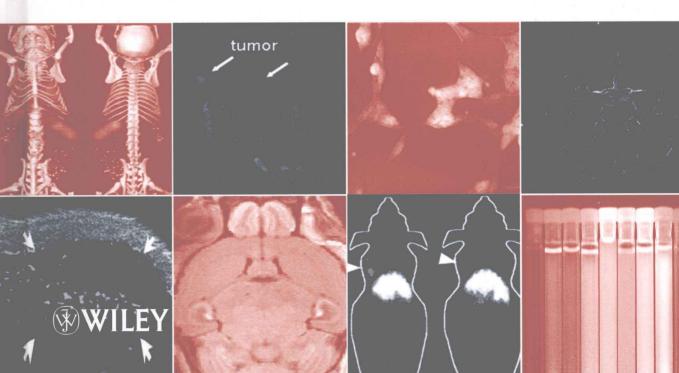


# NANOPLATFORM-BASED MOLECULAR IMAGING

edited by XIAOYUAN CHEN

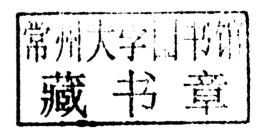


## NANOPLATFORM-BASED MOLECULAR IMAGING

Edited by

#### Xiaoyuan Chen

Laboratory of Molecular Imaging and Nanomedicine National Institute of Biomedical Imaging and Bioengineering National Institutes of Health Bethesda, Maryland





A JOHN WILEY & SONS, INC., PUBLICATION

Copyright © 2011 by John Wiley & Sons, Inc. All rights reserved.

Published by John Wiley & Sons, Inc., Hoboken, New Jersey. Published simultaneously in Canada.

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, scanning, or otherwise, except as permitted under Section 107 or 108 of the 1976 United States Copyright Act, without either the prior written permission of the Publisher, or authorization through payment of the appropriate per-copy fee to the Copyright Clearance Center, Inc., 222 Rosewood Drive, Danvers, MA 01923, (978) 750-8400, fax (978) 750-4470, or on the web at www.copyright.com. Requests to the Publisher for permission should be addressed to the Permissions Department, John Wiley & Sons, Inc., 111 River Street, Hoboken, NJ 07030, (201) 748-6011, fax (201) 748-6008, or online at http://www.wiley.com/go/permission.

Limit of Liability/Disclaimer of Warranty: While the publisher and author have used their best efforts in preparing this book, they make no representations or warranties with respect to the accuracy or completeness of the contents of this book and specifically disclaim any implied warranties of merchantability or fitness for a particular purpose. No warranty may be created or extended by sales representatives or written sales materials. The advice and strategies contained herein may not be suitable for your situation. You should consult with a professional where appropriate. Neither the publisher nor author shall be liable for any loss of profit or any other commercial damages, including but not limited to special, incidental, consequential, or other damages.

For general information on our other products and services or for technical support, please contact our Customer Care Department within the United States at (800) 762-2974, outside the United States at (317) 572-3993 or fax (317) 572-4002.

Wiley also publishes its books in a variety of electronic formats. Some content that appears in print may not be available in electronic formats. For more information about Wiley products, visit our web site at www.wiley.com

#### Library of Congress Cataloging-in-Publication Data:

Nanoplatform-based molecular imaging / edited by Xiaoyuan Chen.

p.; cm.

Includes bibliographical references and index.

ISBN 978-0-470-52115-1

1. Molecular probes. 2. Diagnostic imaging. I. Chen, Xiaoyuan.

[DNLM: 1. Molecular Imaging-methods. 2. Molecular Imaging-trends. 3. Molecular Probes-diagnostic use. 4. Nanoparticles-diagnostic use. 5. Nanotechnology-trends. WN 180 N1865 2010]

QP519.9.M64N36 2010

616.07'54-dc22

2010007984

Printed in Singapore

eBook ISBN: 978-0-470-76703-0 oBook ISBN: 978-0-470-76704-7

10 9 8 7 6 5 4 3 2 1

This book focuses on the rational design of water-soluble, biocompatible nanoparticles for the visualization of the cellular function and follow-up of the molecular processes in living organisms without perturbing them. Molecular imaging probes based on nanotechnology hold great potential in diagnosis, imaging guided intervention, and treatment response monitoring of diseases. This book is logically organized by including the basics of molecular imaging, general strategies of particle synthesis and surface chemistry, applications in computed tomography (CT), optical imaging, magnetic resonance imaging (MRI), ultrasound, multimodality imaging, and theranostics, and finally clinical perspectives of nanoimaging. This comprehensive title provides expert opinions on the latest developments in molecular imaging using nanoparticles. This book consists of 32 chapters and was contributed by nearly 100 authors worldwide, who are among the world's prominent scientists in material science and/or molecular imaging.

Part I consists of Chapters 1–4 Chapter 1 describes the basic principles of molecular imaging, how nanoparticles can be applied to different molecular imaging modalities, and challenges in developing nanoparticle-based molecular imaging probes; Chapter 2 highlights the general strategies to produce narrowly dispersed nanomaterials for molecular imaging; Chapter 3 emphasizes the importance of surface modification to render nanoparticles biocompatible and suitable for molecular imaging applications; and Chapter 4 talks about the toxicity and factors such as size, shape, coating, and surface charge that affect the biodistribution and pharmacokinetics of nanoprobes.

Part II consists of Chapters 5-17 Chapter 5 illustrates the basic principles of CT, the evolution of CT imaging technology, and the rationale for nanoparticle-based CT contrast agents; Chapter 6 describes the advantages of fascinating carbon nanotube field emission X-ray technology over conventional thermionic X-ray tubes that are used in current X-ray imaging systems; Chapter 7 describes the use of unique optical properties of semiconductor quantum dots (QDs) for near-infrared fluorescence imaging in living animals; Chapter 8 introduces macromolecular nanoconstructs such as biopolymers, dendrimers, and liposomes as carriers for fluorophore conjugation and optical imaging; Chapter 9 summarizes recent progress in developing nanoplatforms for Raman imaging of biological systems; Chapter 10 summarizes the work in using single-walled carbon nanotubes (SWNTs) as near-infrared fluorescent sensors for biomolecule detection; Chapter 11 describes the use of micro- and nanoparticles as ultrasound contrast agents; Chapter 12 proposes the use of metal nanoparticles in ultrasound-based photoacoustic and magnetoacoustic imaging modalities; Chapter 13 reports the progress on magnetic resonance imaging (MRI) contrast agents based on inorganic nanoparticles; Chapter 14 emphasizes the use of iron oxide nanoparticles for cellular labeling followed by T2- and T2-weighted MRI; Chapter 15 covers the use of rare earth based nanoparticles for MR imaging as positive contrast agents; Chapter 16 reviews the top-down microfabrication technology to synthesize multispectral MRI contrast agents; and Chapter 17 gives an overview of the strategies to label nanoparticles with radionuclides to study in vivo distribution.

Part III consists of Chapters 18-31 Chapter 18 introduces techniques to incorporate imaging agents into lipoproteins and to reroute lipoproteins to cancer specific epitopes; Chapter 19 exemplifies the use of protein cages such as virus capsids and ferritins as platforms for MRI contrast agents and fluorescent imaging agents; Chapter 20 provides a comprehensive summary of the state-of-the-art of SWNTs for multimodality biomedical imaging applications; Chapter 21 reviews the progress in the controlled synthesis, surface modification, and multimodality imaging applications of multifunctional nanoparticles in recent years; Chapter 22 argues the use of cancer theranostics as a promising new strategy in cancer management, permitting simultaneous cancer diagnosis, drug delivery, and real-time monitoring of therapeutic efficacy; Chapter 23 provides more examples of multifunctional nanoparticles for combined cancer imaging and therapy (theranostics); Chapter 24 describes the recent progress in modifying magnetic nanoparticles for multimodality imaging as well as targeted treatment of a number of diseases; Chapter 25 introduces gold nanocages as contrast agents for optical bioimaging (such as optical and spectroscopic coherence tomography amd photoacoustic tomography) and photothermal treatment; Chapter 26 describes the biological inertness, ease of manufacture and bioconjugation, and presumed lack of toxicity of gold nanoparticles for simultaneous sensing, imaging, and treatment of tumors; Chapter 27 presents the recent developments in the chemistry and photophysics of gold nanorods and their applications toward biological imaging and photothertmally activated therapies; Chapter 28 describes a number of gold core-shell nanostructures for cancer molecular optical imaging, controlled drug delivery, and photothermal ablation therapy; Chapter 29 describes a novel temperature and pH-responsive magnetic nanocarrier that combines tumor targeting and controlled drug release capabilities; Chapter 30 deals with perfluorocarbon nanoparticles as a multidimensional platform for targeted image-guided drug delivery; and Chapter 31 describes the use of radiolabeled nanoparticles and radiolabeled immunonanoparticles for imaging and therapy.

Part IV is the concluding Chapter 32 that highlights some of the nanoparticle-based novel technologies for molecular imaging, diagnosis, and drug delivery formulations. The limitations and future challenges of nanoparticle-based systems are also discussed.

Bethesda, Maryland

XIAOYUAN CHEN

#### ACKNOWLEDGMENTS

The editor thanks the nearly 100 authors throughout the world for their contributions and collaboration on this book project. The editing work of this book was accomplished using a significant amount of the editor's spare time including family time. Therefore the editor also thanks his wife, Michelle Ji, and his daughter, Grace Chen, for their wonderful support and understanding.

#### CONTRIBUTORS

- **Frank Alexis,** Department of Bioengineering, Clemson University, Clemson, South Carolina, USA
- **Leslie Au,** Department of Biomedical Engineering, Washington University, St. Louis, Missouri, USA
- Vaishali Bagalkot, School of Life Sciences, Gwangju Institute of Science and Technology, Gwangju, South Korea
- **R. Bazzi,** Laboratoire Physico-Chimie des Electrolytes, Colloides et Sciences Analytiques, Université Pierre et Marie Curie, Paris, France
- C. Billotey, Laboratoire CREATIS-Animage, Université Claude Bernard, Lyon, France
- **Sébastien Boutry,** Department of General, Organic and Biomedical Chemistry, NMR and Molecular Imaging Laboratory, University of Mons, Mons, Belgium
- J.-L. Bridot, Service de Chimie Générale, Organique et Biomédicale, Laboratoire de RMN et d'Imagerie Moléculaire, Université de Mons-Hainaut, Mons, Belgium
- Weibo Cai, Departments of Radiology and Medical Physics, School of Medicine and Public Health, University of Wisconsin–Madison, and University of Wisconsin Carbone Cancer Center, Madison, Wisconsin, USA
- **Guohua Cao,** Department of Physics and Astronomy, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina, USA
- **Shelton D. Caruthers,** Department of Medicine, Washington University Medical School, St. Louis, Missouri, and Philips Healthcare, Andover, Massachusetts, USA
- **Bo Chen,** Center for Molecular and Functional Imaging, Department of Radiology and Biomedical Imaging, University of California, San Francisco, California, USA
- **Jingyi Chen,** Department of Biomedical Engineering, Washington University, St. Louis, Missouri, USA
- Xiaoyuan Chen, Molecular Imaging Program at Stanford and Bio-X Program, Department of Radiology, Stanford University School of Medicine, Stanford, California, and Laboratory for Molecular Imaging and Nanomedicine, National Institute of Biomedical Imaging and Bioengineering, National Institutes of Health, Bethesda, Maryland, USA
- Claire M. Cobley, Department of Biomedical Engineering, Washington University, St. Louis, Missouri, USA

- Ian R. Corbin, Department of Medical Biophysics, University of Toronto, Toronto, Ontario, and Division of Biophysics and Bioimaging, Ontario Cancer Institute, Toronto, Ontario, Canada
- **Tillmann Cyrus,** Department of Medicine, Washington University Medical School, St. Louis, Missouri, USA
- **Nirupama Deshpande,** Department of Radiology and Molecular Imaging Program at Stanford, Stanford University School of Medicine, Stanford, California, USA
- **Parmeswaran Diagaradjane,** Department of Radiation Oncology, University of Texas M.D. Anderson Cancer Center, Houston, Texas, USA
- **Trevor Douglas,** Department of Chemistry and Biochemistry and Department of Plant Science, Center for Bio-inspired Nanomaterials, Montana State University, Bozeman, Montana, USA
- Luce Vander Elst, Department of General, Organic and Biomedical Chemistry, NMR and Molecular Imaging Laboratory, University of Mons, Mons, Belgium
- **Stanislav Emelianov,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- **Omid C. Farokhzad,** Department of Anesthesiology, Brigham and Women's Hospital, Harvard Medical School, Boston, Massachusetts, USA
- **Fátima Fernández,** Applied Molecular Receptors Group, CIBER de Bioingeniería, Biomateriales y Nanotecnología, IQAC-CSIC, Barcelona, Spain
- **Jinhao Gao,** Molecular Imaging Program at Stanford and Bio-X Program, Department of Radiology, Stanford University School of Medicine, Stanford, California, USA
- Ting Gao, Tyco Electronics Corporation, Menlo Park, California, USA
- **Rui Hao,** Department of Advanced Materials and Nanotechnology, College of Engineering, Peking University, Beijing, China
- **Sven H. Hausner,** Department of Biomedical Engineering, University of California–Davis, Davis, California, USA
- **Kimberly Homan,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- **Hao Hong,** Departments of Radiology and Medical Physics, School of Medicine and Public Health, University of Wisconsin–Madison, Madison, Wisconsin, USA
- **Yanglong Hou,** Department of Advanced Materials and Nanotechnology, College of Engineering, Peking University, Beijing, China
- **Taeghwan Hyeon,** National Creative Research Initiative Center for Oxide Nanocrystalline Materials, and School of Chemical and Biological Engineering, Seoul National University, Seoul, South Korea
- Sangyong Jon, School of Life Sciences, Gwangju Institute of Science and Technology, Gwangju, South Korea

- **Ella Fung Jones,** Center for Molecular and Functional Imaging, Department of Radiology and Biomedical Imaging, University of California, San Francisco, California, USA
- **Kwangmeyung Kim,** Biomedical Research Center, Korea Institute of Science and Technology, Seoul, South Korea
- **Nagesh Kolishetti**, Department of Anesthesiology, Brigham and Women's Hospital, Harvard Medical School, Boston, Massachusetts, USA
- **Alan Koretsky,** Laboratory of Functional and Molecular Imaging, National Institute of Neurological Disorders and Stroke, National Institutes of Health, Bethesda, Maryland, USA
- **Sunil Krishnan,** Department of Radiation Oncology, University of Texas M.D. Anderson Cancer Center, Houston, Texas, USA
- **Ick Chan Kwon**, Biomedical Research Center, Korea Institute of Science and Technology, Seoul, South Korea
- **Gregory M. Lanza**, Department of Medicine, Washington University Medical School, St. Louis, Missouri, USA
- **Timothy Larson,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- **Sophie Laurent,** Department of General, Organic and Biomedical Chemistry, NMR and Molecular Imaging Laboratory, University of Mons, Mons, Belgium
- Seulki Lee, Biomedical Research Center, Korea Institute of Science and Technology, Seoul, South Korea
- **Yueh Z. Lee,** Department of Physics and Astronomy and Department of Radiology, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina, USA
- **Alexei P. Leonov,** Department of Chemistry, Purdue University, West Lafayette, Indiana, USA
- **Chun Li,** Department of Experimental Diagnostic Imaging, University of Texas M.D. Anderson Cancer, Houston, Texas, USA
- Lars Liepold, Department of Chemistry and Biochemistry and Department of Plant Sciences, Center for Bio-Inspired Nanomaterials, Montana State University, Bozeman, Montana, USA
- Pingyu Liu, Palo Alto Unified School District, Palo Alto, California, USA
- **Zhuang Liu**, Institute of Functional Nano & Soft Materials, Soochow University, Suzhou, Jiangsu, China
- **Jianping Lu,** Department of Physics and Astronomy, Curriculum in Applied Sciences and Engineering, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina, USA
- Wei Lu, Department of Experimental Diagnostic Imaging, University of Texas M. D. Anderson Cancer Center, Houston, Texas, USA

- **Srivalleesha Mallidi,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- M.-Pilar Marco, Applied Molecular Receptors Group, CIBER de Bioingeniería, Biomateriales y Nanotecnología, IQAC-CSIC, Barcelona, Spain
- **Jason R. McCarthy,** Center for Molecular Imaging Research, Harvard Medical School and Massachusetts General Hospital, Charlestown, Massachusetts, USA
- **Mohammad Mehrmohammadi**, Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- Marites P. Melancon, Department of Experimental Diagnostic Imaging, University of Texas M. D. Anderson Cancer Center, Houston, Texas, USA
- Mark Michalski, Molecular Imaging Program at Stanford and Bio-X Program, Department of Radiology, Stanford University School of Medicine, Stanford, California, USA
- **R. D. K. Misra**, Center for Structural and Functional Materials, University of Louisiana at Lafayette, Lafayette, Louisiana, USA
- **Pranshu Mohindra**, Department of Radiation Oncology, University of Texas M.D. Anderson Cancer Center, Houston, Texas, USA
- **Robert N. Muller,** Department of General, Organic and Biomedical Chemistry, NMR and Molecular Imaging Laboratory, University of Mons, Mons, Belgium
- **Hyon Bin Na,** National Creative Research Initiative Center for Oxide Nanocrystalline Materials, and School of Chemical and Biological Engineering, Seoul National University, Seoul, South Korea
- **Arutselvan Natarajan,** Department of Radiology and Molecular Imaging Program at Stanford, Stanford University School of Medicine, Stanford, California, USA
- **Kenneth Ng,** Institute of Biomaterials and Biomedical Engineering, University of Toronto, Toronto, Ontario, Canada
- **P. Perriat,** Groupe d'Etudes de Métallurgie Physique et de Physique des Matériaox, Université Claude Bernard, Lyon, France
- **David Pham,** Center for Molecular and Functional Imaging, Department of Radiology and Biomedical Imaging, University of California, San Francisco, California, USA
- Eric M. Pridgen, Department of Chemical Engineering, Massachusetts Institute of Technology, Cambridge, Massachusetts, USA
- **Min Qu,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- C. Rivière, Laboratoire de Physique de la Matière Condensée et Nanostructures, Université de Lyon, Lyon, France
- **Raffaella Rossin,** Department of Biomolecular Engineering, Philips Research Europe, Eindhoven, The Netherlands
- **S. Roux**, Laboratoire de Physico-Chimie des Matériaux Luminescents, Université de Lyon, Lyon, France

- **J.-Pablo Salvador,** Applied Molecular Receptors Group, CIBER de Bioingeniería, Biomateriales y Nanotecnología, IQAC-CSIC, Barcelona, Spain
- **Nuria Sanvicens,** Applied Molecular Receptors Group, CIBER de Bioingeniería, Biomateriales y Nanotecnología, IQAC-CSIC, Barcelona, Spain
- **Anne H. Schmieder,** Department of Medicine, Washington University Medical School, St. Louis, Missouri, USA
- **Konstantin Sokolov,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, and Department of Medical Physics, University of Texas M.D. Anderson Cancer Center, Houston, Texas, USA
- **Michael Strano,** Department of Chemical Engineering, Massachusetts Institute of Technology, Cambridge, Massachusetts, USA
- **O. Tillement,** Laboratoire de Physico-Chimie des Matériaux Luminescents, Université de Lyon, Lyon, France
- Masaki Uchida, Department of Chemistry and Biochemistry and Department of Plant Science, Center for Bio-Inspired Nanomaterials, Montana State University, Bozeman, Montana, USA
- **Bo Wang,** Department of Biomedical Engineering, University of Texas at Austin, Austin, Texas, USA
- Alexander Wei, Department of Chemistry, Purdue University, West Lafayette, Indiana, USA
- Qingshan Wei, Department of Chemistry, Purdue University, West Lafayette, Indiana, USA
- **Ralph Weissleder,** Center for Molecular Imaging Research, Harvard Medical School and Massachusetts General Hospital, Charlestown, Massachusetts, USA
- **Samuel A. Wickline,** Department of Medicine, Washington University Medical School, St. Louis, Missouri, USA
- **Jurgen K. Willmann,** Department of Radiology and Molecular Imaging Program at Stanford, Stanford University School of Medicine, Stanford, California, USA
- **Patrick M. Winter,** Department of Medicine, Washington University Medical School, St. Louis, Missouri, USA
- **Younan Xia,** Department of Biomedical Engineering, Washington University, St. Louis, Missouri, USA
- Jin Xie, Molecular Imaging Program at Stanford and Bio-X Program, Department of Radiology, Stanford University School of Medicine, Stanford, California, and Laboratory for Molecular Imaging and Nanomedicine, National Institute of Biomedical Imaging and Bioengineering, National Institutes of Health, Bethesda, Maryland, USA
- Lei Xing, Department of Radiation Oncology, Stanford University School of Medicine, Stanford, California, USA

- **Yun Xing,** Department of Material Science and Engineering, University of Dayton, Dayton, Ohio, USA
- Bing Xu, Department of Chemistry, Brandeis University, Waltham, Massachusetts, USA
- Mark Young, Department of Chemistry and Biochemistry and Department of Plant Science, Center for Bio-Inspired Nanomaterials, Montana State University, Bozeman, Montana, USA
- Mi Kyung Yu, School of Life Sciences, Gwangju Institute of Science and Technology, Gwangju, South Korea
- **Gary Zabow,** Laboratory of Functional and Molecular Imaging, National Institute of Neurological Disorders and Stroke, National Institutes of Health, Bethesda, Maryland, USA
- **Jingqing Zhang,** Department of Chemical Engineering, Massachusetts Institute of Technology, Cambridge, Massachusetts, USA
- **Ling Zhang,** Center for Molecular and Functional Imaging, Department of Radiology and Biomedical Imaging, University of California, San Francisco, California, USA
- **Gang Zheng,** Department of Medical Biophysics and Institute of Biomaterials and Biomedical Engineering, University of Toronto, Toronto, Ontario, and Division of Biophysics and Bioimaging, Ontario Cancer Institute, Toronto, Ontario, Canada
- Hu Zhou, Community Cancer Center of Roseburg, Roseburg, Oregon, USA
- Otto Zhou, Department of Physics and Astronomy, Curriculum in Applied Sciences and Engineering, and Lineberger Comprehensive Cancer Center, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina, USA

#### **CONTENTS**

Preface	ix
Acknowledgments	xi
Contributors	xiii
PART I BASICS OF MOLECULAR IMAGING AND NANOBIOTECHNOLOGY	
1. Basic Principles of Molecular Imaging Sven H. Hausner	3
2. Synthesis of Nanomaterials as a Platform for Molecular In Jinhao Gao, Jin Xie, Bing Xu, and Xiaoyuan Chen	maging 25
3. Nanoparticle Surface Modification and Bioconjugation Jin Xie, Jinhao Gao, Mark Michalski, and Xiaoyuan Chen	47
4. Biodistribution and Pharmacokinetics of Nanoprobes Nagesh Kolishetti, Frank Alexis, Eric M. Pridgen, and Omid C. Fard	75 okhzad
PART II NANOPARTICLES FOR SINGLE MODALITY MOLECULAR IMAGING	
5. Computed Tomography as a Tool for Anatomical and Mo Pingyu Liu, Hu Zhou, and Lei Xing	lecular Imaging 107
6. Carbon Nanotube X-Ray for Dynamic Micro-CT Imaging Animal Models Otto Zhou, Guohua Cao, Yueh Z. Lee, and Jianping Lu	g of Small 139
7. Quantum Dots for <i>In Vivo</i> Molecular Imaging  Yun Xing	159
8. Biopolymer, Dendrimer, and Liposome Nanoplatforms for Molecular Imaging  David Pham, Ling Zhang, Bo Chen, and Ella Fung Jones	r Optical

9.	Nanoplatforms for Raman Molecular Imaging in Biological Systems Zhuang Liu	197
10.	Single-Walled Carbon Nanotube Near-Infrared Fluorescent Sensors for Biological Systems Jingqing Zhang and Michael S. Strano	217
11.	Microparticle- and Nanoparticle-Based Contrast-Enhanced Ultrasound Imaging Nirupama Deshpande and Jürgen K. Willmann	233
12.	Ultrasound-Based Molecular Imaging Using Nanoagents Srivalleesha Mallidi, Mohammad Mehrmohammadi, Kimberly Homan, Bo Wang, Min Qu, Timothy Larson, Konstantin Sokolov, and Stanislav Emelianov	263
13.	MRI Contrast Agents Based on Inorganic Nanoparticles Hyon Bin Na and Taeghwan Hyeon	279
14.	Cellular Magnetic Labeling with Iron Oxide Nanoparticles Sébastien Boutry, Sophie Laurent, Luce Vander Elst, and Robert N. Muller	309
15.	Nanoparticles Containing Rare Earth Ions: A Tunable Tool for MRI C. Rivière, S. Roux, R. Bazzi, JL. Bridot, C. Billotey, P. Perriat, and O. Tillement	333
16.	Microfabricated Multispectral MRI Contrast Agents Gary Zabow and Alan Koretsky	375
17.	Radiolabeled Nanoplatforms: Imaging Hot Bullets Hitting Their Target Raffaella Rossin	399
PA]	RT III NANOPARTICLE PLATFORMS AS MULTIMODALITY IMAGING AND THERAPY AGENTS	
18.	<b>Lipoprotein-Based Nanoplatforms for Cancer Molecular Imaging</b> <i>Ian R. Corbin, Kenneth Ng, and Gang Zheng</i>	433
19.	Protein Cages as Multimode Imaging Agents  Masaki Uchida, Lars Liepold, Mark Young, and Trevor Douglas	463
20.	Biomedical Applications of Single-Walled Carbon Nanotubes Weibo Cai, Ting Gao, and Hao Hong	481
21.	Multifunctional Nanoparticles for Multimodal Molecular Imaging Yanglong Hou and Rui Hao	529
22.	Multifunctional Nanoparticles for Cancer Theragnosis Seulki Lee, Ick Chan Kwon, and Kwangmeyung Kim	541

CONTENTS	vii
23. Nanoparticles for Combined Cancer Imaging and Therapy Vaishali Bagalkot, Mi Kyung Yu, and Sangyong Jon	565
24. Multimodal Imaging and Therapy with Magnetofluorescent Nanoparticles  Jason R. McCarthy and Ralph Weissleder	593
25. Gold Nanocages: A Multifunctional Platform for Molecular Optical Imaging and Photothermal Treatment Leslie Au, Claire M. Cobley, Jingyi Chen, and Younan Xia	615
26. Theranostic Applications of Gold Nanoparticles in Cancer Parmeswaran Diagaradjane, Pranshu Mohindra, and Sunil Krishnan	639
27. Gold Nanorods as Theranostic Agents Alexander Wei, Qingshan Wei, and Alexei P. Leonov	659
28. Theranostic Applications of Gold Core–Shell Structured Nanoparticles Wei Lu, Marites P. Melancon, and Chun Li	683
29. Magnetic Nanoparticle Carrier for Targeted Drug Delivery: Perspective, Outlook, and Design R. D. K. Misra	709
30. Perfluorocarbon Nanoparticles: A Multidimensional Platform for Targeted Image-Guided Drug Delivery  Gregory M. Lanza, Shelton D. Caruthers, Anne H. Schmieder, Patrick M. Winter, Tillmann Cyrus, and Samuel A. Wickline	725
31. Radioimmunonanoparticles for Cancer Imaging and Therapy  Arutselvan Natarajan	755
PART IV TRANSLATIONAL NANOMEDICINE	
32. Current Status and Future Prospects for Nanoparticle-Based Technology in Human Medicine Nuria Sanvicens, Fátima Fernández, JPablo Salvador, and MPilar Marco	783
Index	815

### BASICS OF MOLECULAR IMAGING AND NANOBIOTECHNOLOGY