

# Regulation of thrombin formation via the protein C pathway in normal and hypercoagulable states

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**Stellingen behorende bij het proefschrift**

**Regulation of Thrombin Formation via the  
Protein C Pathway in  
Normal and Hypercoagulable states**

**Gerry A.F. Nicolaes**

1. It is advisable to perform the functional assessment of APC resistance in the presence of concentrations of protein S and factor Xa that are as low as possible.

Chapter 4, this thesis

2. The observation that coumarin therapy induces skin necrosis in patients with protein S deficiency need not be explained by a similar theory as given for identical observations in protein C deficient patients (i.e. a relatively rapid decrease of coagulation factor concentration, while other vitamin-K dependent factors are relatively still high), but rather be explained by the decrease in protein C concentrations themselves.

Allaart CF and Briët E. Familial Thrombophilia, Chapter 61 In: Bloom AL, Forbes CD, Thomas DP, Tuddenham EGD, eds. Haemostasis and Thrombosis, 3rd ed. Edinburgh: Churchill Livingstone, 1994; 1349-1360

3. The statement that the gaspressure in gascells of doughs, made of 'poor-baking' flour, becomes/is too high, resulting in diffusion of CO<sub>2</sub> to the 'outside', is not correct.

He H en Hosney RC. Factors controlling gas retention in nonheated doughs. Cereal Chemistry 1992; 69: 1-6

4. The suggestion as made by Martin and coworkers that appearance of PS on the external leaflet of the plasma membrane of apoptotic cells is caused by the activation of a specific inside-out PS translocase is not experimentally demonstrated and in full contrast to data published in the same issue of the same journal by Verhoven et al.

Martin SJ, Reutelingsperger CPM, Mc Gahon AJ, Rader JA, van Schie RCAA, LaFace DM and Green DR. Early redistribution of plasma membrane phosphatidylserine is a general feature of apoptosis regardless of the initiating stimulus: inhibition by overexpression of Bcl-2 and Abl J. Exp. Med. 1995; 182: 1543-1556

Verhoven B, Schlegel RA and Williamson P. Mechanisms of phosphatidylserine exposure, a phagocyte recognition signal, on apoptotic T lymphocytes. Abl J. Exp. Med. 1995; 182: 1597-1601

5. Changes in the signal-transduction characteristics of developing megakaryoblasts occur as a result of up- and downregulation of specific G-proteins.

Thesis H. van der Vuurst, 1997, Universiteit Utrecht

6. The effect of playing soft music for one hour, as done by Blombäck et al. in order to allow blood donors to rest and relax before blood donation, is likely to be subject to inter-individual variation which may in some cases have the opposite effect of what was intended.

Blombäck M, Landgren B-M, Stiernholm Y and Andersson O. The effect of progesterone on the haemostatic mechanism. *Thrombosis and Haemostasis* 1997; 77(1): 105-108

7. The hypothesis, stated in several scientific papers, that changes in both the procoagulation factors and profibrinolysis factors balance each other in users of oral contraceptives, so as to maintain a haemostatic balance, lacks an experimental foundation and is therefore false.

Stubblefield PG. The effects on hemostasis of oral contraceptives containing desogestrel. *Am. J. Obstet. Gynecol.* 1993 3(2): 1047-1052

Coata G, Ventura F, Lombardini R, Ciufetti G, Cosmi EM and Di Renzo GC. Effect of low-dose oral triphasic contraceptives on blood viscosity, coagulation and lipid metabolism. *Contraception* 1995; 52: 151-157

8. In contrast to what is reported by Østerud et al., their experiments show that resistance to activated protein C is increased, and not reduced, in women using oral contraceptives.

Østerud B, Robertson R, Åsvang GB, Thijssen F. Resistance to activated protein C is reduced in women using oral contraceptives. *Blood Coagulation and Fibrinolysis* 1994; 5: 853-854

9. Een stelling op de openbare weg dient duidelijk gemarkeerd te worden.

10. De spreuk 'Oost-West, thuis best' zal mogelijk in menig Limburgs huishouden, afhankelijk van de geografische ligging, een relatief begrip worden.

11. Het feit dat sommige harmonie-orkesten zich ook wel "Wind Ensembles" wensen te noemen, pleit niet voor het door deze orkesten voortgebrachte geluid.