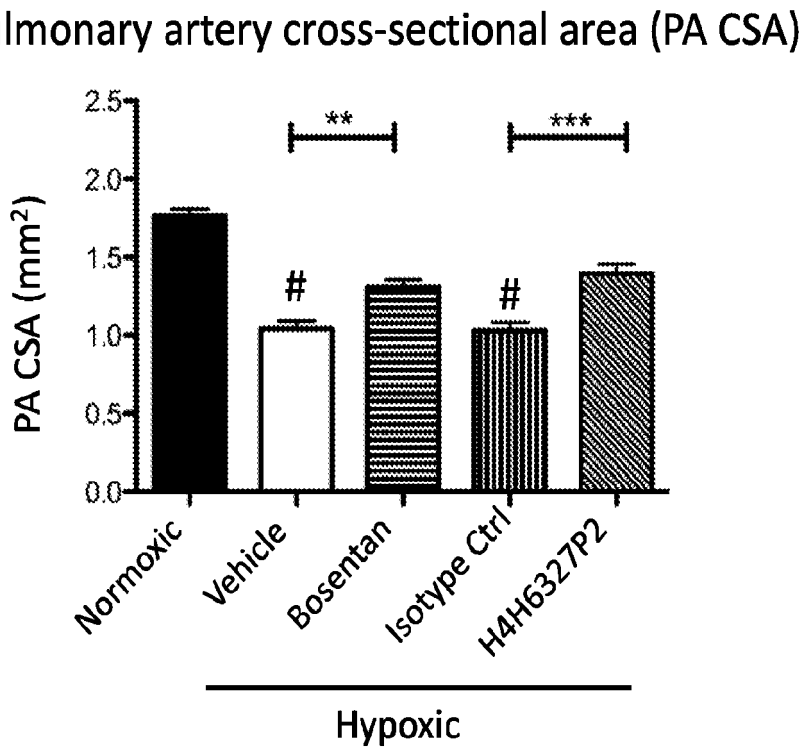

Pulmonary artery cross-sectional area (PA CSA)
PA CSA (mm²)
2.5
2.0
1.5
1.0
0.5
0.0
**
***
#
#
Normoxic
Vehicle
Bosentan
Isotype Ctrl
H4H6327P2
Hypoxic

Figure 1

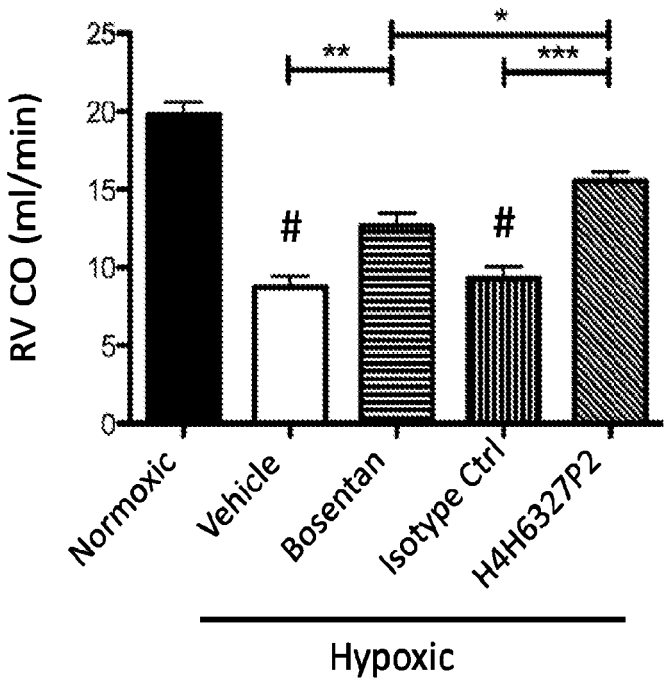

Right ventricular cardiac output (RV CO)
RV CO (ml/min)
25
20
15
10
5
0
**
*
***
#
#
Normoxic
Vehicle
Bosentan
Isotype Ctrl
H4H6327P2
Hypoxic

Figure 2

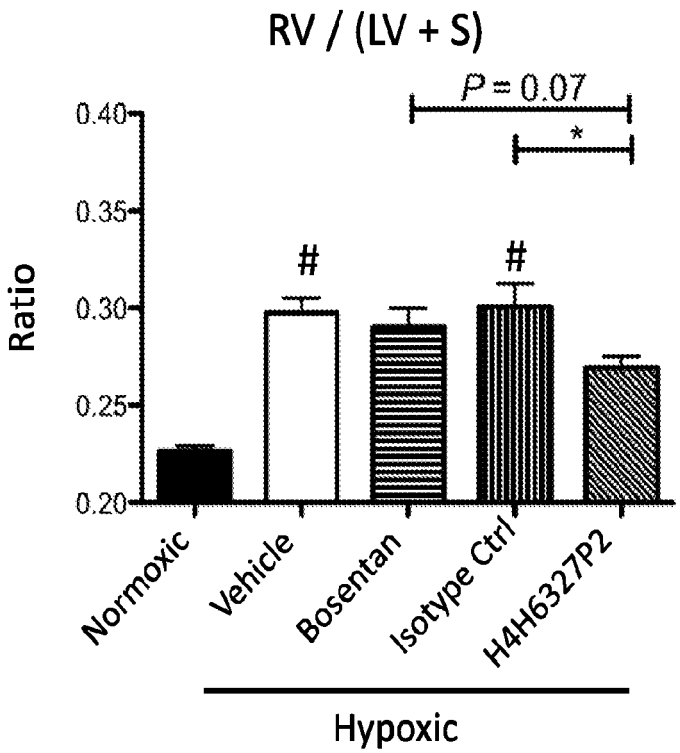

RV / (LV + S)
P = 0.07
*
0.40
0.35
0.30
0.25
0.20
Ratio
#
#
Normoxic
Vehicle
Bosentan
Isotype Ctrl
H4H6327P2
Hypoxic

Figure 3

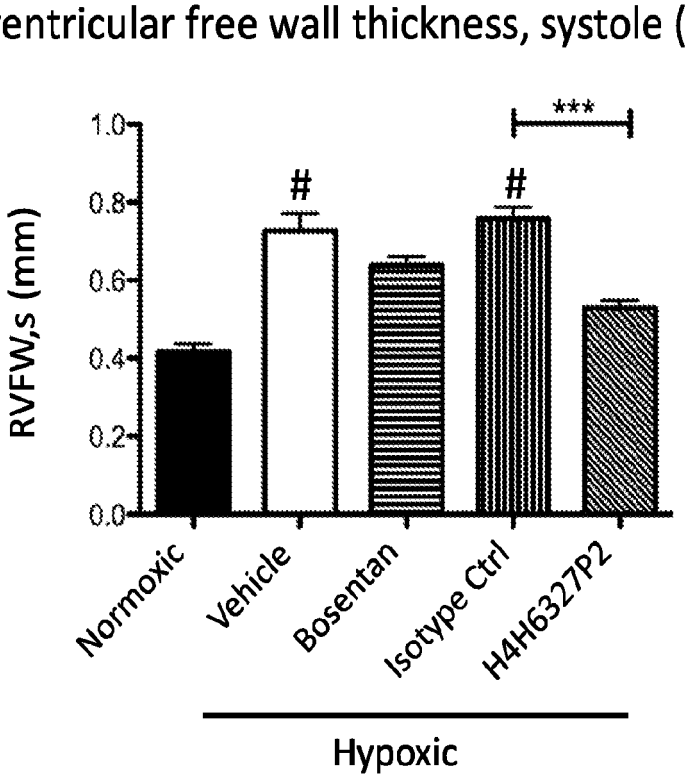

Right ventricular free wall thickness, systole (RVFW,s)
RVFW,s (mm)
1.0
0.8
0.6
0.4
0.2
0.0
#
#
***
Normoxic
Vehicle
Bosentan
Isotype Ctrl
H4H6327P2
Hypoxic

Figure 4

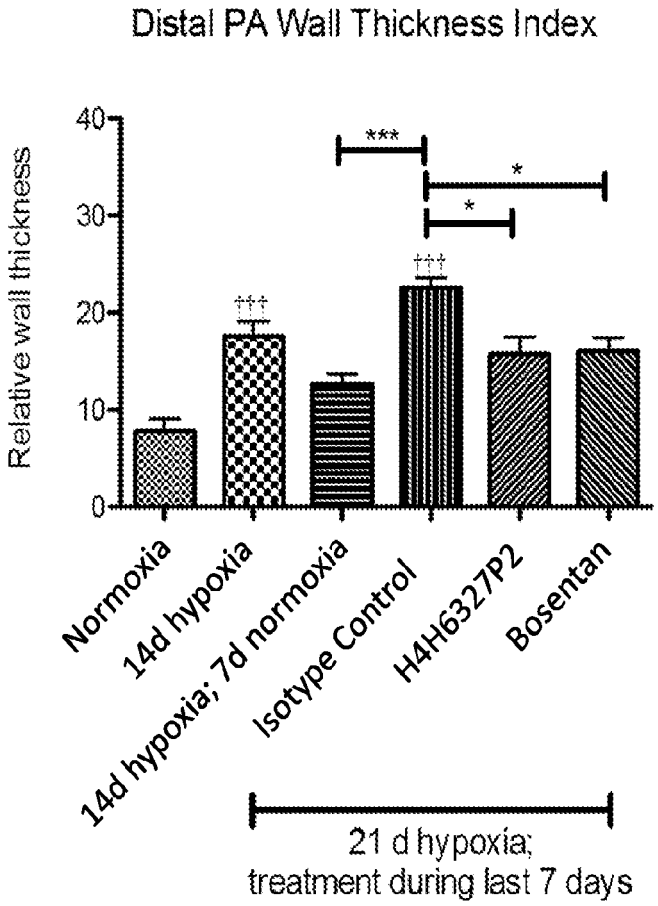

Distal PA Wall Thickness Index
Relative wall thickness
40
30
20
10
0
***
*
*
†††
†††
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 5

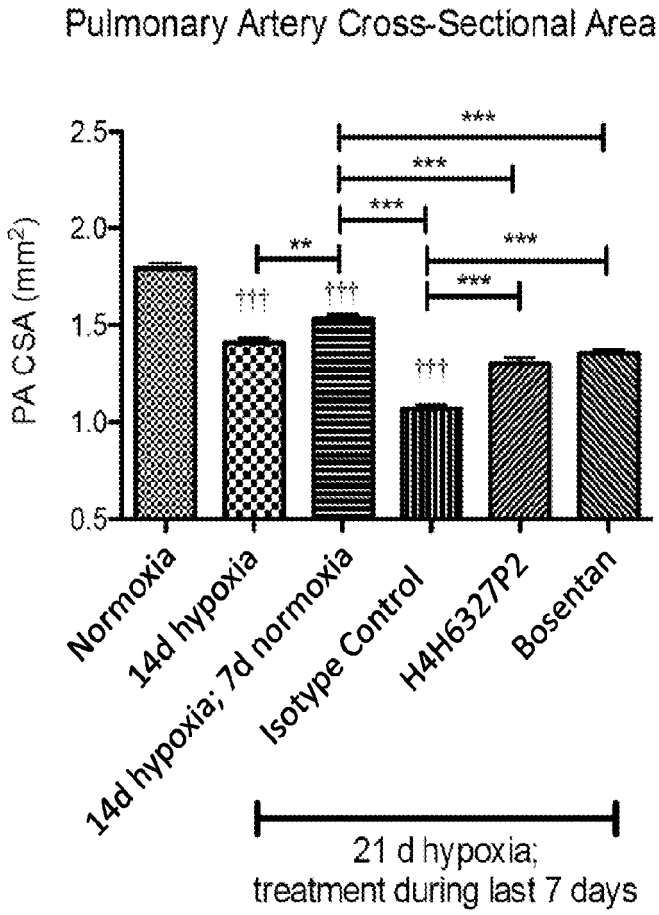

Pulmonary Artery Cross-Sectional Area
PA CSA (mm²)
2.5
2.0
1.5
1.0
0.5
***
***
***
**
***
***
†††
†††
†††
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 6

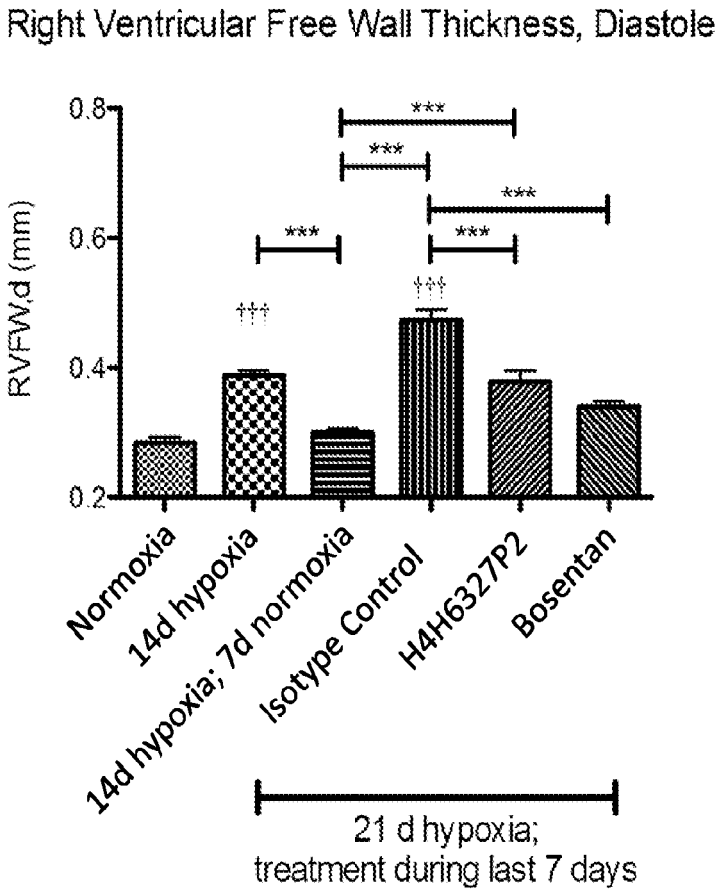

Right Ventricular Free Wall Thickness, Diastole
RVFW,d (mm)
0.8
0.6
0.4
0.2
***
***
***
***
***
†††
†††
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 7

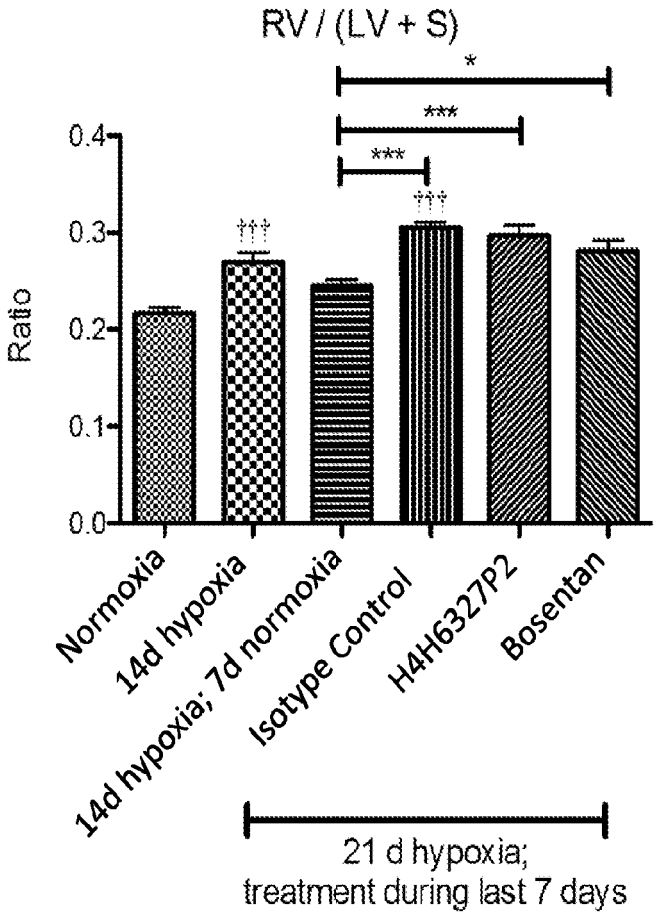

RV / (LV + S)
*
***
***
†††
†††
Ratio
0.4
0.3
0.2
0.1
0.0
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 8

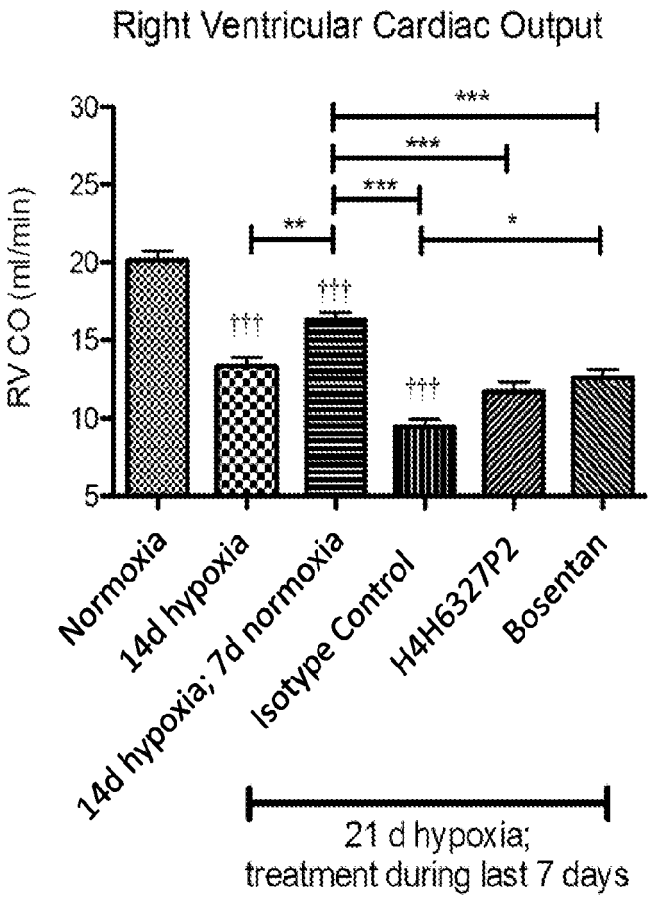

Right Ventricular Cardiac Output
RV CO (ml/min)
30
25
20
15
10
5
***
***
***
**
*
†††
†††
†††
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 9

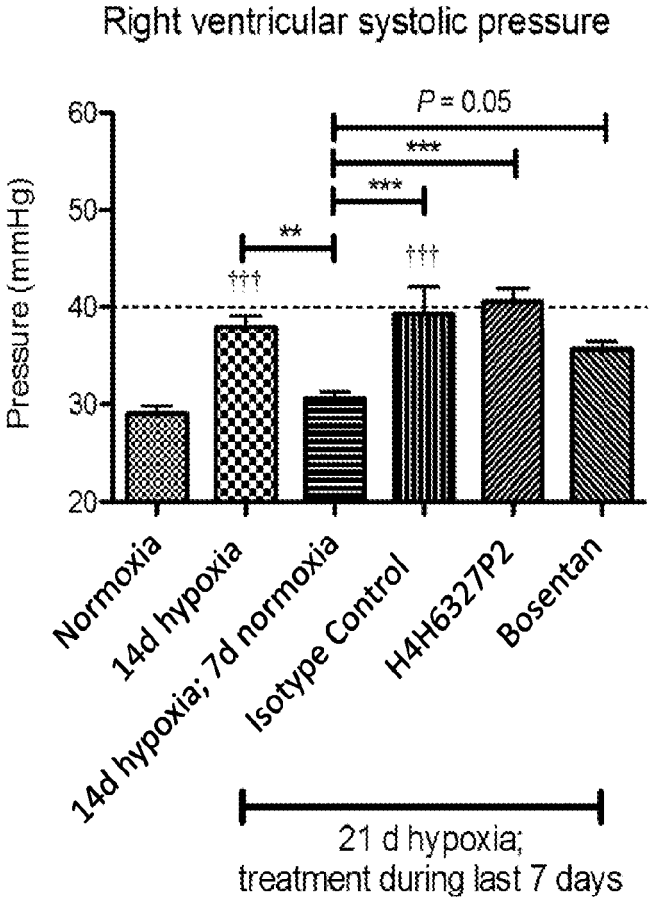

Right ventricular systolic pressure
Pressure (mmHg)
60
50
40
30
20
P = 0.05
***
***
**
†††
†††
Normoxia
14d hypoxia
14d hypoxia; 7d normoxia
Isotype Control
H4H6327P2
Bosentan
21 d hypoxia;
treatment during last 7 days

Figure 10

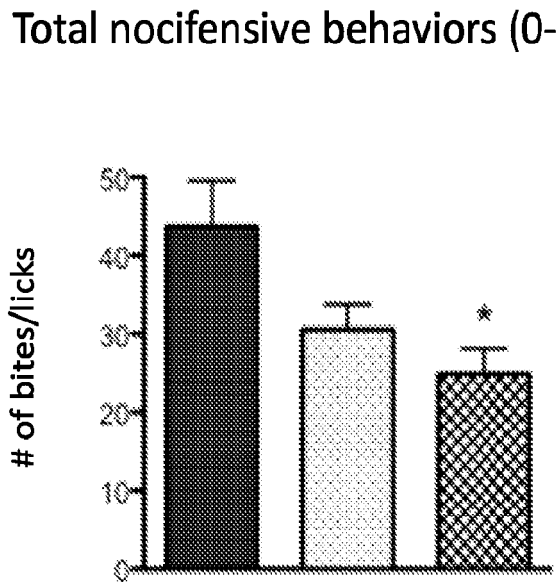

Total nocifensive behaviors (0-60 min)
# of bites/licks
50
40
30
20
10
0
*

Figure 11