

## The unique procoagulant adaptations of pseudonaja textilis venom factor V and factor X Schreuder, M.

## Citation

Schreuder, M. (2022, September 22). *The unique procoagulant adaptations of pseudonaja textilis venom factor V and factor X.* Retrieved from https://hdl.handle.net/1887/3464432

Version:	Publisher's Version
License:	Licence agreement concerning inclusion of doctoral thesis in the Institutional Repository of the University of Leiden
Downloaded from:	https://hdl.handle.net/1887/3464432

**Note:** To cite this publication please use the final published version (if applicable).

Stellingen behorende bij het proefschrift

## The Unique Procoagulant Adaptations of Pseudonaja Textilis Venom Factor V and Factor X

- 1. The dissociation of the factor Va A2 domain following proteolysis by activated protein C is a consequence of the loss of interaction between the A1 and A2 domains this thesis
- 2. The factor Va A2 domain C-terminus contributes to anionic lipid surface-dependent conversion of prothrombin by factor Xa. In Pseudonaja textilis venom factor Va this region has been adapted to facilitate lipid-independent prothrombin conversion by Pseudonaja textilis venom factor Xa – this thesis
- 3. The activation peptide of Pseudonaja textilis venom factor X is modified to shift the conformational equilibrium of factor X towards a more active state, resulting in significant catalytic activity while in zymogen form – this thesis
- 4. A single amino acid substitution can change a protein's characteristics and could provide valuable opportunities for the development of novel therapeutic agents – this thesis
- 5. Proteins do not adopt a single functional conformation but exhibit conformational flexibility resulting in multiple protein states with different characteristics Toso et al., J Biol Chem 2008; Butera et al., Nat Commun, 2020
- 6. With safety concerns remaining regarding the use of andexanet alfa as antidote for direct factor Xa-related major bleeding, factor X-C and F174-substituted factor X variants may potentially provide an alternative specific reversal strategy Ersayin et al., Haematologica, 2017; Connolly et al., N Engl J Med 2019
- 7. Venoms are a valuable source of biological information with significant clinical potential Waheed et al., Curr Med Chem, 2017
- 8. Used throughout several decades of research, biochemistry remains a compelling tool for fundamental and translational research Muczynski et al., Semin Thromb Hemost, 2017
- 9. The workload among academic researchers and staff continues to rise resulting in structural overtime and a very high work pressure. This not only affects the education of students but also the future of academic research with more and more talented researchers choosing a career elsewhere Werkdruk Universiteiten, FNV and VAWO 2019