Fig. 1

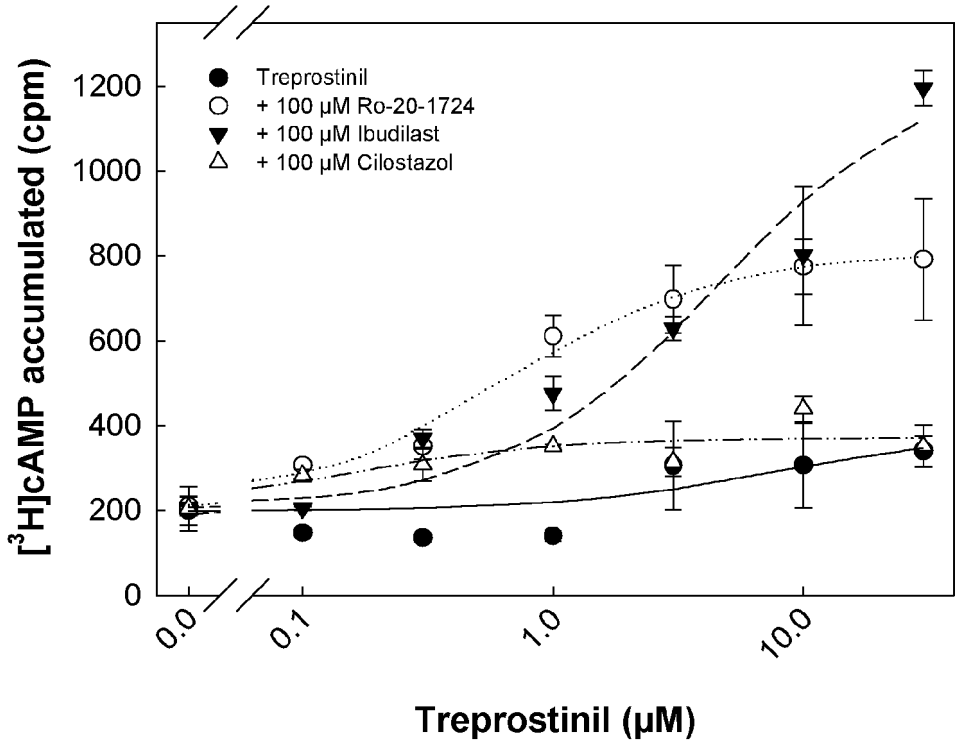

Treprostinil
+ 100 µM Ro-20-1724
+ 100 µM Ibudilast
+ 100 µM Cilostazol
[3H]cAMP accumulated (cpm)
0
200
400
600
800
1000
1200
0.0
0.1
1.0
10.0
Treprostinil (µM)

Fig. 2

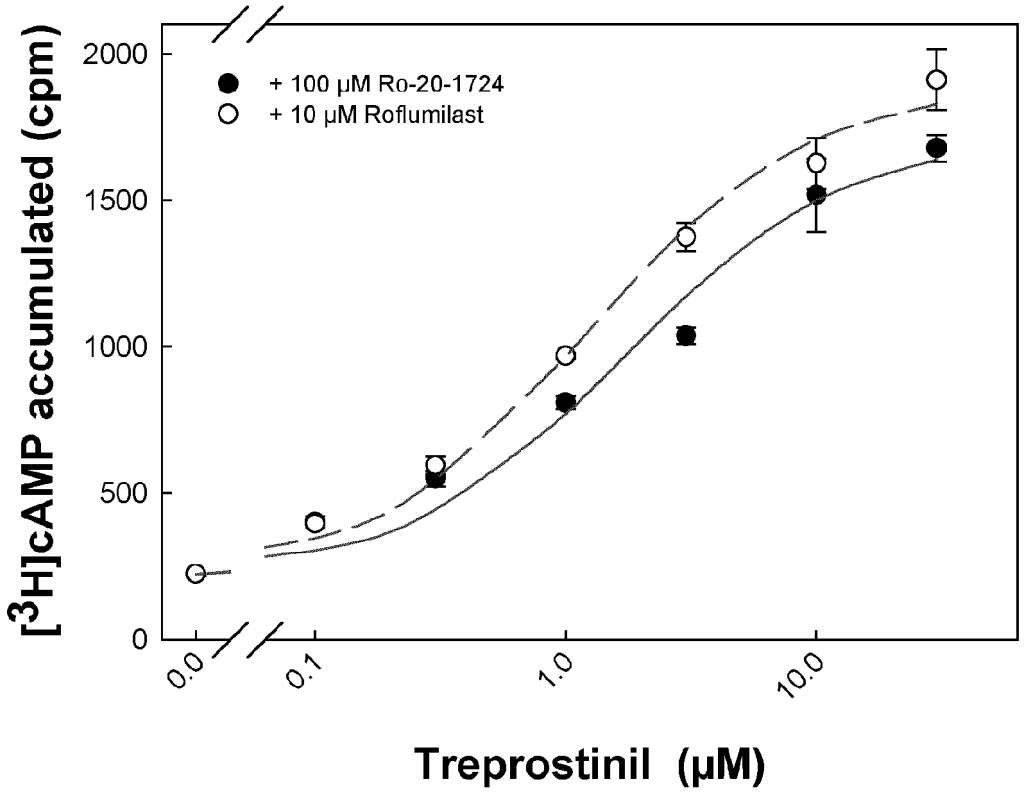

[3H]cAMP accumulated (cpm)
2000
1500
1000
500
0
+ 100 µM Ro-20-1724
+ 10 µM Roflumilast
0.0
0.1
1.0
10.0
Treprostinil (µM)

Fig. 3

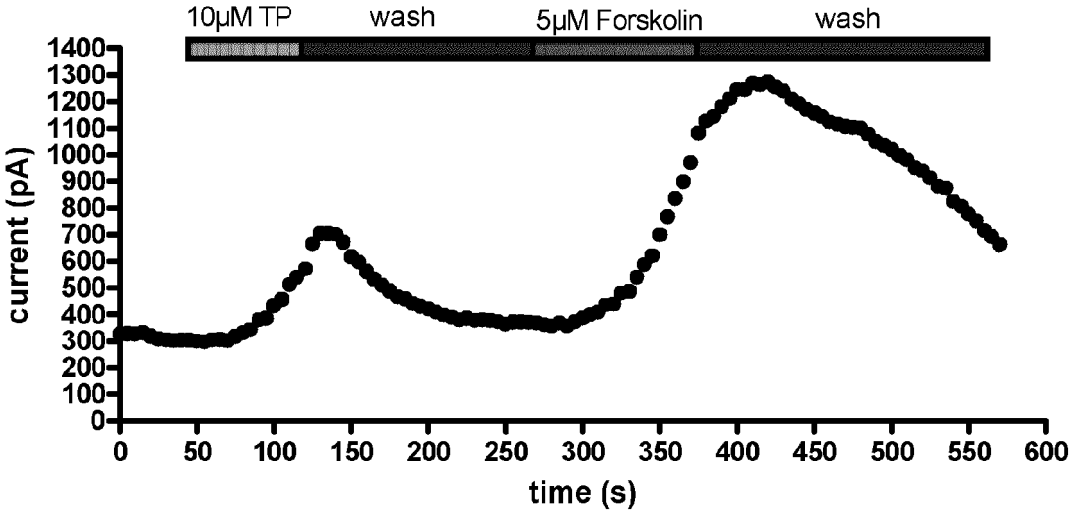

10µM TP
wash
5µM Forskolin
wash
current (pA)
1400
1300
1200
1100
1000
900
800
700
600
500
400
300
200
100
0
0
50
100
150
200
250
300
350
400
450
500
550
600
time (s)

Fig. 4

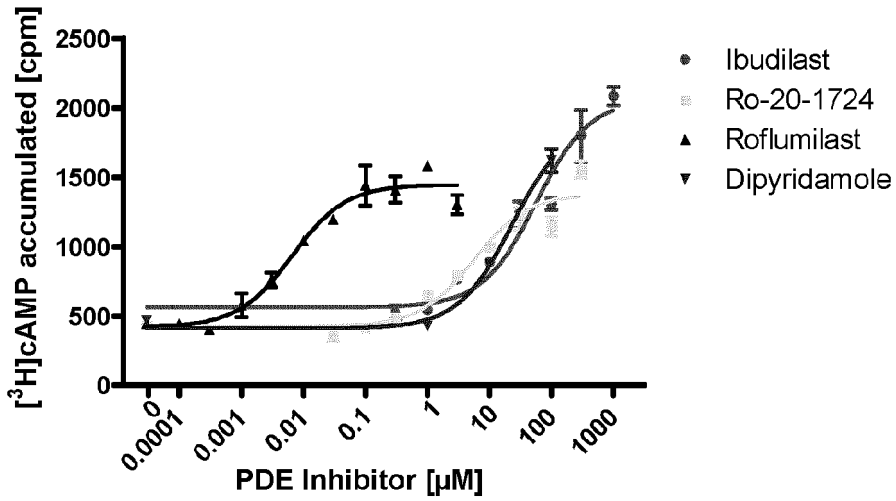

[3H]cAMP accumulated [cpm]
2500
2000
1500
1000
500
0
0
0.0001
0.001
0.01
0.1
1
10
100
1000
PDE Inhibitor [µM]
Ibudilast
Ro-20-1724
Roflumilast
Dipyridamole

Fig. 5

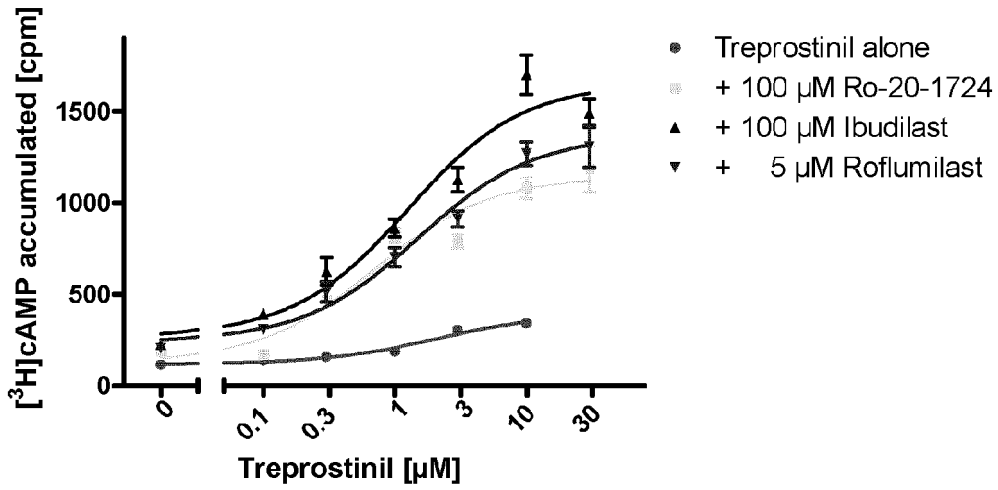
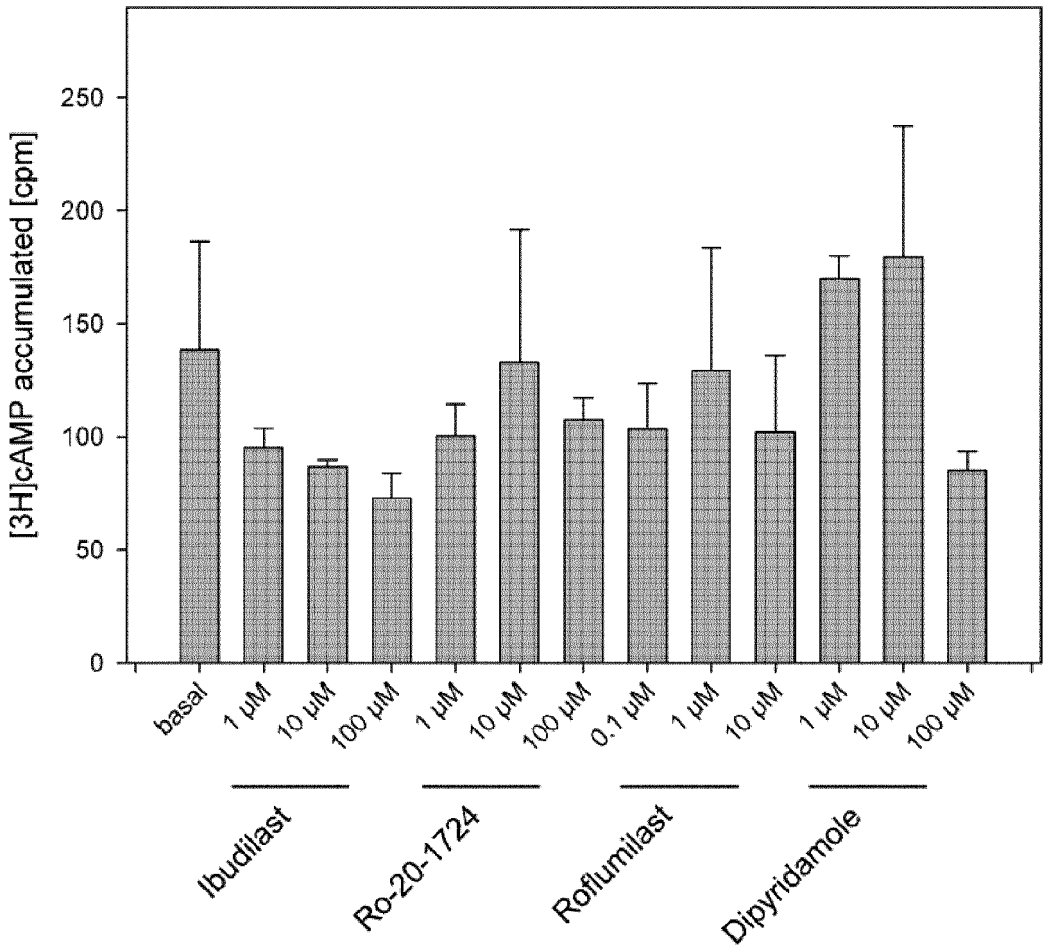

[3H]cAMP accumulated [cpm]
1500
1000
500
0
0
0.1
0.3
1
3
10
30
Treprostinil [µM]
Treprostinil alone
+ 100 µM Ro-20-1724
+ 100 µM Ibudilast
+ 5 µM Roflumilast

Fig. 6


[3H]cAMP accumulated [cpm]
250
200
150
100
50
0
basal
1 µM
10 µM
100 µM
1 µM
10 µM
100 µM
0.1 µM
1 µM
10 µM
1 µM
10 µM
100 µM
Ibudilast
Ro-20-1724
Roflumilast
Dipyridamole