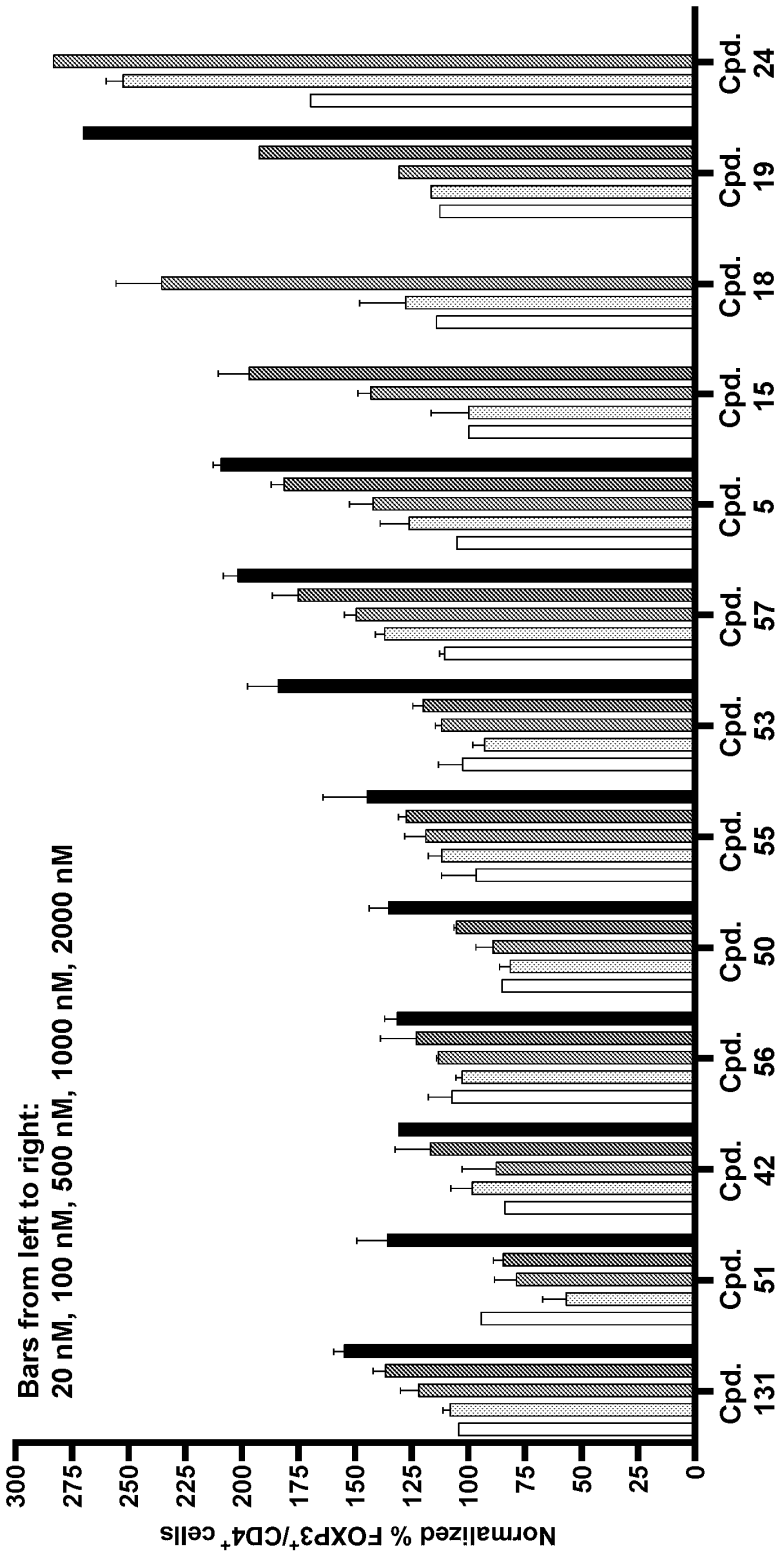
Normalized % FOXP3+/CD4+ cells
300
275
250
225
200
175
150
125
100
75
50
25
0
Bars from left to right:
20 nM, 100 nM, 500 nM, 1000 nM, 2000 nM
Cpd. 131
Cpd. 51
Cpd. 42
Cpd. 56
Cpd. 50
Cpd. 55
Cpd. 53
Cpd. 57
Cpd. 5
Cpd. 15
Cpd. 18
Cpd. 19
Cpd. 24

Figure 1

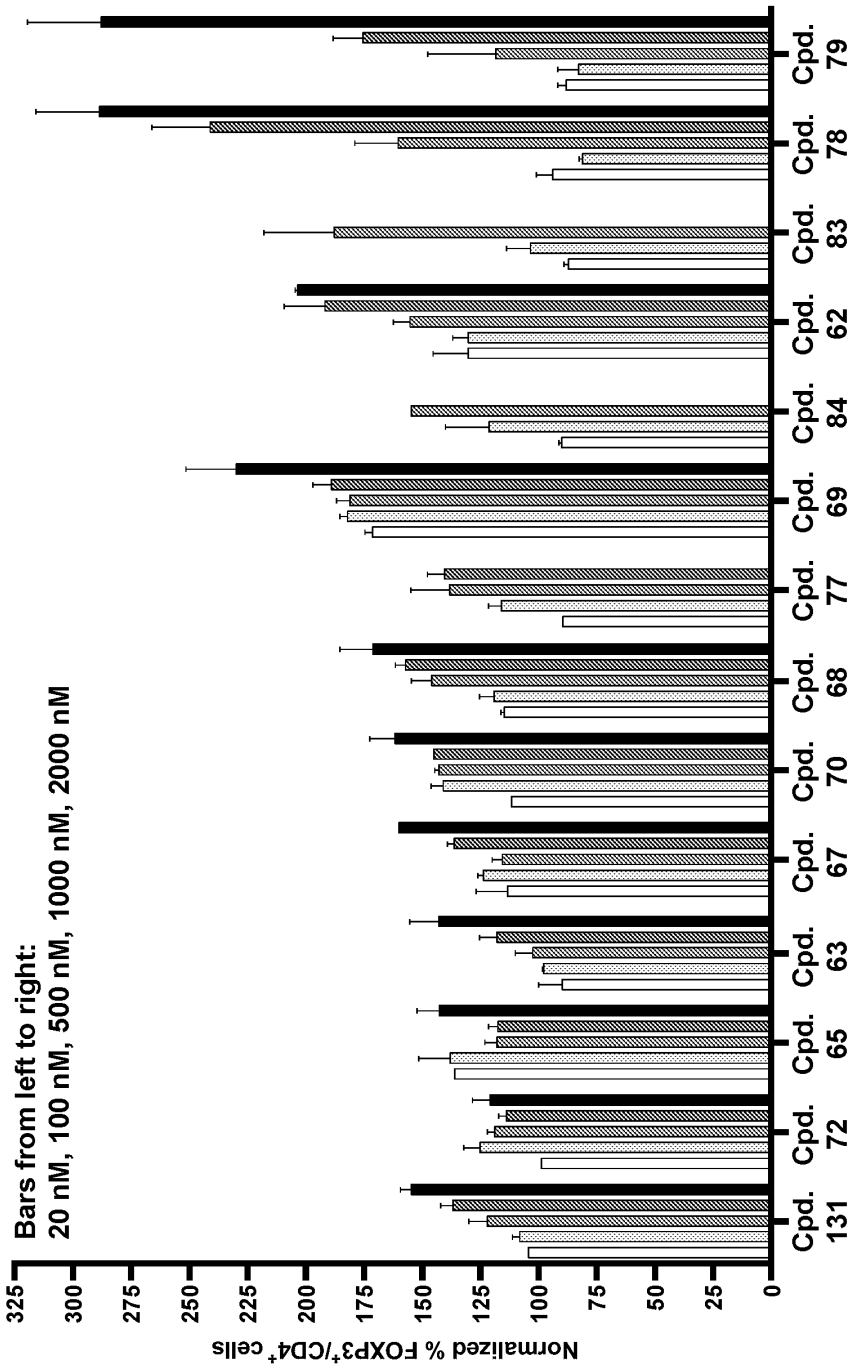
Normalized % FOXP3+/CD4+ cells
325
300
275
250
225
200
175
150
125
100
75
50
25
0
Bars from left to right:
20 nM, 100 nM, 500 nM, 1000 nM, 2000 nM
Cpd. 131
Cpd. 72
Cpd. 65
Cpd. 63
Cpd. 67
Cpd. 70
Cpd. 68
Cpd. 77
Cpd. 69
Cpd. 84
Cpd. 62
Cpd. 83
Cpd. 78
Cpd. 79

Figure 2

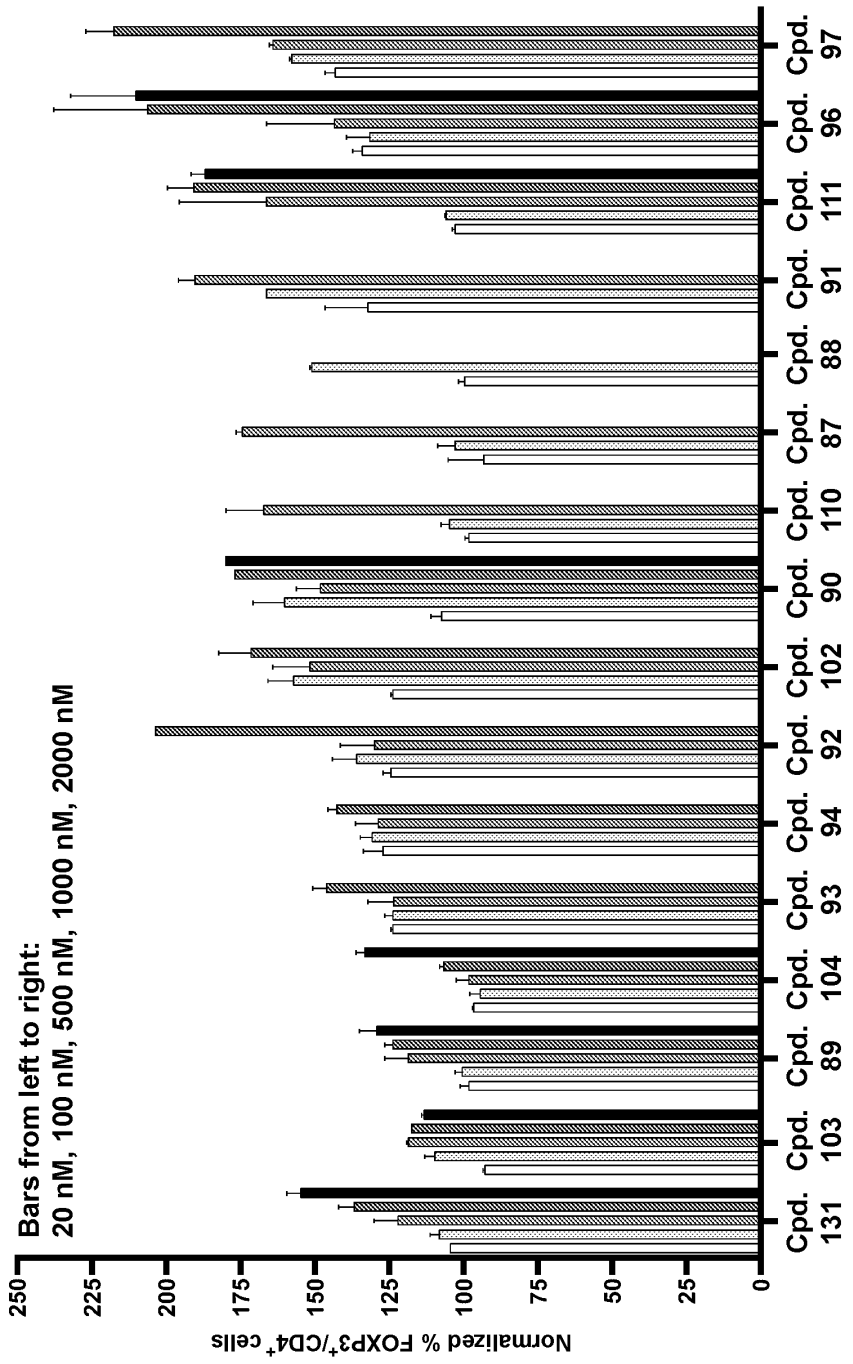
Bars from left to right:
20 nM, 100 nM, 500 nM, 1000 nM, 2000 nM
Normalized % FOXP3+/CD4+ cells
250
225
200
175
150
125
100
75
50
25
0
Cpd. 131
Cpd. 103
Cpd. 89
Cpd. 104
Cpd. 93
Cpd. 94
Cpd. 92
Cpd. 102
Cpd. 90
Cpd. 110
Cpd. 87
Cpd. 88
Cpd. 91
Cpd. 111
Cpd. 96
Cpd. 97

Figure 3

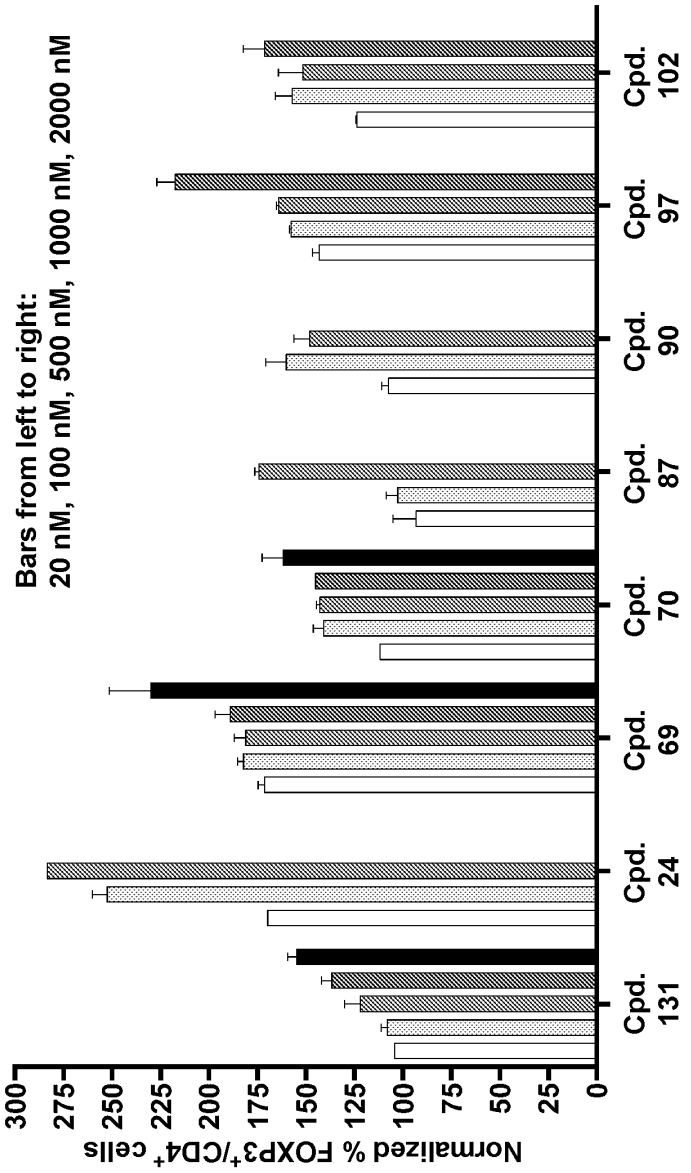
Bars from left to right:
20 nM, 100 nM, 500 nM, 1000 nM, 2000 nM
Normalized % FOXP3+/CD4+ cells
300
275
250
225
200
175
150
125
100
75
50
25
0
Cpd. 131
Cpd. 24
Cpd. 69
Cpd. 70
Cpd. 87
Cpd. 90
Cpd. 97
Cpd. 102

Figure 4

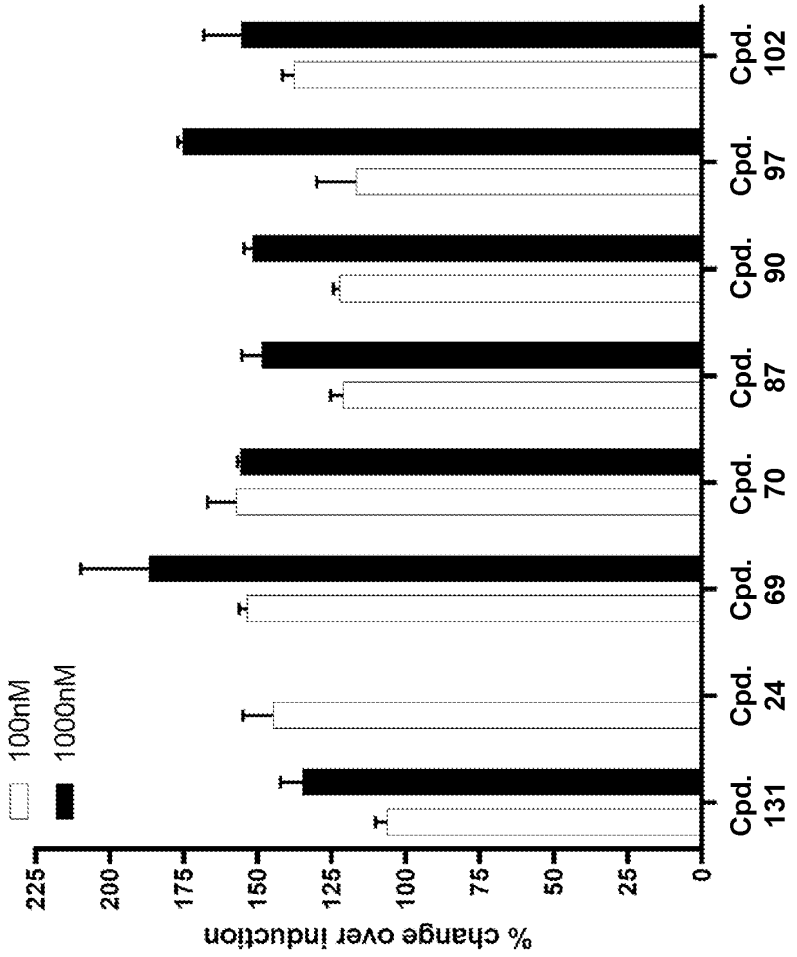
100nM
1000nM
% change over induction
225
200
175
150
125
100
75
50
25
0
Cpd. 131
Cpd. 24
Cpd. 69
Cpd. 70
Cpd. 87
Cpd. 90
Cpd. 97
Cpd. 102

Figure 5

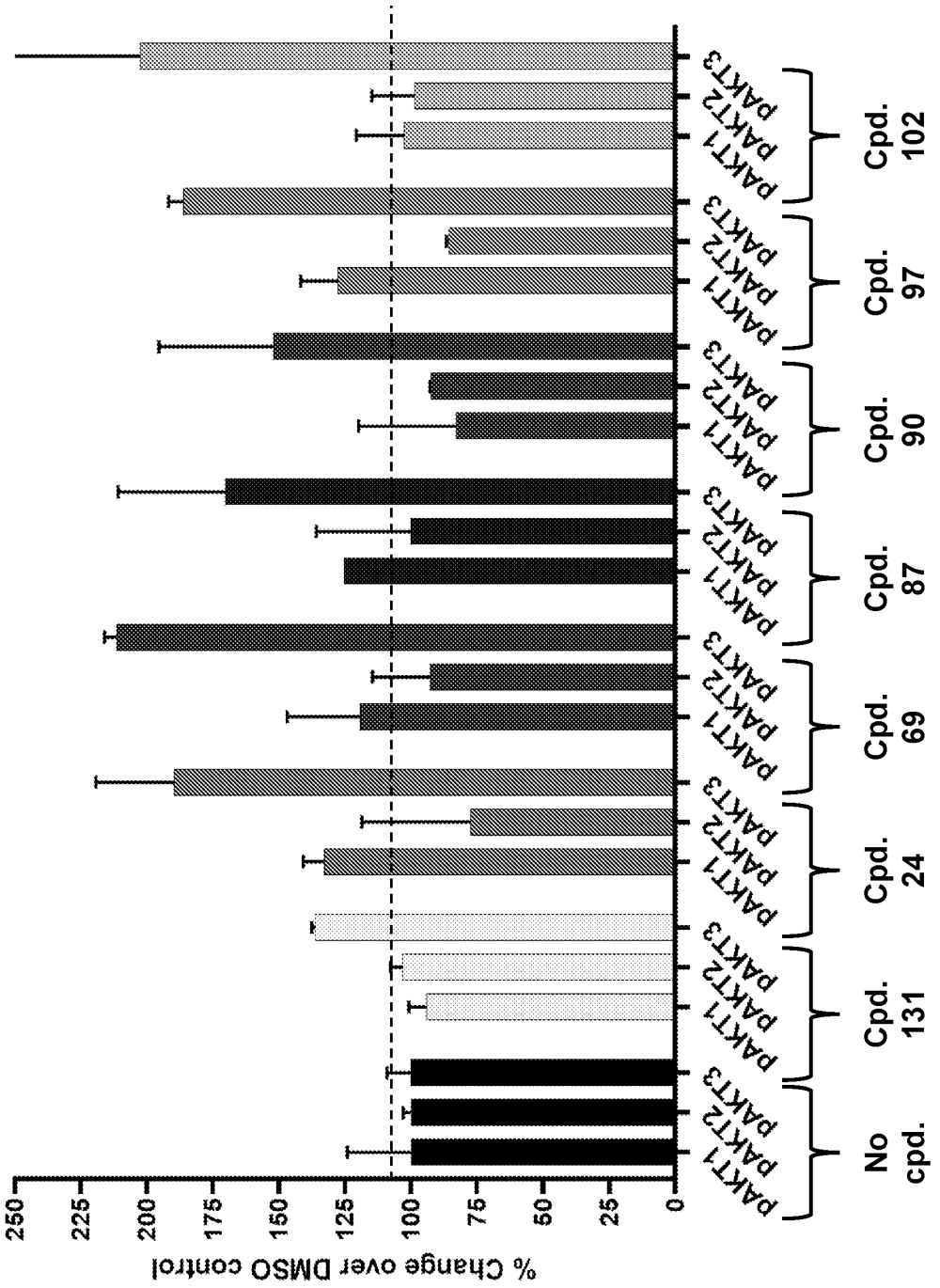
% Change over DMSO control
250
225
200
175
150
125
100
75
50
25
0
No cpd.
Cpd. 131
Cpd. 24
Cpd. 69
Cpd. 87
Cpd. 90
Cpd. 97
Cpd. 102
PAKT1
PAKT2
PAKT3

Figure 6

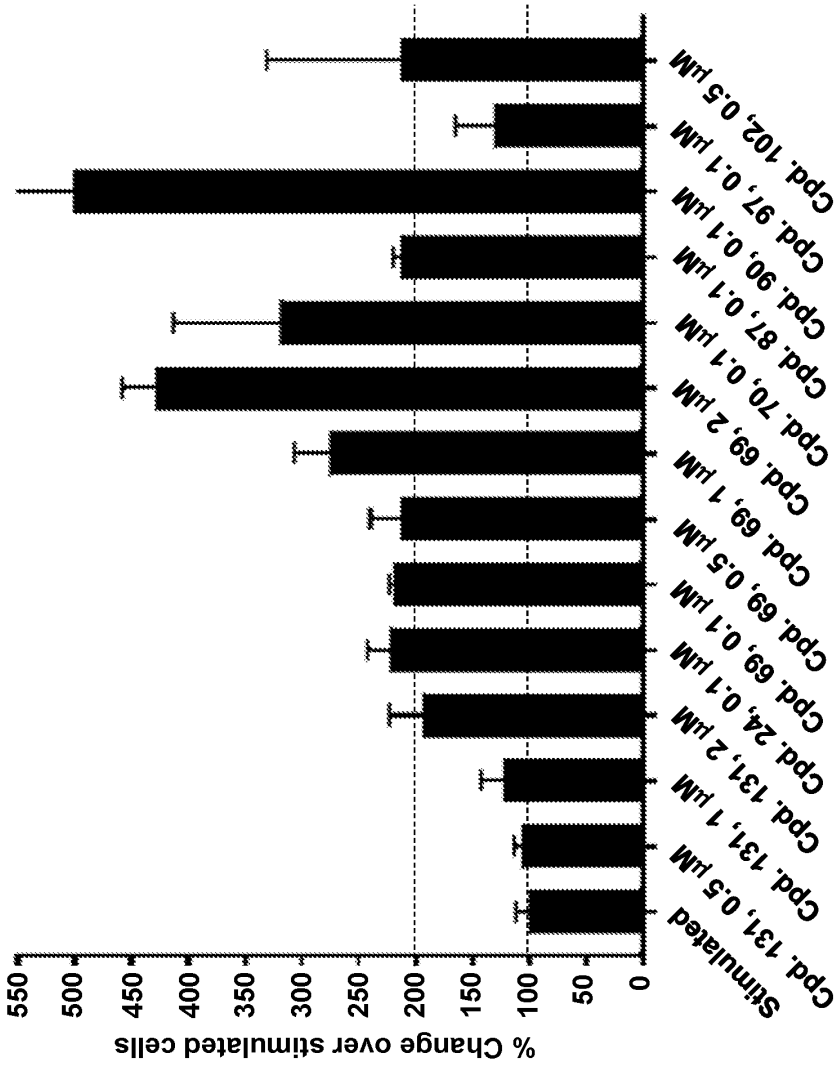
% Change over stimulated cells
550
500
450
400
350
300
250
200
150
100
50
0
Stimulated
Cpd. 131, 0.5 μM
Cpd. 131, 1 μM
Cpd. 131, 2 μM
Cpd. 24, 0.1 μM
Cpd. 69, 0.1 μM
Cpd. 69, 0.5 μM
Cpd. 69, 1 μM
Cpd. 69, 2 μM
Cpd. 70, 0.1 μM
Cpd. 87, 0.1 μM
Cpd. 90, 0.1 μM
Cpd. 97, 0.1 μM
Cpd. 102, 0.1 μM
Cpd. 102, 0.5 μM

Figure 7

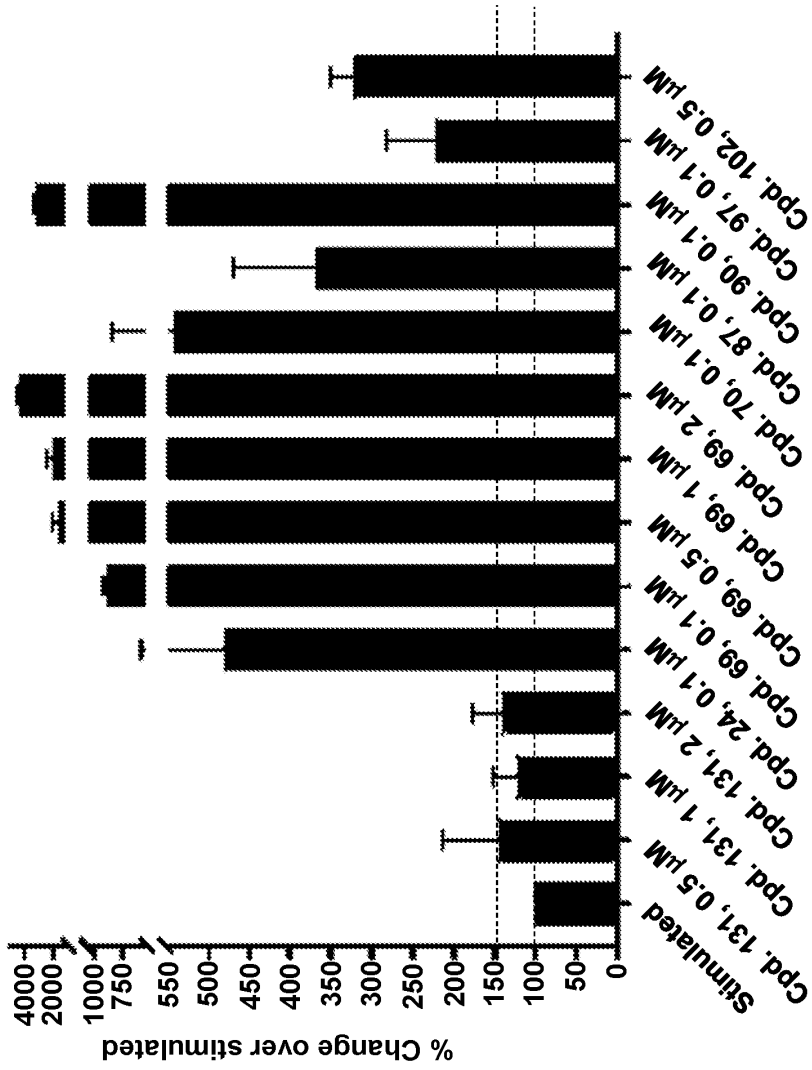
% Change over stimulated
4000
2000
1000
750
550
500
450
400
350
300
250
200
150
100
50
0
Stimulated
Cpd. 131, 0.5 μM
Cpd. 131, 1 μM
Cpd. 131, 2 μM
Cpd. 24, 0.1 μM
Cpd. 69, 0.1 μM
Cpd. 69, 0.5 μM
Cpd. 69, 1 μM
Cpd. 69, 2 μM
Cpd. 70, 0.1 μM
Cpd. 87, 0.1 μM
Cpd. 90, 0.1 μM
Cpd. 97, 0.1 μM
Cpd. 102, 0.5 μM

Figure 8

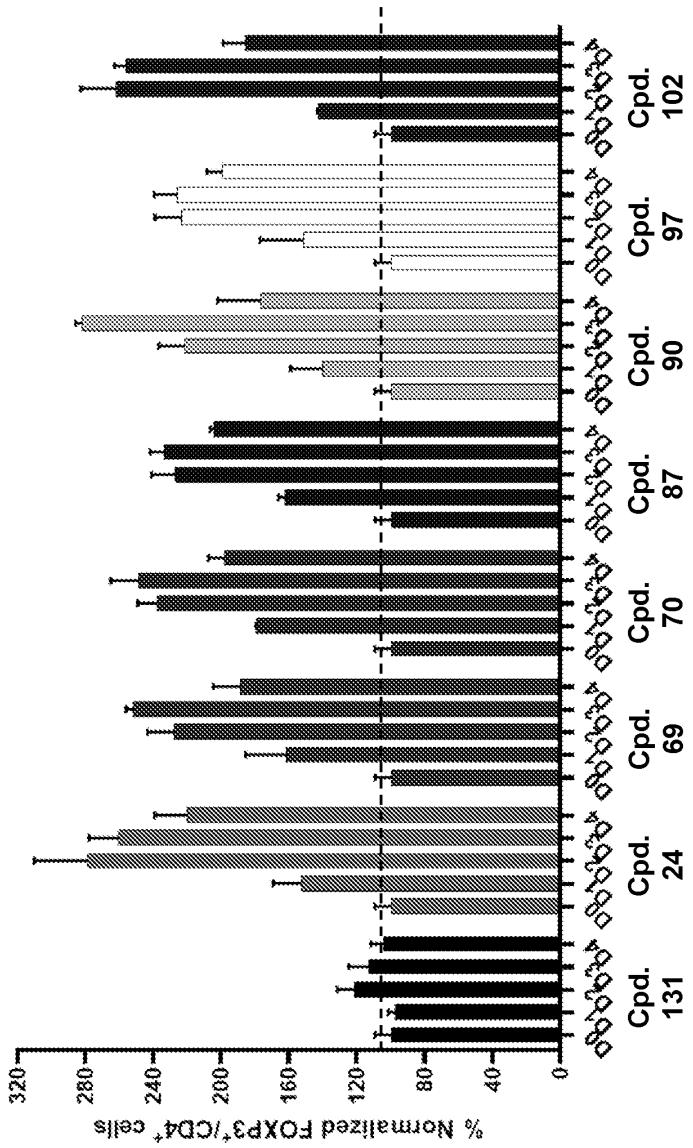
% Normalized FOXP3+/CD4+ cells
320
280
240
200
160
120
80
40
0
Cpd. 131
Cpd. 24
Cpd. 69
Cpd. 70
Cpd. 87
Cpd. 90
Cpd. 97
Cpd. 102

Figure 9

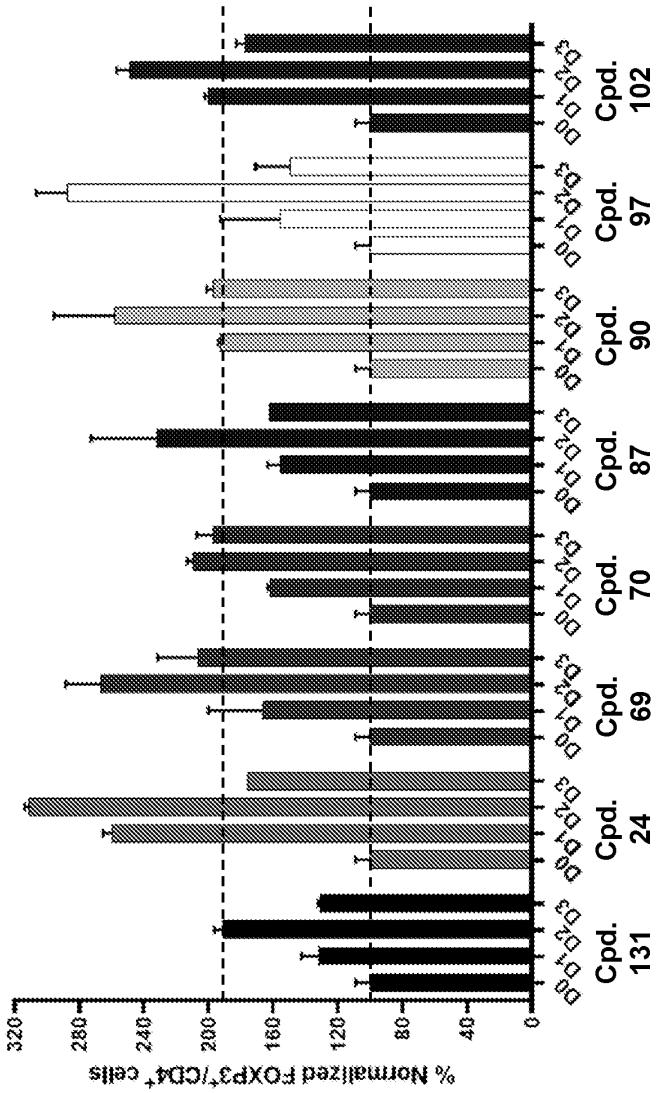
% Normalized FOXP3+/CD4+ cells
0
40
80
120
160
200
240
280
320
0.03
0.10
0.30
1.00
3.00
Cpd. 131
Cpd. 24
Cpd. 69
Cpd. 70
Cpd. 87
Cpd. 90
Cpd. 97
Cpd. 102

Figure 10

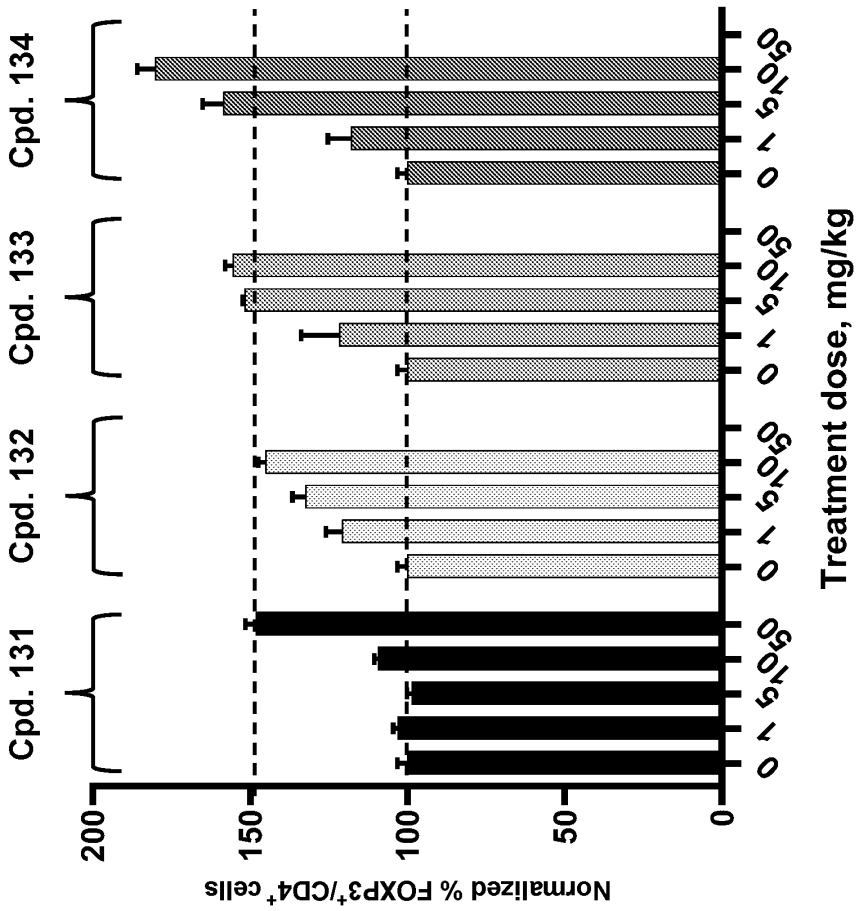
Cpd. 131
Cpd. 132
Cpd. 133
Cpd. 134
0
0.1
0.5
1
5
10
50
100
Treatment dose, mg/kg
0
50
100
150
200
Normalized % FOXP3+/CD4+ cells

Figure 11

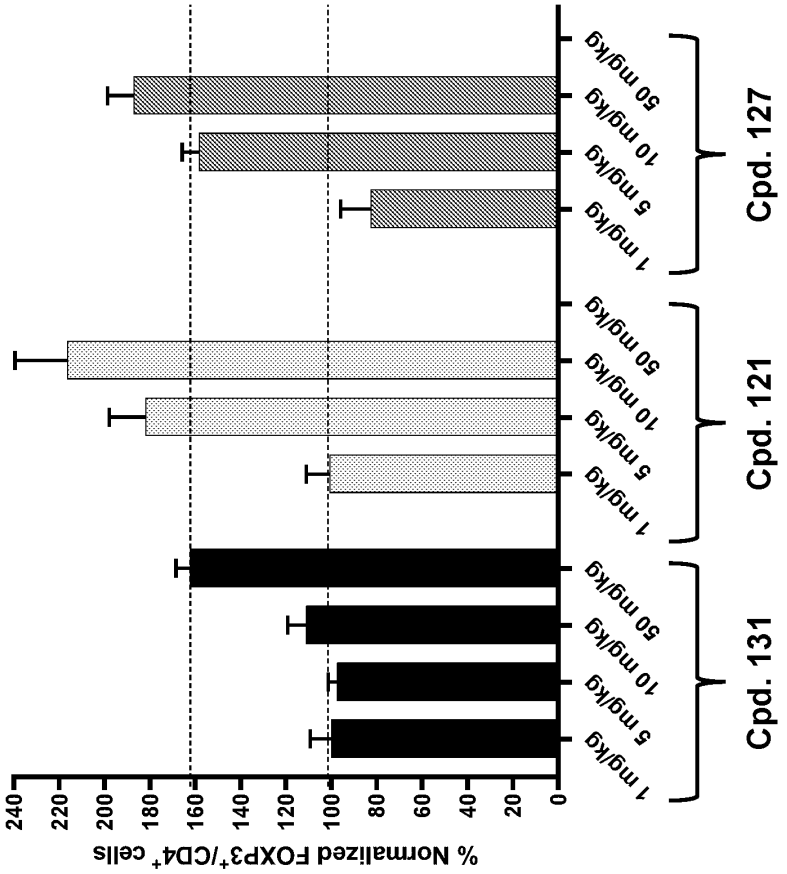
% Normalized FOXP3+/CD4+ cells
240
220
200
180
160
140
120
100
80
60
40
20
0
1 mg/kg
5 mg/kg
10 mg/kg
50 mg/kg
Cpd. 131
1 mg/kg
5 mg/kg
10 mg/kg
50 mg/kg
Cpd. 121
1 mg/kg
5 mg/kg
10 mg/kg
50 mg/kg
Cpd. 127

Figure 12

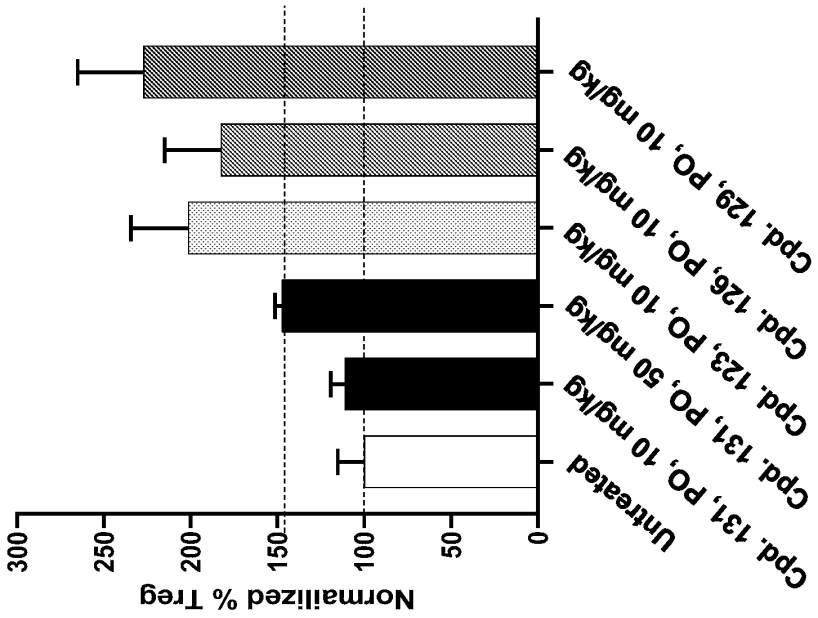
Normalized % Treg
300
250
200
150
100
50
0
Untreated
Cpd. 131, PO, 10 mg/kg
Cpd. 131, PO, 50 mg/kg
Cpd. 123, PO, 10 mg/kg
Cpd. 126, PO, 10 mg/kg
Cpd. 129, PO, 10 mg/kg

Figure 13