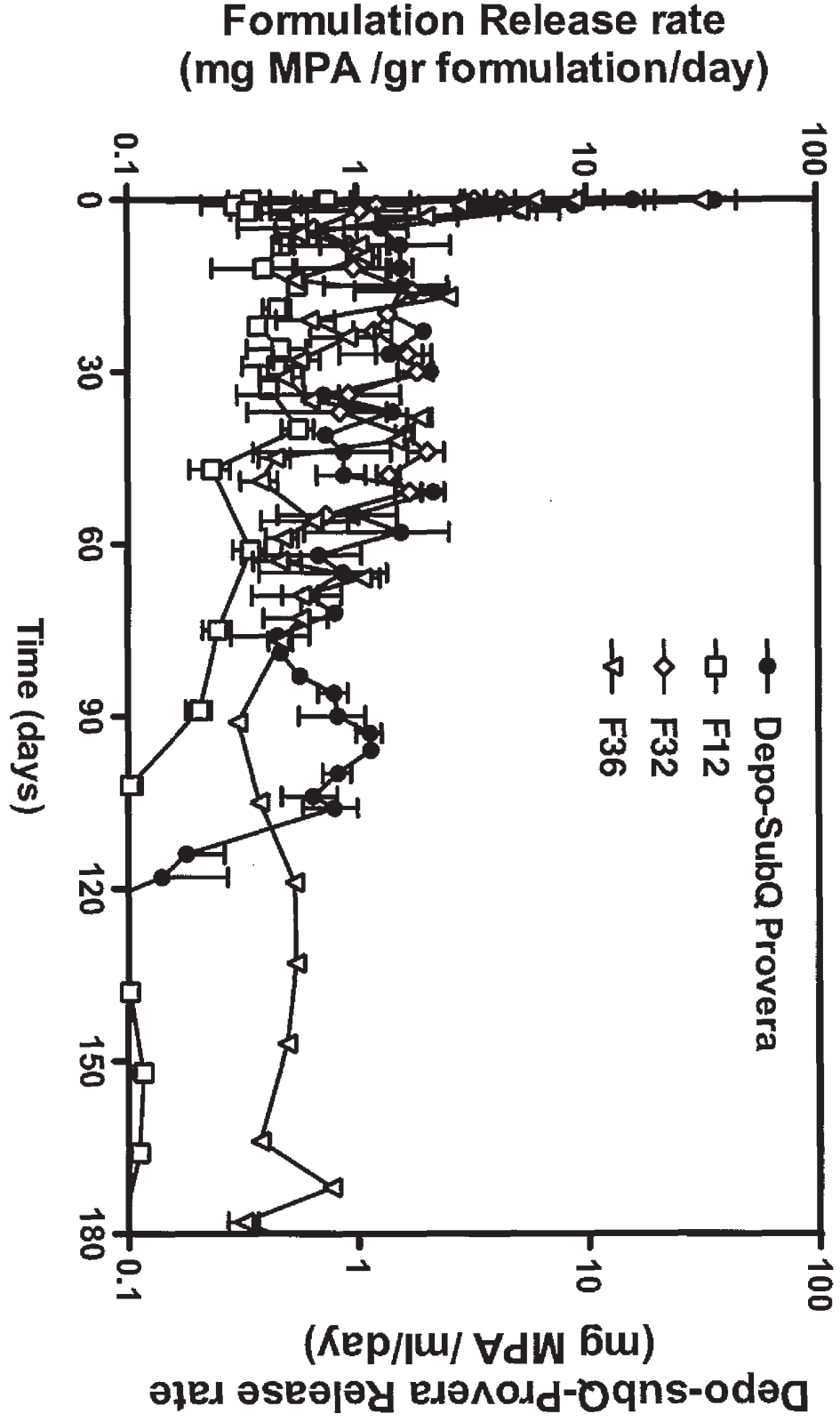

Formulation Release rate
(mg MPA /gr formulation/day)
Depo-subQ-Provera Release rate
(mg MPA /ml/day)
Time (days)
0.1
1
10
100
0
30
60
90
120
150
180
Depo-SubQ Provera
F12
F32
F36

FIGURE 37

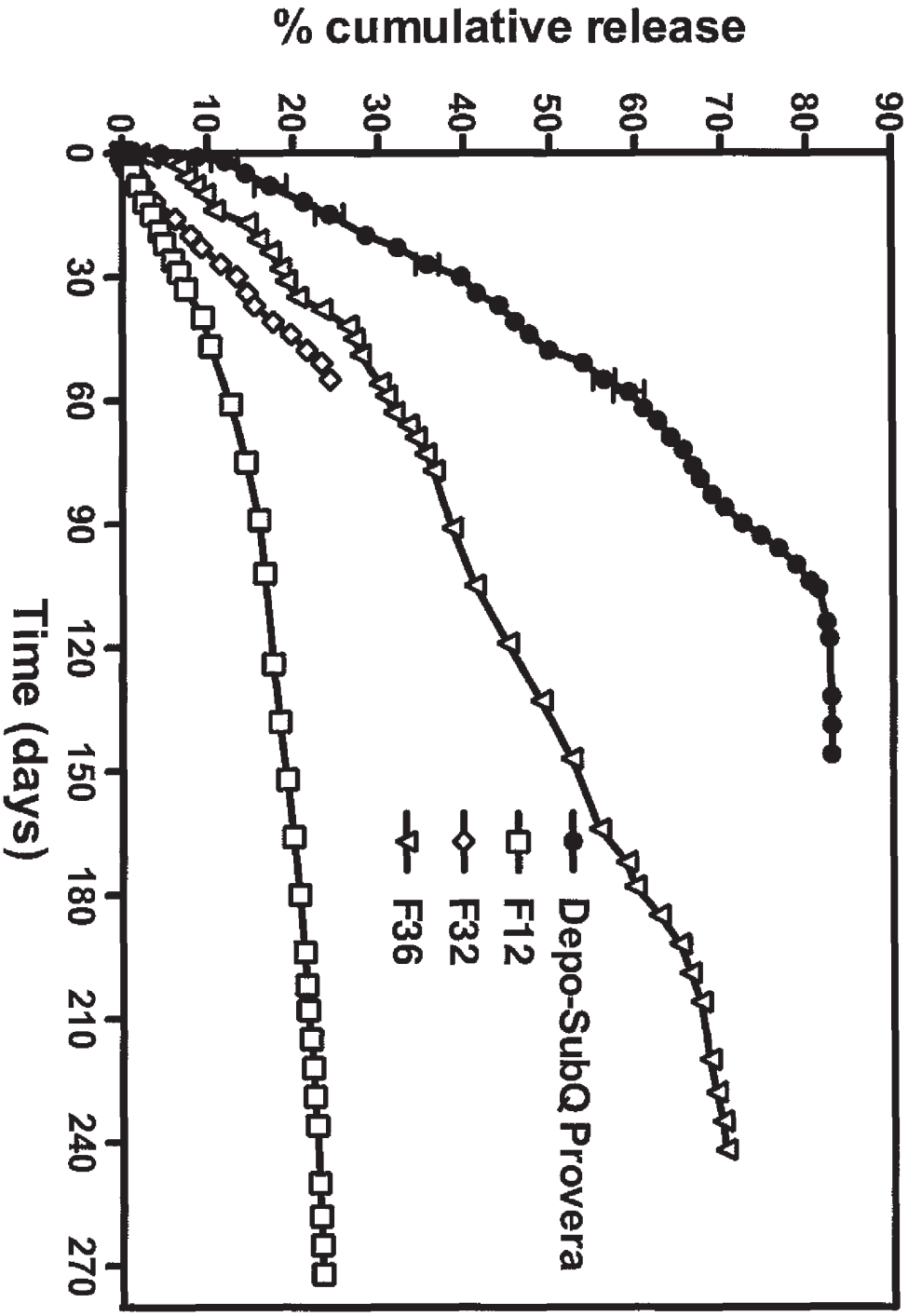

% cumulative release
0
10
20
30
40
50
60
70
80
90
Time (days)
0
30
60
90
120
150
180
210
240
270
Depo-SubQ Provera
F12
F32
F36

FIGURE 38

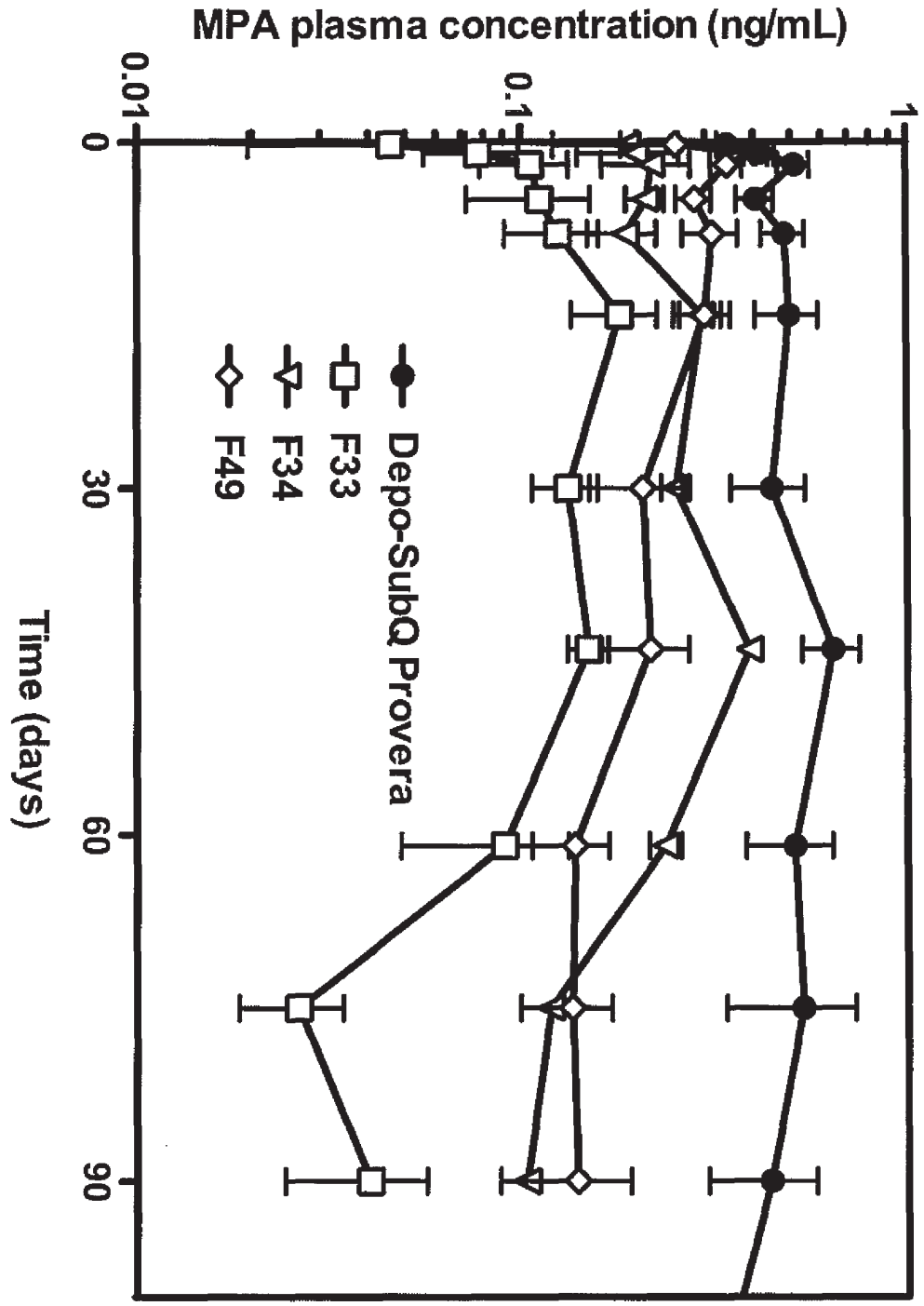

MPA plasma concentration (ng/mL)
0.01
0.1
1
0
30
60
90
Time (days)
Depo-SubQ Provera
F33
F34
F49

FIGURE 39

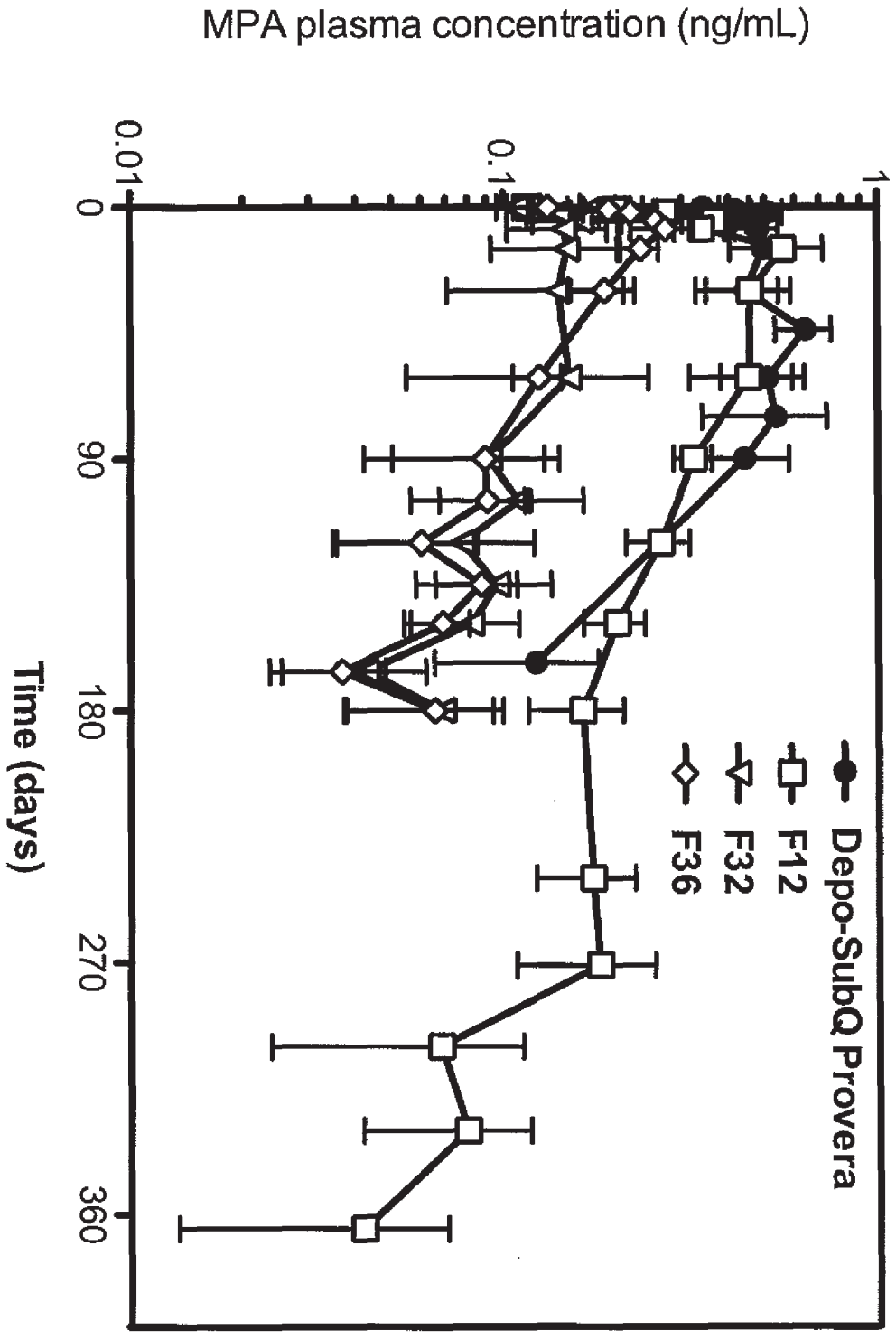

MPA plasma concentration (ng/mL)
0.01
0.1
1
0
90
180
270
360
Time (days)
Depo-SubQ Provera
F12
F32
F36

FIGURE 40

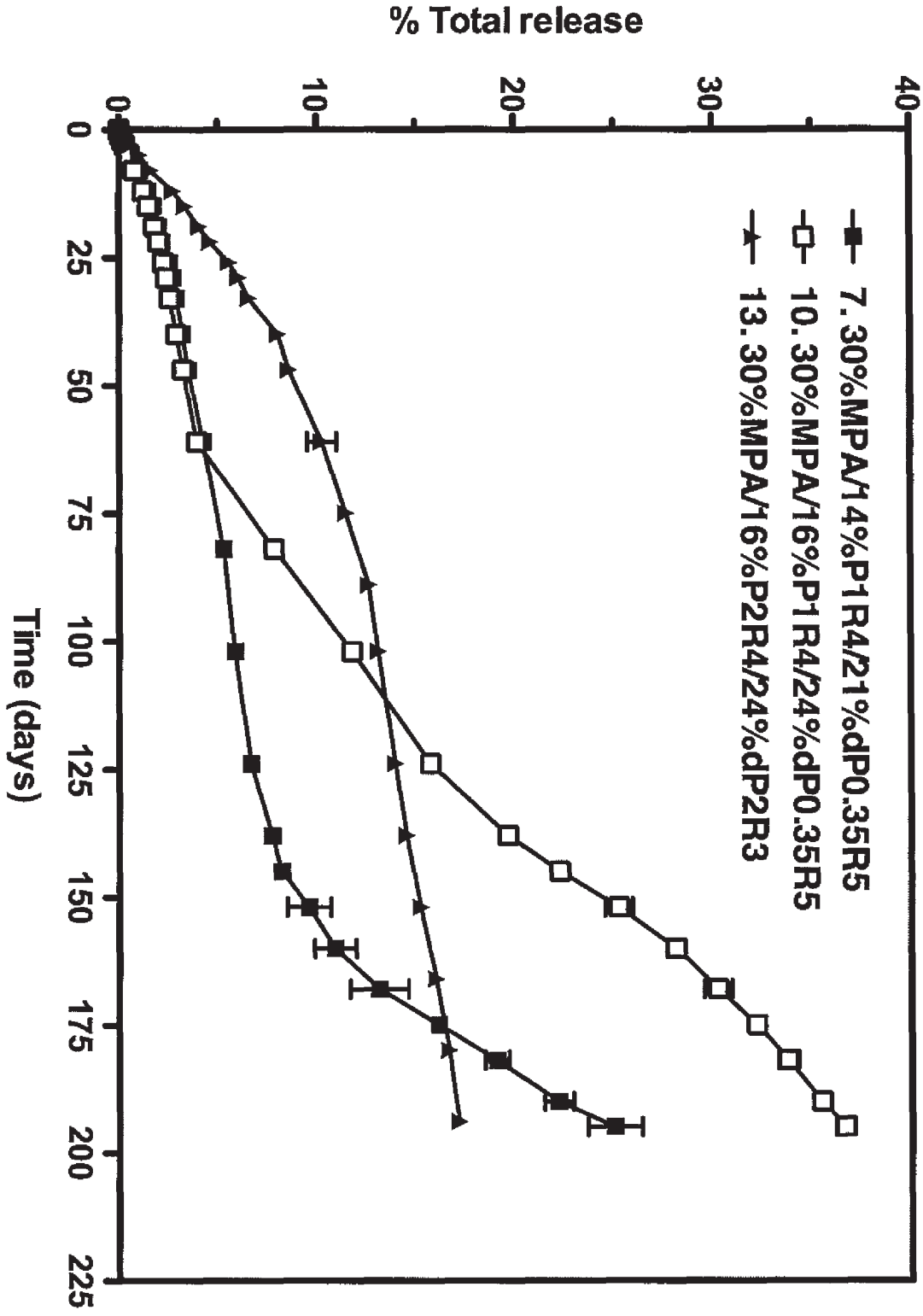

% Total release
0
10
20
30
40
Time (days)
0
25
50
75
100
125
150
175
200
225
7.30%MPA/14%P1R4/21%dP0.35R5
10.30%MPA/16%P1R4/24%dP0.35R5
13.30%MPA/16%P2R4/24%dP2R3

FIGURE 41

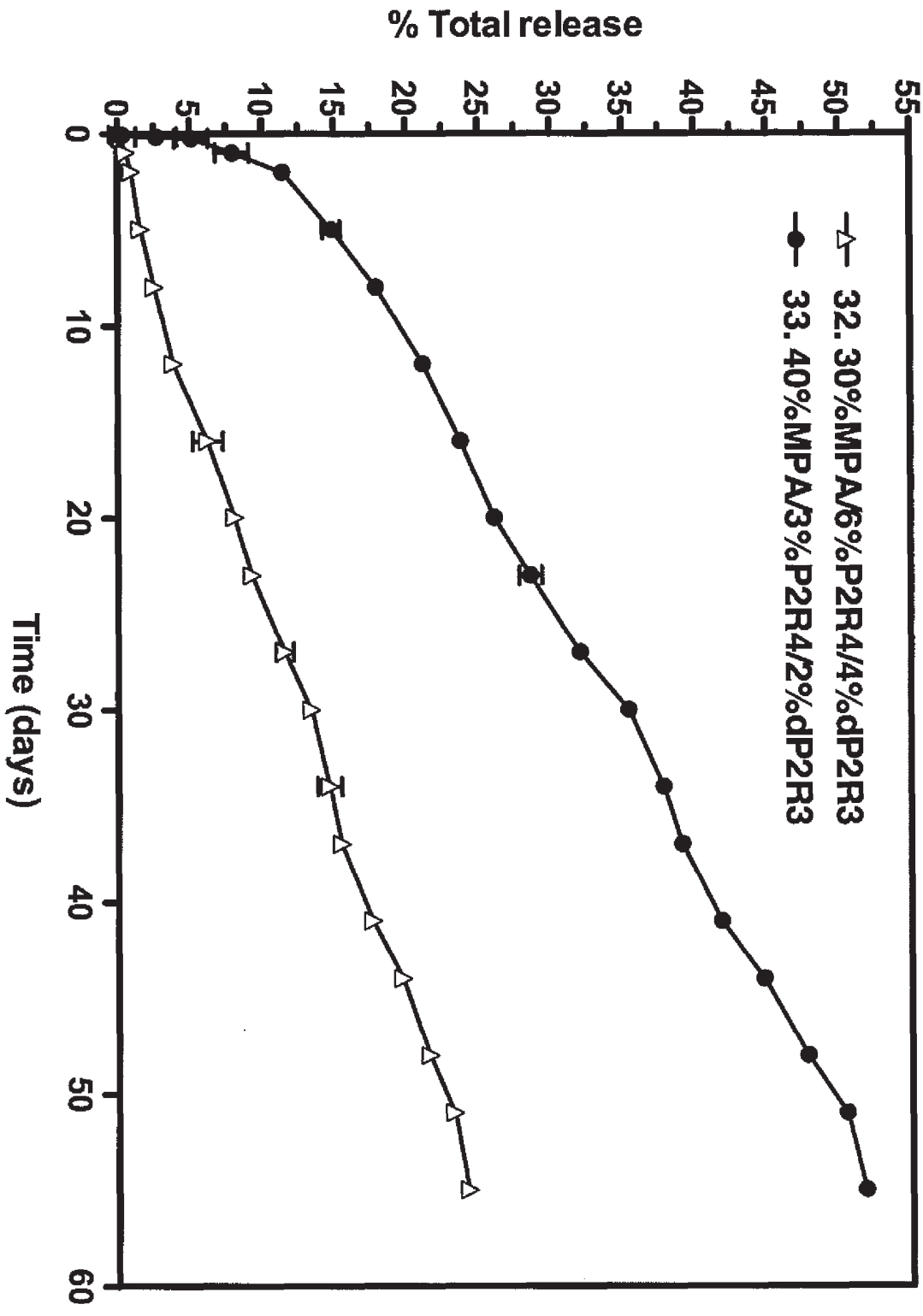

% Total release
0
5
10
15
20
25
30
35
40
45
50
55
Time (days)
0
10
20
30
40
50
60
32. 30%MPA/6%P2R4/4%dP2R3
33. 40%MPA/3%P2R4/2%dP2R3

FIGURE 42

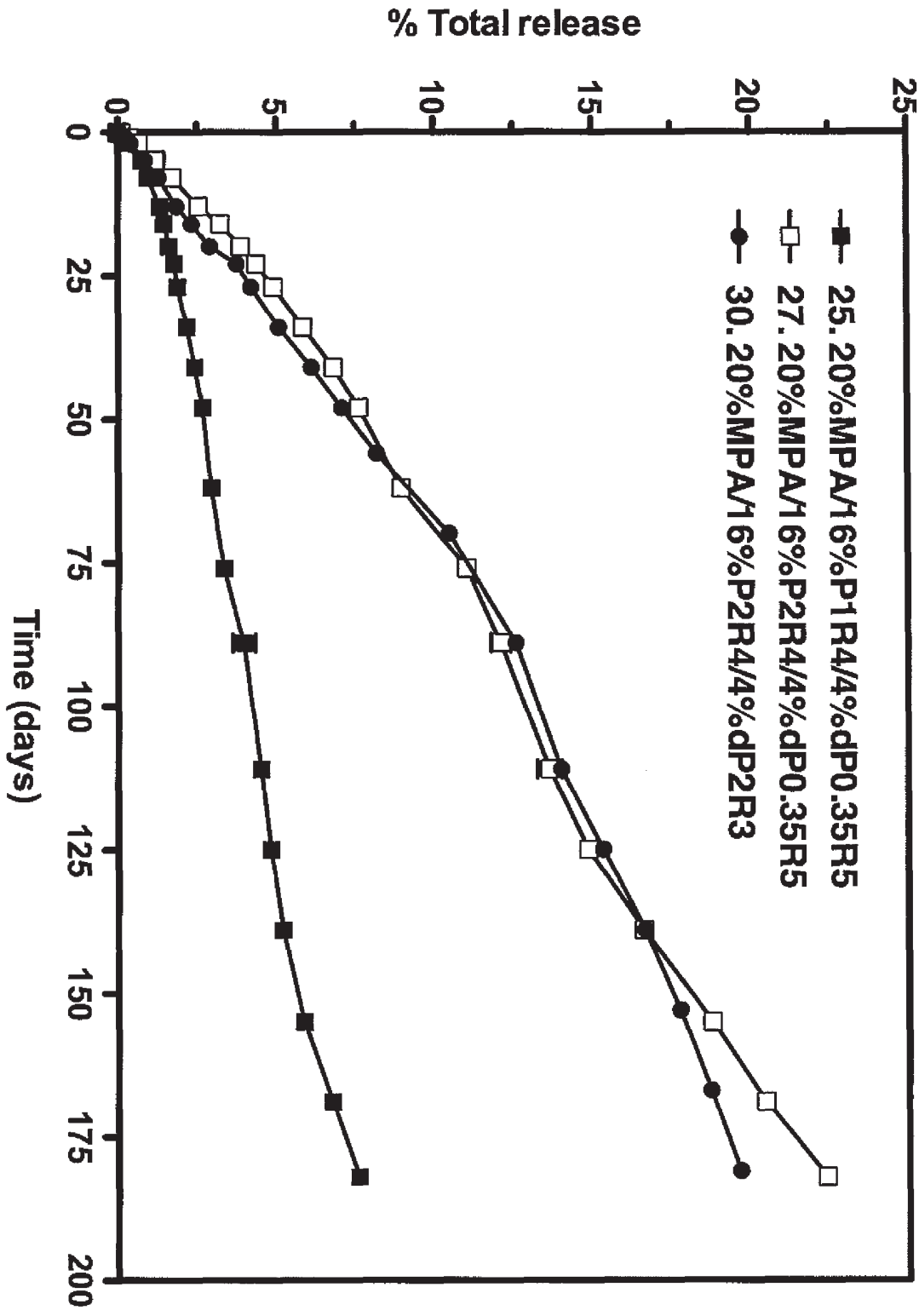

% Total release
0
5
10
15
20
25
Time (days)
0
25
50
75
100
125
150
175
200
25. 20%MPA/16%P1R4/4%dP0.35R5
27. 20%MPA/16%P2R4/4%dP0.35R5
30. 20%MPA/16%P2R4/4%dP2R3

FIGURE 43

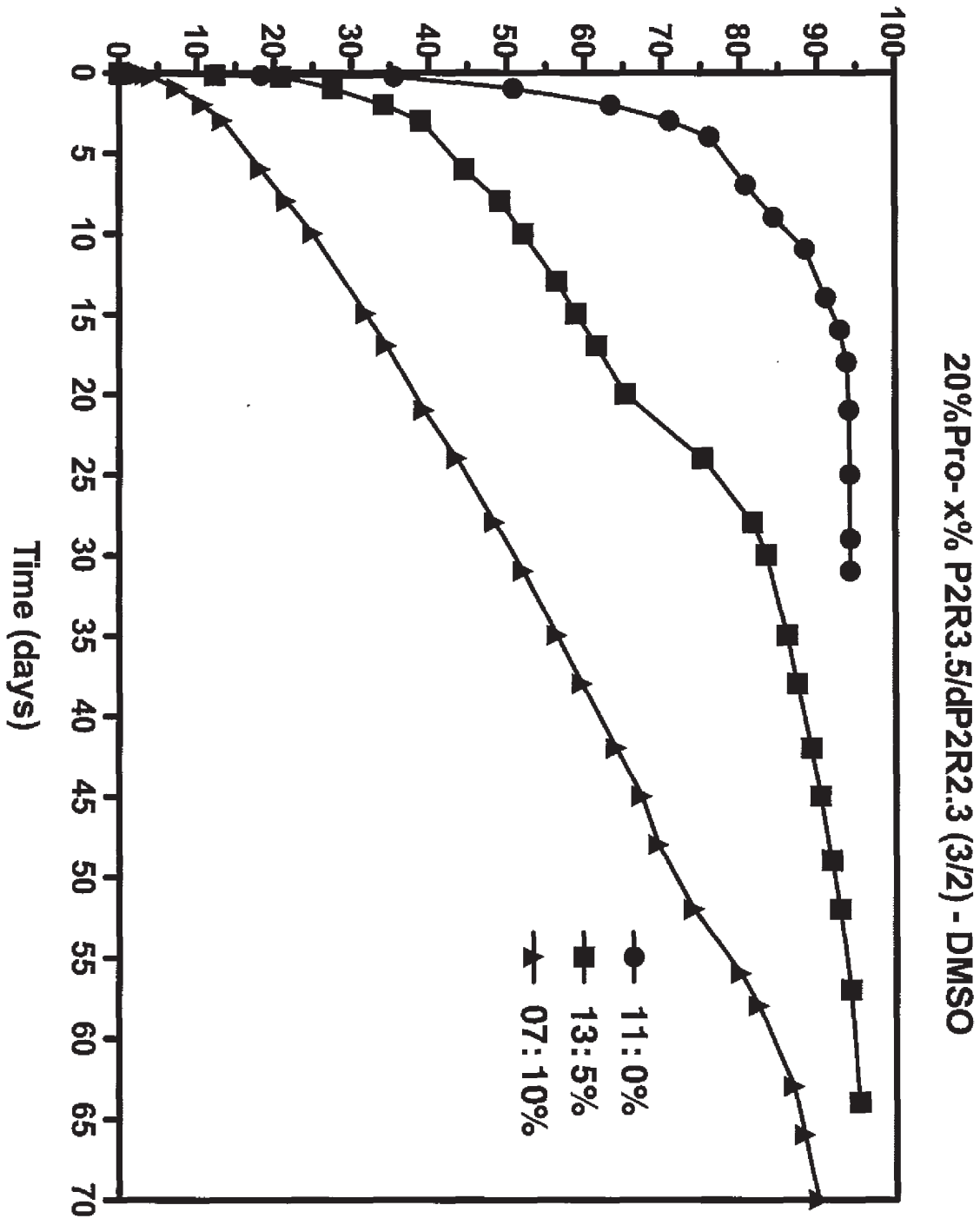

% Total release
0
10
20
30
40
50
60
70
80
90
100
Time (days)
0
5
10
15
20
25
30
35
40
45
50
55
60
65
70
20%Pro- x% P2R3.5/dP2R2.3 (3/2) - DMSO
11:0%
13:5%
07:10%

FIGURE 44

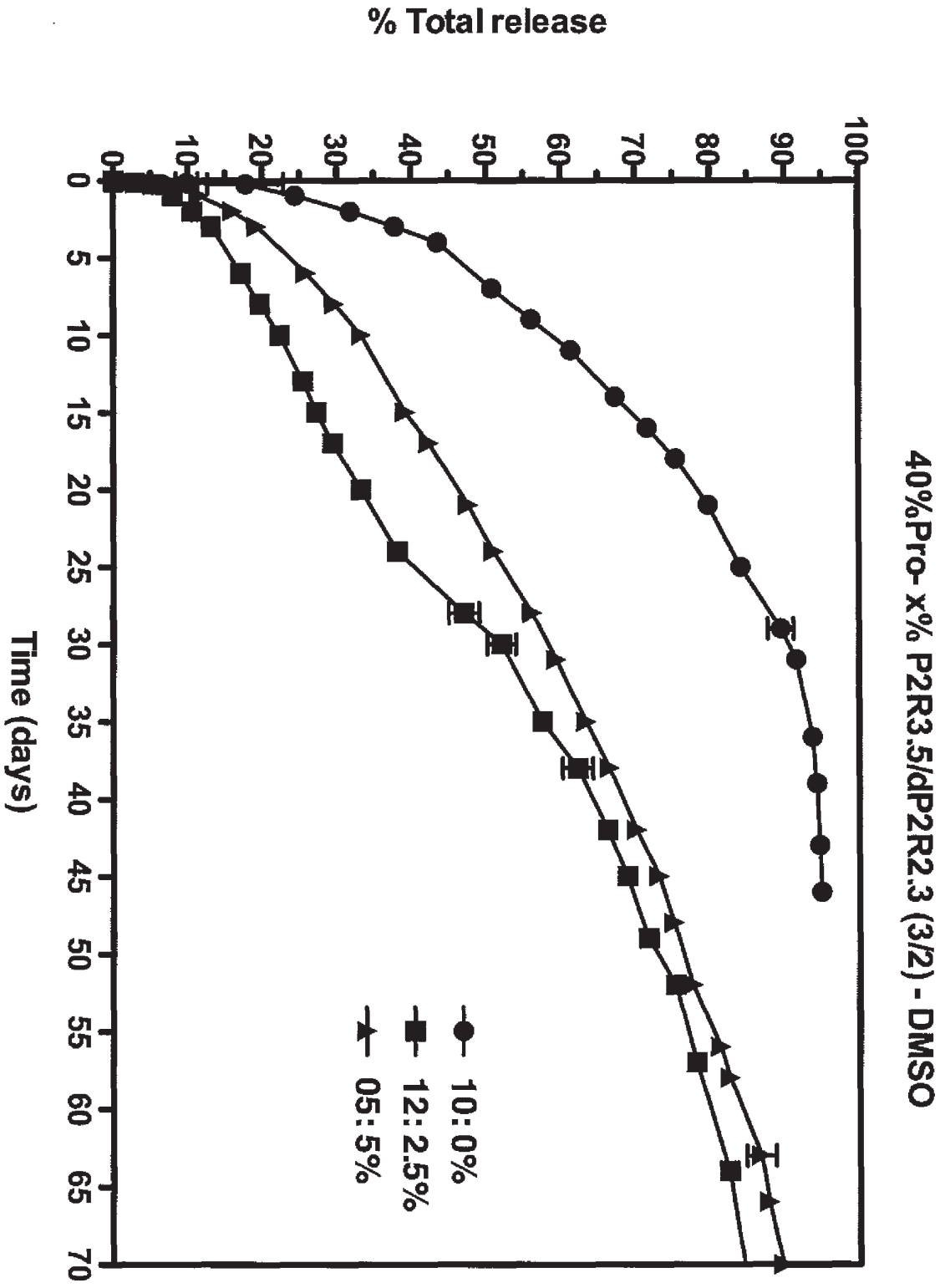

% Total release
0
10
20
30
40
50
60
70
80
90
100
Time (days)
0
5
10
15
20
25
30
35
40
45
50
55
60
65
70
40%Pro- x% P2R3.5/dP2R2.3 (3/2) - DMSO
10: 0%
12: 2.5%
05: 5%

FIGURE 45

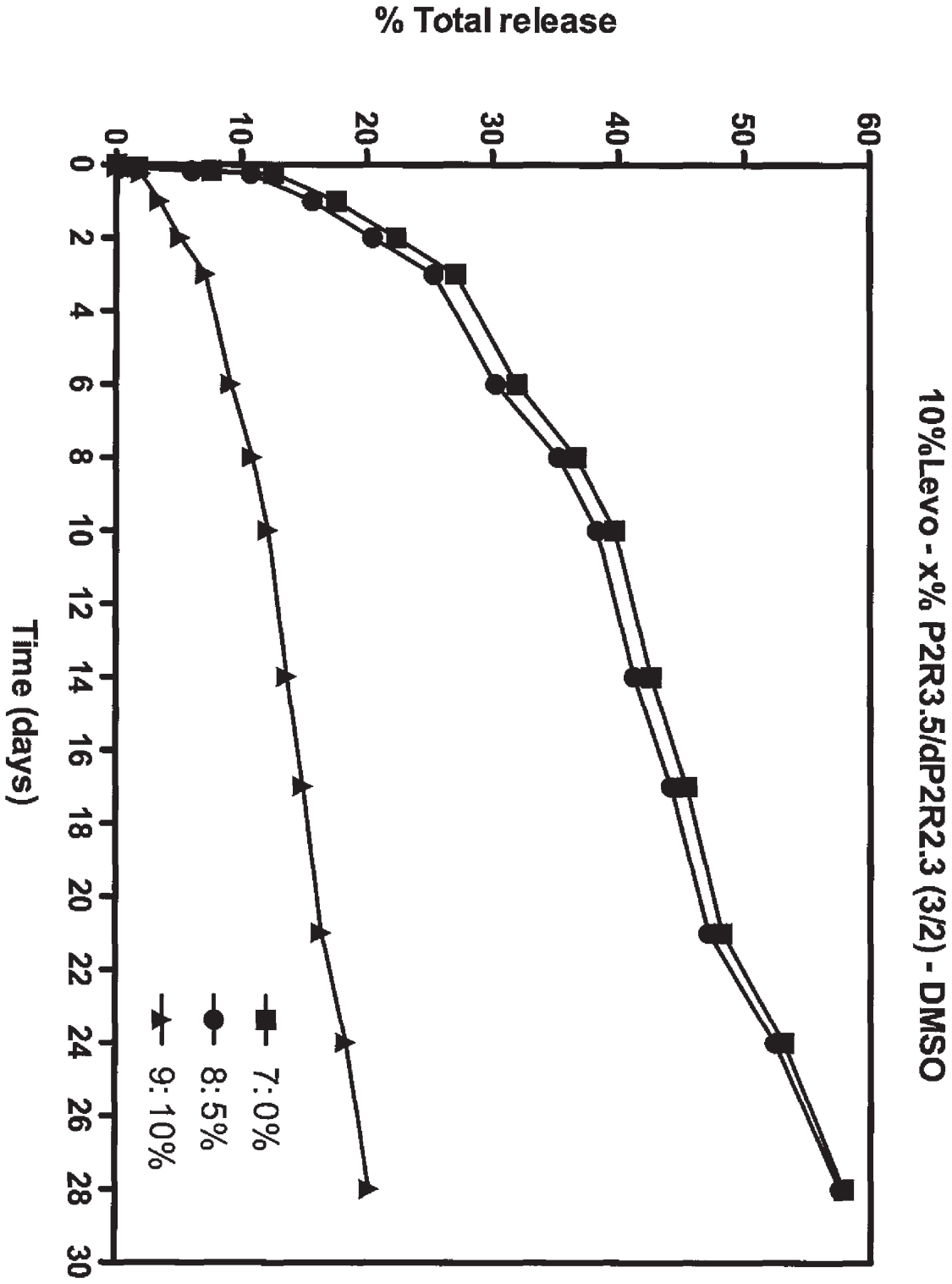

% Total release
0
10
20
30
40
50
60
0
2
4
6
8
10
12
14
16
18
20
22
24
26
28
30
Time (days)
7: 0%
8: 5%
9: 10%
10%Levo - x% P2R3.5/dP2R2.3 (3/2) - DMSO

FIGURE 46

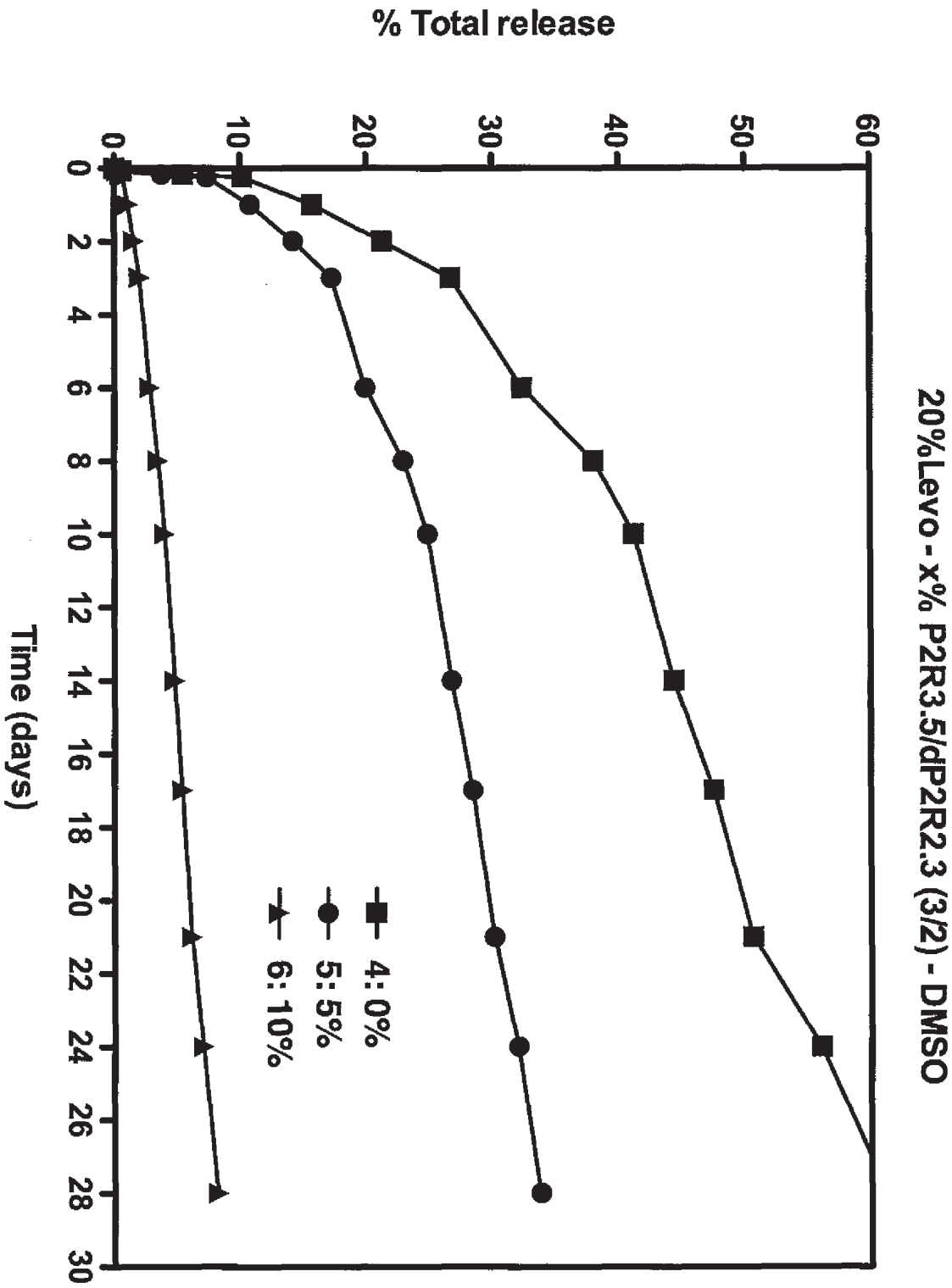

% Total release
0
10
20
30
40
50
60
0
2
4
6
8
10
12
14
16
18
20
22
24
26
28
30
Time (days)
4: 0%
5: 5%
6: 10%
20%Levo - x% P2R3.5/dP2R2.3 (3/2) - DMSO

FIGURE 47

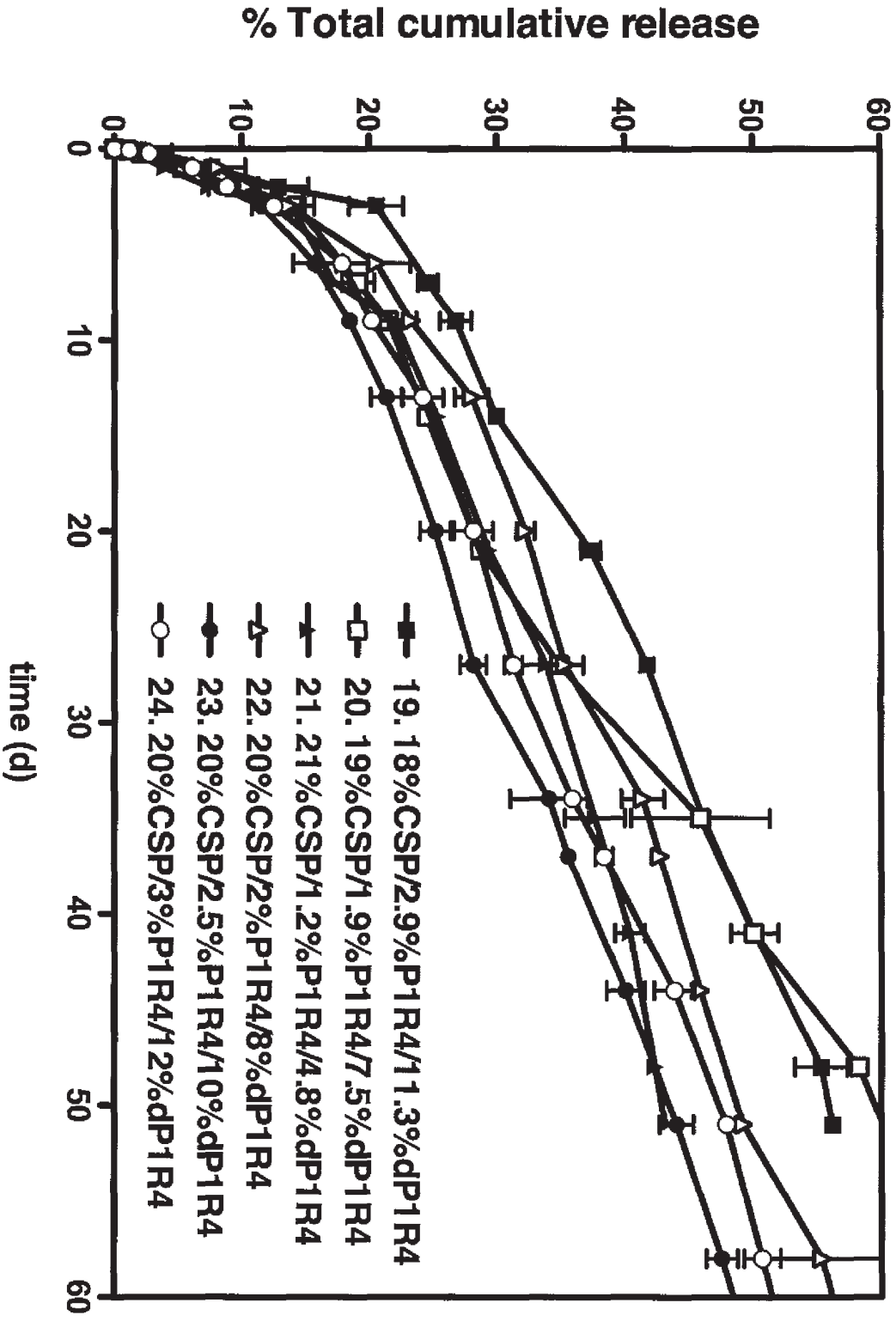

% Total cumulative release
0
10
20
30
40
50
60
time (d)
19. 18%CSP/2.9%P1R4/11.3%dP1R4
20. 19%CSP/1.9%P1R4/7.5%dP1R4
21. 21%CSP/1.2%P1R4/4.8%dP1R4
22. 20%CSP/2%P1R4/8%dP1R4
23. 20%CSP/2.5%P1R4/10%dP1R4
24. 20%CSP/3%P1R4/12%dP1R4

FIGURE 48

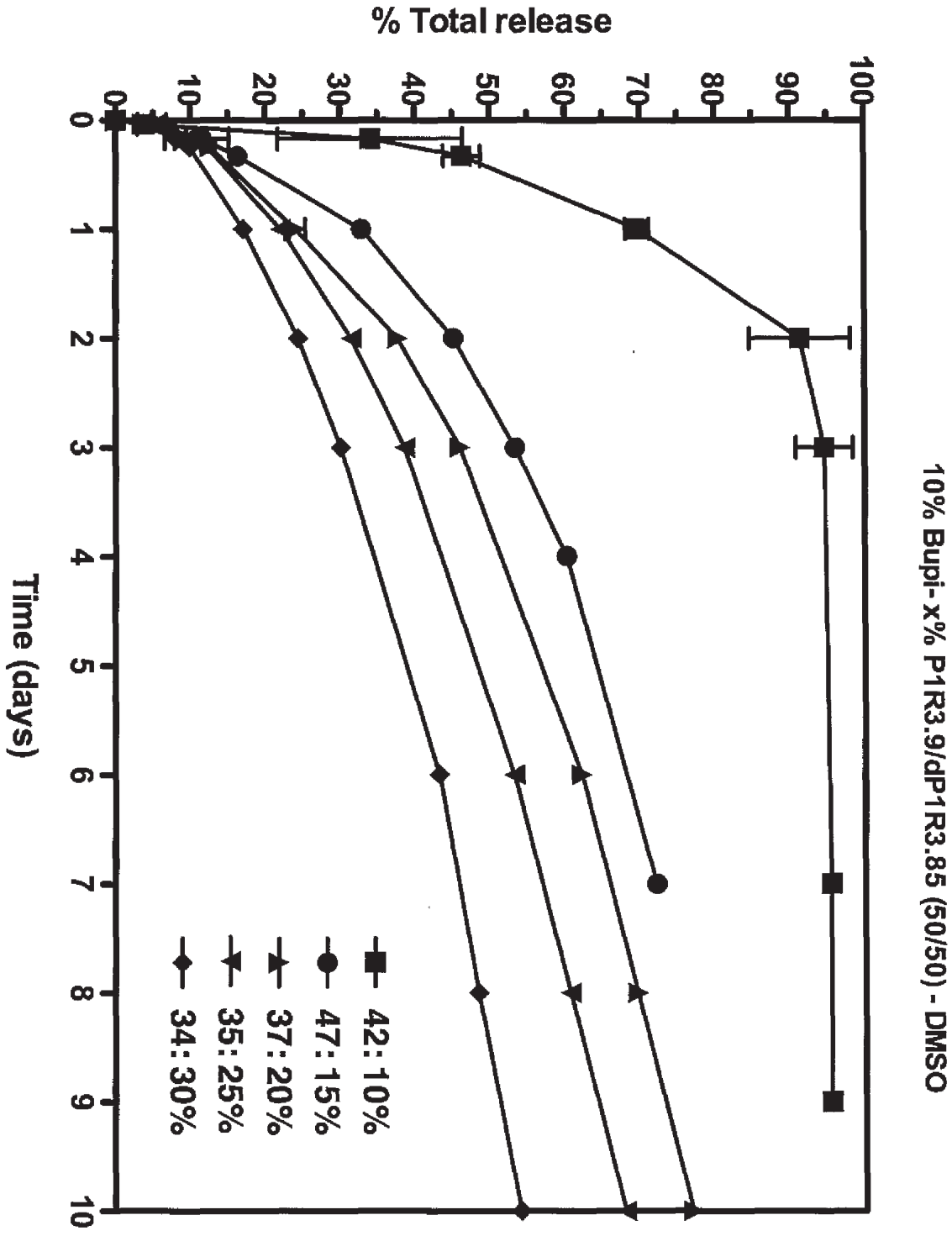

% Total release
0
10
20
30
40
50
60
70
80
90
100
Time (days)
0
1
2
3
4
5
6
7
8
9
10
42:10%
47:15%
37:20%
35:25%
34:30%
10% Bupi- x% P1R3.9/dP1R3.85 (50/50) - DMSO

FIGURE 49