



Biological Identification

DNA Amplification and
Sequencing, Optical Sensing,
Lab-on-chip and Portable
Systems

Edited by R. Paul Schaudies

Woodhead Publishing Series in Electronic and Optical Materials:
Number 59

Biological Identification

DNA Amplification and Sequencing,
Optical Sensing, Lab-on-chip and
Portable Systems

Edited by
R. Paul Schaudies



AMSTERDAM • BOSTON • CAMBRIDGE • HEIDELBERG • LONDON
NEW YORK • OXFORD • PARIS • SAN DIEGO
SAN FRANCISCO • SINGAPORE • SYDNEY • TOKYO

Woodhead Publishing is an imprint of Elsevier



Woodhead Publishing is an imprint of Elsevier
80 High Street, Sawston, Cambridge CB22 3HJ, UK
25 Wyman Street, Waltham, MA 02451, USA
Langford Lane, Kidlington, OX5 1GB, UK

Copyright © 2014 Elsevier Ltd. All rights reserved

Exceptions to the above:

Chapter 14 was prepared by US Government employees; it is therefore in the public domain and cannot be copyrighted. Published by Woodhead Publishing Limited.

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 publisher.

Permissions may be sought directly from Elsevier's Science & Technology Rights Department in Oxford, UK: phone (+44) (0) 1865 843830; fax (+44) (0) 1865 853333; e-mail: permissions@elsevier.com. Alternatively, you can submit your request online by visiting the Elsevier website at <http://elsevier.com/locate/permissions>, and selecting Obtaining permission to use Elsevier material.

Notice

No responsibility is assumed by the publisher for any injury and/or damage to persons or property as a matter of products liability, negligence or otherwise, or from any use or operation of any methods, products, instructions or ideas contained in the material herein. Because of rapid advances in the medical sciences, in particular, independent verification of diagnoses and drug dosages should be made.

British Library Cataloguing-in-Publication Data

A catalogue record for this book is available from the British Library.

Library of Congress Control Number: 2014935097

ISBN 978-0-85709-501-5 (print)

ISBN 978-0-85709-916-7 (online)

For information on all Woodhead Publishing publications
visit our website: <http://store.elsevier.com/>

Typeset by RefineCatch Limited, Bungay, Suffolk

Printed and bound in the United Kingdom



Working together
to grow libraries in
developing countries

www.elsevier.com • www.bookaid.org

Biological Identification

Related titles:

Nanosensors for chemical and biological applications
(ISBN 978-0-85709-660-9)

Semiconductor gas sensors
(ISBN 978-0-85709-236-6)

Smart sensors and MEMS
(ISBN 978-0-85709-502-2)

Contributor contact details

* = main contact

Editor

R. P. Schaudies
GenArraytion Inc.
9700 Great Seneca Hwy Suite 325
Rockville, MD 20850, USA

Email: pschaudies@genarraytion.com

Chapter 1

J. C. Detter*, S. L. Johnson and
P. S. Chain
Los Alamos National Laboratory
Los Alamos, NM 87545, USA

Email: cdetter@lanl.gov; shannonj@
lanl.gov; pchain@lanl.gov

K. A. Bishop-Lilly
Navy Medical Research Center-
Fredrick
Ft Detrick, MD, USA

Email: Kim.Bishop-Lilly@med.navy.mil

H. S. Gibbons
US Army Edgewood Chemical
Biological Center
Aberdeen, MD 21010, USA

Email: henry.s.gibbons.civ@mail.mil

T. D. Minogue
US Army Medical Research of
Infectious Disease
Frederick, MD 21702, USA

Email: timothy.d.minogue.civ@mail.mil

Shanmuga Sozhamannan
Critical Reagents Program
Frederick, MD 21702, USA

Email: shanmuga.sozhamannan.ctr@
mail.mil

E. J. Van Gieson
Defence Threat Reduction Agency
Ft. Belvoir, VA, USA

Email: Eric.VanGieson@DTRA.MIL

I. G. Resnick
IGR Consulting
Park City, UT
USA

Email: gary.resnick2@gmail.com

Chapter 2

W. M. Nelson*, G. W. Long and
L. M. Cockrell
Tetracore, Inc.
9901 Belward Campus Drive, Suite
300
Rockville, MD 20850, USA

Email: wnelson@tetracore.com; glong@tetracore.com

Chapter 3

Y. Tong
New England Biolabs
240 County Road
Ipswich, MA 01938, USA

Email: tong@neb.com; yanhong_t@hotmail.com

Chapter 4

S. A. Dunbar* and S. V. Angeloni
Luminex Corporation
12212 Technology Blvd
Austin, TX 78727, USA

Email: sdunbar@luminexcorp.com;
sangeloni@luminexcorp.com

Chapter 5

P. Skládal
Masaryk University
Department of Biochemistry
Kamenice 5
62500 Brno, Czech Republic

E-mail: skladal@chemi.muni.cz

Chapter 6

S. Dzyadevych
Institute of Molecular Biology
and Genetics
150 Zabolotnogo Str.
Kiev 03680, Ukraine

Email: dzyad@yahoo.com

N. Jaffrezic-Renault*
University of Lyon
Institut des Sciences Analytiques
5 Rue de La Doua
69100 Villeurbanne, France

Email: nicole.jaffrezic@univ-lyon1.fr

Chapter 7

N. Chaniotakis* and M. Fouskaki
Laboratory of Analytical Chemistry
Department of Chemistry
University of Crete
Vasilika Voutes
71003 Heraklion, Crete, Greece

Email: nchan@chemistry.uoc.gr

Chapter 8

H. Becker*, T. Hansen-Hagge and
C. Gärtner
microfluidic ChipShop GmbH
Stockholmer Str. 20
D-07747 Jena
Germany

Email: hb@microfluidic-chipshop.com;
Thomas.hansen-hagge@microfluidic-chipshop.com; claudia.gaertner@microfluidic-chipshop.com

Chapter 9

H. J. Harmon* and A. Oliver
Department of Physics
Oklahoma State University
Stillwater
OK74078
USA

Email: james.harmon@okstate.edu

Chapter 10

T. Globus*, B. Gelmont and I. Sizov
Department of Electrical and
Computer Engineering
University of Virginia
351 McCormick Road
Charlottesville, VA 22904-4743,
USA

Email: tg9a@virginia.edu; gb7k@
virginia.edu

Chapter 11

T. J. Ronningen, J. M. Schuetter,
J. L. Wightman, A. Murdock and
A. P. Bartko*
Battelle Memorial Institute
Columbus, OH 43201, USA

Email: bartkoa@battelle.org

Chapter 12

H. S. Lee, C. R. Prasad and
R. M. Serino*
Science & Engineering Services,
Inc. (SESI)
6992 Columbia Gateway Drive
Columbia, MD 21046, USA

Email: rserino@sesi-md.com

Chapter 13

C. Bradburne
Johns Hopkins University
Applied Physics Laboratory
Applied Biology Branch
Asymmetric Operations Department
11100 Johns Hopkins Road
Laurel, MD 20723-6099, USA

Email: Chris.Bradburne@jhuapl.edu

Chapter 14

R. E. Jabbour and A. P. Snyder*
US Army Edgewood Chemical
Biological Center
Aberdeen Proving Ground
MD 21010, USA

Email: Arnold.p.snyder.civ@mail.mil

Woodhead Publishing Series in Electronic and Optical Materials

- 1 **Circuit analysis**
J. E. Whitehouse
- 2 **Signal processing in electronic communications: For engineers and mathematicians**
M. J. Chapman, D. P. Goodall and N. C. Steele
- 3 **Pattern recognition and image processing**
D. Luo
- 4 **Digital filters and signal processing in electronic engineering: Theory, applications, architecture, code**
S. M. Bozic and R. J. Chance
- 5 **Cable engineering for local area networks**
B. J. Elliott
- 6 **Designing a structured cabling system to ISO 11801: Cross-referenced to European CENELEC and American Standards**
Second edition
B. J. Elliott
- 7 **Microscopy techniques for materials science**
A. Clarke and C. Eberhardt
- 8 **Materials for energy conversion devices**
Edited by C. C. Sorrell, J. Nowotny and S. Sugihara
- 9 **Digital image processing: Mathematical and computational methods**
Second edition
J. M. Blackledge
- 10 **Nanolithography and patterning techniques in microelectronics**
Edited by D. Bucknall
- 11 **Digital signal processing: Mathematical and computational methods, software development and applications**
Second edition
J. M. Blackledge
- 12 **Handbook of advanced dielectric, piezoelectric and ferroelectric materials: Synthesis, properties and applications**
Edited by Z.-G. Ye
- 13 **Materials for fuel cells**
Edited by M. Gasik
- 14 **Solid-state hydrogen storage: Materials and chemistry**
Edited by G. Walker

- 15 **Laser cooling of solids**
S. V. Petrushkin and V. V. Samartsev
- 16 **Polymer electrolytes: Fundamentals and applications**
Edited by C. A. C. Sequeira and D. A. F. Santos
- 17 **Advanced piezoelectric materials: Science and technology**
Edited by K. Uchino
- 18 **Optical switches: Materials and design**
Edited by S. J. Chua and B. Li
- 19 **Advanced adhesives in electronics: Materials, properties and applications**
Edited by M. O. Alam and C. Bailey
- 20 **Thin film growth: Physics, materials science and applications**
Edited by Z. Cao
- 21 **Electromigration in thin films and electronic devices: Materials and reliability**
Edited by C.-U. Kim
- 22 ***In situ* characterization of thin film growth**
Edited by G. Koster and G. Rijnders
- 23 **Silicon-germanium (SiGe) nanostructures: Production, properties and applications in electronics**
Edited by Y. Shiraki and N. Usami
- 24 **High-temperature superconductors**
Edited by X. G. Qiu
- 25 **Introduction to the physics of nanoelectronics**
S. G. Tan and M. B. A. Jalil
- 26 **Printed films: Materials science and applications in sensors, electronics and photonics**
Edited by M. Prudenziati and J. Hormadaly
- 27 **Laser growth and processing of photonic devices**
Edited by N. A. Vainos
- 28 **Quantum optics with semiconductor nanostructures**
Edited by F. Jahnke
- 29 **Ultrasonic transducers: Materials and design for sensors, actuators and medical applications**
Edited by K. Nakamura
- 30 **Waste electrical and electronic equipment (WEEE) handbook**
Edited by V. Goodship and A. Stevels
- 31 **Applications of ATILA FEM software to smart materials: Case studies in designing devices**
Edited by K. Uchino and J.-C. Debus
- 32 **MEMS for automotive and aerospace applications**
Edited by M. Kraft and N. M. White
- 33 **Semiconductor lasers: Fundamentals and applications**
Edited by A. Baranov and E. Tournie
- 34 **Handbook of terahertz technology for imaging, sensing and communications**
Edited by D. Saeedkia
- 35 **Handbook of solid-state lasers: Materials, systems and applications**
Edited by B. Denker and E. Shklovsky
- 36 **Organic light-emitting diodes (OLEDs): Materials, devices and applications**
Edited by A. Buckley

- 37 **Lasers for medical applications: Diagnostics, therapy and surgery**
Edited by H. Jelínková
- 38 **Semiconductor gas sensors**
Edited by R. Jaaniso and O. K. Tan
- 39 **Handbook of organic materials for optical and (opto)electronic devices: Properties and applications**
Edited by O. Ostroverkhova
- 40 **Metallic films for electronic, optical and magnetic applications: Structure, processing and properties**
Edited by K. Barmak and K. Coffey
- 41 **Handbook of laser welding technologies**
Edited by S. Katayama
- 42 **Nanolithography: The art of fabricating nanoelectronic and nanophotonic devices and systems**
Edited by M. Feldman
- 43 **Laser spectroscopy for sensing: Fundamentals, techniques and applications**
Edited by M. Baudelet
- 44 **Chalcogenide glasses: Preparation, properties and applications**
Edited by J.-L. Adam and X. Zhang
- 45 **Handbook of MEMS for wireless and mobile applications**
Edited by D. Uttamchandani
- 46 **Subsea optics and imaging**
Edited by J. Watson and O. Zielinski
- 47 **Carbon nanotubes and graphene for photonic applications**
Edited by S. Yamashita, Y. Saito and J. H. Choi
- 48 **Optical biomimetics: Materials and applications**
Edited by M. Large
- 49 **Optical thin films and coatings**
Edited by A. Piegari and F. Flory
- 50 **Computer design of diffractive optics**
Edited by V. A. Soifer
- 51 **Smart sensors and MEMS: Intelligent devices and microsystems for industrial applications**
Edited by S. Nihtianov and A. Luque
- 52 **Fundamentals of femtosecond optics**
S. A. Kozlov and V. V. Samartsev
- 53 **Nanostructured semiconductor oxides for the next generation of electronics and functional devices: Properties and applications**
S. Zhuilykov
- 54 **Nitride semiconductor light-emitting diodes (LEDs): Materials, technologies and applications**
Edited by J. J. Huang, H. C. Kuo and S. C. Shen
- 55 **Sensor technologies for civil infrastructures**
Volume 1: Sensing hardware and data collection methods for performance assessment
Edited by M. Wang, J. Lynch and H. Sohn
- 56 **Sensor technologies for civil infrastructures**
Volume 2: Applications in structural health monitoring
Edited by M. Wang, J. Lynch and H. Sohn

- 57 **Graphene: Properties, preparation, characterisation and devices**
Edited by V. Skákalová and A. B. Kaiser
- 58 **Silicon-on-insulator (SOI) technology**
Edited by O. Kononchuk and B.-Y. Nguyen
- 59 **Biological identification: DNA amplification and sequencing, optical sensing, lab-on-chip and portable systems**
Edited by R. P. Schaudies
- 60 **High performance silicon imaging: Fundamentals and applications of CMOS and CCD sensors**
Edited by D. Durini
- 61 **Nanosensors for chemical and biological applications: Sensing with nanotubes, nanowires and nanoparticles**
Edited by K. C. Honeychurch
- 62 **Composite magnetoelectrics: Materials, structures, and applications**
G. Srinivasan, S. Priya and N. Sun
- 63 **Quantum information processing with diamond: Principles and applications**
Edited by S. Praver and I. Aharonovich
- 64 **Advances in non-volatile memory and storage technology**
Edited by Y. Nishi
- 65 **Laser surface engineering: Processes and applications**
Edited by J. Lawrence, C. Dowding, D. Waugh and J. Griffiths
- 66 **Power ultrasonics: A handbook of materials, design and applications of high power ultrasound transducers**
Edited by J. A. Gallego-Juárez
- 67 **Advances in delay-tolerant networks (DTNs): Architectures, routing and challenges**
Edited by J. J. P. C. Rodrigues
- 68 **Handbook of flexible organic electronics: Materials, manufacturing and applications**
Edited by S. Logothetidis
- 69 **Machine-to-machine (M2M) communications: Architecture, performance and applications**
Edited by C. Anton-Haro and M. Dohler
- 70 **Ecological design of smart home networks: Technologies, social impact and sustainability**
Edited by N. Saito and D. Menga
- 71 **Industrial tomography: Systems and applications**
Edited by M. Wang
- 72 **Vehicular communications and networks: Architectures, protocols, operation and deployment**
Edited by W. Chen
- 73 **Modeling, characterization, and production of nanomaterials: Electronics, photonics and energy applications**
Edited by V. Tewary and Y. Zhang
- 74 **Reliability characterisation of electrical and electronic systems**
Edited by J. Swingler
- 75 **Handbook of industrial wireless sensor networks: Monitoring, control and automation**
Edited by R. Budampati and S. Kolavennu

- 76 **Epitaxial growth of complex metal oxides: Techniques, properties and applications**

Edited by G. Koster and G. Rijnders

- 77 **Semiconductor nanowires: Materials, synthesis, characterization and applications**

Edited by J. Arbiol and Q. Xiong

The need to accurately identify and characterize biological organisms in a rapid fashion pervades almost all aspects of our life. The safety of the life essentials of food and water requires that we be able to identify and eliminate contamination at all levels of collection and distribution. Zoonotic infections, introduced either deliberately or through natural exposure, pose a cross-species threat to human life. The rise of emerging infectious diseases exacerbated by widespread international travel adds the requirement for not only rapid discovery and test distribution, but also improved treatment methods. Hospitals have become reservoirs of infectious organisms, leading to the now pervasive term ‘hospital acquired infections’ (HAI). HAIs are accounting for a growing percentage of infections leading to sepsis and frequently death. Then there is concern over the deliberate release of infectious organisms into a densely populated environment. The US Government has spent significant resources on the BioWatch program designed to protect our major population centers from a biological warfare attack.

The diversity of biological microorganisms presents a challenge almost as great as their ability to defend themselves against our attempts to kill them, or at least inhibit their replication. The diversity of replication mechanisms and the ability to use host mechanisms for replication only complicate our attempts to selectively inhibit the invader. All of the diverse approaches for the identification and characterization of microbiological organisms share the requirement to differentiate the target from the environment it is part of. This may involve attempts to isolate the target from the environment, frequently termed ‘sample preparation’. Sample preparation is the ultimate driver for sensitivity, as the material must be extracted and concentrated and transferred into a matrix that is compatible with the identification strategy. The ability to rapidly and accurately identify microorganisms, especially those that cause disease in man, is paramount in all regions where people live. This work provides highlights of a multitude of scientific approaches to both identifying and characterizing biological material by exploiting the characteristics that are unique to an organism and differentiate it from its environment.

Part I of this book concentrates on the genetic elements of DNA and RNA for the identification and characterization of biological organisms. The DNA or RNA from an organism provides the blueprint from which activities are derived. The presence of a specific genetic element does not guarantee expression, but the

absence of a specific gene does indicate an absence of activity associated with that genetic element. The chapters begin with Next Generation Sequencing (NGS) in complex matrices and how these technical approaches identify both known and previously undiscovered organisms. While NGS is becoming much more affordable and the data analysis time is decreasing, there is still a pressing need to ask and answer much simpler and specific questions using the amplification of specific smaller regions of nucleic acids. The following chapters include classical polymerase chain reaction methods and isothermal amplifications as well as bead arrays for the analysis of amplified nucleic acid fragments. The need to make identification systems portable resulted in Part II of the book.

Part II, Lab-on-chip and portable systems for biodetection and analysis, is a collection of diverse approaches and technologies, not limited to nucleic acid signatures, to make systems efficient and smaller. Small systems require different approaches to fluid movement than bench-top systems, as well as unique power requirements. Smaller assay volumes require highly efficient sample preparation and detection methods because the small volume frequently requires higher concentrations than larger-volume detection approaches. This second part of the publication provides an in-depth overview of multiple approaches for specific signal generation based on the structural recognition of biological signatures. This section is followed by Part III, which focuses on optical approaches for recognition and identification.

Optical identification methods either detect modifications after an optically active molecule interacts with the target or take advantage of unique electromagnetic spectral properties of the organism itself. Most optical approaches have the advantage of being non-destructive and thereby preserving the sample for either culture or further analytical examination. The final section of the book describes a unique approach to sample preparation utilizing charge and mass differences to concentrate material of interest. The final chapter documents some very promising approaches for biological identification based on mass spectrometry.

I would personally like to thank all of the authors, as well as the individuals at Woodhead Publishing for their professionalism and dedication in producing this work.

Contents

<i>Contributor contact details</i>	<i>xi</i>
<i>Woodhead Publishing Series in Electronic and Optical Materials</i>	<i>xv</i>
<i>Preface</i>	<i>xxi</i>
Part I Technology for DNA and RNA analysis of pathogens	1
1 Nucleic acid sequencing for characterizing infectious and/or novel agents in complex samples	3
J. C. DETTER and S. L. JOHNSON, Los Alamos National Laboratory, USA, K. A. BISHOP-LILLY, Navy Medical Research Center-Frederick, USA, P. S. CHAIN, Los Alamos National Laboratory, USA, H. S. GIBBONS, US Army Edgewood Chemical Biological Center, USA, T. D. MINOGUE, US Army Medical Research of Infectious Disease, USA, S. SOZHAMANNAN, Critical Reagents Program, USA, E. J. VAN GIESON, Defense Threat Reduction Agency, USA and I. G. RESNICK, IGR Consulting, USA	
1.1 Pathogen sequencing and applications in public health and biosecurity	3
1.2 Next-generation sequencing (NGS) technologies and the sequencing landscape	15
1.3 Characterization of known pathogens	24
1.4 Discovery of novel agents	30
1.5 Future trends	38
1.6 Acknowledgments	43
1.7 References	43
2 Multiplexed, lateral flow, polymerase chain reaction (PCR) techniques for biological identification	54
W. M. NELSON, G. W. LONG and L. M. COCKRELL, Tetracore, Inc., USA	
2.1 Introduction	54

vi	Contents	
2.2	Real-time PCR: development and description	57
2.3	Considerations when developing a real-time PCR assay	63
2.4	Real-time PCR instrument platforms	64
2.5	References	67
3	Isothermal amplification of specific sequences Y. TONG, New England Biolabs, USA	69
3.1	Introduction	69
3.2	Melting temperature (T_m) estimation and categories of isothermal amplification technologies	70
3.3	Isothermal amplification based on DNA polymerases	74
3.4	Isothermal amplification based on RNA polymerases	84
3.5	Future prospects	85
3.6	References	87
4	Bead array technologies for genetic disease screening and microbial detection S. A. DUNBAR and S. V. ANGELONI, Luminex Corporation, USA	93
4.1	Introduction	93
4.2	Luminex [®] xMAP [®] Technology	94
4.3	Illumina VeraCode	108
4.4	NanoString nCounter	111
4.5	Applications	115
4.6	Conclusion	121
4.7	References	122
Part II	Lab-on-chip and portable systems for biodetection and analysis	129
5	Electrochemical detection for biological identification P. SKLÁDAL, Masaryk University, Czech Republic	131
5.1	Introduction	131
5.2	Electrochemical techniques for bioanalysis	132
5.3	Electrochemical biosensors for pathogens	141
5.4	Conclusions	146
5.5	References	147
6	Conductometric biosensors S. DZYADEVYCH, National Academy of Sciences of Ukraine, Ukraine and N. JAFFREZIC-RENAULT, University of Lyon, France	153
6.1	Introduction	153
6.2	Conductometry in enzyme catalysis	154
6.3	Conductometric enzyme biosensors based on direct analysis – I: Biosensors for biomedical applications	155