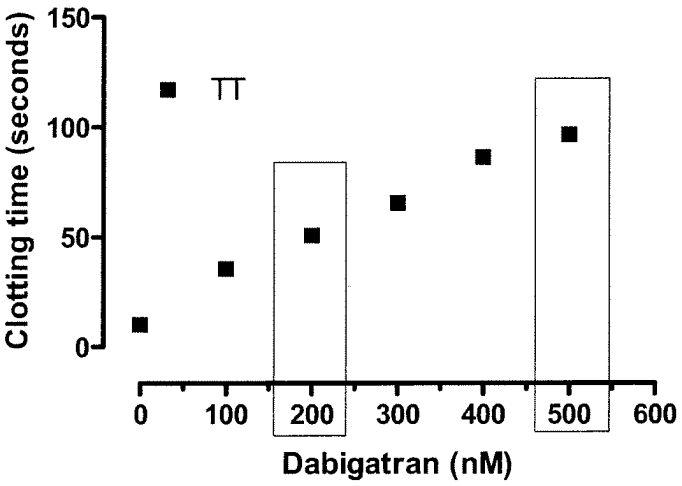

Clotting time (seconds)
150
100
50
0
TT
0
100
200
300
400
500
600
Dabigatran (nM)

Figure 1

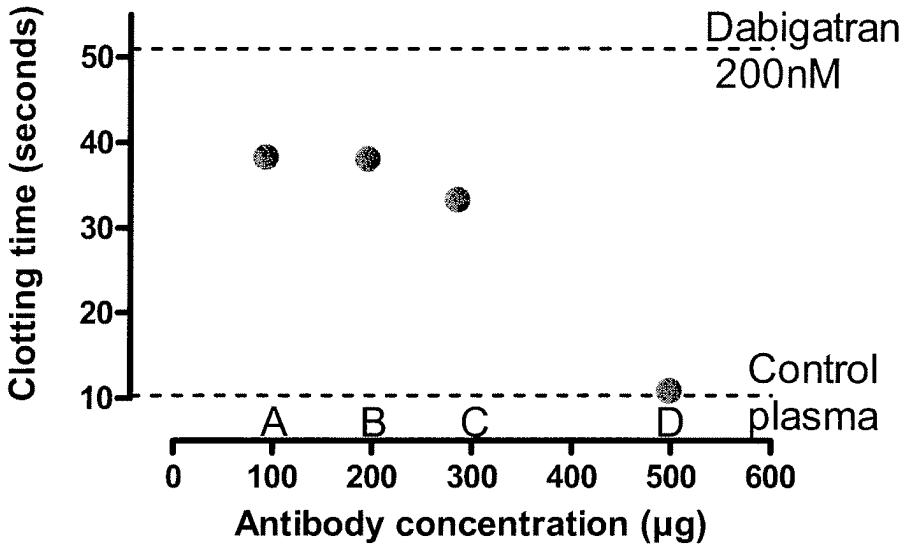

Clotting time (seconds)
50
40
30
20
10
Dabigatran
200nM
Control
plasma
A
B
C
D
0
100
200
300
400
500
600
Antibody concentration (µg)

Figure 2

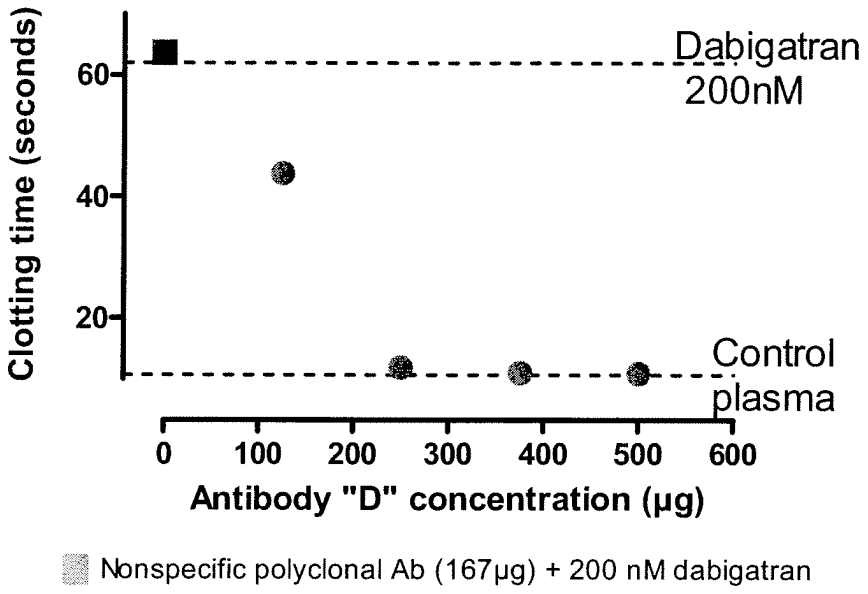

Clotting time (seconds)
60
40
20
Dabigatran
200nM
Control
plasma
0
100
200
300
400
500
600
Antibody "D" concentration (µg)
Nonspecific polyclonal Ab (167µg) + 200 nM dabigatran

Figure 3

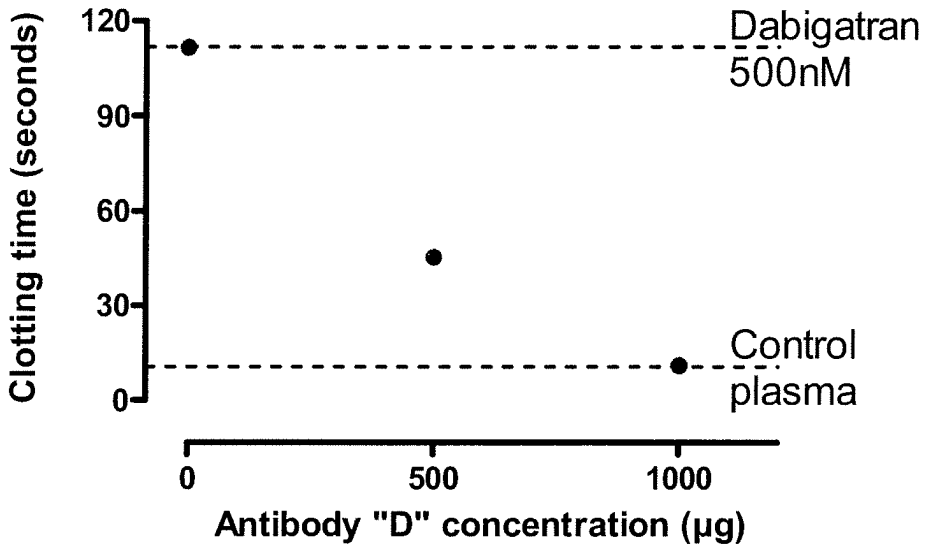

120
90
60
30
0
Clotting time (seconds)
Dabigatran
500nM
Control
plasma
0
500
1000
Antibody "D" concentration (µg)

FIGURE 4

Figure 5

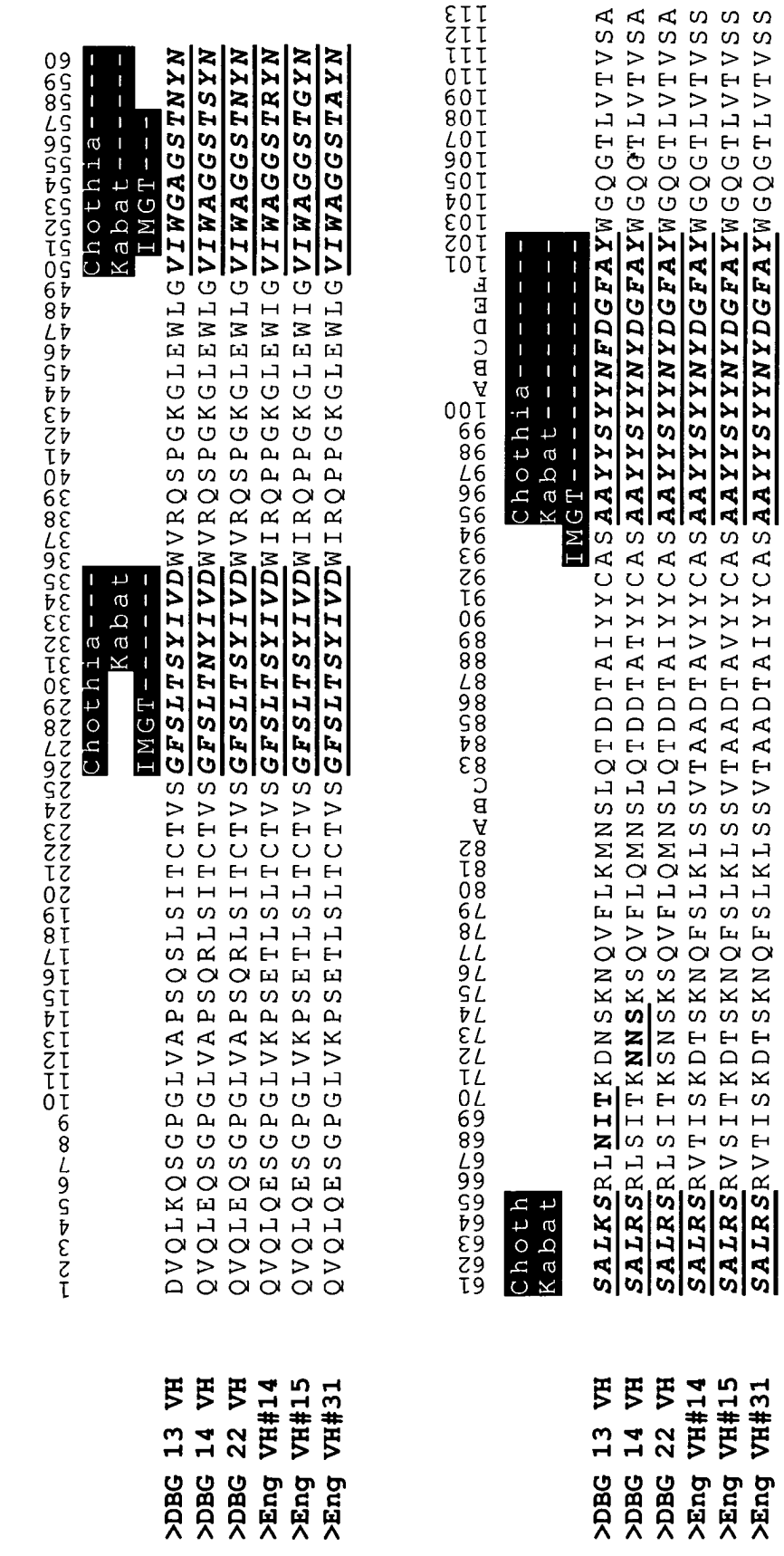

Chothia----
Kabat------
IMGT---
>DBG 13 VH DVQLKQSGPGLVAPSQSLSITCTVSGFSLTSYIVDWVRQSPGKGLEWLGVIWGAGGSTNYN
>DBG 14 VH QVQLEQSGPGLVAPSQRLSITCTVSGFSLTNYIVDWVRQSPGKGLEWLGVIWAGGSTSYN
>DBG 22 VH QVQLEQSGPGLVAPSQRLSITCTVSGFSLTSYIVDWVRQSPGKGLEWLGVIWAGGSTNYN
>Eng VH#14 QVQLQESGPGLVKPSETLSLTCTVSGFSLTSYIVDWIRQPPGKGLEWIGVIWAGGSTRYN
>Eng VH#15 QVQLQESGPGLVKPSETLSLTCTVSGFSLTSYIVDWIRQPPGKGLEWIGVIWAGGSTGYN
>Eng VH#31 QVQLQESGPGLVKPSETLSLTCTVSGFSLTSYIVDWIRQPPGKGLEWLGVIWAGGSTAYN
Chothia----------
Kabat------------
IMGT-------------
>DBG 13 VH SALKSRLNITKDNSKNQVFLKMNSLQTDDTAIYYCASAAYYSYYNFDGFAYWGQGTLVTVSA
>DBG 14 VH SALRSRLSITKNNSKSQVFLQMNSLQTDDTATYYCASAAYYSYYNYDGFAYWGQGTLVTVSA
>DBG 22 VH SALRSRLSITKSNSKSQVFLQMNSLQTDDTAIYYCASAAYYSYYNYDGFAYWGQGTLVTVSA
>Eng VH#14 SALRSRVTISKDTSKNQFSLKLSSVTAADTAVYYCASAAYYSYYNYDGFAYWGQGTLVTVSS
>Eng VH#15 SALRSRVSITKDTSKNQFSLKLSSVTAADTAVYYCASAAYYSYYNYDGFAYWGQGTLVTVSS
>Eng VH#31 SALRSRVTISKDTSKNQFSLKLSSVTAADTAIYYCASAAYYSYYNYDGFAYWGQGTLVTVSS

Figure 6

Chothia----------
Kabat----------
IMGT-------
>DBG 13 VK DIVMTQTPLTLSVTIGQPASISCKSSQSLLYTNGKTYLYWLLQRPGQSPKRLIY
>DBG 14 VK DVVLTQTPLTLSVTIGQPASISCKSSQSLLYTNGKSYLYWLLQRPGQSPKRLIY
>DBG 22 VK DVVMTQTPLTLSVTIGQPASISCKSSQSLLYTNGKTYLYWLLQRPGQSPKRLIY
>Eng VK#11 DVVMTQSPLSLPVTLGQPASISCKSSQSLLYTDGKTYLYWFLQRPGQSPKRLIY
>Eng VK#17 DVVMTQSPLSLPVTLGQPASISCKSSQSLLYTDGKTYLYWFQQRPGQSPKRLIY
>Eng VK#18 DVVMTQSPLSLPVTLGQPASISCKSSQSLLYTDGKTYLYWFLQRPGQSPRRLIY
Chothia--
Kabat----
IMGT-----
>DBG 13 VK LVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYYCLQSTHFPHTFGGGTKLEIK
>DBG 14 VK LVSKLDSGVPDRFSGSGSGTDFTLKISGVEAEDVGIYYCLQSTHFPHTFGGGTKLEIK
>DBG 22 VK LVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDVGIYYCLQSTHFPHTFGGGTKLEIK
>Eng VK#11 LVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDVGIYYCLQSTHFPHTFGGGTKVEIK
>Eng VK#17 LVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDVGIYYCLQSTHFPHTFGGGTKVEIK
>Eng VK#18 LVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDVGVYYCLQSTHFPHTFGGGTKVEIK

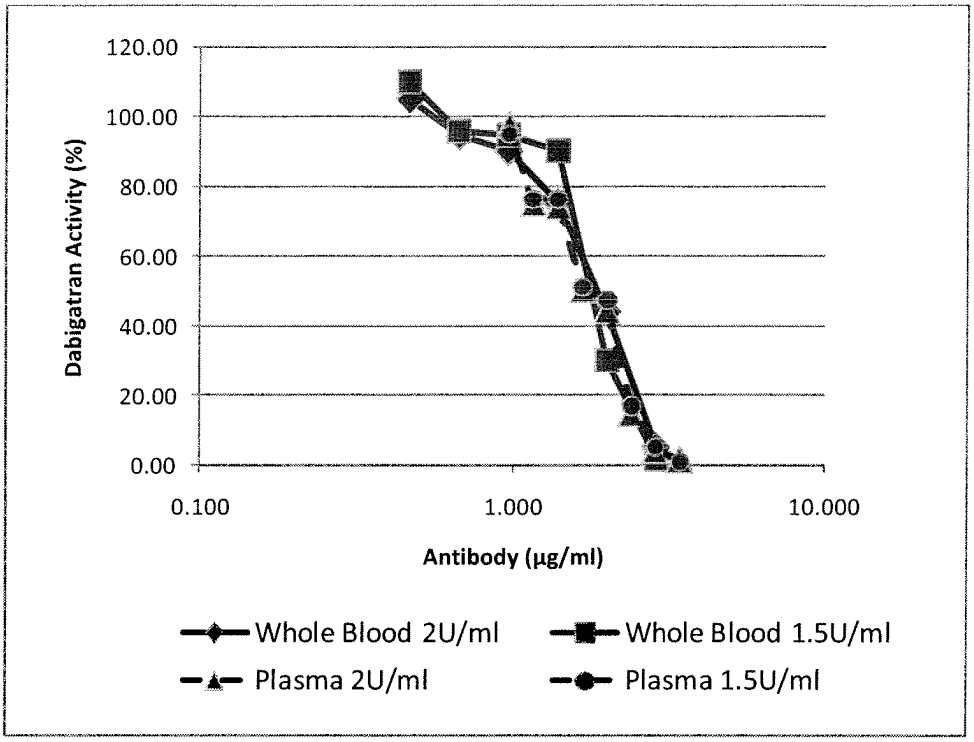

120.00
100.00
80.00
60.00
40.00
20.00
0.00
Dabigatran Activity (%)
0.100
1.000
10.000
Antibody (µg/ml)
Whole Blood 2U/ml
Whole Blood 1.5U/ml
Plasma 2U/ml
Plasma 1.5U/ml

FIGURE 7

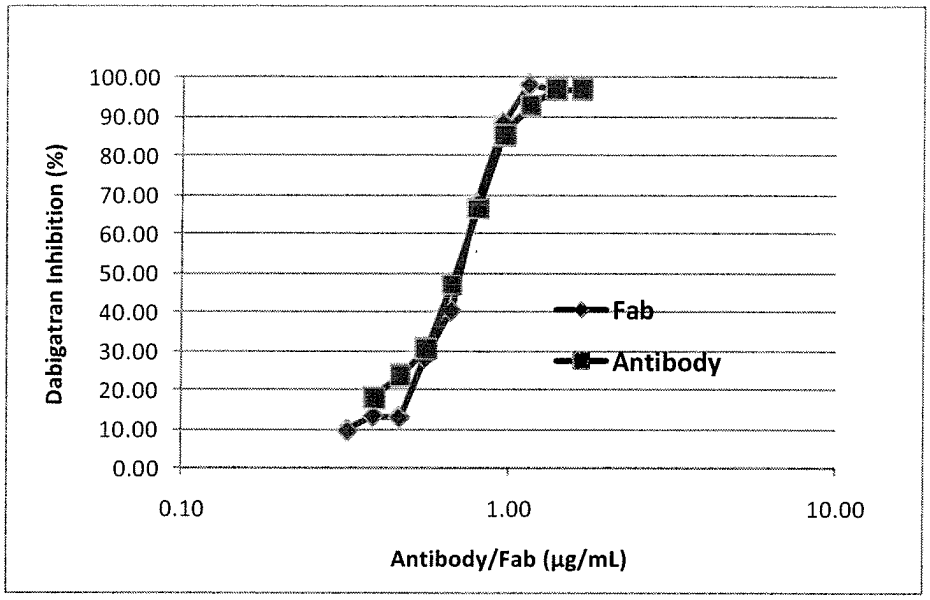

100.00
90.00
80.00
70.00
60.00
50.00
40.00
30.00
20.00
10.00
0.00
Dabigatran Inhibition (%)
0.10
1.00
10.00
Antibody/Fab (µg/mL)
Fab
Antibody

FIGURE 8

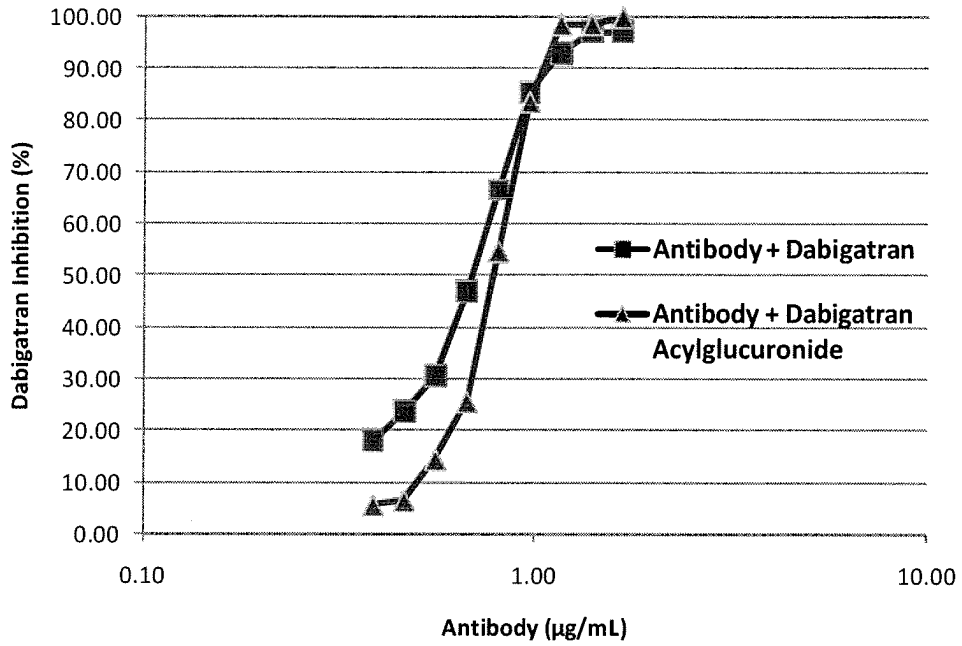

100.00
90.00
80.00
70.00
60.00
50.00
40.00
30.00
20.00
10.00
0.00
Dabigatran Inhibition (%)
0.10
1.00
10.00
Antibody (µg/mL)
Antibody + Dabigatran
Antibody + Dabigatran Acylglucuronide

FIGURE 9

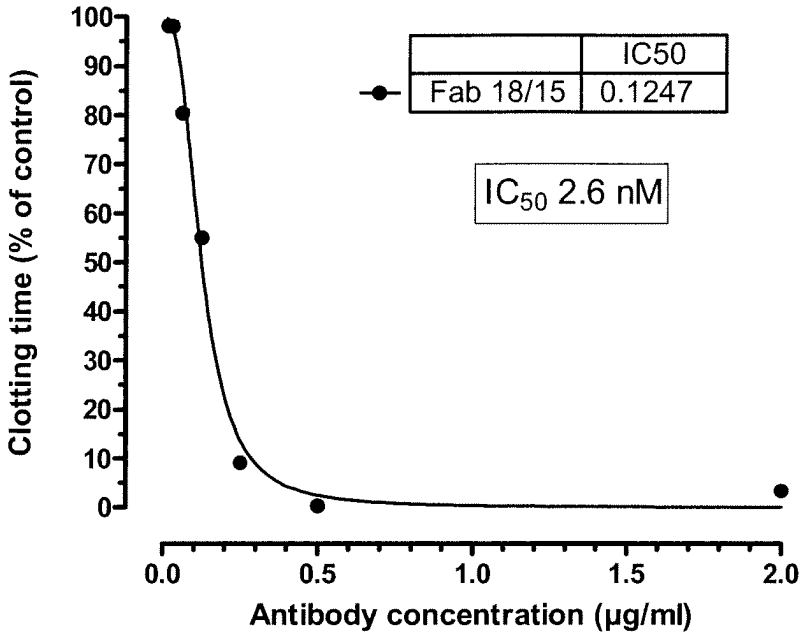
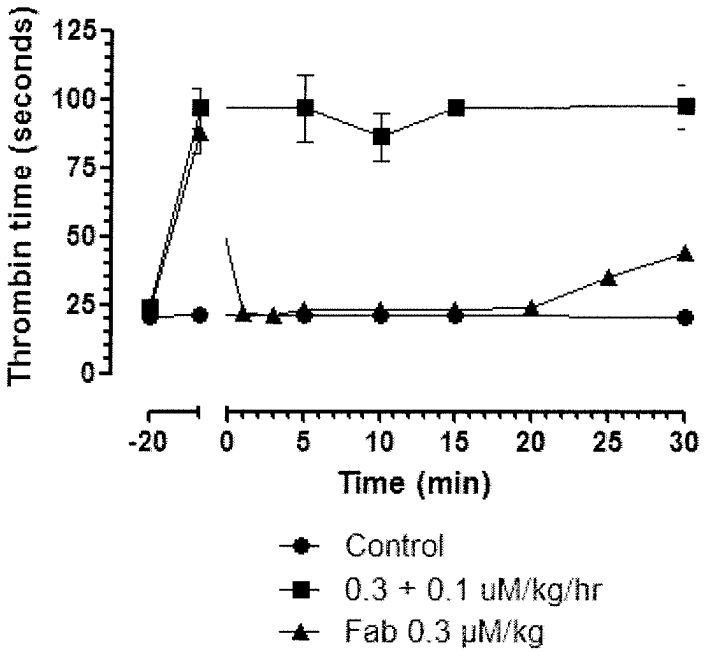
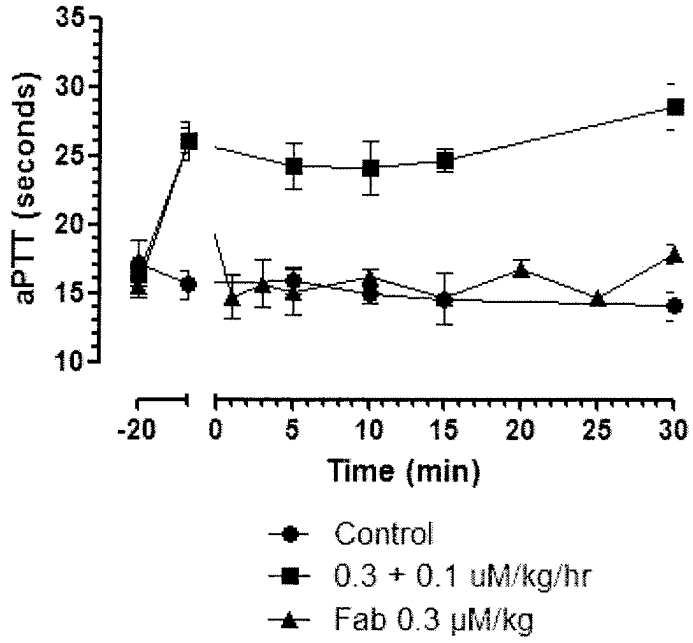

Clotting time (% of control)
100
90
80
70
60
50
40
30
20
10
0
0.0
0.5
1.0
1.5
2.0
Antibody concentration (µg/ml)
IC50
Fab 18/15
0.1247
IC50 2.6 nM

Figure 10



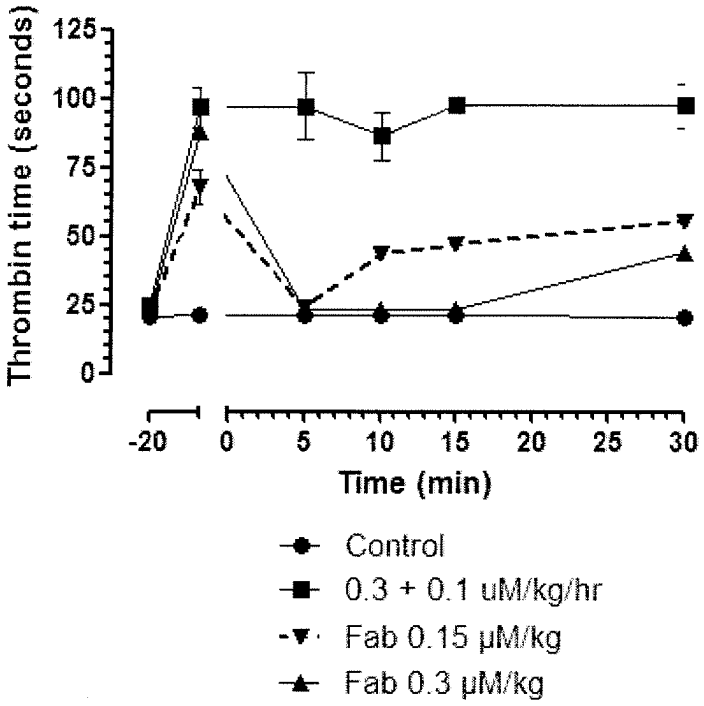
Dabigatran: bolus (0.3 µM/kg) + infusion (0.1 µM/kg/hr)
Thrombin time: 3.0 U/ml thrombin used in the assay
Data expressed as mean ± SE, n=4.

Figure 11



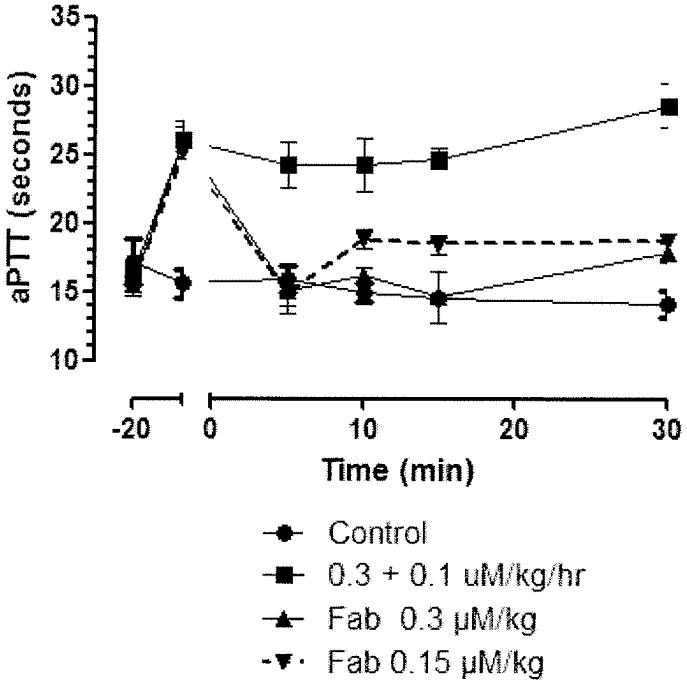
Dabigatran: Bolus (0.3 µM/kg) + infusion (0.1 µM/kg/hr)
Data expressed as mean ± SE, n=4.

Figure 12



Dabigatran bolus (0.3 µM/kg) + infusion (0.1 µM/kg/hr)
Thrombin time: 3.0 U/ml thrombin used in the assay
Data expressed as mean ± SE, n=4

Figure 13



Dabigatran bolus (0.3 µM/kg) + infusion (0.1 µM/kg/hr)
Data expressed as mean ± SE, n=4

Figure 14