immuno biology6

THE IMMUNE SYSTEM IN HEALTH AND DISEASE



LARM Find authenticated court documents without watermarks at <u>docketalarm.com</u>.

DOCKET

Δ

immuno biology 5

THE IMMUNE SYSTEM IN HEALTH AND DISEASE

Charles A. Janeway, Jr.

Yale University School of Medicine

Paul Travers

Anthony Nolan Research Institute, London

Mark Walport

Imperial College School of Medicine, London

Mark J. Shlomchik

Yale University School of Medicine



Vice President: Text Editors: Managing Editor: Editorial Assistant: Managing Production Editor: Production Assistant: New Media Editor: Copyeditor: Indexer: Illustration and Layout: Manufacturing: Denise Schanck Penelope Austin, Eleanor Lawrence Sarah Gibbs Mark Ditzel Emma Hunt Angela Bennett Michael Morales Len Ceglelka Liza Furnival Blink Studio, London Marion Morrow, Rory MacDonald

© 2001 by Garland Publishing.

All rights reserved. No part of this publication may be reproduced, stored in a retrieval system or transmitted in any form or by any means—electronic, mechanical, photocopying, recording, or otherwise—without the prior written permission of the copyright holder.

Distributors:

Inside North America: Garland Publishing, 29 West 35th Street, New York, NY 10001-2299. Inside Japan: Nankodo Co. Ltd., 42-6, Hongo 3-Chrome, Bunkyo-ku, Tokyo, 113-8410, Japan. Outside North America and Japan: Churchill Livingstone, Robert Stevenson House, 1–3 Baxter's Place, Leith Walk, Edinburgh, EH1 3AF.

ISBN 0 8153 3642 X (paperback) Garland

ISBN 0 4430 7096 9 (paperback) Churchill Livingstone ISBN 0 4430 7099 7 (paperback) International Student Edition

Library of Congress Cataloging-in-Publication Data

Immunobiology : the immune system in health and disease / Charles A. Janeway, Jr. ... [et al.].-- 5th ed. p. cm. Includes bibliographical references and index.

ISBN 0-8153-3642-X (pbk.) 1. Immunology. 2. Immunity, I. Janeway, Charles. II. Title.

QR181 .1454 2001 616.07'9--dc21

DOCKE

Δ

R

Μ

Δ

2001016039

This book was produced using QuarkXpress 4.11 and Adobe Illustrator 9.0

Published by Garland Publishing, a member of the Taylor & Francis Group, 29 West 35th Street, New York, NY 10001-2299.

Printed in the United States of America. 15 14 13 12 11 10 9 8 7 6 5 4 3 2

Find authenticated court documents without watermarks at docketalarm.com.

CONTENTS

ix.

PARTI	AN INTRODUCTION TO IMMUNOBIOLOGY AND INNATE IMMUNIT	Υ
Chapter 1	Basic Concepts In Immunology	1
Chapter 2	Innate Immunity	35
PART II	THE RECOGNITION OF ANTIGEN	
Chapter 3	Antigen Recognition by B-cell and T-cell Receptors	93
Chapter 4	The Generation of Lymphocyte Antigen Receptors	123
Chapter 5	Antigen Presentation to T Lymphocytes	155
PART III	THE DEVELOPMENT OF MATURE LYMPHOCYTE RECEPTOR REPERTOIRES	
Chapter 6	Signaling Through Immune System Receptors	187
Chapter 7	The Development and Survival of Lymphocytes	221
PART IV	THE ADAPTIVE IMMUNE RESPONSE	
Chapter 8	T Cell-Mediated Immunity	295
Chapter 9	The Humoral Immune Response	341
Chapter 10	Adaptive Immunity to Infection	381
PART V	THE IMMUNE SYSTEM IN HEALTH AND DISEASE	
Chapter 11	Failures of Host Defense Mechanisms	425
Chapter 12	Allergy and Hypersensitivity	471
Chapter 13	Autoimmunity and Transplantation	501
Chapter 14	Manipulation of the Immune Response	553
Afterword	Evolution of the Immune System: Past, Present, and Future,	
	by Charles A. Janeway, Jr.	597
Appendix I	Immunologists' Toolbox	613
Appendix II	CD Antigens	661
Appendix III	Cytokines and their Receptors	677
Appendix IV	Chemokines and their Receptors	680
Appendix V	Immunological Constants	681
Biographies		682
Glossary		683
Index		708

DOCKET

ALARM

Find authenticated court documents without watermarks at <u>docketalarm.com</u>.

Immunologists' Toolbox

APPENDIX I

Immunization.

Natural adaptive immune responses are normally directed at antigens borne by pathogenic microorganisms. The immune system can, however, also be induced to respond to simple nonliving antigens, and experimental immunologists have focused on the responses to these simple antigens in developing our understanding of the immune response. The deliberate induction of an immune response is known as **immunization**. Experimental immunizations are routinely carried out by injecting the test antigen into the animal or human subject. The route, dose, and form in which antigen is administered can profoundly affect whether a response occurs and the type of response that is produced, and are considered in Sections A-1–A-4. The induction of protective immune responses against common microbial pathogens in humans is often called **vaccination**, although this term is correctly only applied to the induction of immune responses against smallpox by immunizing with the cross-reactive cowpox virus, vaccinia (see Chapter 14).

To determine whether an immune response has occurred and to follow its course, the immunized individual is monitored for the appearance of immune reactants directed at the specific antigen. Immune responses to most antigens elicit the production of both specific antibodies and specific effector T cells. Monitoring the antibody response usually involves the analysis of relatively crude preparations of **antiserum** (plural: **antisera**). The **serum** is the fluid phase of clotted blood, which, if taken from an immunized individual, is called antiserum because it contains specific antibodies against the immunizing antigen as well as other soluble serum proteins. To study immune responses mediated by T cells, blood lymphocytes or cells from lymphoid organs such as the spleen are tested; T-cell responses are more commonly studied in experimental animals than in humans.

Any substance that can elicit an immune response is said to be **immunogenic** and is called an **immunogen**. There is a clear operational distinction between an immunogen and an antigen. An **antigen** is defined as any sustance that can bind to a specific antibody. All antigens therefore have the potential to elicit specific antibodies, but some need to be attached to an immunogen in order to do so. This means that although all immunogens are antigens, not all antigens are immunogenic. The antigens used most frequently in experimental immunology are proteins, and antibodies to proteins are of enormous utility in experimental biology and medicine. Purified proteins are, however, not always highly immunogenic and to provoke an immune response have to be administered with an adjuvant (see Section A-4). Carbohydrates, nucleic acids, and other types of molecule are all potential antigens, but will often only induce an immune response if attached to a protein carrier. Thus, the immunogenicity of protein antigens determines the outcome of virtually every immune response.

DOCKET



Explore Litigation Insights

Docket Alarm provides insights to develop a more informed litigation strategy and the peace of mind of knowing you're on top of things.

Real-Time Litigation Alerts



Keep your litigation team up-to-date with **real-time** alerts and advanced team management tools built for the enterprise, all while greatly reducing PACER spend.

Our comprehensive service means we can handle Federal, State, and Administrative courts across the country.

Advanced Docket Research



With over 230 million records, Docket Alarm's cloud-native docket research platform finds what other services can't. Coverage includes Federal, State, plus PTAB, TTAB, ITC and NLRB decisions, all in one place.

Identify arguments that have been successful in the past with full text, pinpoint searching. Link to case law cited within any court document via Fastcase.

Analytics At Your Fingertips



Learn what happened the last time a particular judge, opposing counsel or company faced cases similar to yours.

Advanced out-of-the-box PTAB and TTAB analytics are always at your fingertips.

API

Docket Alarm offers a powerful API (application programming interface) to developers that want to integrate case filings into their apps.

LAW FIRMS

Build custom dashboards for your attorneys and clients with live data direct from the court.

Automate many repetitive legal tasks like conflict checks, document management, and marketing.

FINANCIAL INSTITUTIONS

Litigation and bankruptcy checks for companies and debtors.

E-DISCOVERY AND LEGAL VENDORS

Sync your system to PACER to automate legal marketing.

