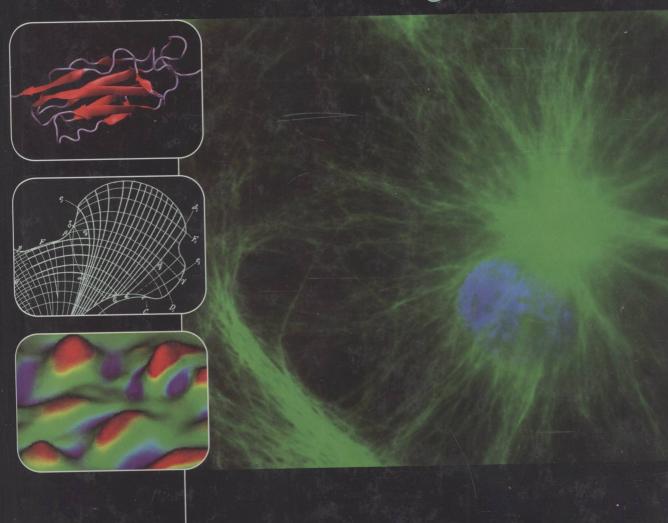
CAMBRIDGE TEXTS IN BIOMEDICAL FNGINEFRING

Introductory Biomechanics

From Cells to Organisms



C. Ross Ethier and Craig A. Simmons

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Introductory Biomechanics

From Cells to Organisms

C. Ross Ethier and Craig A. Simmons University of Toronto, Canada







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Introductory Biomechanics From Cells to Organisms

Introductory Biomechanics is a new, integrated text written specifically for engineering students. It provides a broad overview of this important branch of the rapidly growing field of bioengineering. A wide selection of topics is presented, ranging from the mechanics of single cells to the dynamics of human movement. No prior biological knowledge is assumed and in each chapter, the relevant anatomy and physiology are first described. The biological system is then analyzed from a mechanical viewpoint by reducing it to its essential elements, using the laws of mechanics, and then linking mechanical insights back to biological function. This integrated approach provides students with a deeper understanding of both the mechanics and the biology than that obtained from qualitative study alone. The text is supported by a wealth of illustrations, tables, and examples, a large selection of suitable problems and many current references, making it an essential textbook for any biomechanics course.

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To my family, who make it all worthwhile. c. Ross ethier

To Deborah, and to my parents, who inspired my love of learning. CRAIG A. SIMMONS

About the cover

The cover contains images that together represent the broad scope of modern biomechanics. The figures are as follows:

- Main image: A fluorescent immunohistochemical image of an endothelial cell
 isolated from the surface of a pig aortic heart valve and grown in culture. Within
 the cell, the nucleus is stained blue and vimentin filaments are stained green.
 Vimentin is an intermediate filament protein of the cellular cytoskeleton that
 plays an important role in cellular mechanics.
- Left top: An intermediate stage from a simulation of the forced unfolding of repeats 4 and 5 of chain A of the protein filamin. Filamin is an actin cross-linking protein and therefore plays a role in the biomechanics of the cytoskeleton. The simulation was based on the crystal structure of part of filamin [1], and was carried out in NAMD [2] and visualized using the VMD package [3]. (Image courtesy of Mr. Blake Charlebois.)
- Left middle: A sketch by the Swiss anatomist Hermann von Meyer of the orientation of trabecular bone in the proximal human femur. This sketch was accompanied in the original article by a sketch of the principal stress trajectories in a crane having a shape similar to the femur. Together these sketches are believed to have inspired "Wolff's Law" of bone remodelling. From [4].
- Left lower: The distribution of mass transfer rates from flowing blood to cultured vascular endothelial cells. The contoured quantity (the Sherwood number) was computed by first measuring the topography of the endothelial cells using atomic force microscopy and then solving the convection-diffusion equation in the blood flowing over the cells. Mass transfer from blood to endothelial cells is important in cell-cell signalling. (Image courtesy of Mr. Ji Zhang.)

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Preface

For some years, we have taught an introductory course in biomechanics within the Department of Mechanical and Industrial Engineering at the University of Toronto. We have been unable to find a textbook suitable for the purpose of introducing engineers and others having a "hard science" background to the field of biomechanics. That is not to say that excellent books on biomechanics do not exist; in fact, there are many. However, they are typically at a level that is too advanced for an introductory course, or they cover too limited a subset of topics for purposes of an introductory course.

This book represents an attempt to fill this void. It is not meant to be an extensive treatise on any particular branch of biomechanics, but rather to be an introduction to a wide selection of biomechanics-related topics. Our hope is that it will aid the student in his or her introduction to the fascinating world of bioengineering, and will lead some to pursue the topic in greater detail.

In writing this book, we have assumed that the reader has a background in engineering and mathematics, which includes introductory courses in dynamics, statics, fluid mechanics, thermodynamics, and solid mechanics. No prior knowledge of biology, anatomy, or physiology is assumed, and in fact every section begins with a review of the relevant biological background. Each chapter then emphasizes identification and description of the essential aspects of the related biomechanics problems. Because of the introductory nature of this book, this has led in some cases to a great deal of simplification, but in all instances, we have tried to maintain a firm link to "biological reality."

We wish to thank Professor David F. James, of the Department of Mechanical and Industrial Engineering, University of Toronto. He first developed the introductory course in biomechanical engineering at the University and his course notes provided the inspiration for parts of this book. Professors James E. Moore Jr. and Takami Yamaguchi provided important material for Ch. 1. We have benefited greatly from interactions with our students, who sometimes are the best teachers, and our colleagues and mentors.

We shall be most grateful to students who, upon discovering errors in the text, bring them to our attention.

Contents

About the cover		<i>page</i> xii	
Pr	eface		xv
1	Introduction		1
	1.1	A brief history of biomechanics	3
	1.2	An outline of this book	12
		References	15
2	Cellular biomechanics		18
	2.1	Introduction to eukaryotic cellular architecture	18
	2.2	The cell's energy system	22
	2.3	Overview of the cytoskeleton	23
		2.3.1 Actin filaments	25
		2.3.2 Intermediate filaments	28
		2.3.3 Microtubules	28
	2.4	Cell–matrix interactions	29
	2.5	Methods to measure the mechanical properties of cells	
		and biomolecules	35
		2.5.1 Atomic force microscopy	35
		2.5.2 Optical trapping ("optical tweezers")	41
		2.5.3 Magnetic bead microrheometry	42
		2.5.4 Micropipette aspiration	43
	2.6	Models of cellular biomechanical behavior	53
		2.6.1 Lumped parameter viscoelastic model of the cell	54
		2.6.2 Tensegrity model of the cytoskeleton	60
		2.6.3 Modeling actin filaments as a foam	69
		2.6.4 Computational model of a chondrocyte in its matrix	72
	2.7	Mechanotransduction: how do cells sense and respond to	
		mechanical events?	76
		2.7.1 Mechanoreceptors	76
		2.7.2 Intracellular signal transduction	78
		2.7.2 Callular response to machanical signals	97

	2.8	Techniques for mechanical stimulation of cells	80
		2.8.1 Compressive loading	81
		2.8.2 Stretching	82
		2.8.3 Fluid flow	86
	2.9	Summary of mechanobiological effects on cells in	
		selected tissues	90
		2.9.1 Endothelial cells in the vascular system	91
		2.9.2 Smooth muscle cells in vascular tissue	93
		2.9.3 Chondrocytes in articular cartilage	94
		2.9.4 Osteoblasts and osteocytes in bone	95
	2.10	Problems	99
		References	111
3	Her	modynamics	119
	3.1	Blood rheology	119
		3.1.1 Blood composition	121
		3.1.2 Relationship between blood composition and rheology	124
		3.1.3 Constitutive equation for blood	129
	3.2	Large artery hemodynamics	130
		3.2.1 Physical characteristics of blood flow patterns in vivo	130
		3.2.2 Steady blood flow at low flow rates	133
		3.2.3 Unsteady flow in large vessels	138
	3.3	Blood flow in small vessels	142
		3.3.1 Fahraeus-Lindqvist effect	143
		3.3.2 "Inverse" Fahraeus-Lindqvist effect	146
	3.4	Problems	147
		References	161
4	The	e circulatory system	164
	4.1	Anatomy of the vasculature	164
	4.2	The heart	169
		4.2.1 Gross anatomy of the heart	171
		4.2.2 Qualitative description of cardiac pumping	172
		4.2.3 Cardiac pumping power and ventricular function	175
	4.3	Arterial pulse propagation	179
		4.3.1 Systolic and diastolic pressure	179
		4.3.2 Windkessel model	180
		4.3.3 Arterial wall structure and elasticity	183
		4.3.4 Elastic waves	186
		4.3.5 Pressure–flow relationships: purely oscillatory flow	194

		4.3.6 Pressure-flow relationships: mean flow effects	197
		4.3.7 Pressure–flow relationships: deviations from ideality	198
	4.4	The capillaries	204
		4.4.1 Capillary filtration: the experiments of Landis	207
		4.4.2 Osmotic pressure	209
		4.4.3 Quantitative analysis of capillary leakage	211
	4.5	The veins	212
	4.6	Scaling of hemodynamic variables	213
	4.7	Problems	218
		References	235
5	The	interstitium	240
	5.1	Interstitial fluid flow	240
		5.1.1 Darcy's law	241
		5.1.2 Clearance of edema	242
	5.2	Problems	248
		References	249
6	Ocı	ılar biomechanics	250
	6.1	Ocular anatomy	250
	6.2	Biomechanics of glaucoma	251
		6.2.1 Tonometry	252
		6.2.2 Drainage of aqueous humor in normal and glaucomatous eyes	257
		6.2.3 Aqueous humor circulation in the anterior chamber	263
		6.2.4 Optic nerve head biomechanics	264
	6.3	Ocular blood flow	271
	6.4	Problems	274
		References	276
7	The	respiratory system	282
	7.1	Gross anatomy	282
		7.1.1 The conducting airways and pulmonary vasculature	282
		7.1.2 Associated structures	285
	7.2	Biomechanics of breathing	287
	7.3	Lung elasticity and surface tension effects	288
	7.4	Mass transfer	294
		7.4.1 Blood-side acinar mass transfer	295
		7.4.2 Air-side acinar mass transfer	307
		7.4.3 Whole lung mass transfer	308
	7.5	Particle transport in the lung	313

	7.6	Problems	316
		References	329
8	Musc	eles and movement	332
	8.1	Skeletal muscle morphology and physiology	333
		8.1.1 Isotonic versus isometric contraction	339
	8.2	Muscle constitutive modeling	343
	8.3	Whole muscle mechanics	351
		8.3.1 Parallel versus pinnate muscle types	351
	8.4	Muscle/bone interactions	353
		8.4.1 Foreleg motion in two species	353
		8.4.2 Flexion of the elbow	355
		8.4.3 Biomechanics of the knee	365
	8.5	Problems	369
		References	376
9	Skel	etal biomechanics	379
	9.1	Introduction to bone	379
	9.2	Composition and structure of bone	382
		9.2.1 Cortical bone	384
		9.2.2 Trabecular bone	384
	9.3	Biomechanical properties of cortical and trabecular bone	387
		9.3.1 Cortical bone mechanics	388
		9.3.2 Trabecular bone mechanics	389
		9.3.3 Trabecular bone mechanics: density dependence	390
		9.3.4 Trabecular bone mechanics: unit cell models	393
	9.4	Bone fracture and failure mechanics	395
		9.4.1 Fast fracture	397
		9.4.2 Fatigue fracture	403
	9.5	Functional adaptation and mechanobiology	407
	9.6	The design of bone	409
	9.7	Introduction to soft connective tissues	411
	9.8	Structure of collagen	412
	9.9	Structure of ligament, tendon, and cartilage	414
		9.9.1 Ligament	414
		9.9.2 Tendon	416 418
	9.10	9.9.3 Cartilage Biomechanical properties of ligament, tendon, and cartilage	419
	7.10	9.10.1 Structural properties	420
		9.10.2 Material properties	423
		9.10.3 Material properties: tension	425

		9.10.4 Material properties: compression	426
		9.10.5 Material properties: viscoelasticity	428
		9.10.6 Material properties: biphasic mixture theory of cartilage	435
	9.11	Problems	436
		References	439
10	Terre	estrial locomotion	444
	10.1	Jumping	444
		10.1.1 Standing jump	444
		10.1.2 Running jumps	448
	10.2	Description of walking and running	451
		10.2.1 Walking	451
		10.2.2 Running	459
	10.3	Gait analysis	461
		10.3.1 Kinematics	463
		10.3.2 Anthropometry	468
		10.3.3 Kinetics	471
	10.4	Problems	480
		References	487
Арр	endix	The electrocardiogram	489
Inde		<u> </u>	498

Color plate section between pages 118 and 119

1 Introduction

Biomechanics is a branch of the field of bioengineering, which we define as the application of engineering principles to biological systems. Most bioengineering is applied to humans, and in this book the primary emphasis will be on *Homo sapiens*. The bioengineer seeks to understand basic physiological processes, to improve human health via applied problem solving, or both. This is a difficult task, since the workings of the body are formidably complex. Despite this difficulty, the bioengineer's contribution can be substantial, and the rewards for success far outweigh the difficulties of the task.

Biomechanics is the study of how physical forces interact with living systems. If you are not familiar with biomechanics, this might strike you as a somewhat esoteric topic, and you may even ask yourself the question: Why does biomechanics matter? It turns out that biomechanics is far from esoteric and plays an important role in diverse areas of growth, development, tissue remodeling and homeostasis. Further, biomechanics plays a central role in the pathogenesis of some diseases, and in the treatment of these diseases. Let us give a few specific examples:

- How do your bones "know" how big and strong to be so that they can support your weight and deal with the loads imposed on them? Evidence shows that the growth of bone is driven by mechanical stimuli [1]. More specifically, mechanical stresses and strains induce bone cells (osteoblasts and osteoclasts) to add or remove bone just where it is needed. Because of the obvious mechanical function played by bone, it makes good sense to use mechanical stress as the feedback signal for bone growth and remodeling. But biomechanics also plays a "hidden" regulatory role in other growth processes, as the next example will show.
- How do our arteries "know" how big to be so that they can deliver just the right amount of blood to their distal capillary beds? There is good evidence that this is determined in large part by the mechanical stress exerted on the artery wall by flowing blood. Endothelial cells lining the inner arterial surface sense this shear stress and send signals to cells deeper in the artery wall to direct the remodeling of the artery so as to enlarge or reduce its caliber [2].

- What about biomechanics in everyday life? Probably the most obvious application of biomechanics is in locomotion (walking, running, jumping), where our muscles generate forces that are transferred to the ground by bones and soft connective tissue. This is so commonplace that we rarely think about it, yet the biomechanics of locomotion is remarkably complex (watch a baby learning to walk!) and still incompletely understood.
- Locomotion happens on many scales, from whole organisms all the way down to individual cells. Unicellular organisms must be able to move so as to gather nutrients, and they have evolved a variety of clever strategies to accomplish this task [3]. In multicellular organisms, the ability of single cells to move is essential in processes such as repair of wounds, capture of foreign pathogens, and tissue differentiation. Force generation at the cellular level is a fascinating topic that is the subject of much active research.
- Cells can generate forces, but just as importantly, they can sense and respond to forces. We alluded to this above in the examples of bone remodeling and arterial caliber adjustment, but it is not only endothelial and bone cells that can sense forces. In fact, the ability of mechanical stress to elicit a biological response in cells seems to be the rule rather than the exception, and some cells are exquisitely specialized for just this task. One remarkable example is the hair cells in the ear. These cells have bundles of thin fibers (the *stereocilia*) that protrude from the apical cell surface and act as sensitive accelerometers; as a result, the hair cells are excited by sound-induced vibrations in the inner ear. This excitation produces electrochemical signals that are conducted by the auditory nerve to the auditory centers in the brain in a process that we call hearing [4,5].¹
- The examples above show that biomechanics is important in homeostasis and normal function. Unfortunately, biomechanics also plays a role in some diseases. One example is glaucoma, an ocular disease that affects about 65 million people worldwide [6]. Normally the human eye is internally pressurized, a fact that you can verify by gently touching your eye through the closed eyelid. In most forms of glaucoma, the pressure in the eye becomes elevated to pathological levels, and the resulting extra biomechanical load somehow damages the optic nerve, eventually leading to blindness [7]. A second example is atherosclerosis, a common arterial disease in which non-physiological stress distributions on endothelial cells promote the disease process [8].
- What about biomechanics in the treatment of disease and dysfunction? There are obvious roles in the design of implants that have a mechanical function,

Actually, the function of the hair cells is even more amazing than it first appears. The outer hair cells are active amplifiers, changing their shape in response to mechanical stimulation and thus generating sounds. The net effect is to apply a frequency-selective boost to incoming sounds and hence improve the sensitivity of the ear.

such as total artificial hips [9], dental implants [10], and mechanical heart valves [11]. In the longer term, we expect to treat many diseases by implanting engineered replacement tissue into patients. For tissues that have a mechanical function (e.g., heart valves, cartilage), there is now convincing evidence that application of mechanical load to the tissue while it is being grown is essential for proper function after implantation. For example, heart valves grown in a bioreactor incorporating flow through the valve showed good mechanical properties and function when implanted [12]. Cartilage subjected to cyclic shearing during growth was stiffer and could bear more load than cartilage grown without mechanical stimulation [13]. We expect that biomechanics will become increasingly important in tissue engineering, along the way leading to better fundamental understanding of how cells respond to stresses.

The above examples should give a flavor of the important role that biomechanics plays in health and disease. One of the central characteristics of the field is that it is highly interdisciplinary: to be called biomechanics, there must be elements of both mechanics and biology (or medicine). Advances in the field occur when people can work at the frontier of these two areas, and accordingly we will try to give both the "bio" and the "mechanics" due consideration in this book.

Another characteristic feature of biomechanics is that the topic is fairly broad. We can get a sense of just how broad it is by looking at some of the professional societies that fall under the heading of biomechanics. For example, in Japan alone, at least six different professional societies cover the field of biomechanics.² Obviously we cannot, in a single book, go into detail in every topic area within such a broad field. Therefore, we have given an introduction to a variety of topics, with the hope of whetting readers' appetites.

1.1 A brief history of biomechanics

We can learn more about the field of biomechanics by looking at its history. In one sense, biomechanics is a fairly young discipline, having been recognized as an independent subject of enquiry with its own body of knowledge, societies, journals, and conferences for only around 30–40 years. For example, the "Biomechanics and Human Factors Division" (later to become the "Bioengineering Division") of the American Society of Mechanical Engineering was established in late 1966. The International Society of Biomechanics was founded August 30, 1973; the European

² These are the Japanese Society of Biomechanics, the Bioengineering Division of the Japan Society of Mechanical Engineers, the Japan Society of Medical Electronics and Biological Engineering, the Association of Oromaxillofacial Biomechanics, the Japanese Society for Clinical Biomechanics and the Japanese Society of Biorheology.