

The General Synthetic Scheme for the Preparation of Intermediates

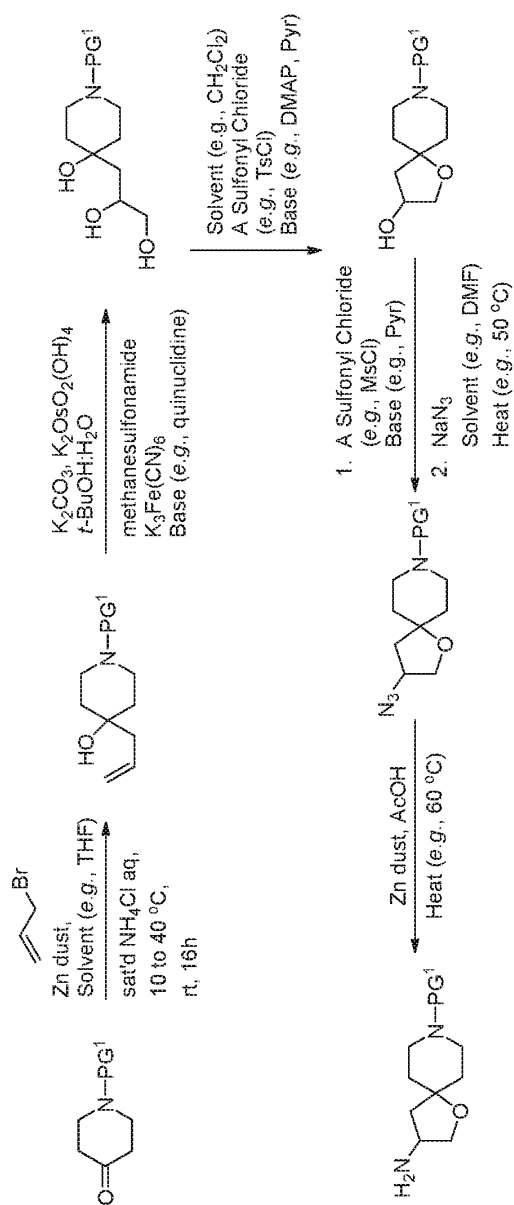


FIGURE 1

The General Synthetic Scheme for the Preparation of Intermediates

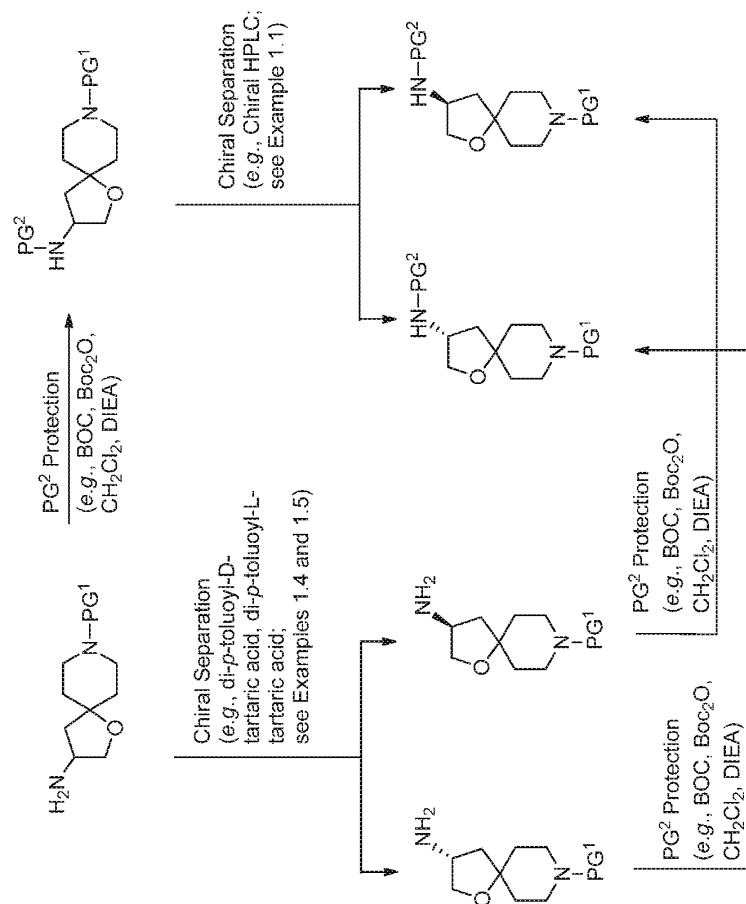


FIGURE 2

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

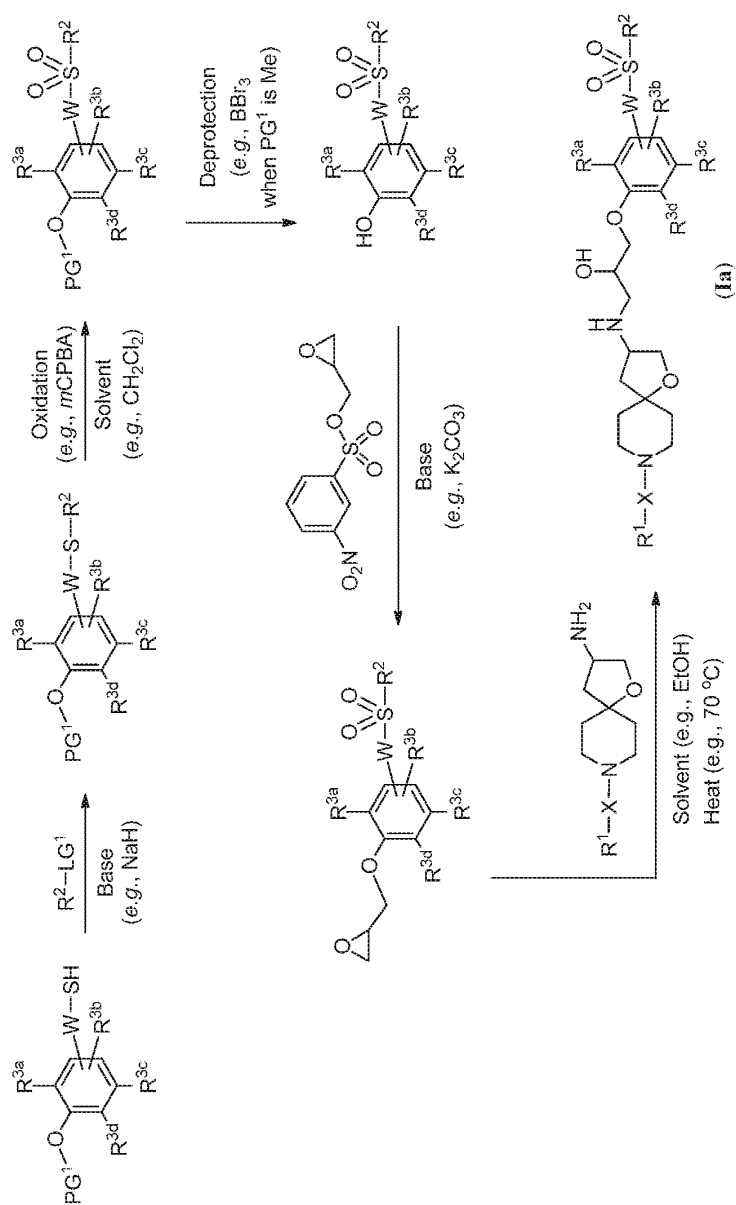


FIGURE 3

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

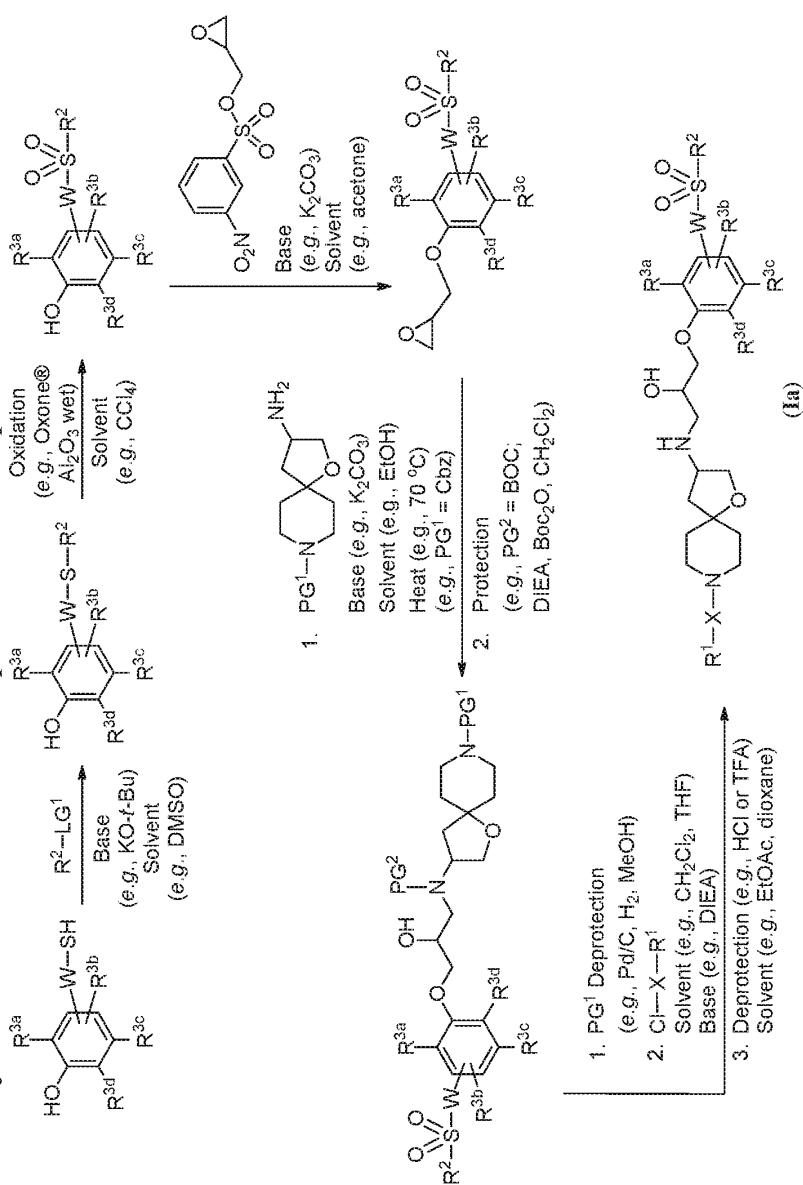


FIGURE 4

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

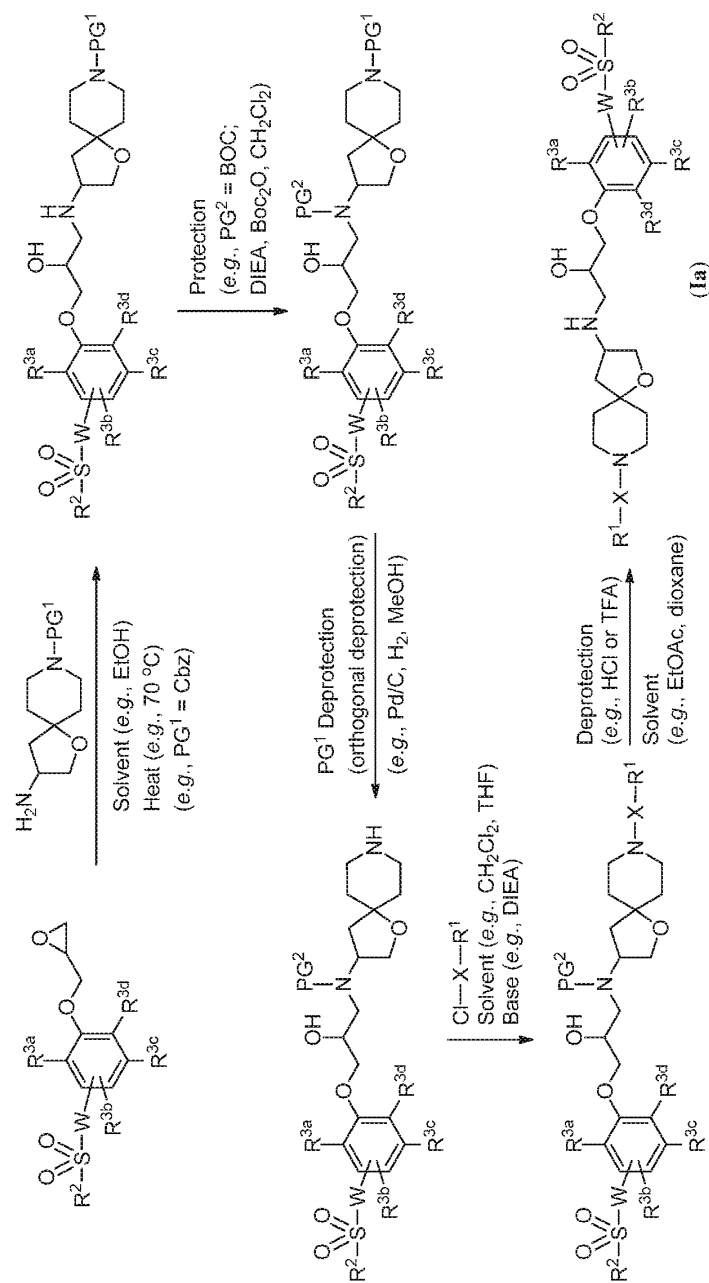


FIGURE 5

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

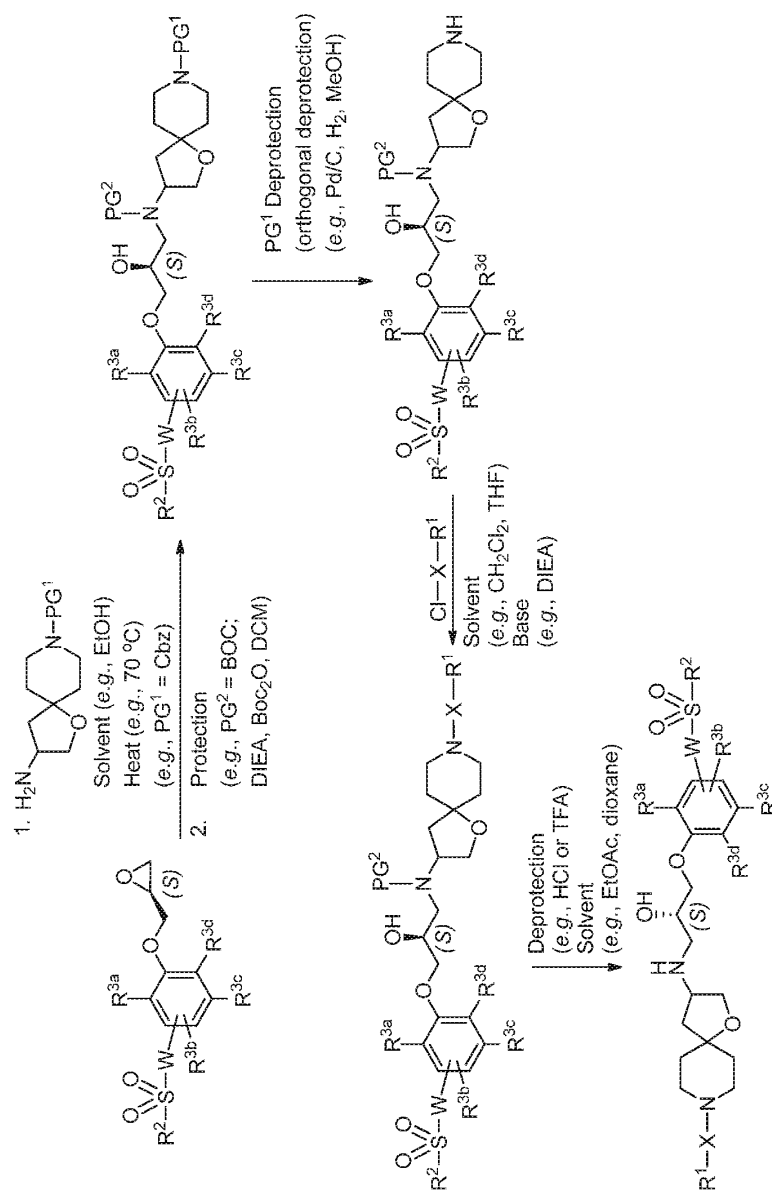


FIGURE 6

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

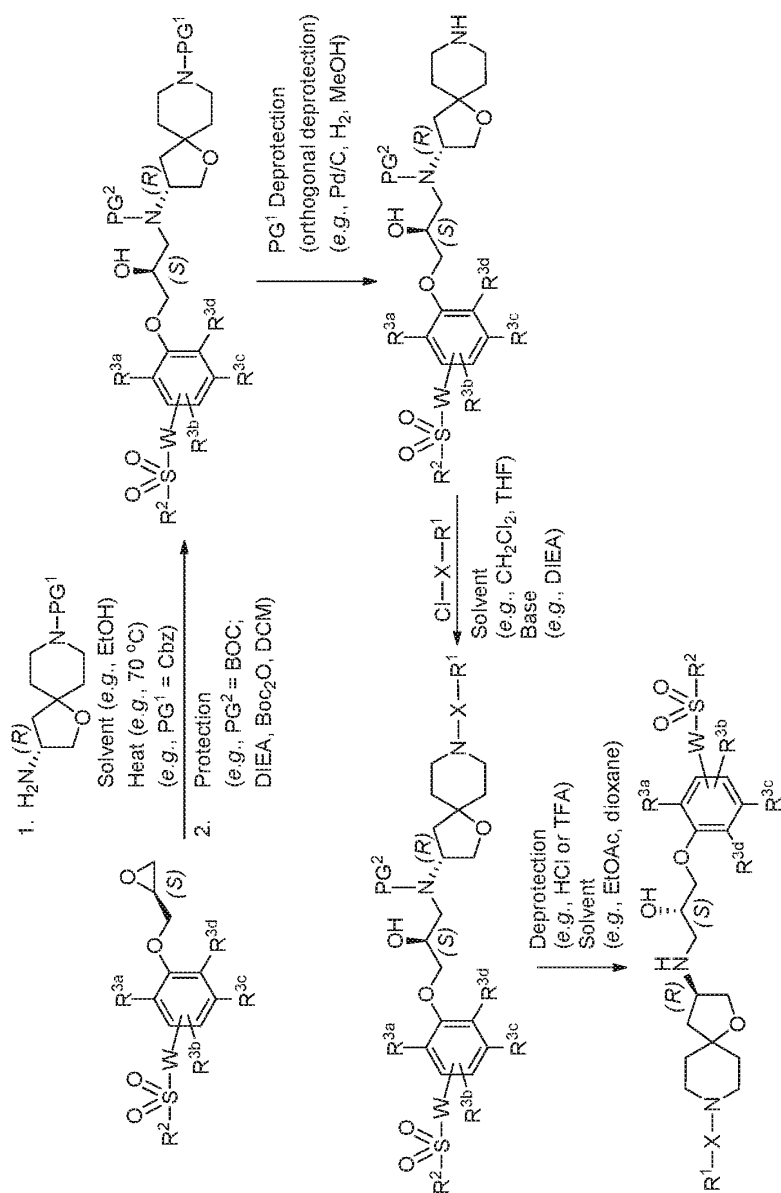


FIGURE 7

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

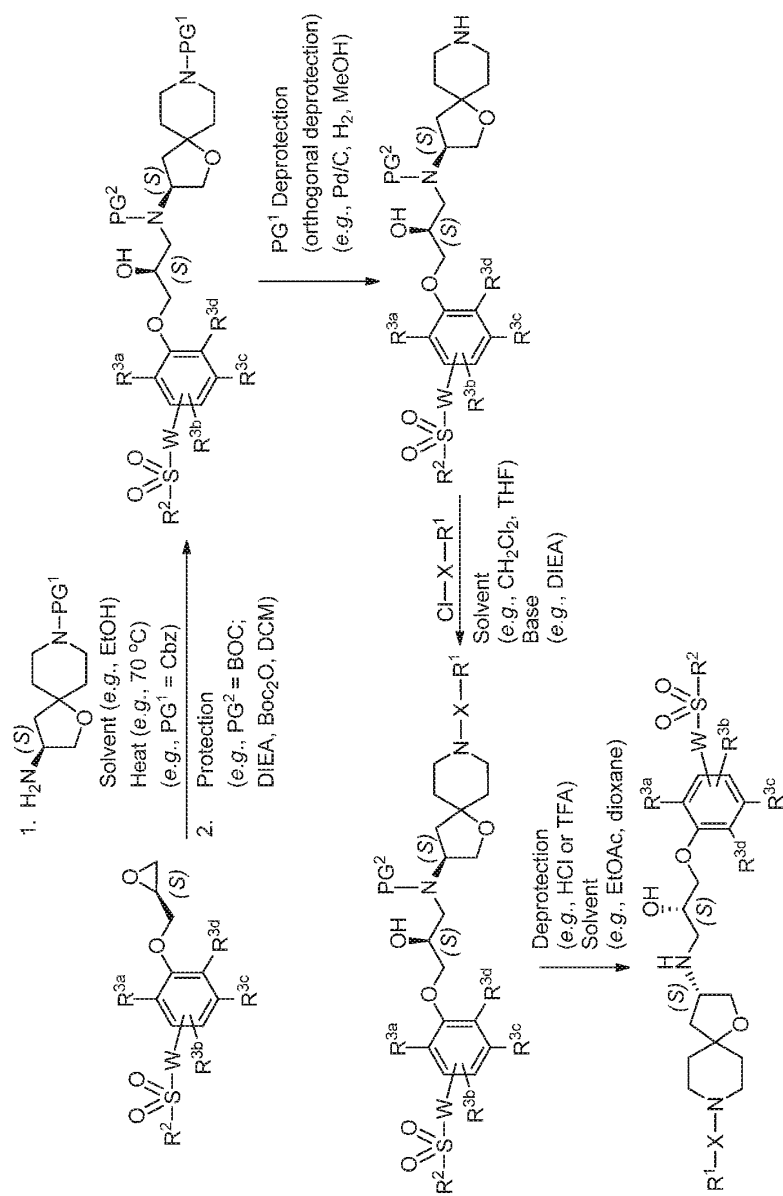


FIGURE 8

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

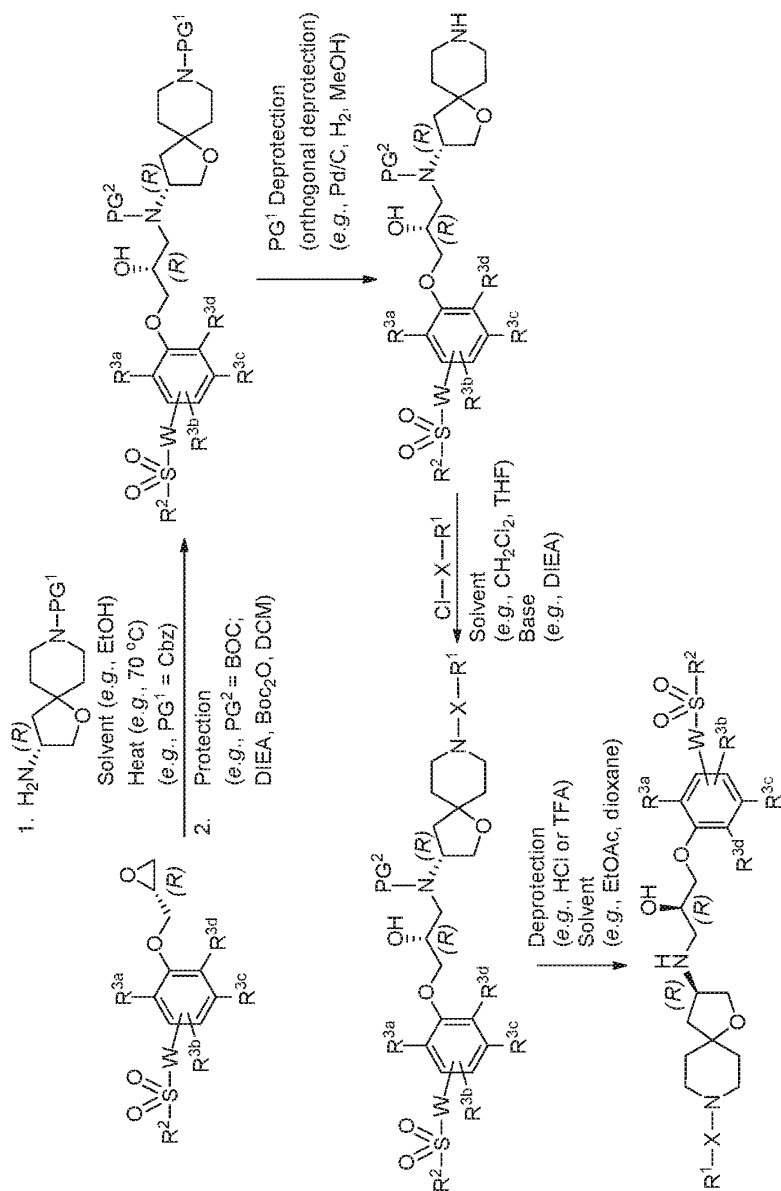


FIGURE 9

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

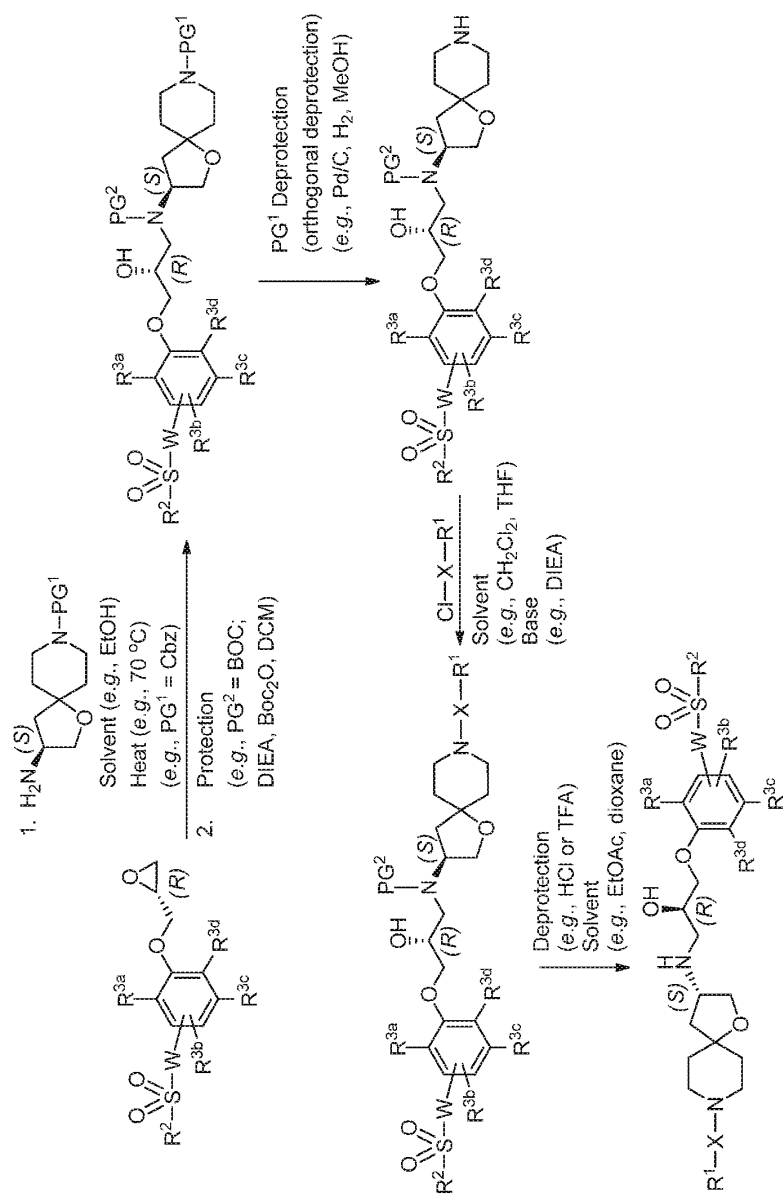


FIGURE 10

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

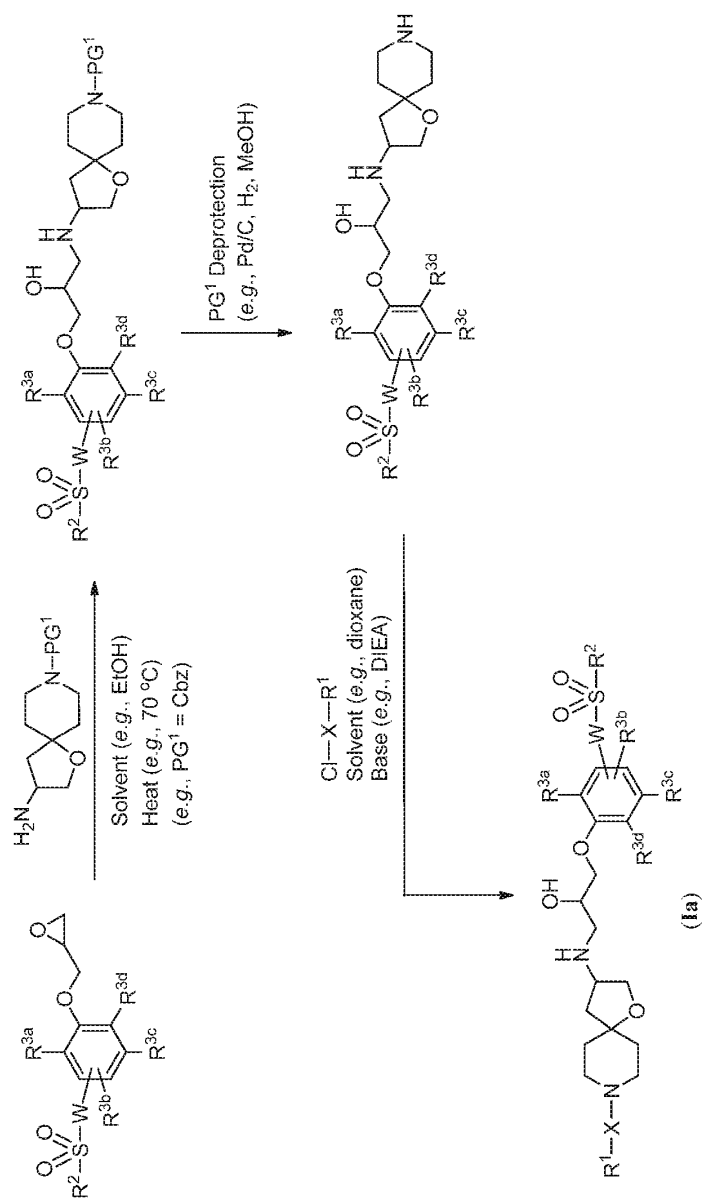


FIGURE 11

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

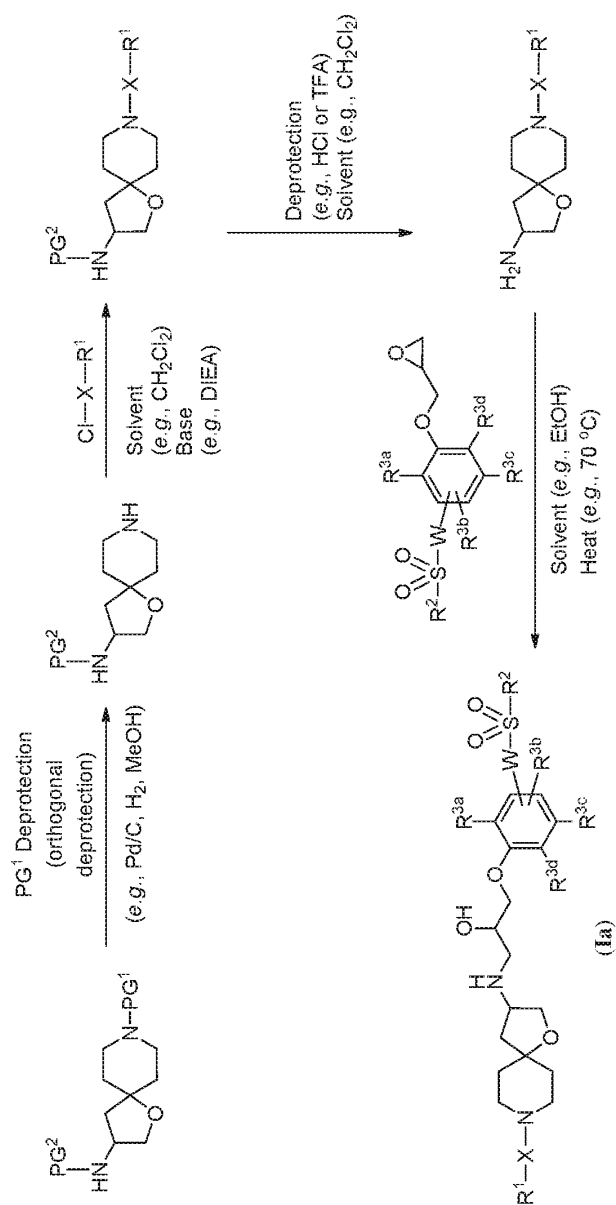


FIGURE 12

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

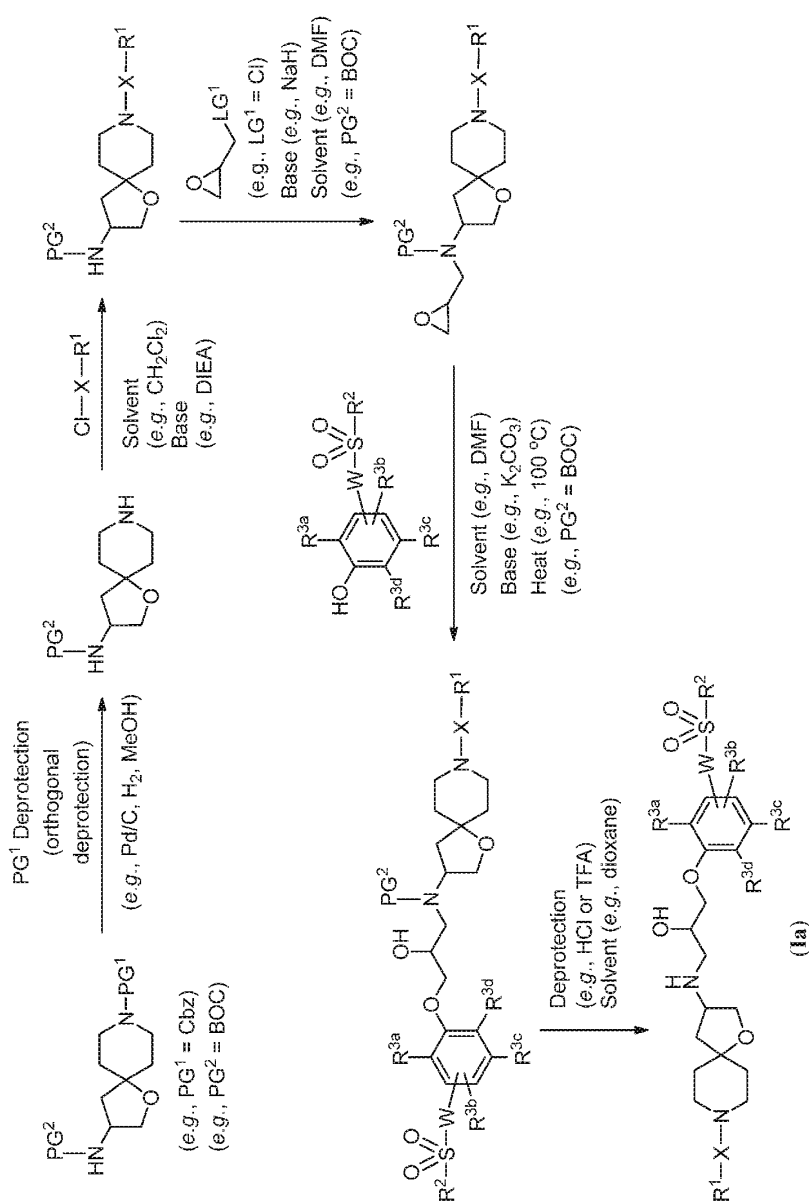


FIGURE 13

General Synthetic Scheme for the Preparation of Compounds of the Present Invention

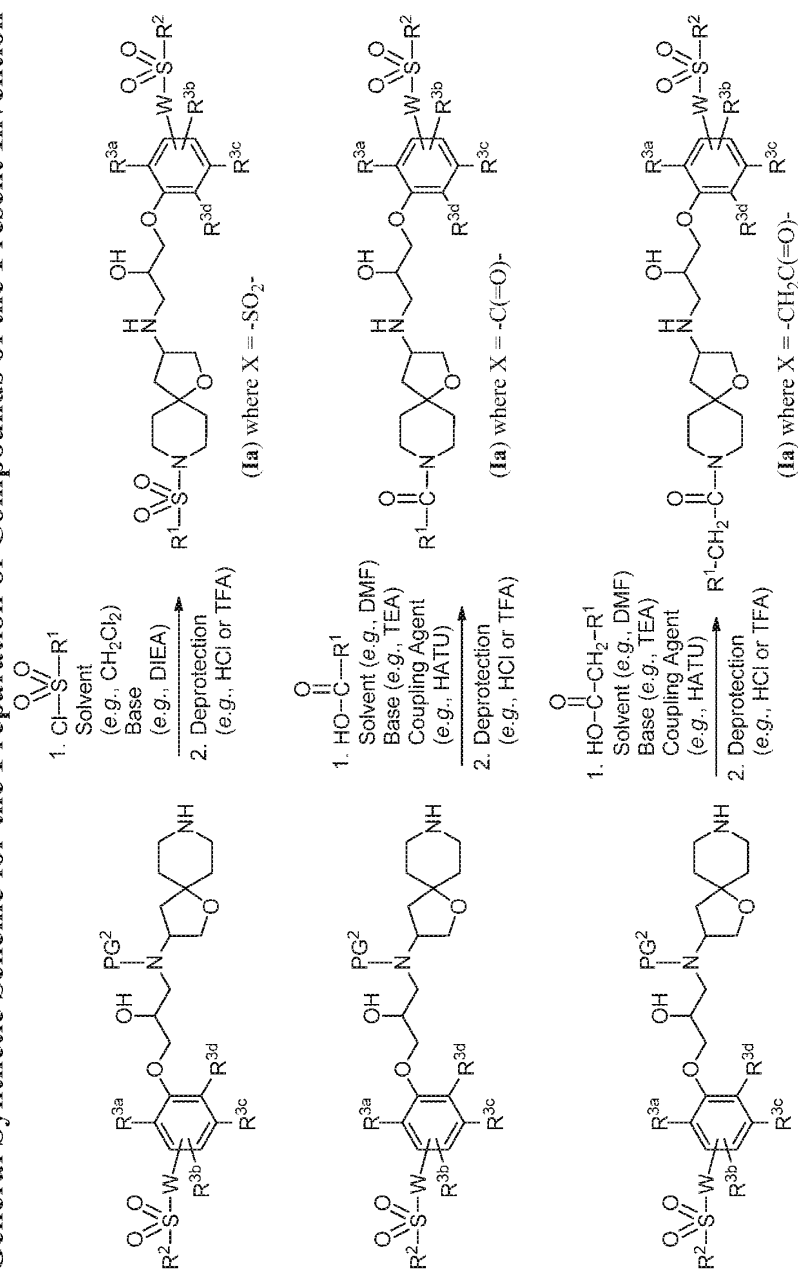


FIGURE 14

The General Synthetic Scheme for the Preparation of Certain Intermediates

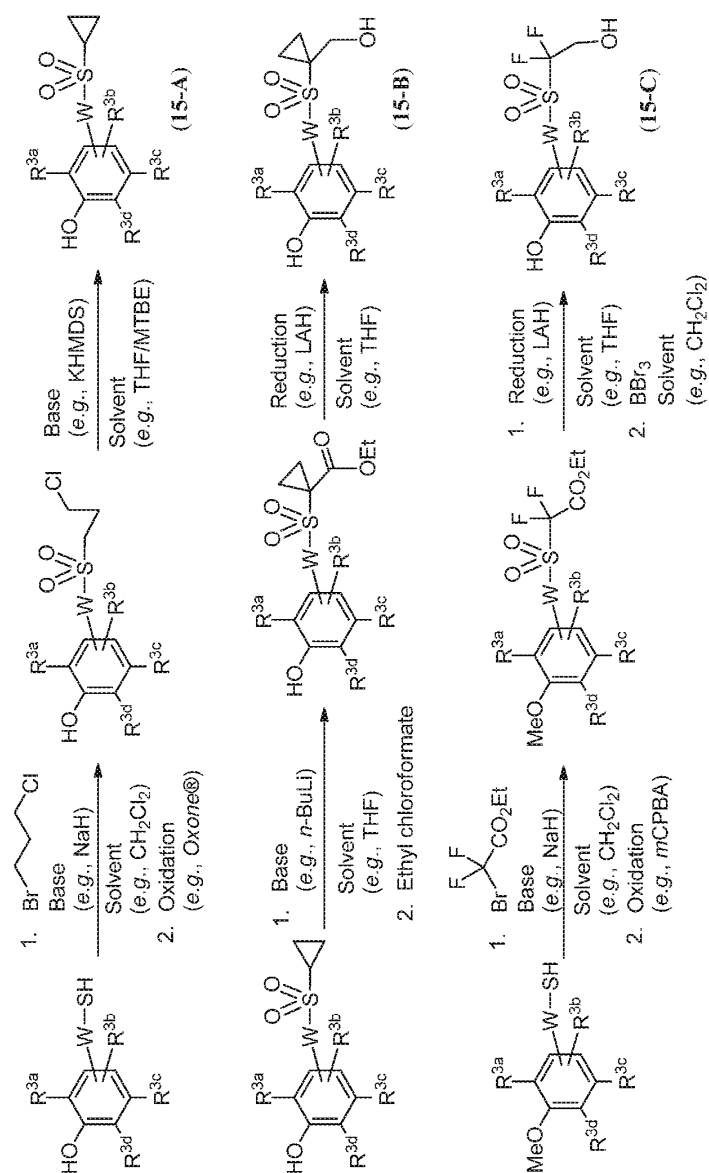


FIGURE 15

The General Synthetic Scheme for the Preparation of Certain Intermediates

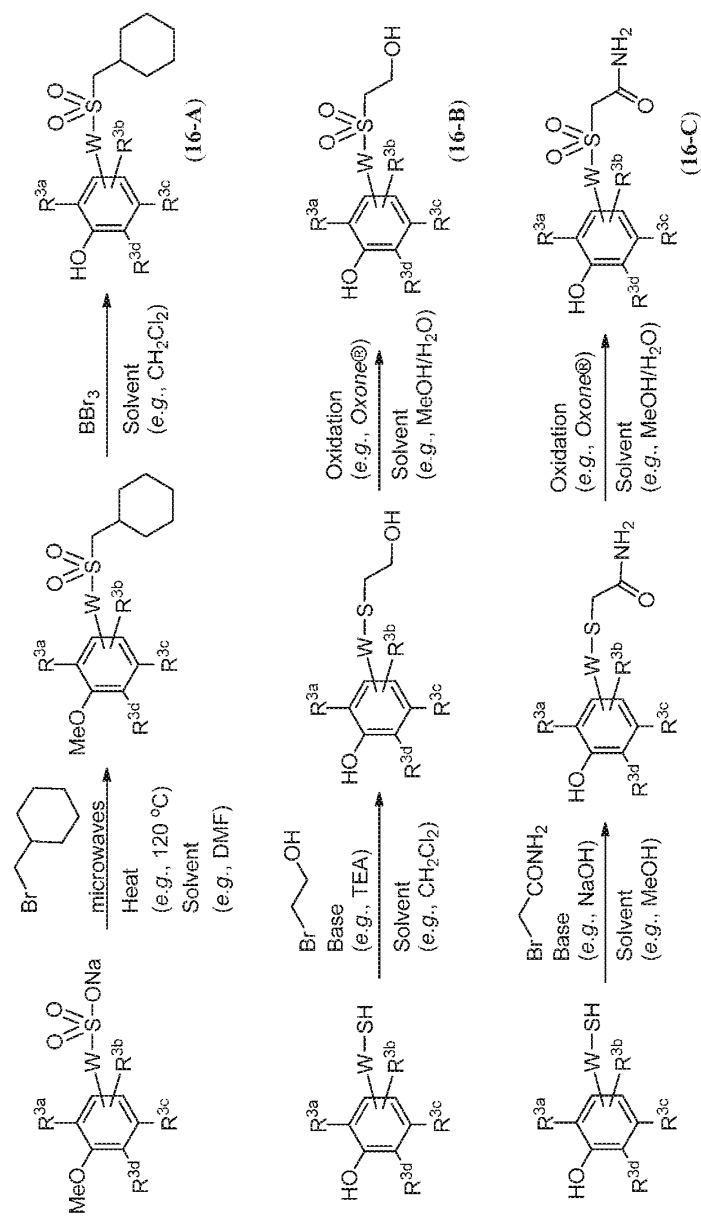


FIGURE 16

The General Synthetic Scheme for the Preparation of Certain Intermediates

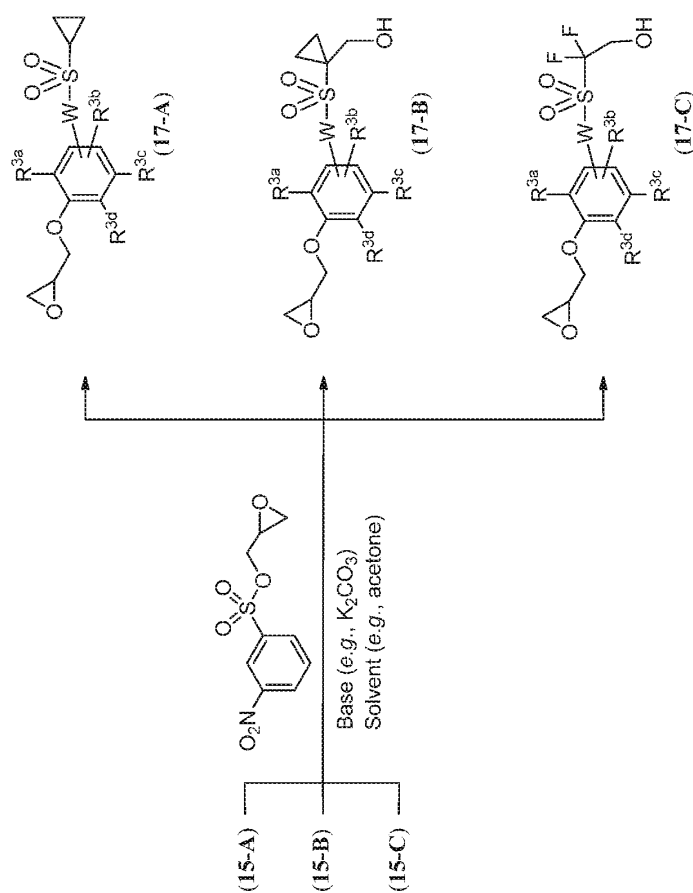


FIGURE 17

The General Synthetic Scheme for the Preparation of Certain Intermediates

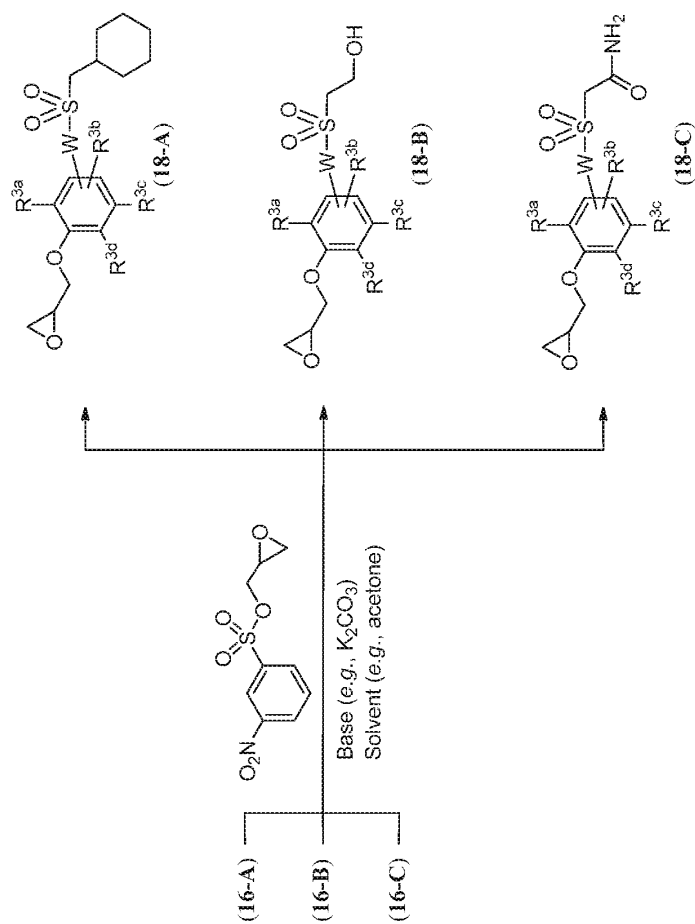


FIGURE 18

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

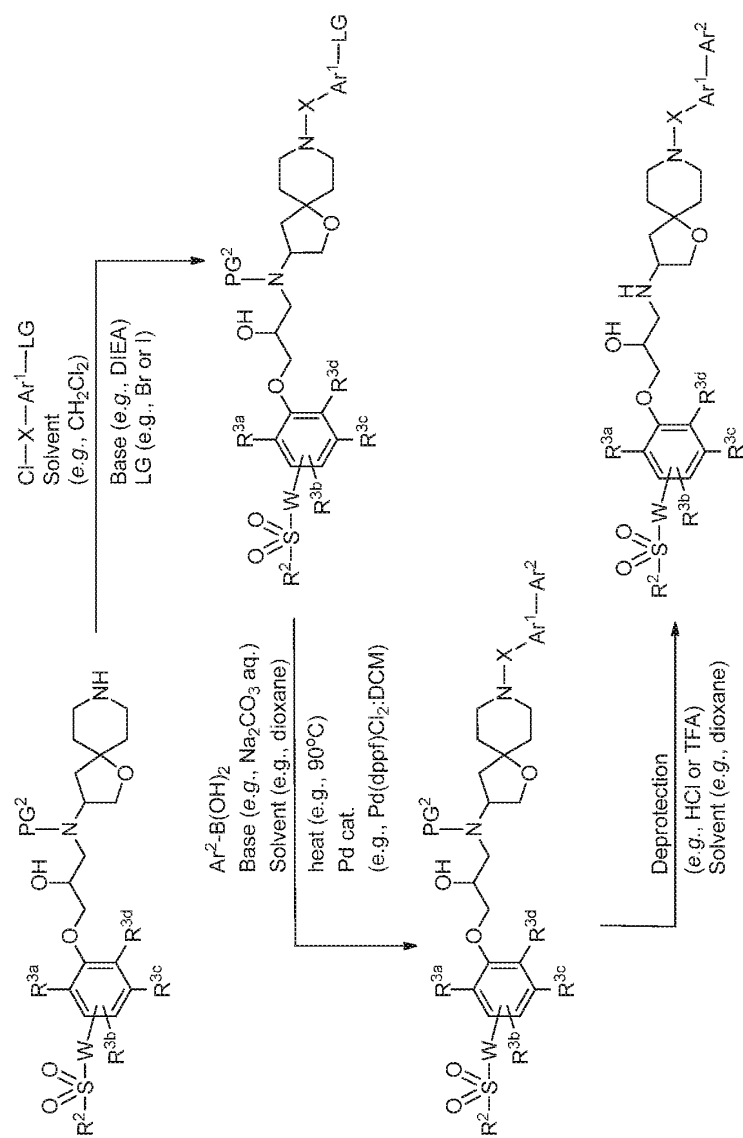


FIGURE 19

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

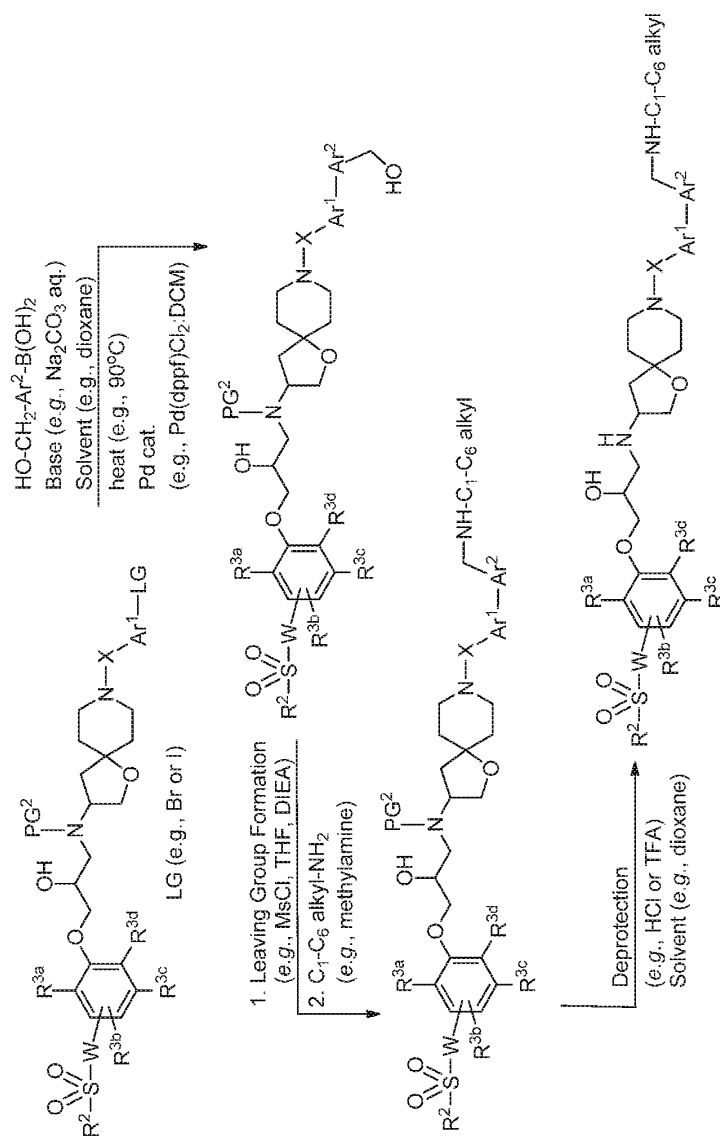


FIGURE 20

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

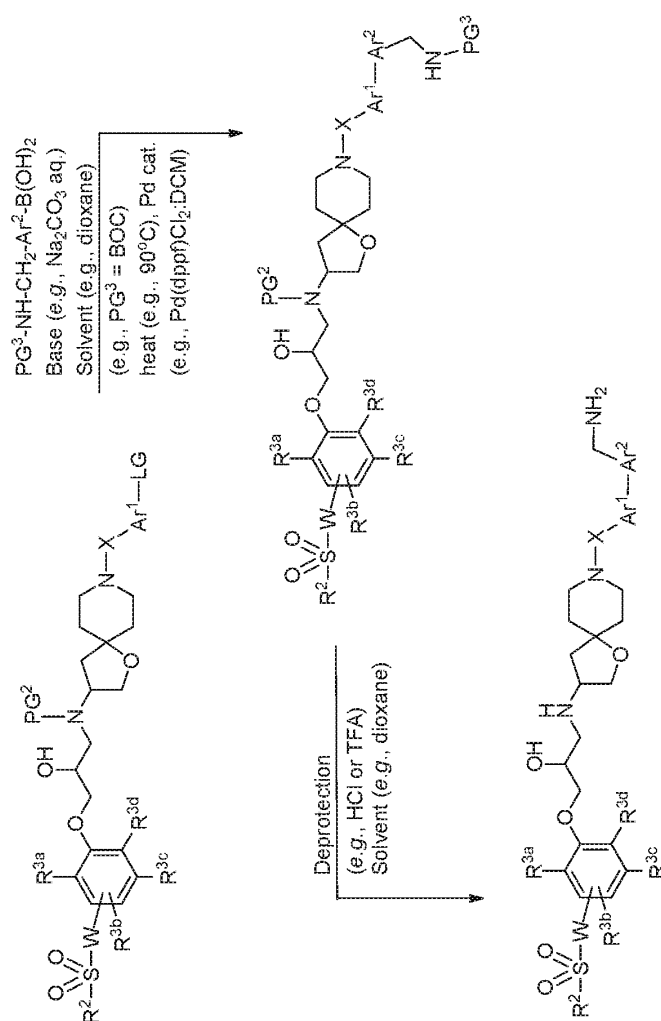


FIGURE 21

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

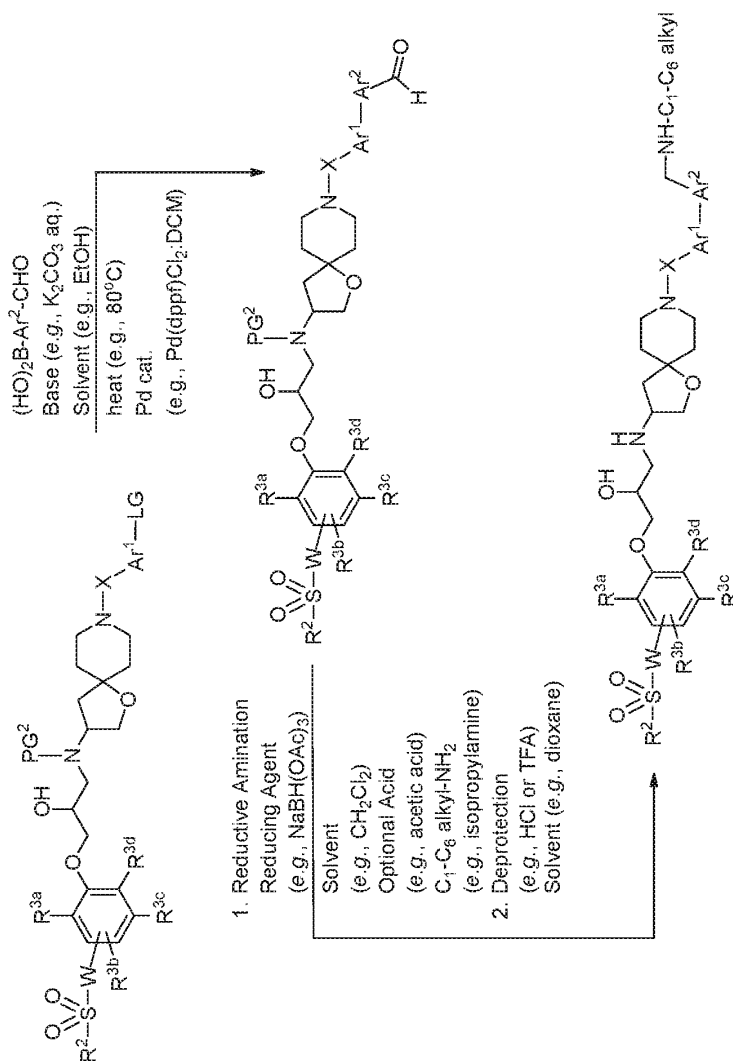


FIGURE 22

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

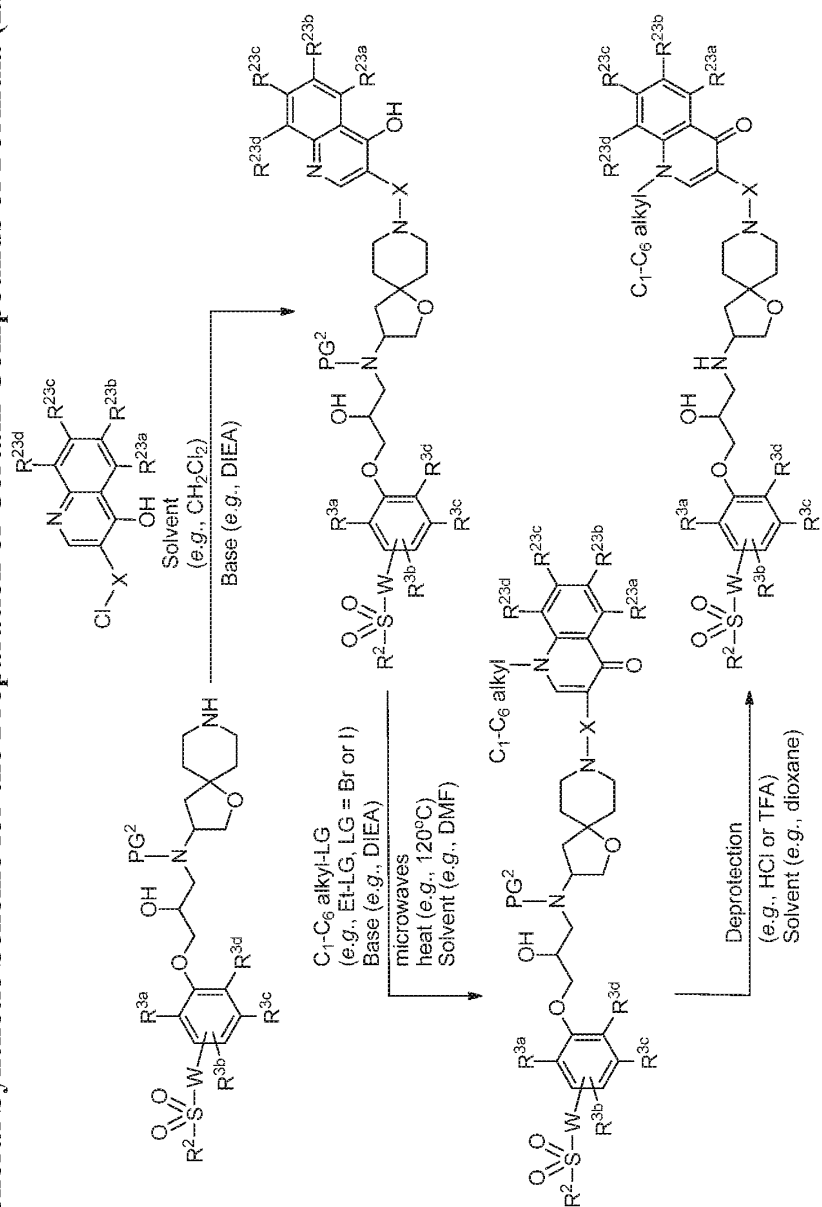


FIGURE 23

General Synthetic Scheme for the Preparation of Certain Compounds of Formula (Ia)

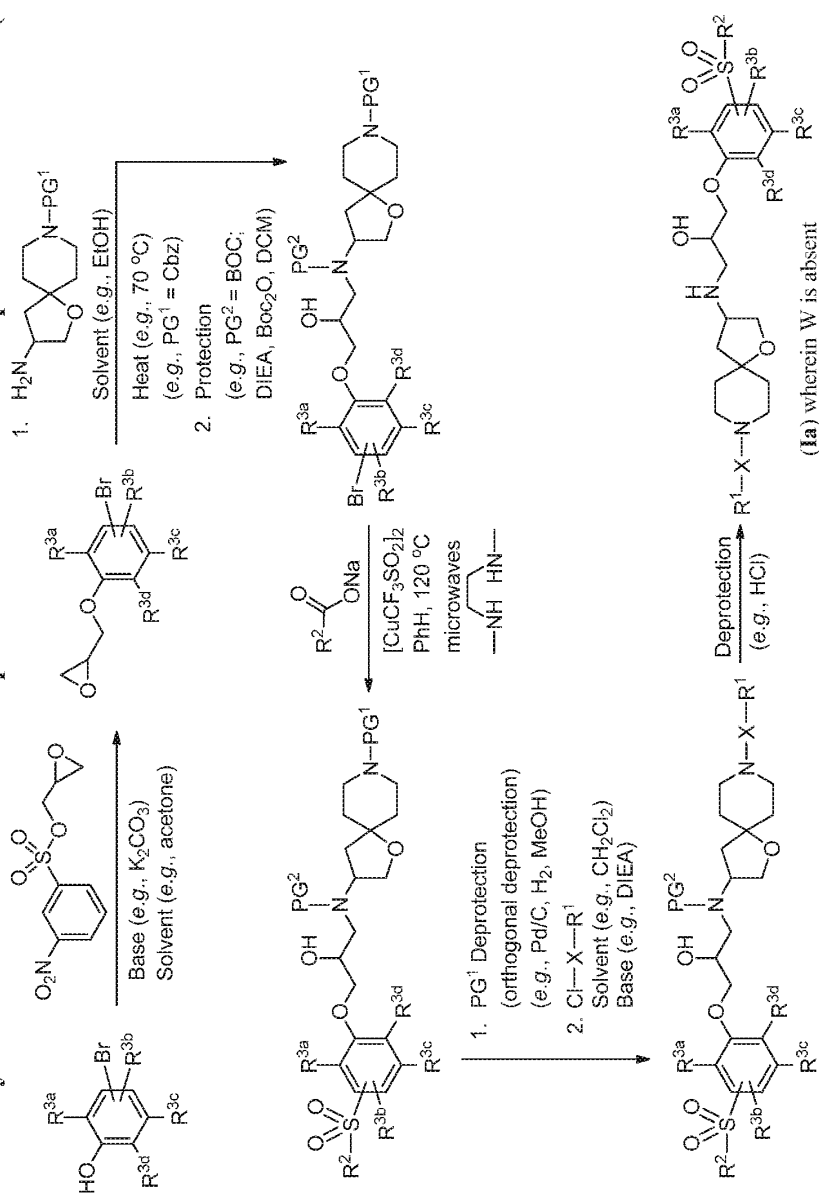


FIGURE 24

Scheme for Administration of Test Compounds in Normal Rats in Example 4

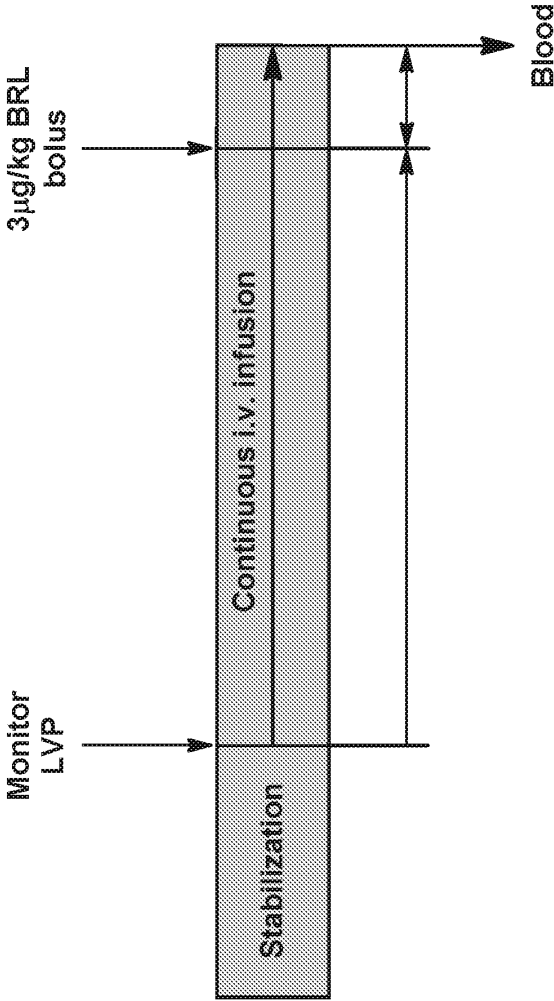


FIGURE 25

Scheme for Administration of Test Compounds in Heart Failure Rats in Example 4

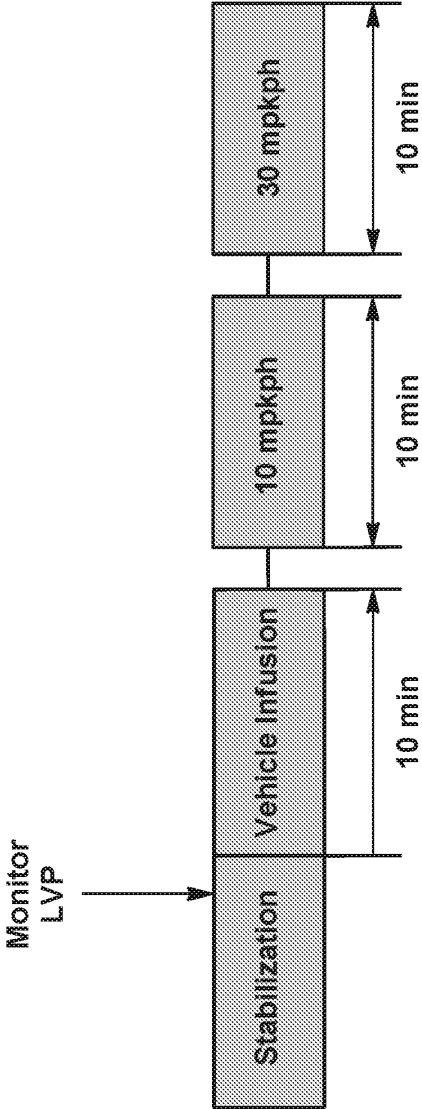


FIGURE 26

Inhibition of Negative Effects of BRL on LVP in Normal Rats
Following Administration of Compound 123

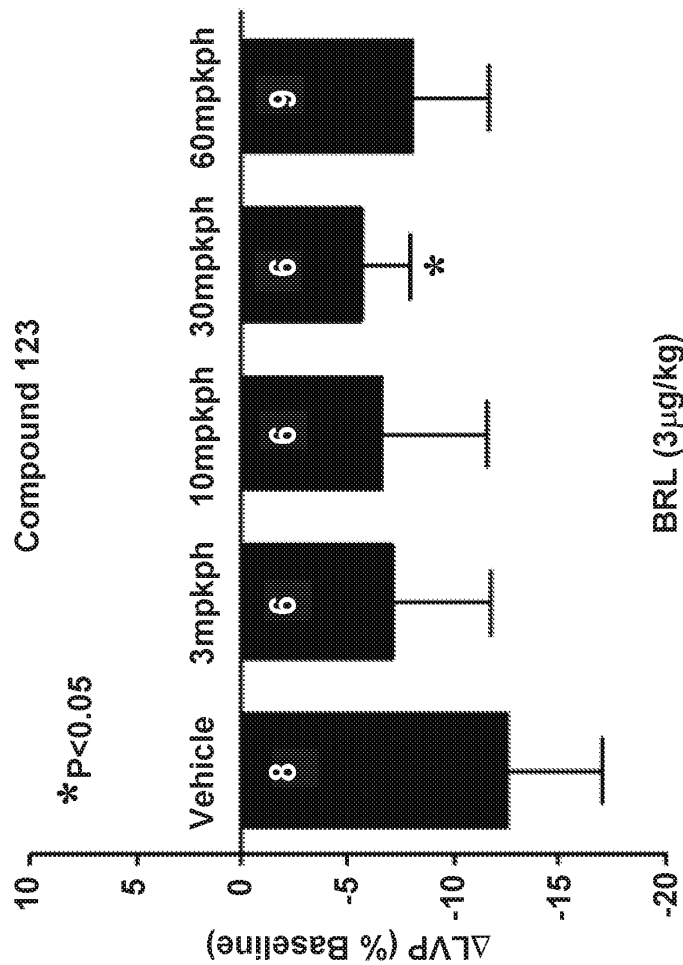
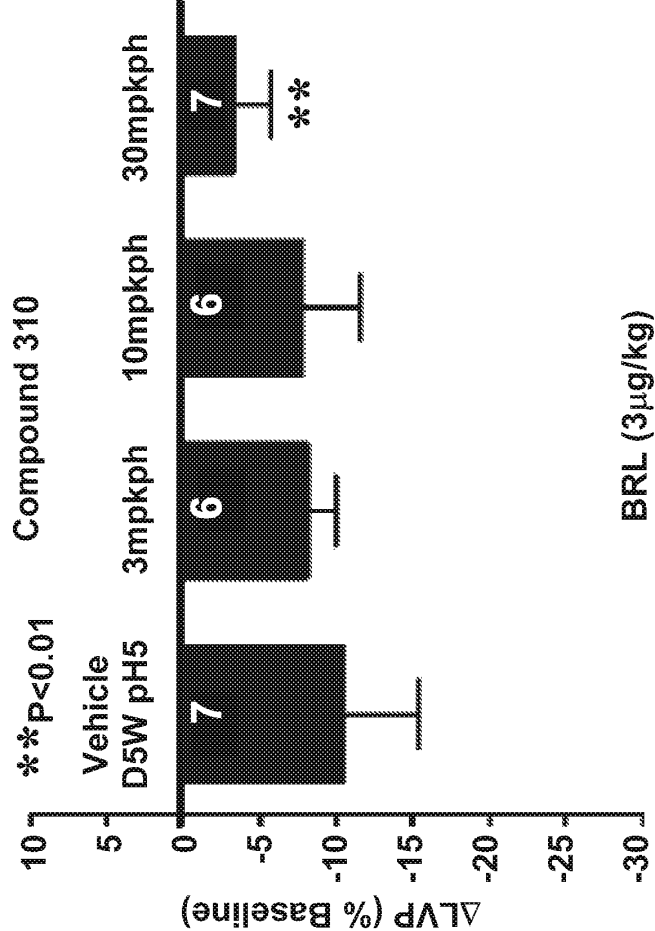


FIGURE 27

Inhibition of Negative Effects of BRL on LVP in Normal Rats
Following Administration of Compound 310



BRL (3 μ g/kg)

FIGURE 28

Inhibition of Negative Effects of BRL on LVP in Normal Rats
Following Administration of Compound 163

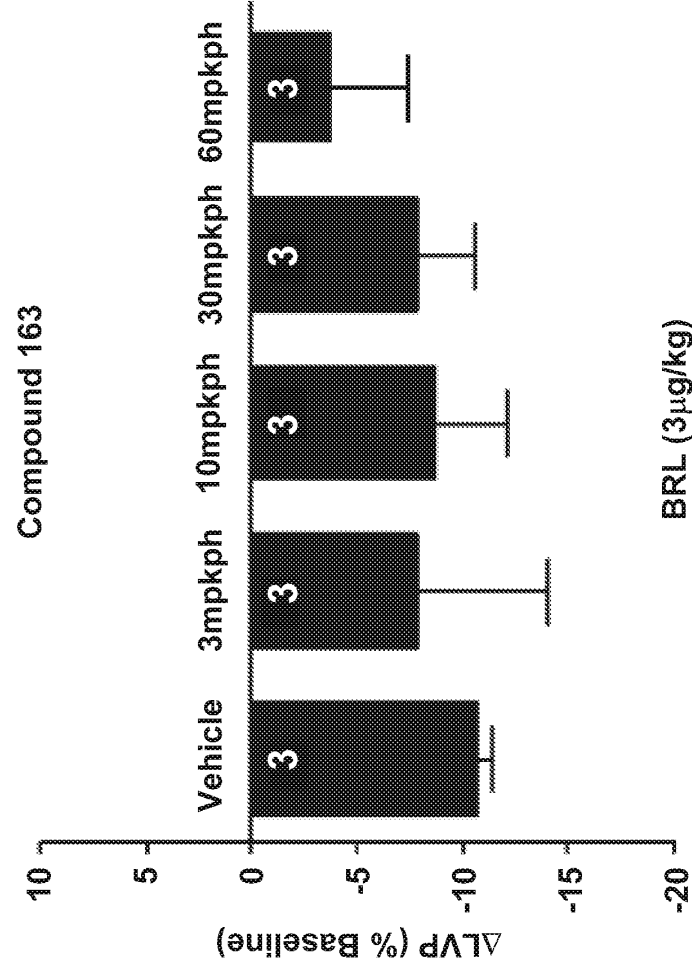


FIGURE 29

Inhibition of Negative Effects of BRL on LVP in Normal Rats
Following Administration of Compound 154

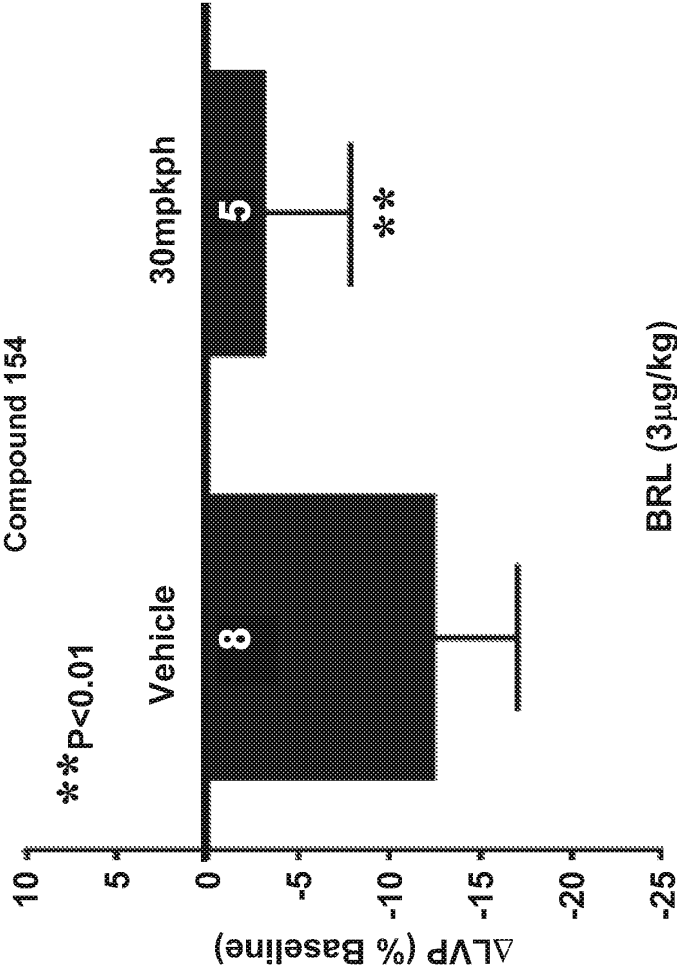


FIGURE 30

Inhibition of Negative Effects of BRL on LVP in Normal Rats
Following Administration of Compound 88

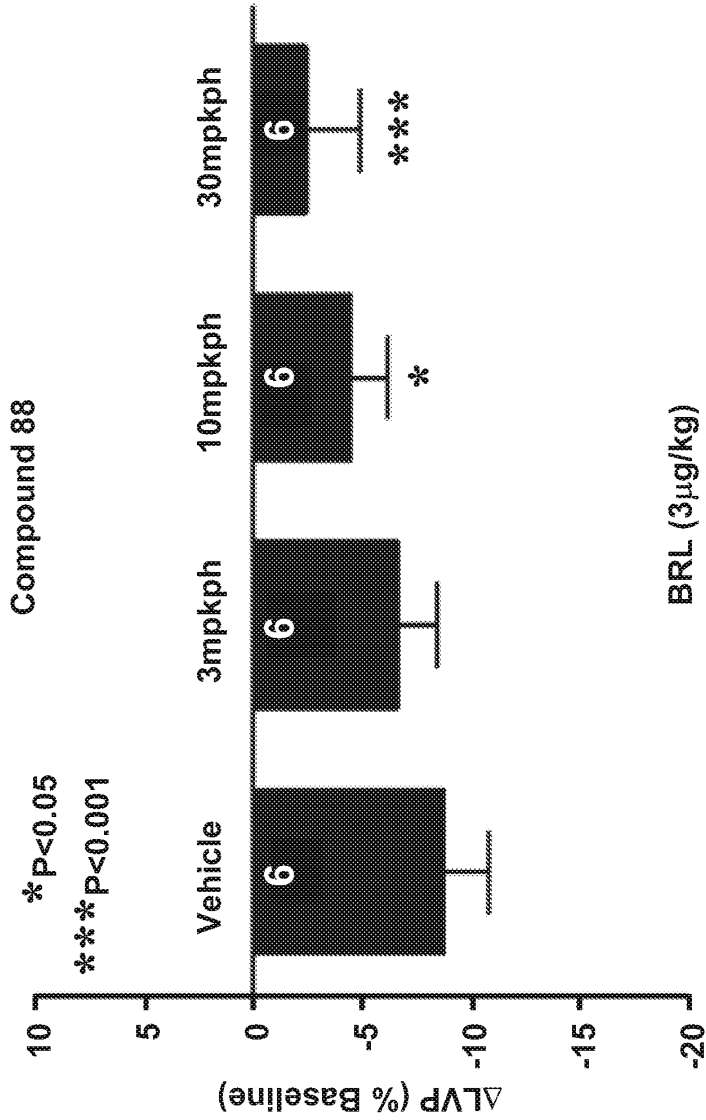


FIGURE 31

Inhibition of Negative Effects of BRL on LVP in a Rat with heart failure
Following Administration of Compound 88

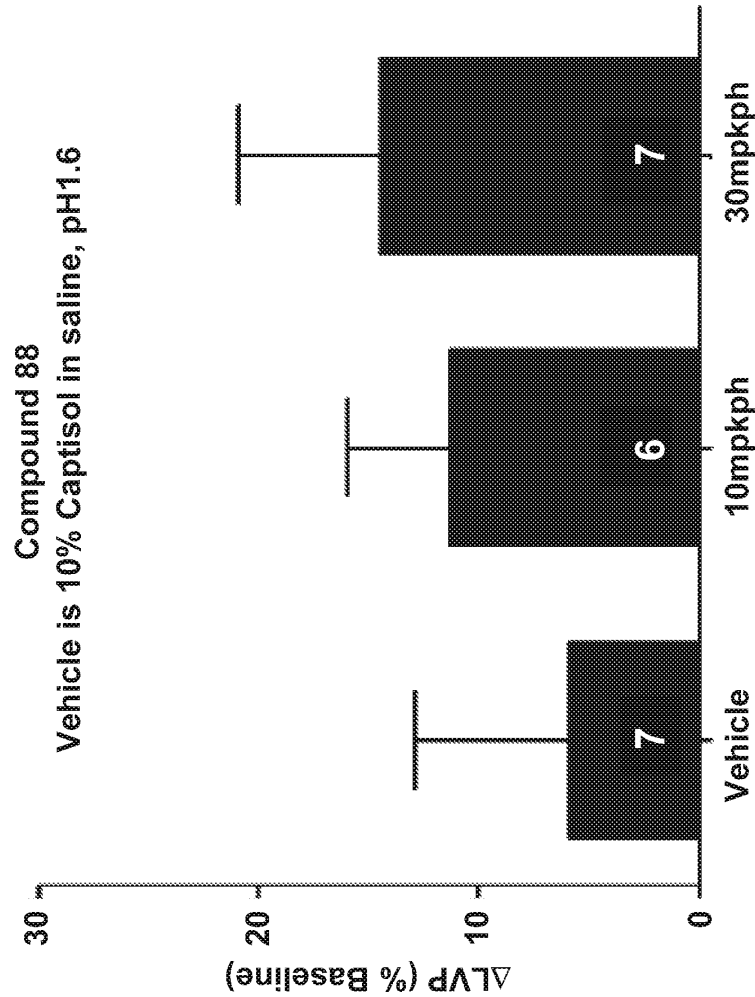


FIGURE 32

Effect of Compound 88 in a Dog Prior to Heart Failure

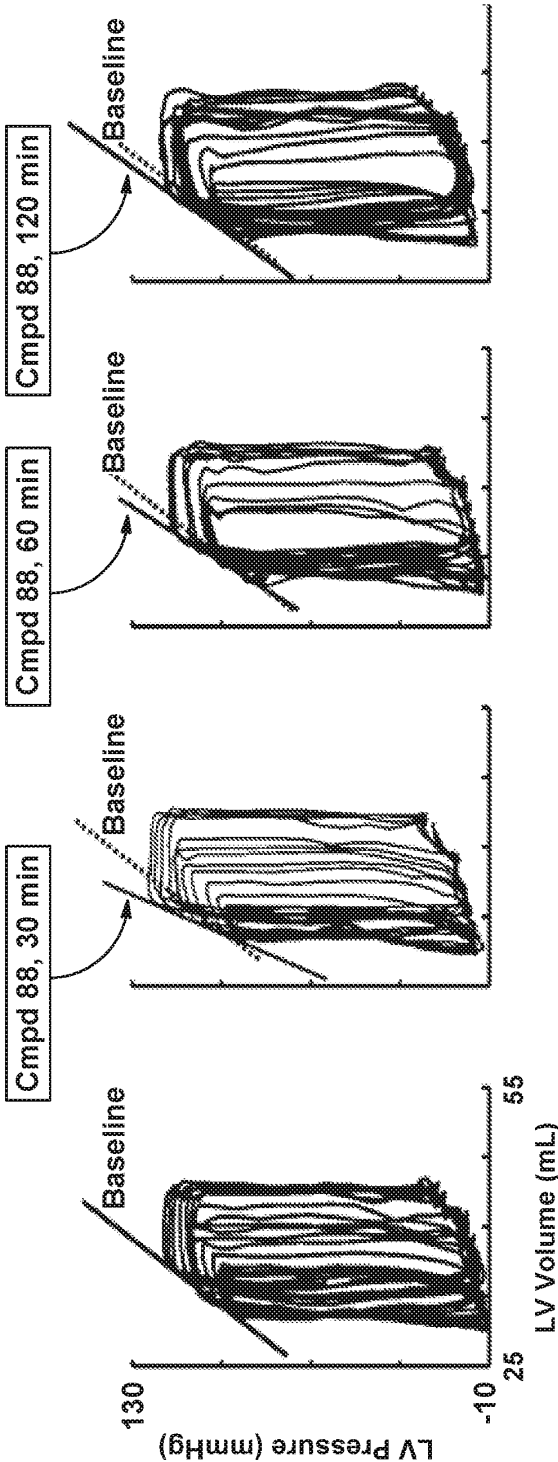


FIGURE 33A

Effect of Compound 88 in a Dog Following Heart Failure

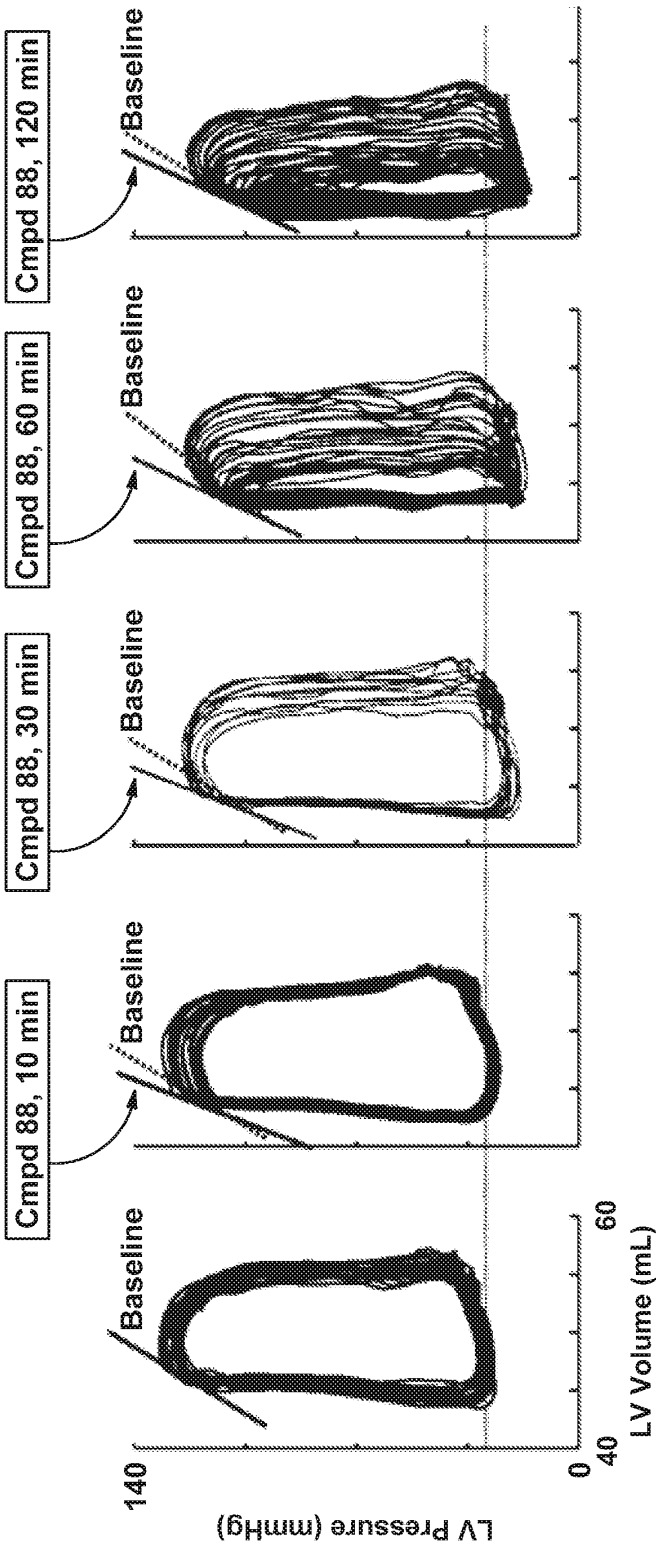


FIGURE 33B

Effect of Compound 88 on Hemodynamics and LV Contractility
in Normal Dogs (n=4)

Hemodynamic Parameters	Normal Control	After Compound 88		
	Baseline	30 min	60 min	120 min
Heart rate (beats/min)	108±8	109±7	105±8	108±7
Maximum dP/dt (mmHg/sec)	2320±213	2368±147	2240±139	2204±139
Minimum dP/dt (mmHg/sec)	-1993±143	-2053±109	-1973±109	-1944±98
Stroke Volume (mL)	16.1±2.3	16.8±3.1	17.5±3.3	16.7±2.7
LV end-diastolic pressure (mmHg)	6.4±2.7	6.0±1.4	7.2±1.6	6.9±2.3
LV end-systolic pressure (mmHg)	101±8.4	99±3.8	100±5.8	98±3.4
Minimum LV pressure (mmHg)	1.7±2.5	-0.8±2.0	0.3±2.1	-0.8±1.9
LV end-diastolic volume (mL)	50.3±9.3	50.7±5.1	51.2±4.2	51.6±5.0
LV end-systolic volume (mL)	34.2±4.8	33.9±3.5	33.7±3.1	34.9±4.0
Time constant of relaxation (msec)	27.9±2.0	25.8±2.6	26.7±2.0	26.7±1.9
E _{ES} (mmHg/mL)	6.3±1.4	7.6 ±1.0	7.1±0.9	6.4±1.2
M _{SW} (mmHg)	73.7±6.3	80.2±5.6	79.5±5.3	72.3±5.3

Values are means ± SD (n = 4)
E_{ES}: slope of linear P_{ES}-V_{ES} relation
M_{SW}: slope of SW-V_{ED} relation

FIGURE 34