Figure 1

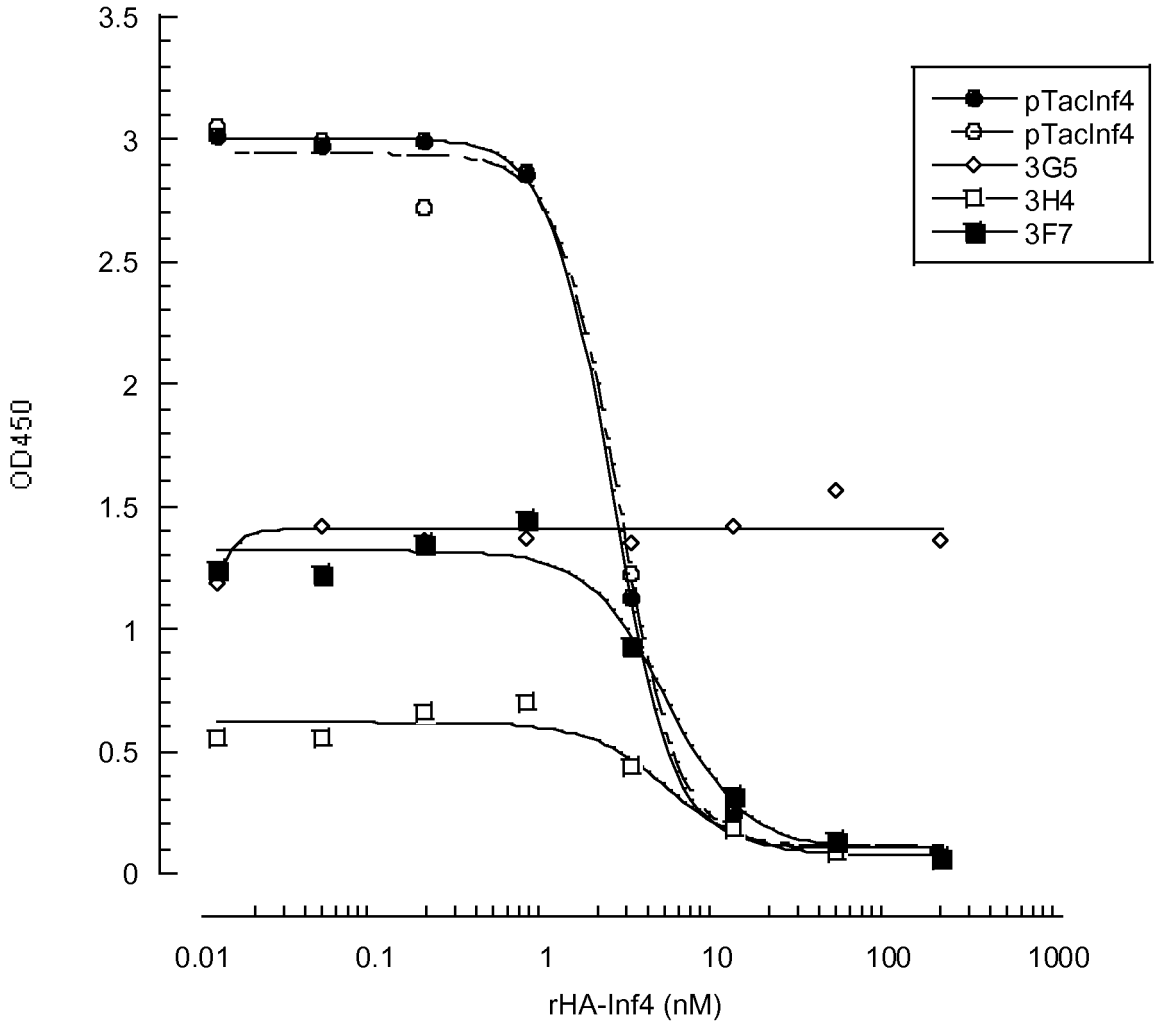

3.5
3
2.5
2
1.5
1
0.5
0
OD450
0.01
0.1
1
10
100
1000
rHA-Inf4 (nM)
pTacInf4
pTacInf4
3G5
3H4
3F7

Figure 2

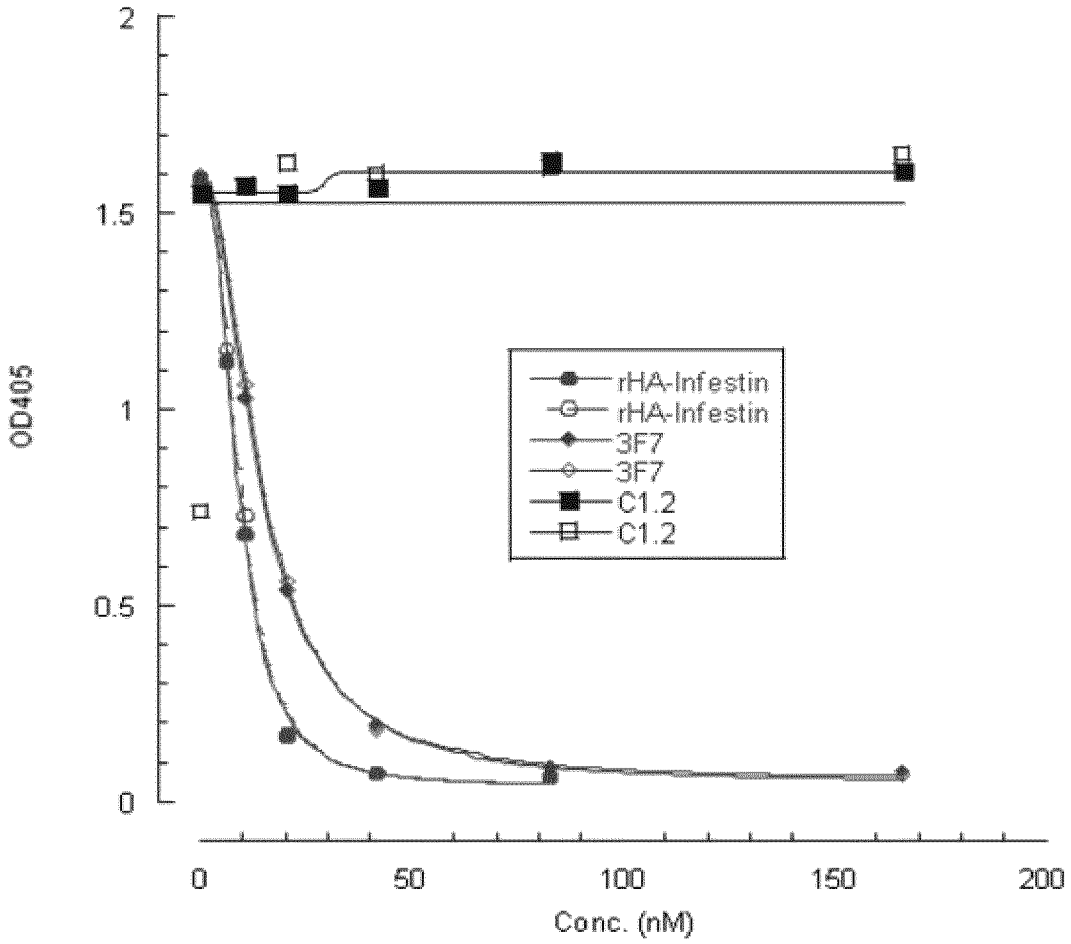

2
1.5
1
0.5
0
OD405
0
50
100
150
200
Conc. (nM)
rHA-Infestin
rHA-Infestin
3F7
3F7
C1.2
C1.2

Figure 3

3F7 H1

EVQLLESGGGLVQPGGSLRLSCAASGFTFxxxxxxWVRQAPGKGLEWVSGIRPSGGTTVYAD

SVKGRFTISRDNSKNTLYLQMNSLRAEDTAVYYCARALPRSGYLISPHYYYYALDVWGQGTT

VTVSSASTK

3F7 H2

EVQLLESGGGLVQPGGSLRLSCAASGFTFSKYIMQWVRQAPGKGLEWVSGIxxxxxxxTVYAD

SVKGRFTISRDNSKNTLYLQMNSLRAEDTAVYYCARALPRSGYLISPHYYYYALDVWGQGTT

VTVSSASTK

3F7 H3.1

EVQLLESGGGLVQPGGSLRLSCAASGFTFSKYIMQWVRQAPGKGLEWVSGIRPSGGTTVYAD

SVKGRFTISRDNSKNTLYLQMNSLRAEDTAVYYCARAxxxxxxxLISPHYYYYALDVWGQGTT

VTVSSASTK

3F7 H3.2

EVQLLESGGGLVQPGGSLRLSCAASGFTFSKYIMQWVRQAPGKGLEWVSGIRPSGGTTVYAD

SVKGRFTISRDNSKNTLYLQMNSLRAEDTAVYYCARALPRSGxxxxxxxYYYYALDVWGQGTT

VTVSSASTK

Figure 4

3F7 L1

QSVLTQPPSASGTPGQRVTISCSGSSxxxxxxYVYWYQQLPGTAPKLLIYSNNQRPSGVPDR

FSGSKSGTSASLAISGLRSEDEADYYCAAWDASLRGVFGGGTKLTVLGQPKAAPSVTL

3F7 L3.1

QSVLTQPPSASGTPGQRVTISCSGSSSNIGRNYVYWYQQLPGTAPKLLIYSNNQRPSGVPDR

FSGSKSGTSASLAISGLRSEDEADYYCxxxxxxLRGVFGGGTKLTVLGQPKAAPSVTL

3F7 L3.2

QSVLTQPPSASGTPGQRVTISCSGSSSNIGRNYVYWYQQLPGTAPKLLIYSNNQRPSGVPDR

FSGSKSGTSASLAISGLRSEDEADYYCAAWDxxxxxxFGGGTKLTVLGQPKAAPSVTL

Figure 5

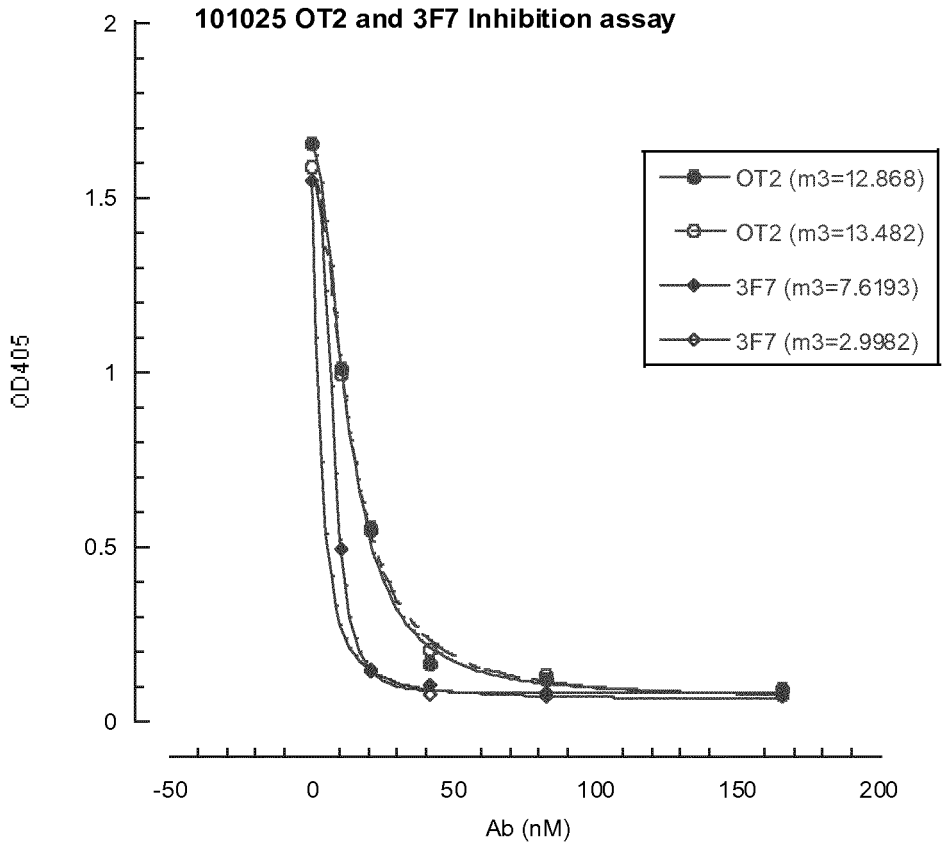

101025 OT2 and 3F7 Inhibition assay
OD405
2
1.5
1
0.5
0
-50
0
50
100
150
200
Ab (nM)
OT2 (m3=12.868)
OT2 (m3=13.482)
3F7 (m3=7.6193)
3F7 (m3=2.9982)

Figure 6

A

Human/mouse/rat FXIIa catalytic domain alignment

```
                                                    *
Mouse   VVGGLVALPGSHPYIAALYWGNNFCAGSLIAPCWVLTAAHCLQNRPAPEELTVVLGQDRH  414
Rat     VVGGLVALPGSHPYIAALYWGDSFCAGSLIDPCWVLTAAHCLQKRPAPEELTVVLGQDRH  414
Human   VVGGLVALRGAHPYIAALYWGHSFCAGSLIAPCWVLTAAHCLQDRPAPEDLTVVLGQERR  432
                           !        !            !

                                    *
Mouse   NQSCEWCQTLAVRSYRLHEGFSSITYQHDLALLRLQESKTNSCAILSPHVQPVCLPSGAA  474
Rat     NQSCERCQTLAVHSYRLHEGFSSKTYQHDLALLRLR-GRKNSCAILSPHVQPVCLPSSAA  473
Human   NHSCEPCQTLAVRSYRLHEAFSPVSYQHDLALLRLQEDADGSCALLSPYVQPVCLPSGAA  492
             !      !          !           !!!!!                !

Mouse   PPSETVLCEVAGWGHQFEGAEEYSTFLQEAQVPFIALDRCSNSNVHGDAILPGMLCAGFL  534
Rat     PPSETVLCEVAGWGHQFEGAEEYATFLQEAQVPFISLDRCSSSNVHGDAILPGMLCAGFL  533
Human   RPSETTLCQVAGWGHQFEGAEEYASFLQEAQVPFLSLERCSAPDVHGSSILPGMLCAGFL  552
                                                  !

                  *
Mouse   EGGTDACQGDSGGPLVCEEGTAEHQLTLRGVISWGSGCGDRNKPGVYTDVANYLAWIQKH  594
Rat     EGGADACQGDSGGPLVCDEGVTERQLTLRGVISWGSGCGDRNKPGVYTDVANYLDWIQEH  593
Human   EGGTDACQGDSGGPLVCEDQAAERRLTLQGIISWGSGCGDRNKPGVYTDVAYYLAWIREH  612
           !             !  !!                                  !

Mouse   IAS  597
Rat     TAF  596
Human   TVS  615
```

Figure 6

B

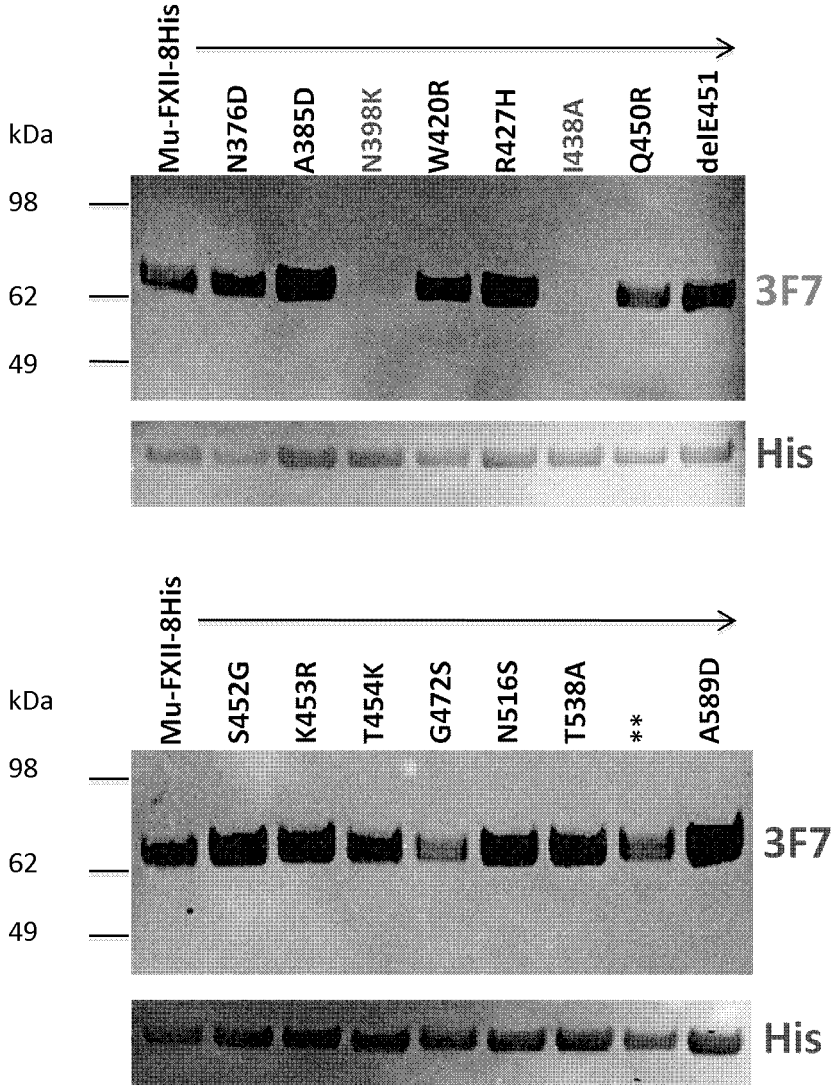

Mu-FXII-8His
N376D
A385D
N398K
W420R
R427H
I438A
Q450R
delE451
kDa
98
62
49
3F7
His
Mu-FXII-8His
S452G
K453R
T454K
G472S
N516S
T538A
**
A589D
kDa
98
62
49
3F7
His

Figure 7

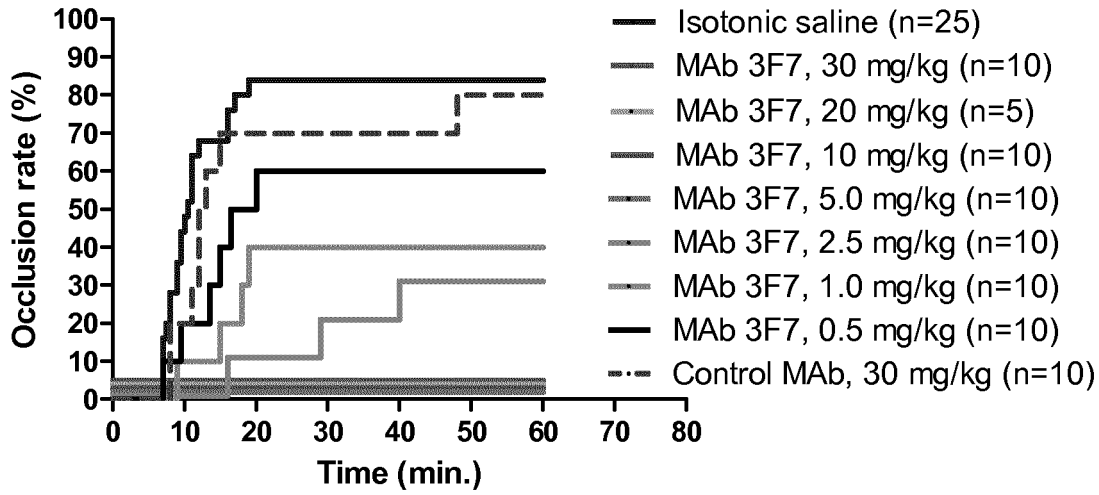


Figure 8

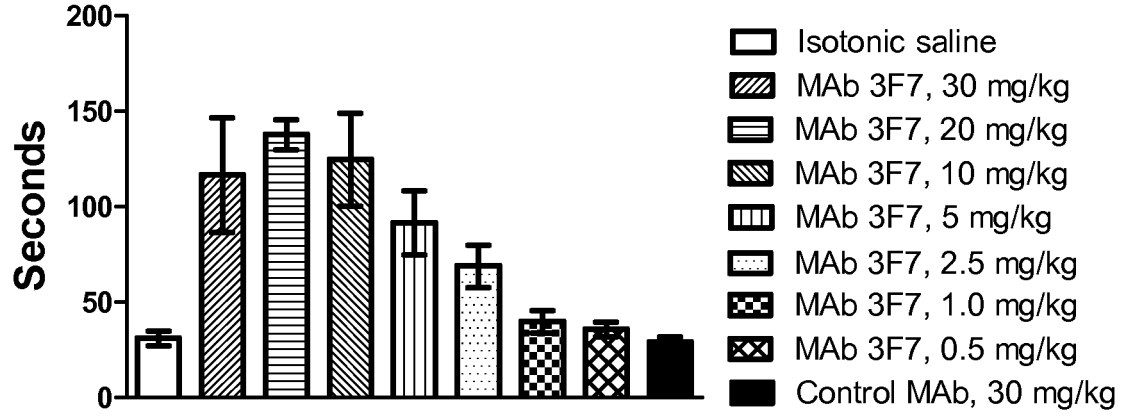


Figure 9

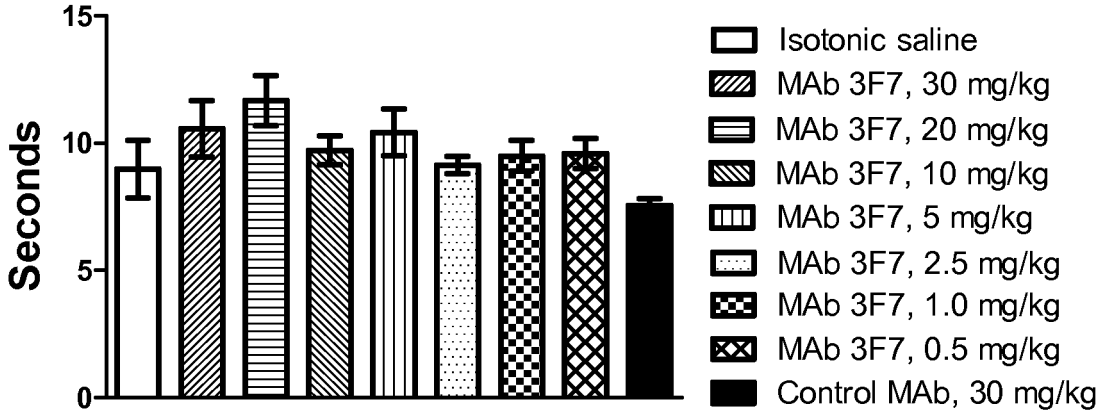


Figure 10

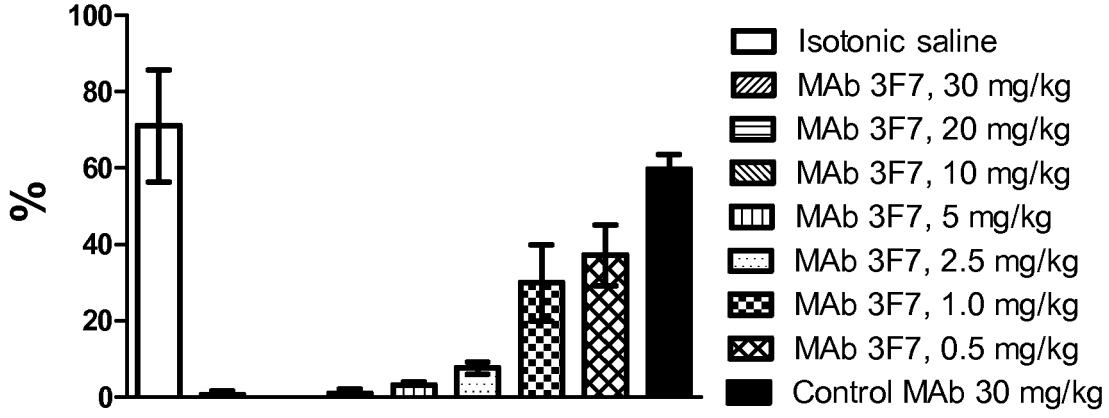


Figure 11

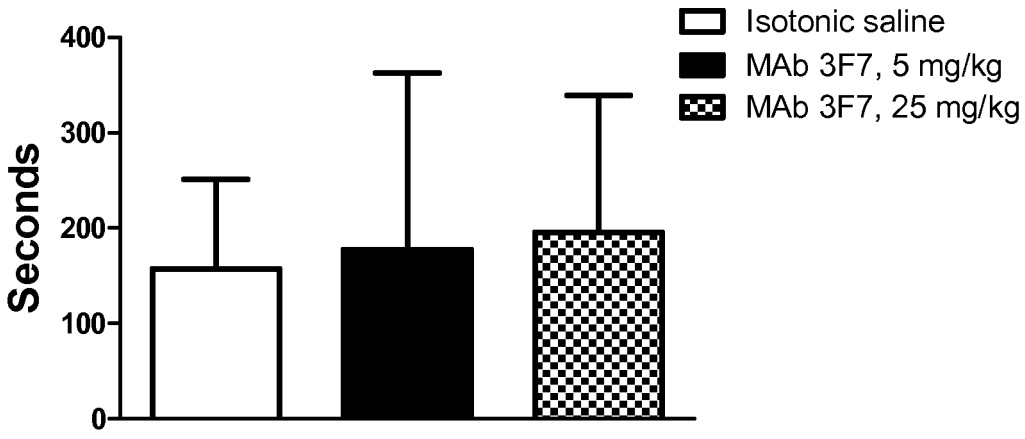


Figure 12

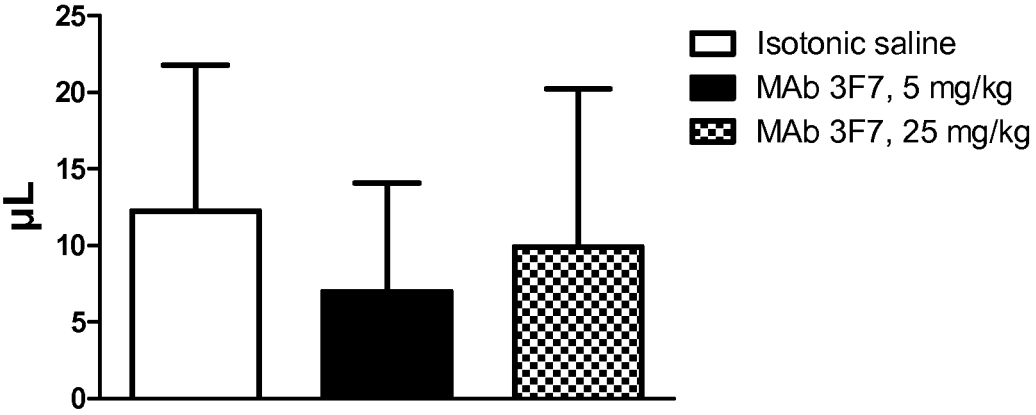


Figure 13

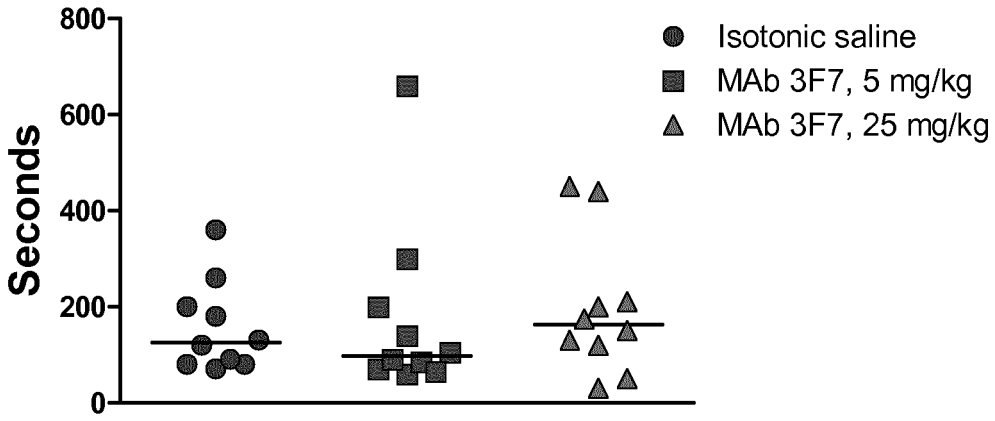


Figure 14

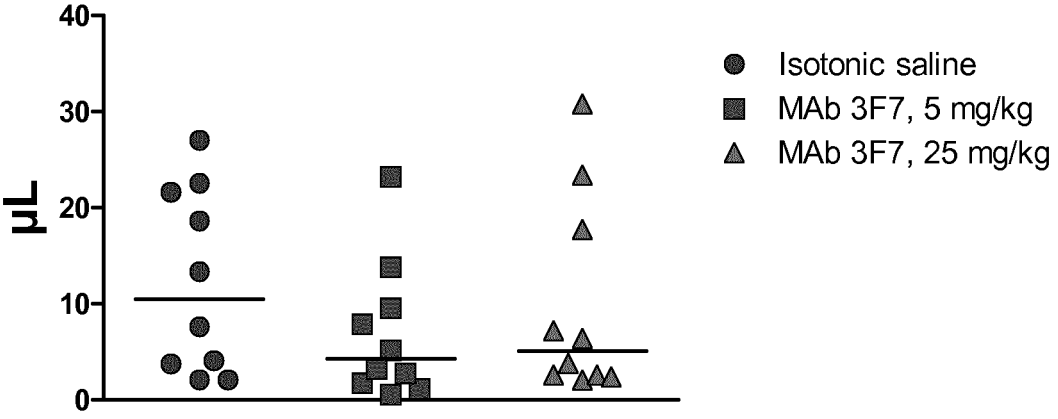


Figure 15

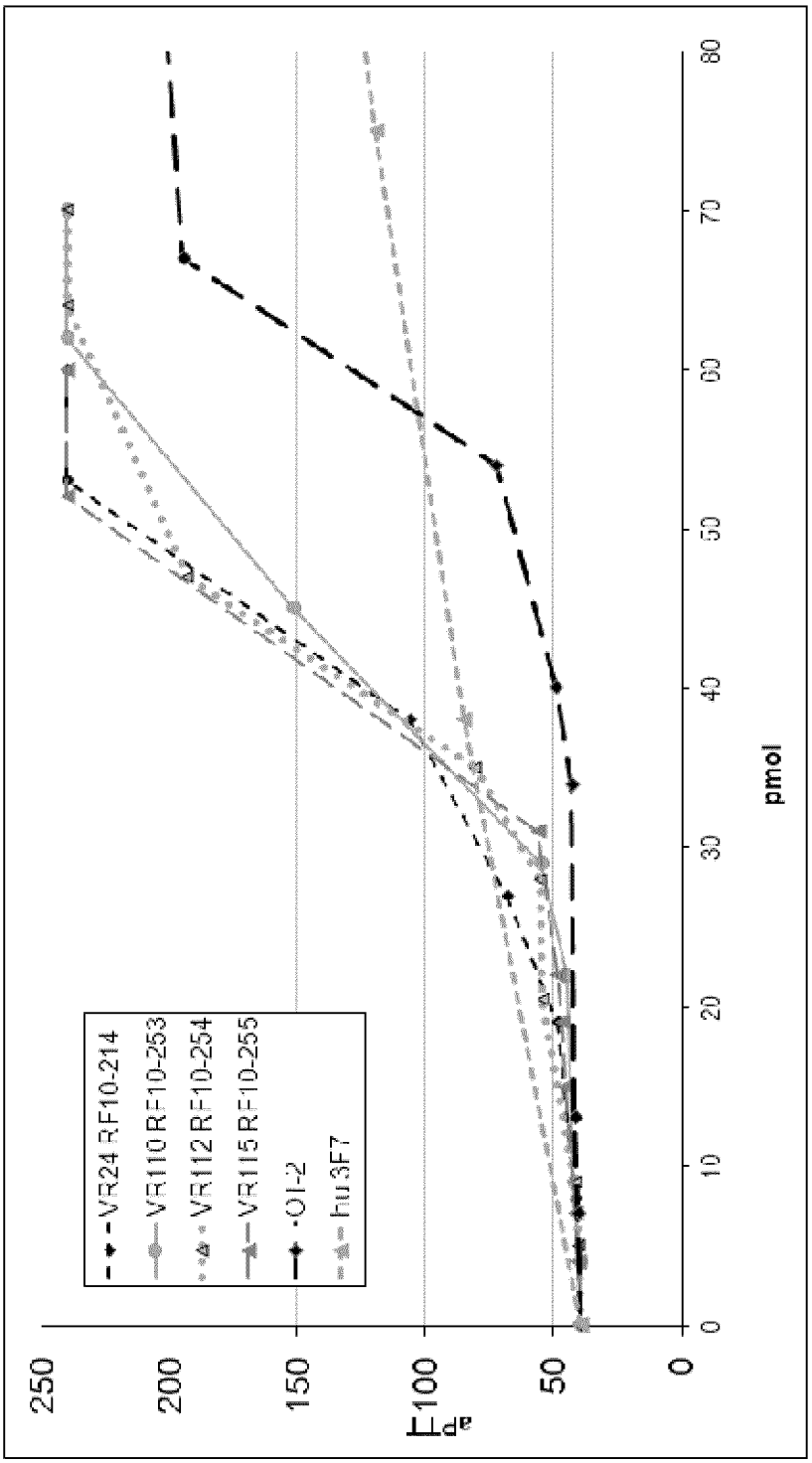
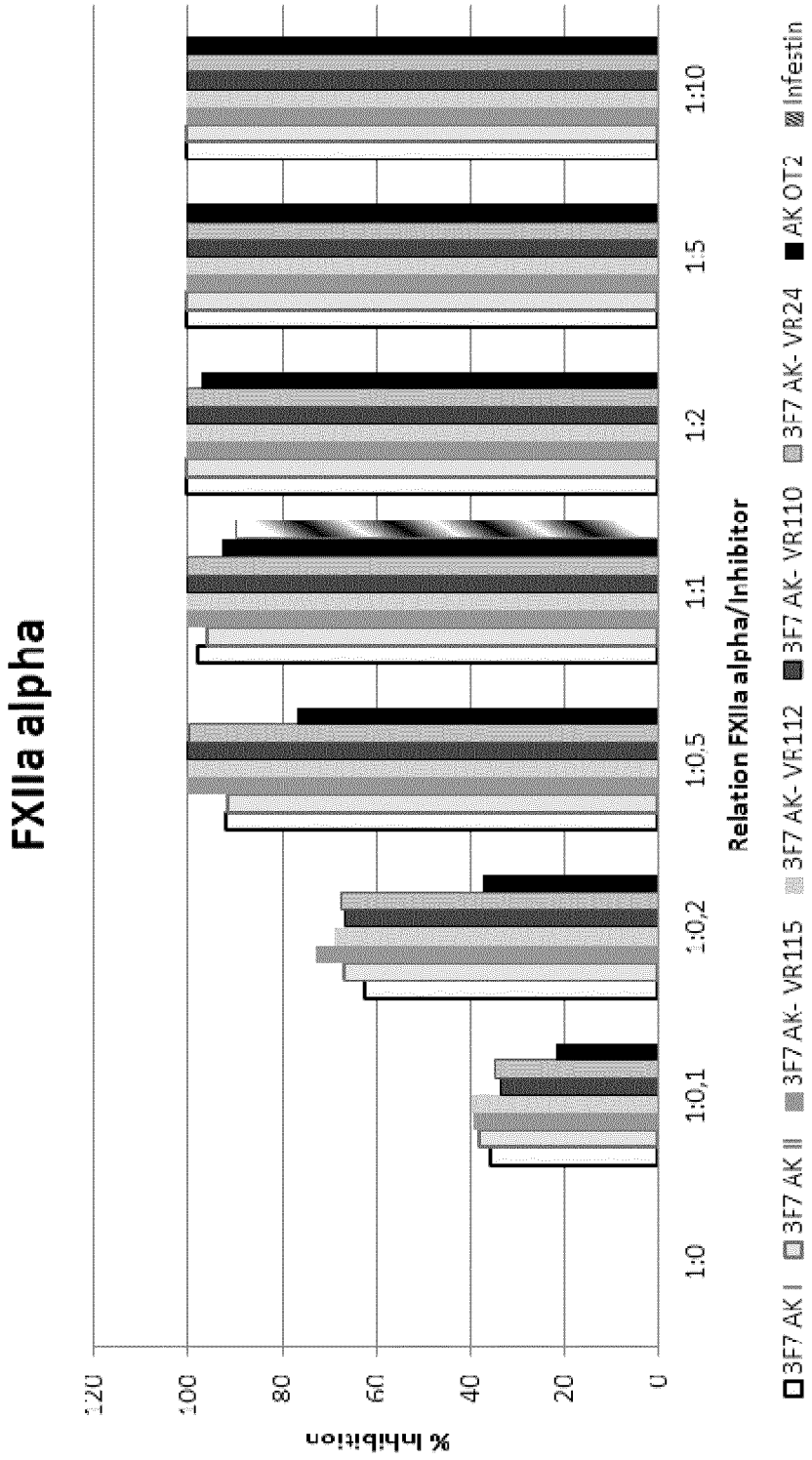

VR24 RF10-214
VR110 RF10-253
VR112 RF10-254
VR115 RF10-255
OI-2
hu3F7
250
200
150
100
50
0
aPTT
0
10
20
30
40
50
60
70
80
pmol

Figure 16


FXIIa alpha
% Inhibition
120
100
80
60
40
20
0
1:0
1:0,1
1:0,2
1:0,5
1:1
1:2
1:5
1:10
Relation FXIIa alpha/Inhibitor
3F7 AK I
3F7 AK II
3F7 AK- VR115
3F7 AK- VR112
3F7 AK- VR110
3F7 AK- VR24
AK OT2
Infestin