

Stratum corneum model membranes : molecular organization in relation to skin barrier function Groen, D.

Citation

Groen, D. (2011, October 25). *Stratum corneum model membranes : molecular organization in relation to skin barrier function*. Retrieved from https://hdl.handle.net/1887/17978

Version:	Corrected Publisher's Version
License:	<u>Licence agreement concerning inclusion of doctoral</u> <u>thesis in the Institutional Repository of the University</u> <u>of Leiden</u>
Downloaded from:	https://hdl.handle.net/1887/17978

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

Stratum corneum model membranes: molecular organization in relation to skin barrier function

The investigations described in this thesis were performed at the division of Drug Delivery Technology of the Leiden/Amsterdam Center for Drug Research (LACDR), Leiden University, Leiden, The Netherlands. The studies described in this thesis were financially supported by a grant from the Technology Foundation (STW, project LGP 7503). The printing of this thesis was financially supported by STW, LACDR and the Leiden University library.

Cover image: Gothic bell tower of the church of St-André in Grenoble, France (1298).

Printed by Wöhrmann Print Service, Zutphen, The Netherlands

Stratum corneum model membranes: molecular organization in relation to skin barrier function

Proefschrift

ter verkrijging van de graad van Doctor aan de Universiteit Leiden, op gezag van Rector Magnificus prof. mr. P.F. van der Heijden, volgens besluit van het College voor Promoties te verdedigen op dinsdag 25 oktober 2011 klokke 15:00

door

Daniël Groen geboren te Den Haag in 1979

Promotiecommissie

Promotor:

Prof. Dr. J.A. Bouwstra

Overige leden:

Prof. Dr. M. DanhofProf. Dr. W. JiskootProf. Dr. W.J. BrielsProf. Dr. M.J. LawrenceProf. Dr. J.P. Abrahams

Stellingen

Behorende bij het proefschrift

Stratum corneum model membranes: molecular organization in relation to skin barrier function

- 1. The barrier function of the skin is fundamental to life on dry land. (D. Attenborough, Life on Earth, Little, Brown and Company, Boston, 1980)
- 2. Understanding the permeability barrier function of the skin is important for rational design of transdermal drug delivery systems as well as for our understanding the etiology, and possible treatment, of a range of skin diseases in which barrier function is compromised. (*J.R. Hill and P.W. Wertz, BBA 1616, 2003, p121-126*)
- 3. The stratum corneum substitute, prepared with synthetic lipids only, closely mimics the stratum corneum lipid organization and can replace human stratum corneum in permeability studies. (*This thesis*)
- 4. Because of its adjustable lipid composition, the stratum corneum substitute is a suitable model to mimic the lipid organization in stratum corneum of diseased skin. (*This thesis*)
- 5. Whether an orthorhombic or hexagonal lipid packing is present in stratum corneum, is of less importance for a proper barrier function than the presence of a proper lamellar organization. *(This thesis)*
- 6. The presence of water is not required to form a proper lipid organization in a lipid model, mimicking that in human stratum corneum.
- 7. Many papers in the scientific literature on percutaneous absorption represent a complex blend of physicochemical theory and physiological practicalities. (Brian W. Barry, Dermatological Formulations: Percutaneous Absorption, Marcel Dekker inc, New York, 1983)
- 8. If you wish to make an apple pie from scratch, you must first invent the universe. (*Garl Sagan, Cosmos, 1980, p. 218.*)
- 9. The propositions in a thesis are a rudimentary organ that lost its function many years ago.

Table of contents

Chapter 1	Introduction, aim and outline of this thesis	1
Part I		
Chapter 2	Two new methods for preparing a unique stratum corneum substitute	29
Chapter 3	Is an orthorhombic lateral packing and a proper lamellar organization important for the skin barrier function?	61
Chapter 4	Investigating the barrier function of skin lipid models with varying compositions	91
Part II		
Chapter 5	Model membranes prepared with ceramide EOS, cholesterol and free fatty acids form a very unique lamellar phase	121
Chapter 6	New insights into the stratum corneum lipid organization by x-ray diffraction analysis	151
Chapter 7	Disposition of ceramide in model lipid membranes determined by neutron diffraction	175
Chapter 8	Summary and perspectives	205