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Gerd Keiser

# Biophotonics

Concepts to Applications



Springer

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*To  
Ching-yun, Nishla, Kai, and Neyla  
for their loving patience and encouragement*

# Preface

The discipline of biophotonics or biomedical optics has undergone a fascinating journey in the past several decades and it is still growing rapidly. As the name *biophotonics* implies, the application of this field to biological disciplines is based on the wide range of photonics technologies, which involve the generation, detection, and control of photons for enabling functions such as transferring or processing information, analyzing material characteristics, sensing or measuring changes in physical parameters, and modifying material characteristics. Basically biophotonics deals with the interaction between light and biological material. The resulting interactions can be used in almost all biomedical areas for basic life science research and for biomedical diagnosis, therapy, monitoring, imaging, and surgery.

Owing to the importance of biophotonics to all aspects of human health, it is essential that a wide range of biomedical researchers, healthcare professionals, clinical technicians, and biomedical engineers have a good understanding of biophotonics and its applications. To address the attainment and implementation of these skills, this book provides the basic material for a one-semester entry-level course in the fundamentals and applications of biophotonics technology for senior or postgraduate students. It also will serve well as a working reference or as a short-course textbook for biomedical researchers, practicing physicians, healthcare professionals, clinical technicians, and biomedical engineers and technicians dealing with the design, development, and application of photonics components and instrumentation to biophotonics issues.

In Chap. 1–5 the sequence of topics takes the reader systematically from the underlying principles of light and biology, through the fundamentals of optical fiber light guiding, and then through optical sources and photodetection methods. Next, the topics in Chap. 6–10 address the concepts of light–tissue interactions, various optical probes and photonics sensing techniques, the principles of microscopy and spectroscopy, and biophotonics imaging modalities. The final chapter discusses advanced techniques and developments such as optical trapping, miniaturized instruments, single nanoparticles detection, and optogenetics procedures. By mastering these fundamental topics the reader will be prepared not only to contribute to

current biomedical photonics disciplines, but also to understand quickly any further technology developments for future enhanced biophotonics developments.

The background required to study the book is that of typical senior-level science and engineering students. This background includes introductory biology and chemistry, calculus, and basic concepts of electromagnetism and optics as presented in a freshman physics course. To assist readers in learning the material and applying it to practical situations, 104 worked examples are given throughout the text. A collection of 129 homework problems is included to test the readers' comprehension of the material covered, and to extend and elucidate the text.

The articles and books cited as references in the text were selected from among numerous texts and thousands of papers in the literature relating to the material covered in each chapter. Because biophotonics brings together research, development, and application efforts from many different scientific, medical, and engineering disciplines, these references are a small sample of the major contributions to biophotonics. A number of these references are review papers and provide a starting point for delving deeper into any given topic.

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Newton, USA

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# Contents

<b>1</b>	<b>Overview of Biophotonics . . . . .</b>	<b>1</b>
1.1	What Is Biophotonics? . . . . .	2
1.2	Diverse Applications . . . . .	6
1.3	Biophotonics Spectral Windows. . . . .	8
1.4	Light Absorption . . . . .	11
1.5	Signal Attenuation . . . . .	12
1.6	Structures of Biological Cells and Tissues. . . . .	14
1.6.1	Macromolecules. . . . .	15
1.6.2	Biological Cells . . . . .	17
1.6.3	Biological Tissues and Organs. . . . .	19
1.7	Summary . . . . .	20
1.8	Problems. . . . .	21
	References . . . . .	22
<b>2</b>	<b>Basic Principles of Light . . . . .</b>	<b>25</b>
2.1	Lightwave Characteristics . . . . .	26
2.1.1	Monochromatic Waves . . . . .	27
2.1.2	Pulsed Plane Waves . . . . .	29
2.2	Polarization . . . . .	29
2.2.1	Linear Polarization . . . . .	31
2.2.2	Elliptical and Circular Polarization . . . . .	33
2.3	Quantized Photon Energy and Momentum . . . . .	35
2.4	Reflection and Refraction . . . . .	37
2.4.1	Snells' Law . . . . .	38
2.4.2	The Fresnel Equations . . . . .	40
2.4.3	Diffuse Reflection . . . . .	43
2.5	Interference . . . . .	44
2.6	Optical Coherence . . . . .	45
2.7	Lightwave-Molecular Dipole Interaction . . . . .	46

2.8	Summary . . . . .	48
2.9	Problems. . . . .	49
	References . . . . .	51
<b>3</b>	<b>Optical Fibers for Biophotonics Applications. . . . .</b>	<b>53</b>
3.1	Light Guiding Principles in Conventional Optical Fibers. . . . .	54
3.1.1	Ray Optics Concepts . . . . .	57
3.1.2	Modal Concepts. . . . .	60
3.1.3	Mode Field Diameter . . . . .	63
3.2	Graded-Index Optical Fibers . . . . .	64
3.2.1	Core Index Structure . . . . .	64
3.2.2	Graded-Index Numerical Aperture . . . . .	65
3.2.3	Cutoff Condition in Graded-Index Fibers. . . . .	66
3.3	Performance Characteristics of Generic Optical Fibers . . . . .	66
3.3.1	Attenuation Versus Wavelength . . . . .	66
3.3.2	Bend-Loss Insensitivity. . . . .	67
3.3.3	Mechanical Properties. . . . .	67
3.3.4	Optical Power-Handling Capability. . . . .	68
3.4	Conventional Solid-Core Fibers . . . . .	68
3.5	Specialty Solid-Core Fibers . . . . .	69
3.5.1	Photosensitive Optical Fiber . . . . .	70
3.5.2	Fibers Resistant to UV-Induced Darkening . . . . .	71
3.5.3	Bend Insensitive Fiber . . . . .	72
3.5.4	Polarization-Maintaining Fiber. . . . .	73
3.6	Double-Clad Fibers . . . . .	75
3.7	Hard-Clad Silica Fibers. . . . .	76
3.8	Coated Hollow-Core Fibers. . . . .	77
3.9	Photonic Crystal Fibers. . . . .	78
3.10	Plastic Fibers. . . . .	79
3.11	Side-Emitting or Glowing Fibers . . . . .	80
3.12	Middle-Infrared Fibers . . . . .	81
3.13	Optical Fiber Bundles. . . . .	83
3.14	Summary . . . . .	84
3.15	Problems. . . . .	85
	References . . . . .	87
<b>4</b>	<b>Fundamentals of Light Sources . . . . .</b>	<b>91</b>
4.1	Radiometry . . . . .	93
4.1.1	Optical Flux and Power . . . . .	94
4.1.2	Irradiance or Exposure Rate . . . . .	94
4.1.3	Radiant Intensity . . . . .	95
4.1.4	Radiant Exposure or Radiant Fluence . . . . .	96
4.1.5	Radiance. . . . .	96
4.2	Arc Lamps . . . . .	97

4.3	Light-Emitting Diodes . . . . .	98
4.3.1	LED Operation and Structures . . . . .	99
4.3.2	LED Wavelengths and Device Uses . . . . .	101
4.3.3	Modulation of an LED . . . . .	102
4.4	Lasers for Biophotonics . . . . .	105
4.4.1	Basic Laser Actions . . . . .	106
4.4.2	Laser Diodes . . . . .	108
4.4.3	Solid-State Lasers . . . . .	111
4.4.4	Gas Lasers . . . . .	113
4.4.5	Optical Fiber Lasers . . . . .	114
4.5	Superluminescent Diodes . . . . .	114
4.6	Summary . . . . .	115
4.7	Problems. . . . .	115
	References . . . . .	117
<b>5</b>	<b>Fundamentals of Optical Detectors . . . . .</b>	<b>119</b>
5.1	The <i>pin</i> Photodetector. . . . .	120
5.2	Avalanche Photodiodes. . . . .	127
5.3	Photodiode Noises . . . . .	128
5.3.1	Signal-to-Noise Ratio . . . . .	128
5.3.2	Noise Sources . . . . .	129
5.3.3	Noise-Equivalent Power and Detectivity . . . . .	131
5.3.4	Comparisons of Photodiodes . . . . .	132
5.4	Multichannel Detectors . . . . .	133
5.4.1	CCD Array Technology . . . . .	133
5.4.2	CMOS Array Technology . . . . .	134
5.5	Photomultiplier Tubes. . . . .	136
5.6	Optical Filters . . . . .	138
5.7	Optical Couplers and Optical Circulators. . . . .	141
5.8	Summary . . . . .	143
5.9	Problems. . . . .	143
	References . . . . .	145
<b>6</b>	<b>Light-Tissue Interactions . . . . .</b>	<b>147</b>
6.1	Reflection and Refraction Applications . . . . .	149
6.1.1	Refraction in Ophthalmology. . . . .	150
6.1.2	Specular Reflection . . . . .	150
6.1.3	Diffuse Reflection . . . . .	153
6.2	Absorption . . . . .	154
6.2.1	Absorption Characteristics. . . . .	154
6.2.2	Absorption in Biological Tissues . . . . .	157
6.3	Scattering . . . . .	160
6.3.1	Elastic Scattering . . . . .	162
6.3.2	Rayleigh Scattering . . . . .	165
6.3.3	Anisotropy Factor . . . . .	167
6.3.4	Inelastic (Raman) Scattering . . . . .	168

6.4	Scattering with Absorption . . . . .	169
6.5	Light-Tissue Interaction Mechanisms . . . . .	170
6.5.1	Photobiomodulation . . . . .	174
6.5.2	Photochemical Interaction . . . . .	177
6.5.3	Thermal Interaction . . . . .	179
6.5.4	Photoablation . . . . .	183
6.5.5	Plasma-Induced Photoablation . . . . .	186
6.5.6	Photodisruption . . . . .	187
6.6	Formation of Speckles . . . . .	188
6.7	Fluorescence Basics . . . . .	189
6.8	Summary . . . . .	190
6.9	Problems. . . . .	192
	References . . . . .	193
<b>7</b>	<b>Optical Probes and Biosensors . . . . .</b>	<b>197</b>
7.1	Overview of Biosensors and Probes . . . . .	198
7.2	Optical Fiber Probe Configurations. . . . .	200
7.3	Optical Fiber Tip Geometries . . . . .	206
7.4	Optical Sensors . . . . .	209
7.4.1	Biorecognition Optical Fiber Sensors . . . . .	209
7.4.2	ELISA . . . . .	210
7.4.3	Sensors Based on Optical Fiber Movements . . . . .	211
7.4.4	Microbending Fiber Sensors . . . . .	214
7.5	Interferometric Sensors . . . . .	216
7.5.1	Mach-Zehnder Interferometer . . . . .	217
7.5.2	Michelson Interferometer . . . . .	219
7.5.3	Sagnac Interferometer . . . . .	220
7.6	Photonic Crystal Fiber Biosensors . . . . .	221
7.6.1	Interferometry Sensing Methods . . . . .	221
7.6.2	Liquid Infiltration Sensor . . . . .	222
7.7	Fiber Bragg Grating Sensors . . . . .	223
7.7.1	Smart-Bed FBG System . . . . .	223
7.7.2	Distributed FBG-Based Catheter Sensor . . . . .	224
7.8	Surface Plasmon Resonance Biosensors . . . . .	225
7.9	Optical Fiber Nanoprobes . . . . .	226
7.10	Summary . . . . .	227
7.11	Problems. . . . .	227
	References . . . . .	229
<b>8</b>	<b>Microscopy . . . . .</b>	<b>233</b>
8.1	Concepts and Principles of Microscopy . . . . .	234
8.1.1	Viewing and Illumination Techniques . . . . .	234
8.1.2	Observation Methods . . . . .	237
8.1.3	Numerical Aperture . . . . .	240

8.1.4	Field of View . . . . .	241
8.1.5	Depth of Field . . . . .	242
8.2	Resolution and Diffraction Limit . . . . .	244
8.3	Confocal Microscopy . . . . .	247
8.4	Fluorescence Microscopy . . . . .	249
8.5	Multiphoton Microscopy . . . . .	251
8.6	Raman Microscopy . . . . .	253
8.7	Light Sheet Fluorescence Microscopy . . . . .	254
8.8	Super-Resolution Fluorescence Microscopy . . . . .	255
8.9	Summary . . . . .	255
8.10	Problems . . . . .	256
	References . . . . .	257
<b>9</b>	<b>Spectroscopic Methodologies</b> . . . . .	259
9.1	Fluorescence Spectroscopy . . . . .	261
9.2	FRET/FLIM . . . . .	263
9.2.1	Förster Resonance Energy Transfer . . . . .	263
9.2.2	Fluorescence Lifetime Imaging Microscopy . . . . .	266
9.3	Fluorescence Correlation Spectroscopy . . . . .	269
9.4	Elastic Scattering Spectroscopy . . . . .	273
9.5	Diffuse Correlation Spectroscopy . . . . .	275
9.6	Raman Spectroscopy . . . . .	276
9.7	Surface Enhanced Raman Scattering Spectroscopy . . . . .	279
9.8	Coherent Anti-stokes Raman Scattering Spectroscopy . . . . .	280
9.9	Stimulated Raman Scattering Spectroscopy . . . . .	282
9.10	Photon Correlation Spectroscopy . . . . .	282
9.11	Fourier Transform Infrared Spectroscopy . . . . .	284
9.12	Brillouin Scattering Spectroscopy . . . . .	285
9.13	Summary . . . . .	286
9.14	Problems . . . . .	286
	References . . . . .	288
<b>10</b>	<b>Optical Imaging Procedures</b> . . . . .	291
10.1	Optical Coherence Tomography . . . . .	292
10.1.1	Time Domain OCT . . . . .	294
10.1.2	Spectral Domain OCT . . . . .	301
10.1.3	Swept Source OCT . . . . .	301
10.2	Endoscopy . . . . .	303
10.2.1	Basic Endoscopy . . . . .	304
10.2.2	Minimally Invasive Surgery . . . . .	305
10.2.3	Tethered Capsule Endomicroscopy . . . . .	307
10.3	Laser Speckle Imaging . . . . .	307
10.4	Optical Coherence Elastography . . . . .	310
10.5	Photoacoustic Tomography . . . . .	312
10.6	Hyperspectral Imaging . . . . .	316

10.7	Summary . . . . .	317
10.8	Problems. . . . .	317
	References . . . . .	319
<b>11</b>	<b>Biophotonics Technology Applications</b> . . . . .	323
11.1	Optical Manipulation . . . . .	324
11.2	Miniaturized Analyses Tools . . . . .	329
11.2.1	Lab-on-a-Chip Technology . . . . .	329
11.2.2	Lab-on-Fiber Concept. . . . .	331
11.3	Microscope in a Needle . . . . .	332
11.4	Single Nanoparticle Detection . . . . .	333
11.5	Neurophotonics . . . . .	334
11.6	Summary . . . . .	335
11.7	Problems. . . . .	335
	References . . . . .	336
	<b>Erratum to: Biophotonics</b> . . . . .	E1
	<b>Index</b> . . . . .	339

## About the Author

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# Abbreviations

ADP	Adenosine diphosphate
AMP	Adenosine monophosphate
APD	Avalanche photodiode
ATP	Adenosine triphosphate
BFP	Blue fluorescent protein
CARS	Coherent anti-Stokes Raman spectroscopy
CCD	Charge-coupled device
CFP	Cyan fluorescent protein
CMOS	Complementary metal-oxide-semiconductor
CW	Continuous wave
DCF	Double-clad fiber
DCS	Diffuse correlation spectroscopy
DFB	Distributed feedback (laser)
DIC	Differential interference contrast
DNA	Deoxyribonucleic acid
DOF	Depth of field
DPSS	Diode-pumped solid-state
DRS	Diffuse reflectance spectroscopy
ELISA	Enzyme-linked immunosorbent assay
ESS	Elastic scattering spectroscopy
eV	Electron volt
FBG	Fiber Bragg grating
FCS	Fluorescence correlation spectroscopy
FD-OCT	Fourier domain OCT
FEL	Free-electron laser
FIDA	Fluorescence intensity distribution analysis
FLIM	Fluorescence lifetime imaging microscopy
FN	Field number
FOV	Field of view
FRET	Fluorescence resonance energy transfer
FRET	Förster resonance energy transfer

FTIR	Fourier transform infrared
FWHM	Full-width half-maximum
FWM	Four-wave mixing
GFP	Green fluorescent protein
GRIN	Graded-index
HCPCF	Hollow-core photonic crystal fiber
HCS	Hard-clad silica
HMFG	Heavy metal fluoride glasses
HRCT	High-resolution computed tomography
HSI	Hyperspectral imaging
IR	Infrared
IRIS	Interferometric reflectance imaging sensor
LED	Light-emitting diode
LITT	Laser-induced interstitial thermotherapy
LLLT	Low-level light therapy (photobiomodulation)
LOC	Lab-on-a-chip
LP	Linearly polarized
LPG	Long-period grating
LSFM	Light sheet fluorescence microscopy
LSPR	Localized surface plasmon resonance
LSS	Light scattering spectroscopy
MFD	Mode-field diameter
MI	Michelson interferometer
MMF	Multimode fiber
MOSFET	Metal-oxide-semiconductor field-effect transistor
MRI	Magnetic resonance imaging
MZI	Mach–Zehnder interferometer
NA	Numerical aperture
Nd:YAG	Neodymium:yttrium aluminum garnet
NEP	Noise equivalent power
NIR	Near-infrared
NO	Nitric oxide
OCE	Optical coherence elastography
OCT	Optical coherence tomography
OSA	Optical spectrum analyzer
PALM	Photo activated localization microscopy
PAT	Photoacoustic tomography
PCF	Photonic crystal fiber
PCH	Photon counting histograms
PCS	Photon correlation spectroscopy
PDL	Polarization dependent loss
PDT	Photodynamic therapy
PLA	Percutaneous laser ablation
PMMA	Polymethylmethacrylate
PMT	Photomultiplier tube

POC	Point of care
POF	Plastic optical fiber
POF	Polymer optical fiber
PSF	Point-spread function
QCL	Quantum cascade laser
RET	Resonance energy transfer
RI	Refractive index
RNA	Ribonucleic acid
ROS	Reactive oxygen species
SD-OCT	Spectral domain OCT
SERS	Surface-enhanced Raman scattering
SFG	Sum-frequency generation
SHG	Second-harmonic generation
SM	Single-mode
SMF	Single-mode fiber
SNR	Signal-to-noise ratio
SPR	Surface plasmon resonance
SRG	Stimulated Raman gain
SRI	Surrounding refractive index
SRL	Stimulated Raman loss
SRS	Stimulated Raman scattering
SSIM	Saturated structured illumination microscopy
SS-OCT	Swept source OCT
STED	Stimulated emission depletion microscopy
STORM	Stochastic optical reconstruction microscopy
TD-OCT	Time domain OCT
TE	Transverse electric
TFF	Thin film filter
THG	Third-harmonic generation
TPEF	Two-photon excitation fluorescence
TRT	Thermal relaxation time
UV	Ultraviolet
VCSEL	Vertical cavity surface-emitting laser
WDL	Wavelength dependent loss
YAG	Yttrium aluminum garnet
YFP	Yellow fluorescent protein