surfactant science series - volume 62

VESICLES



edited by Morton Rosoff

VESICLES

edited by Morton Rosoff

Professor Emeritus Long Island University Brooklyn, New York

Library of Congress Cataloging-in-Publication Data

Vesicles / edited by Morton Rosoff.

p. cm. — (Surfactant science series; v. 62)

Includes index.

ISBN 0-8247-9603-9 (hardcover: alk. paper)

1. Liposomes. 2. Surface chemistry. I. Rosoff, Morton. II. Series

RS201.L55V47 1996

574.87'4—dc20

96-15201

CIP

The publisher offers discounts on this book when ordered in bulk quantities. For more information, write to Special Sales/Professional Marketing at the address below.

This book is printed on acid-free paper.

Copyright © 1996 by Marcel Dekker, Inc. All Rights Reserved.

Neither this book nor any part may be reproduced or transmitted in any form or by any means, electronic or mechanical, including photocopying, microfilming, and recording, or by any information storage and retrieval system, without permission in writing from the publisher.

Marcel Dekker, Inc.

270 Madison Avenue, New York, New York 10016

Current printing (last digit):

10987654321

PRINTED IN THE UNITED STATES OF AMERICA

SURFACTANT SCIENCE SERIES

CONSULTING EDITORS

MARTIN J. SCHICK

FREDERICK M. FOWKES

Consultant

(1915–1990)

New York, New York

- 1. Nonionic Surfactants, *edited by Martin J. Schick* (see also Volumes 19, 23, and 60)
- Solvent Properties of Surfactant Solutions, edited by Kozo Shinoda (see Volume 55)
- 3. Surfactant Biodegradation, R. D. Swisher (see Volume 18)
- Cationic Surfactants, edited by Eric Jungermann (see also Volumes 34, 37, and 53)
- Detergency: Theory and Test Methods (in three parts), edited by W. G. Cutler and R. C. Davis (see also Volume 20)
- Emulsions and Emulsion Technology (in three parts), edited by Kenneth J. Lissant
- Anionic Surfactants (in two parts), edited by Wamer M. Linfield (see Volume 56)
- 8. Anionic Surfactants: Chemical Analysis, edited by John Cross (out of print)
- Stabilization of Colloidal Dispersions by Polymer Adsorption, Tatsuo Sato and Richard Ruch (out of print)
- Anionic Surfactants: Biochemistry, Toxicology, Dermatology, edited by Christian Gloxhuber (see Volume 43)
- 11. Anionic Surfactants: Physical Chemistry of Surfactant Action, edited by E. H. Lucassen-Reynders (out of print)
- 12. Amphoteric Surfactants, edited by B. R. Bluestein and Clifford L. Hilton (see Volume 59)
- 13. Demulsification: Industrial Applications, Kenneth J. Lissant (out of print)
- 14. Surfactants in Textile Processing, Arved Datyner
- 15. Electrical Phenomena at Interfaces: Fundamentals, Measurements, and Applications, edited by Ayao Kitahara and Akira Watanabe
- 16. Surfactants in Cosmetics, edited by Martin M. Rieger (out of print)
- Interfacial Phenomena: Equilibrium and Dynamic Effects, Clarence A. Miller and P. Neogi
- 18. Surfactant Biodegradation: Second Edition, Revised and Expanded, R. D. Swisher
- 19. Nonionic Surfactants: Chemical Analysis, edited by John Cross
- Detergency: Theory and Technology, edited by W. Gale Cutler and Erik Kissa

- 21. Interfacial Phenomena in Apolar Media, edited by Hans-Friedrich Eicke and Geoffrev D. Parfitt
- 22. Surfactant Solutions: New Methods of Investigation, edited by Raoul Zana
- 23. Nonionic Surfactants: Physical Chemistry, edited by Martin J. Schick
- 24. Microemulsion Systems, edited by Henri L. Rosano and Marc Clausse
- 25. Biosurfactants and Biotechnology, edited by Naim Kosaric, W. L. Caims, and Neil C. C. Gray
- 26. Surfactants in Emerging Technologies, edited by Milton J. Rosen
- 27. Reagents in Mineral Technology, edited by P. Somasundaran and Brij M. Moudgil
- 28. Surfactants in Chemical/Process Engineering, edited by Darsh T. Wasan, Martin E. Ginn, and Dinesh O. Shah
- 29. Thin Liquid Films, edited by I. B. Ivanov
- 30. Microemulsions and Related Systems: Formulation, Solvency, and Physical Properties, edited by Maurice Bourrel and Robert S. Schechter
- 31. Crystallization and Polymorphism of Fats and Fatty Acids, edited by Nissim Garti and Kiyotaka Sato
- 32. Interfacial Phenomena in Coal Technology, edited by Gregory D. Botsaris and Yuli M. Glazman
- 33. Surfactant-Based Separation Processes, edited by John F. Scamehom and Jeffrey H. Harwell
- 34. Cationic Surfactants: Organic Chemistry, edited by James M. Richmond
- Alkylene Oxides and Their Polymers, F. E. Bailey, Jr., and Joseph V. Koleske
- 36. Interfacial Phenomena in Petroleum Recovery, edited by Norman R. Morrow
- 37. Cationic Surfactants: Physical Chemistry, edited by Donn N. Rubingh and Paul M. Holland
- 38. Kinetics and Catalysis in Microheterogeneous Systems, edited by M. Grätzel and K. Kalyanasundaram
- 39. Interfacial Phenomena in Biological Systems, edited by Max Bender
- 40. Analysis of Surfactants, Thomas M. Schmitt
- 41. Light Scattering by Liquid Surfaces and Complementary Techniques, edited by Dominique Langevin
- 42. Polymeric Surfactants, Irja Piirma
- 43. Anionic Surfactants: Biochemistry, Toxicology, Dermatology. Second Edition, Revised and Expanded, edited by Christian Gloxhuber and Klaus Künstler
- 44. Organized Solutions: Surfactants in Science and Technology, edited by Stig E. Friberg and Björn Lindman
- 45. Defoaming: Theory and Industrial Applications, edited by P. R. Garrett
- 46. Mixed Surfactant Systems, edited by Keizo Ogino and Masahiko Abe
- 47. Coagulation and Flocculation: Theory and Applications, edited by Bohuslav Dobiáš
- 48. Biosurfactants: Production Properties Applications, edited by Naim Kosaric
- 49. Wettability, edited by John C. Berg
- 50. Fluorinated Surfactants: Synthesis Properties Applications, Erik Kissa
- 51. Surface and Colloid Chemistry in Advanced Ceramics Processing, edited by Robert J. Pugh and Lennart Bergström

- 52. Technological Applications of Dispersions, edited by Robert B. McKay
- 53. Cationic Surfactants: Analytical and Biological Evaluation, edited by John Cross and Edward J. Singer
- 54. Surfactants in Agrochemicals, Tharwat F. Tadros
- 55. Solubilization in Surfactant Aggregates, edited by Sherril D. Christian and John F. Scamehom
- 56. Anionic Surfactants: Organic Chemistry, edited by Helmut W. Stache
- 57. Foams: Theory, Measurements, and Applications, edited by Robert K. Prudhomme and Saad A. Khan
- 58. The Preparation of Dispersions in Liquids, H. N. Stein
- 59. Amphoteric Surfactants: Second Edition, edited by Eric G. Lomax
- 60. Nonionic Surfactants: Polyoxyalkylene Block Copolymers, edited by Vaughn M. Nace
- 61. Emulsions and Emulsion Stability, edited by Johan Sjöblom
- 62. Vesicles, edited by Morton Rosoff

ADDITIONAL VOLUMES IN PREPARATION

Applied Surface Thermodynamics, edited by A. W. Neumann and Jan K. Spelt

Liquid Detergents, edited by Kuo-Yann Lai

Detergents in the Environment, edited by M. J. Schwuger

Preface

The Pythagorean view of the formation of the universe from the commingling of the elements—earth, air, fire, and water—notably omits oil. Oil and water are the Yin and Yang—in molecular terms, the spatial separation of dichotomous groups or segments, hydrophilic and oleophilic—underlying the formation of multitudes of organized assemblies or microphases.

Monolayers, multilayers, micelles, microemulsions, bilayers, and vesicles are an important set of interrelated states of matter. In particular, two-dimensional closed membranes or liposomes, consisting of a bilayer(s) of phospholipid molecules surrounding an aqueous core(s), form a reservoir for encapsulation and a diffusion barrier. Vesicles exhibit many basic and applied phenomena, serve as a model for the fundamental unit of biological structure—the cell membrane—and are uniquely suited to bridge the biological and materials sciences. From an academic viewpoint, they are represented at the growing tips of such active topics as self-assembly, self-organization, and complexity.

Reviews can serve a variety of functions, not the least of which is providing literature references sufficient to outline the development of a subject as well as its most recent results. For newcomers who need to learn more about a field, reviews fulfill a high form of education not available in formal courses. For the experienced researcher there is the stimulation of critical surveys and fresh ideas. Finally, for the authors, expository writing about a research area is a genuine form of scientific activity (recall that Mendeleev discovered the periodic table while planning an elementary chemistry textbook).

With the widening applications of vesicles and investigations to characterize the systems themselves, a multiauthored comprehensive review would be encyclopedic. Rather, this volume presents selected topics as signposts indicating the iv Preface

scope and direction of vesicle research. According to the "80/20" rule of the Pareto principle, named after the nineteenth-century Italian economist Vilfredo Pareto, 80% of the value of a group of items is generally concentrated in only 20% of the items. Hence, the average reader will probably be interested in only 20% of the topics. Nevertheless, it is hoped that a synthesis of ideas drawn from different (but related) subjects will create new and fruitful general conceptions. Often overlooked is the serendipitous breakthrough of a given idea when transplanted to another context. Undoubtedly, scattered among the diverse chapters lie solutions in search of problems.

The book is divided into four sections covering some main topics in vesicle research: physical and surface chemistry; methods; drug delivery; and diverse applications.

I thank the contributors for making this book possible and hope that it will be opened with expectation and closed with profit.

Morton Rosoff

Contributors

Francesco Castelli Department of Chemistry, University of Catania, Catania, Italy

Eric Claassen Division of Immunological and Infectious Diseases, TNO-PG, Leiden, and Department of Immunology, Erasmus University, Rotterdam, The Netherlands

Guy Duportail Laboratoire de Biophysique, Centre de Recherches Pharmaceutiques, Université Louis Pasteur, Illkirch, France

Gary Fujii Gene Delivery, NeXstar Pharmaceuticals, Inc., San Dimas, California

Oleg V. Gerasimov Department of Chemistry, Purdue University, West Lafayette, Indiana

Tibor Hianik Department of Biophysics and Chemical Physics, Comenius University, Bratislava, Slovak Republic

Tohru Inoue Department of Chemistry, Fukuoka University, Fukuoka, Japan

Shigeyuki Komura Department of Mechanical System Engineering, Kyushu Institute of Technology, Iizuka, Japan

viii Contributors

Hironobu Kunieda Department of Physical Chemistry, Yokohama National University, Yokohama, Japan

Danilo D. Lasic MegaBios Corporation, Burlingame, California

Panagiotis Lianos Engineering Science Department, University of Patras, Patras, Greece

Robert C. MacDonald Department of Biochemistry, Molecular Biology, and Cell Biology, Northwestern University, Evanston, Illinois

Rimona Margalit Department of Biochemistry, Tel Aviv University, Tel Aviv, Israel

Grant Meng Faculty of Pharmaceutical Sciences, University of British Columbia, Vancouver, British Columbia, Canada

David Needham Department of Mechanical Engineering and Materials Science and the Center for Cellular and Biosurface Engineering, Duke University, Durham, North Carolina

Angela Ottová-Leitmannová Department of Microelectronics, Slovak Technical University, Bratislava, Slovak Republic

Rodney Pearlman MegaBios Corporation, Burlingame, California

Vijay Rajagopalan Chemical Center, University of Lund, Lund, Sweden

Antonio Raudino Department of Chemistry, University of Catania, Catania, Italy

Yuanjin Rui Department of Chemistry, Purdue University, West Lafayette, Indiana

Masahiko Sisido Department of Bioengineering Science, Okayama University, Okayama, Japan

David H. Thompson Department of Chemistry, Purdue University, West Lafayette, Indiana

H. Ti Tien Department of Physiology, Michigan State University, East Lansing, Michigan

Contributors ix

Colin Tilcock Faculty of Pharmaceutical Sciences, University of British Columbia, Vancouver, British Columbia, Canada

Deepank Utkhede Faculty of Pharmaceutical Sciences, University of British Columbia, Vancouver, British Columbia, Canada

John H. van Zanten Department of Chemical Engineering, Johns Hopkins University, Baltimore, Maryland

Doncho V. Zhelev Department of Mechanical Engineering and Materials Science, Duke University, Durham, North Carolina

Contents

Preface iii Contributors vii

1 Physical and Surface Chemistry 1

- The Relationship and Interactions Between Lipid Bilayers Vesicles and Lipid Monolayers at the Air/Water Interface 3 Robert C. MacDonald
- Physical and Chemical Aspects of Liposomes and Some of Their Applications 49
 Tibor Hianik, Angela Ottová-Leitmannová, and H. Ti Tien
- 3. Formation and Structure of Reverse Vesicles 79 Hironobu Kunieda and Vijay Rajagopalan
- Thermodynamics and Kinetics of Inhomogeneous Distribution of Lipid Membrane Components 105 Antonio Raudino and Francesco Castelli
- 5. Interaction of Surfactants with Phospholipid Vesicles 151

 Tohru Inoue
- 6. Shape Fluctuations of Vesicles 197 Shigeyuki Komura

vi Contents

II Methods 237

- Characterization of Vesicles and Vesicular Dispersions via Scattering Techniques 239
 John H. van Zanten
- Fluorescence Probing of Vesicles Using Pyrene and Pyrene Derivatives 295
 Guy Duportail and Panagiotis Lianos
- The Mechanochemistry of Lipid Vesicles Examined by Micropipet Manipulation Techniques 373
 David Needham and Doncho V. Zhelev

III Drug Delivery 445

- 10. Liposomes in Drug Delivery 447

 Danilo D. Lasic
- 11. Liposomes and Lipidic Particles in Gene Therapy 477

 Danilo D. Lasic and Rodney Pearlman
- Liposomal Amphotericin B (AmBisome): Realization of the Drug Delivery Concept 491 Gary Fujii
- 13. Vesicles as Topical Drug Delivery Systems 527 *Rimona Margalit*

IV Diverse Applications 561

- 14. Photoenergy Harvesting on Two-Dimensional Vesicular Assemblies 563

 Masahiko Sisido
- 15. Vesicles as Imaging Agents 593

 Colin Tilcock, Deepank Utkhede, and Grant Meng
- 16. Liposomes in In Vivo Immunology 649

 Eric Claassen
- Triggered Release from Liposomes Mediated by Physically and Chemically Induced Phase Transitions 679 Oleg V. Gerasimov, Yuanjin Rui, and David H. Thompson

Index 747

I

Physical and Surface Chemistry

The papers in this section represent some of the research being devoted to basic vesicular properties.

Studies of lipid monolayers considered as half a vesicle bilayer provide insight into the structure and dynamics of vesicles unobtainable through other techniques. By reducing the dimensionality of the system, monolayer studies allow for simplification of complex thermodynamics. The first chapter deals with the relationship between lipid bilayers in the form of vesicles and monolayers at the air-water interface.

In Chapter 2, a wide survey of liposomal topics is presented ranging from the use of molecular acoustics for studying the mechanical properties of liposomes and protein-lipid interactions to vesicle fusion, photochemistry, and energy transfer.

The next paper, Chapter 3, describes the correlation between the phase behavior of vesicles and microemulsions. The importance of the HLB of the surfactant for the formation of reverse vesicles is discussed and the methods for preparing reverse vesicles by dispersion or spontaneous transition are described.

Chapter 4 reviews the phenomenon of phase separation in bilayers. The formation of lipid and protein domains and its consequences, particularly for biology, are examined from the viewpoints of thermodynamics and kinetics.

Interaction of surfactants or amphiphiles with vesicles is important for the solubilization of bilayer membranes as well as their preparation. Additionally, surfactant vesicle interaction serves as a model for the stability of vesicles in the presence of foreign molecules. These matters and the effect of surfactants in the sublytic concentration range are taken up in Chapter 5.

The next contribution, Chapter 6, treats the statistical mechanics of membrane conformational fluctuations. Both fluid and polymerized vesicles are discussed. Shape instabilities may play a role in biological processes such as endocytosis and cell recognition.

1

The Relationship and Interactions Between Lipid Bilayers Vesicles and Lipid Monolayers at the Air/Water Interface

ROBERT C. MACDONALD Department of Biochemistry, Molecular Biology, and Cell Biology, Northwestern University, Evanston, Illinois

I.	Inti	roduction	4
II.	For	ces Operating in Monolayers and Bilayers	5
	A.		
		tension of the air/water interface by surface active molecules	5
	В.	Surface tension is the resultant of internal pressures and	
		tensions	8
	C.		
	_	in concert, to produce the macroscopic surface pressure	9
	D.	With the exception of an oil/air tension, the forces operating	
		in bilayers are essentially the same as those in monolayers	10
III.	Sur	face Thermodynamics Relevant to the Study of Monolayers	
		Bilayers	11
	A.	Basic thermodynamic relationships	11
	В.	The chemical potential depends on surface tension (pressure)	
		and surface concentration	12
	C.	Monolayer surface pressure can be expressed in terms of	
		surface phase water activity	14
	D.	The chemical potential of lipid molecules in monolayers	15
	E.	The chemical potential of lipid molecules in bilayers	16
	F.	Bilayers exhibit an elastic response around the equilibrium area	
		per molecule, whereas monolayers tend to expand at all areas	16
IV.	For	mation of Monolayers from Vesicles of Natural Membranes	19
V.	For	rmation of Monolayers from Lipid Vesicles	19

4 MacDonald

VI.	from Vesicles	26
VII.	Effects of Vesicle Size on the Magnitude of the Equilibrium Monolayer Pressure	27
VIII.	Influences of the Lipid-Phase State on the Monolayer Formed at the Surface of Vesicle Dispersions	
IX.	The Contact Between Bilayers and Monolayers at the Air/Water Interface	31
X.	Interactions of Vesicles with Solid Surfaces	33
XI.	Determining When Monolayers Have the Same Surface Pressure as Bilayers: Methods Based on Penetration by Enzymes and	
	Other Exogenous Molecules	34
	A. Observations on enzymes	34
	B. The theoretical analysis of surface pressure on enzyme	
	activities	36
	C. Penetration by small molecules	39
XII.	Determining When Monolayers Have the Same Surface Pressure as Bilayers: Methods Not Involving Penetration	39
XIII.	The Relationship Between the Organization of Lipid Molecules in Monolayers and in Bilayers	42
	References	46

I. INTRODUCTION

There are several reasons for inquiring into the relationship between lipid bilayers in the form of vesicles and monolayers at macroscopic interfaces, particularly at the air/water interface.* Historically, the most important and perhaps still foremost reason is that monolayers at the air/water interface allow measurements of interaction of membrane-active molecules with an array of lipids that approximates a bilayer membrane in ways that are either not possible or are very difficult with the bilayer itself. The most common such measurement is penetration by the foreign molecule of the monolayer, the extent of which is easily monitored either as an increase in surface pressure of the monolayer at constant area or an increase in area of the monolayer at constant surface pressure. These measurements can be used to

^{*}Although a bilayer is obviously composed of two monolayers, the latter term, unless specified otherwise, implies a monolayer of lipid at the air/water interface.