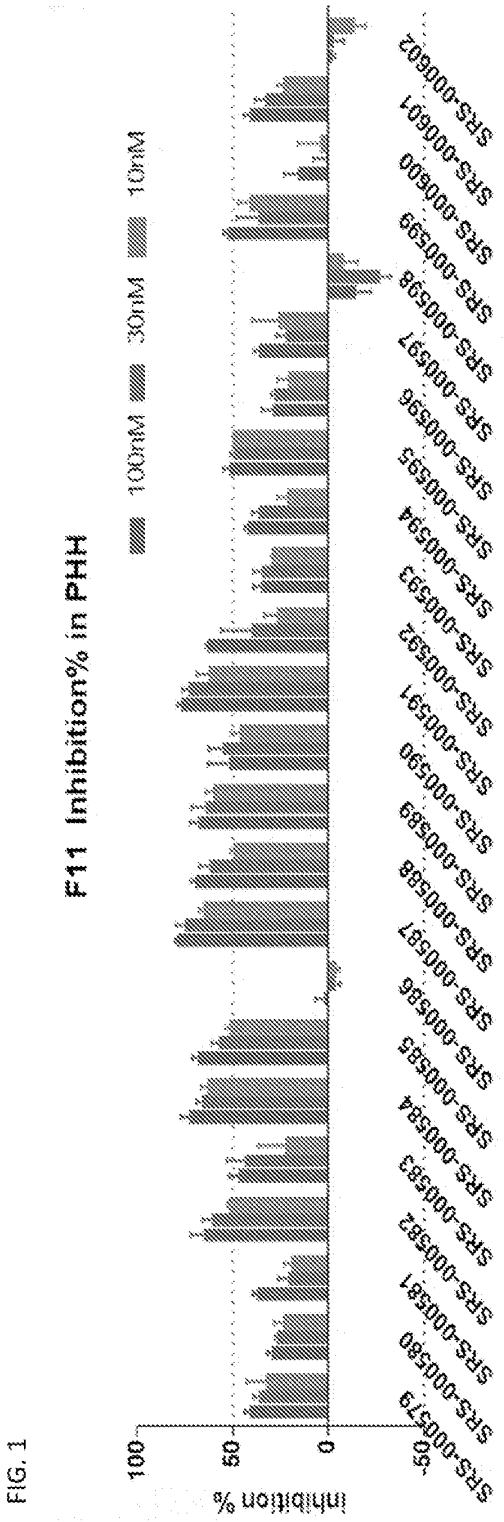

F11 Inhibition% in PHH
100nM
30nM
10nM
Inhibition %
100
50
0
-50
SRS-000579
SRS-000580
SRS-000581
SRS-000582
SRS-000583
SRS-000584
SRS-000585
SRS-000586
SRS-000587
SRS-000588
SRS-000589
SRS-000590
SRS-000591
SRS-000592
SRS-000593
SRS-000594
SRS-000595
SRS-000596
SRS-000597
SRS-000598
SRS-000599
SRS-000600
SRS-000601
SRS-000602

FIG. 1

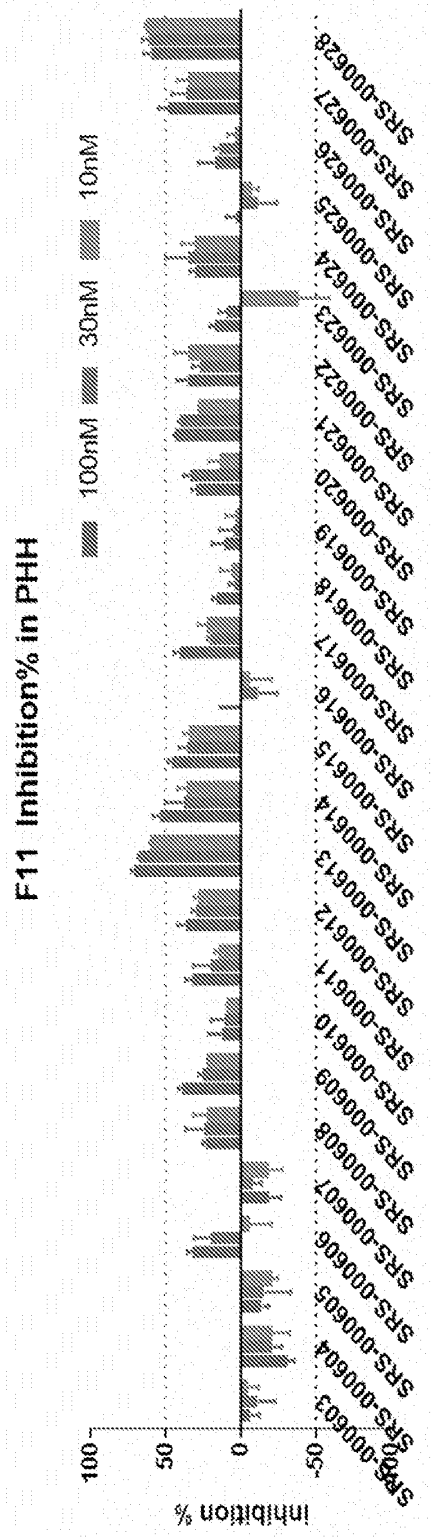

F11 Inhibition% in PHH
100nM
30nM
10nM
inhibition %
100
50
0
-50
SRS-000603
SRS-000604
SRS-000605
SRS-000606
SRS-000607
SRS-000608
SRS-000609
SRS-000610
SRS-000611
SRS-000612
SRS-000613
SRS-000614
SRS-000615
SRS-000616
SRS-000617
SRS-000618
SRS-000619
SRS-000620
SRS-000621
SRS-000622
SRS-000623
SRS-000624
SRS-000625
SRS-000626
SRS-000627
SRS-000628

FIG. 2

FIG. 3

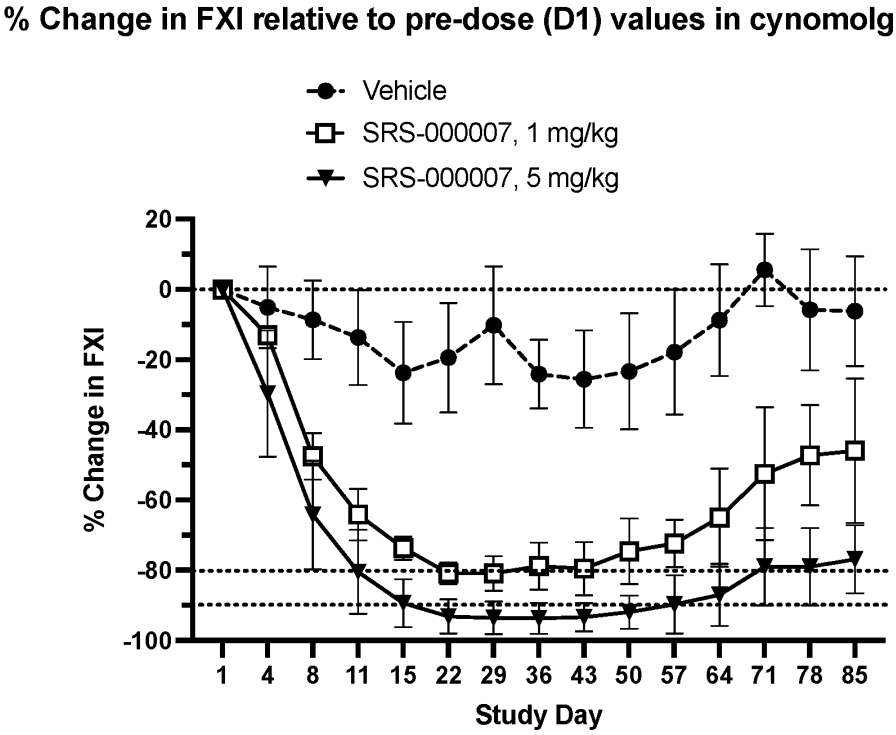

% Change in FXI relative to pre-dose (D1) values in cynomolgus monkeys
Vehicle
SRS-000007, 1 mg/kg
SRS-000007, 5 mg/kg
% Change in FXI
20
0
-20
-40
-60
-80
-100
1
4
8
11
15
22
29
36
43
50
57
64
71
78
85
Study Day

FIG. 4

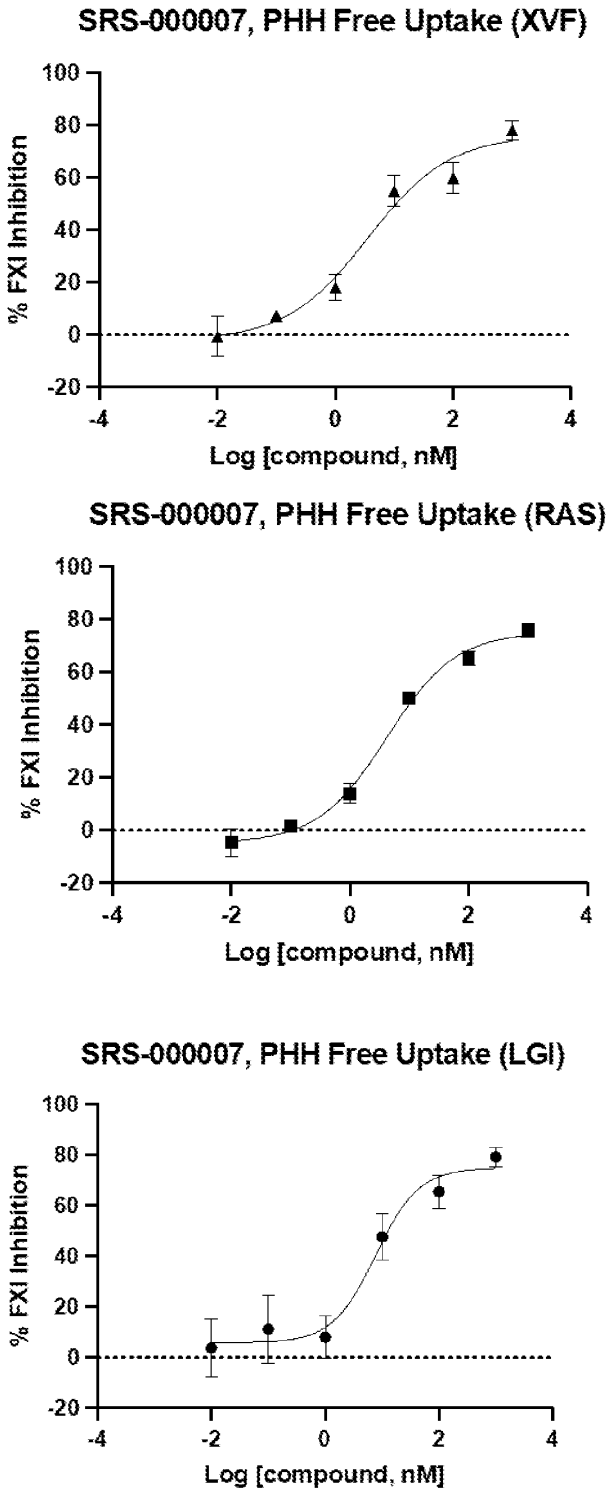

SRS-000007, PHH Free Uptake (XVF)
% FXI Inhibition
100
80
60
40
20
0
-20
-4
-2
0
2
4
Log [compound, nM]
SRS-000007, PHH Free Uptake (RAS)
% FXI Inhibition
100
80
60
40
20
0
-20
-4
-2
0
2
4
Log [compound, nM]
SRS-000007, PHH Free Uptake (LGI)
% FXI Inhibition
100
80
60
40
20
0
-20
-4
-2
0
2
4
Log [compound, nM]

FIG. 5

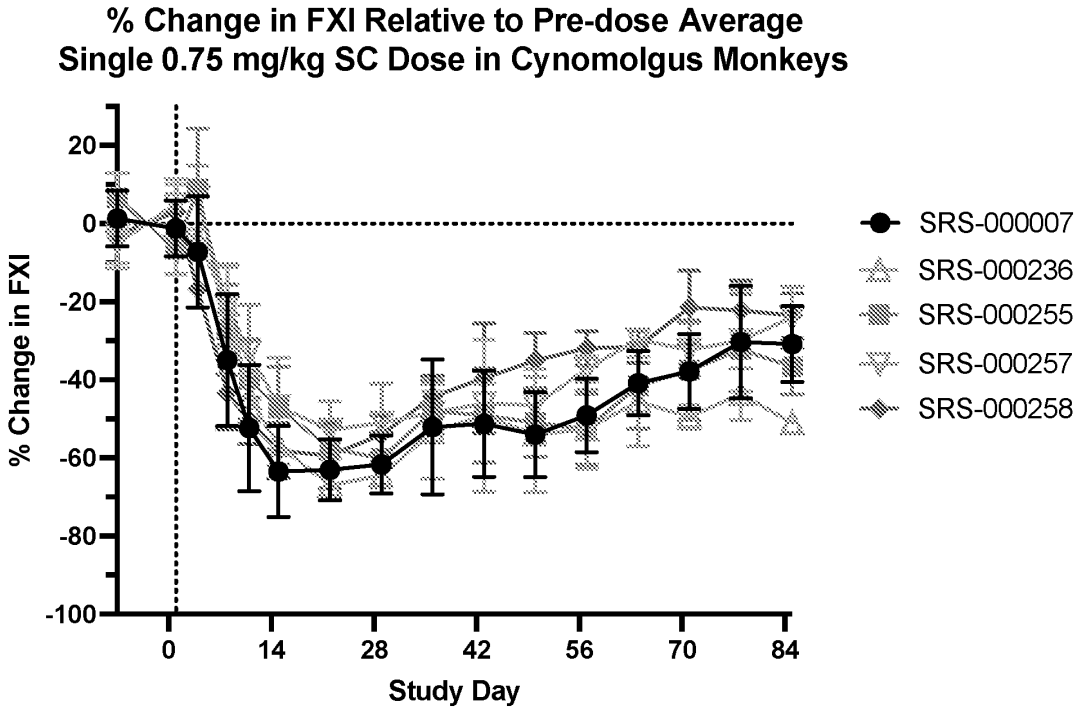

% Change in FXI Relative to Pre-dose Average
Single 0.75 mg/kg SC Dose in Cynomolgus Monkeys
% Change in FXI
20
0
-20
-40
-60
-80
-100
0
14
28
42
56
70
84
Study Day
SRS-000007
SRS-000236
SRS-000255
SRS-000257
SRS-000258

FIG. 6

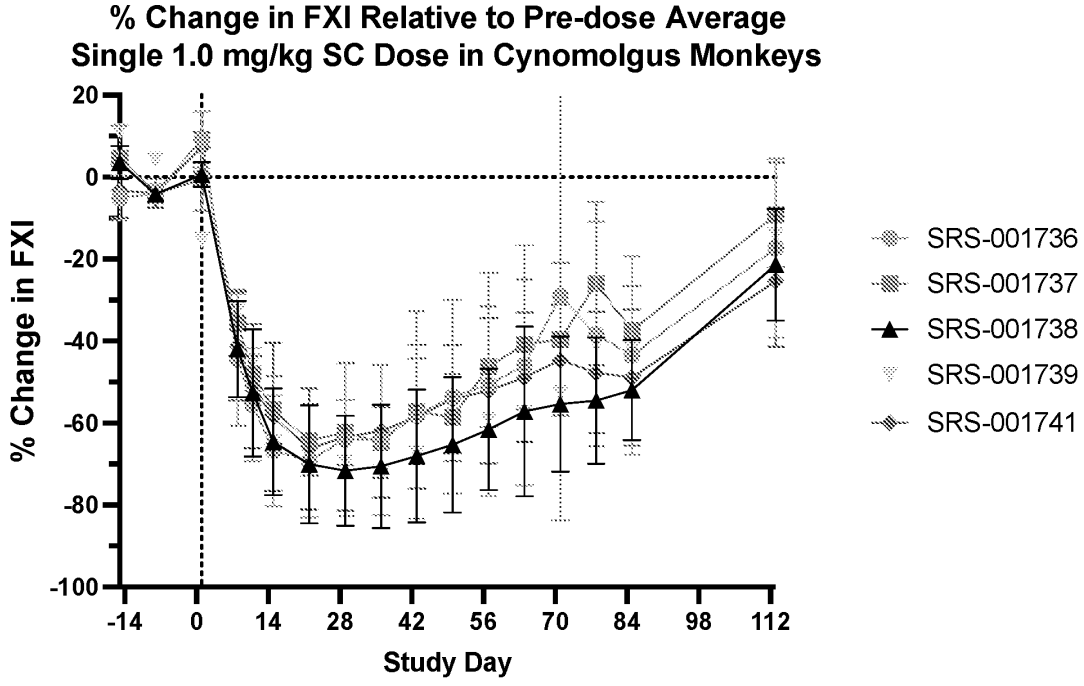

% Change in FXI Relative to Pre-dose Average
Single 1.0 mg/kg SC Dose in Cynomolgus Monkeys
% Change in FXI
20
0
-20
-40
-60
-80
-100
-14
0
14
28
42
56
70
84
98
112
Study Day
SRS-001736
SRS-001737
SRS-001738
SRS-001739
SRS-001741

FIG. 7

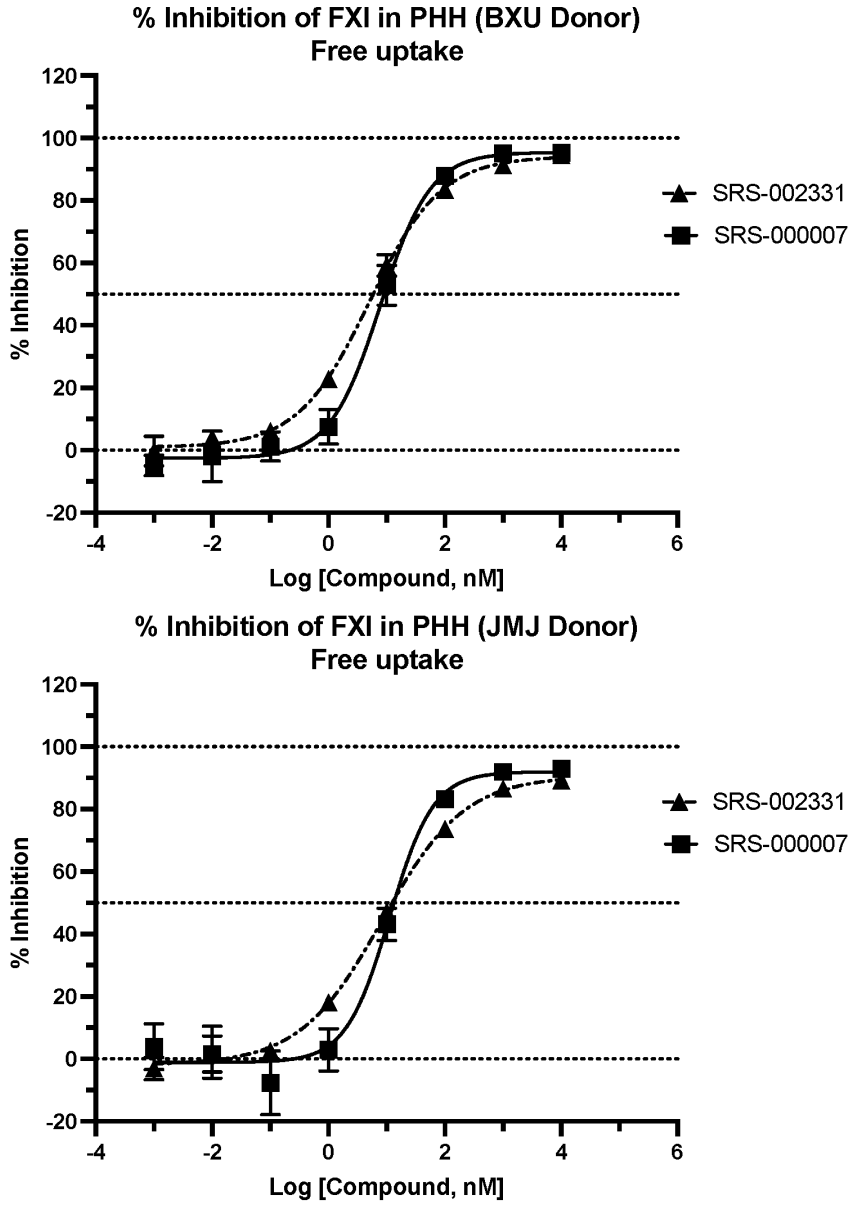

% Inhibition of FXI in PHH (BXU Donor)
Free uptake
% Inhibition
Log [Compound, nM]
SRS-002331
SRS-000007
% Inhibition of FXI in PHH (JMJ Donor)
Free uptake
% Inhibition
Log [Compound, nM]
SRS-002331
SRS-000007

FIG. 8

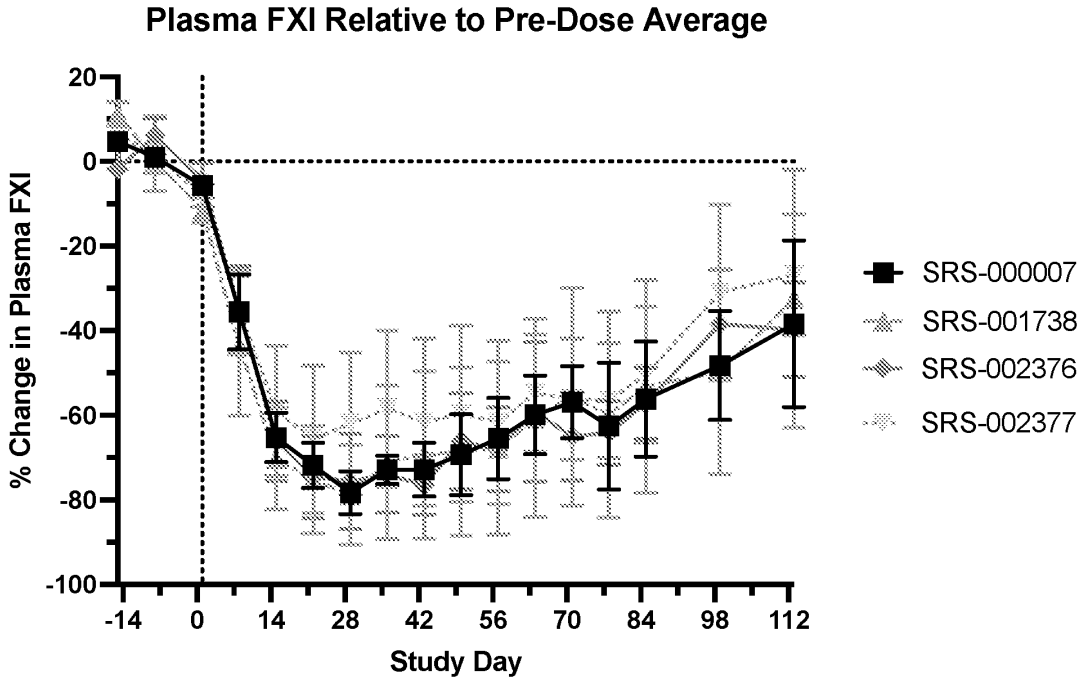
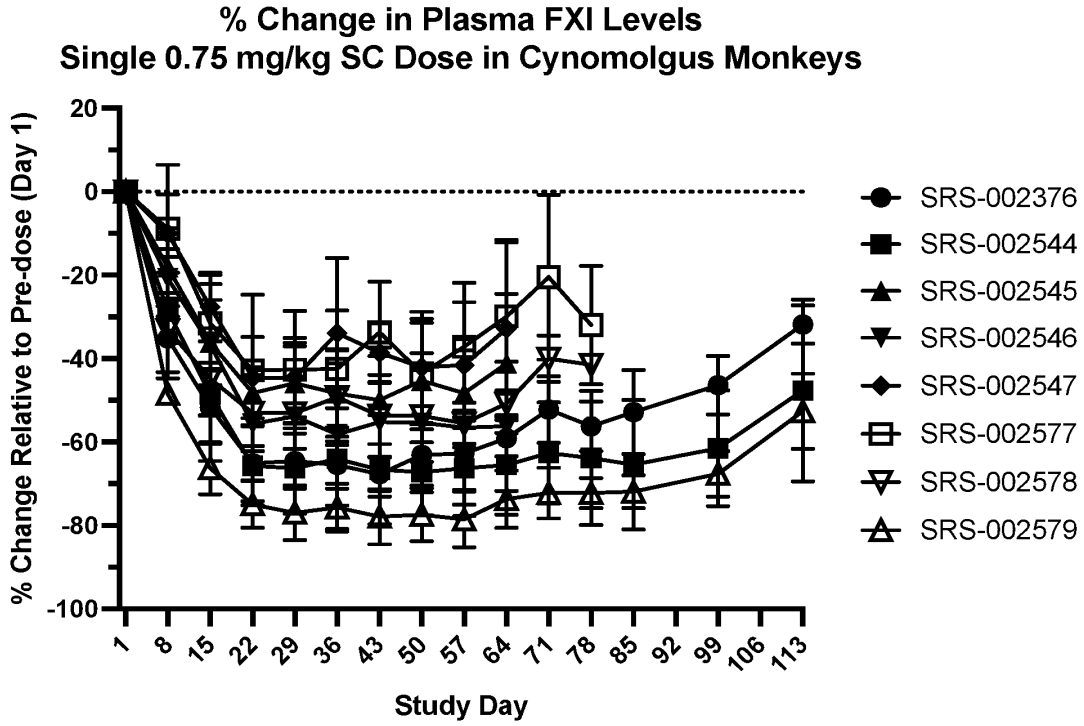

Plasma FXI Relative to Pre-Dose Average
% Change in Plasma FXI
20
0
-20
-40
-60
-80
-100
-14
0
14
28
42
56
70
84
98
112
Study Day
SRS-000007
SRS-001738
SRS-002376
SRS-002377

FIG. 9


% Change in Plasma FXI Levels
Single 0.75 mg/kg SC Dose in Cynomolgus Monkeys
% Change Relative to Pre-dose (Day 1)
20
0
-20
-40
-60
-80
-100
1
8
15
22
29
36
43
50
57
64
71
78
85
92
99
106
113
Study Day
SRS-002376
SRS-002544
SRS-002545
SRS-002546
SRS-002547
SRS-002577
SRS-002578
SRS-002579