DUKE **BIOMEDICAL ENGINEERING BME** Pratt School of Engineering

| about | |
|-----------------------|--|
| people | |
| - faculty | |
| - recent publications | |
| - courses | |
| - staff | |
| research | |
| events | |
| contacts | |
| news | |
| bme home | |
| pratt home | |
| duke home | |

undergrads

grads

industry

employment

MORTON H FRIEDMAN, PROFESSOR

Dr. Friedman's research interests focus on the role of mechanical forces in the localization and evolution of vascular disease, particularly atherosclerosis. Specific areas of research are the influence of adaptation on vascular biology and physiology, the influence of shear stress gradients and harmonic content on the vascular endothelium, and the influence of coronary artery geometry and dynamics on plaque formation and rupture.

The first of these explores the effect of changes in blood flow on the permeability of the arterial wall to large molecules, and the expression and distribution of proteins that might mediate the permeability response. Our interest in permeability is motivated by the fact that a key initial step in the development of atherosclerosis is the accumulation of cholesterol-carrying



lipoproteins in the vessel wall. In many areas of the vasculature, the blood flow changes during the day, as, for instance, during digestion or exercise. Our hypothesis is that these changes cause transient increases in permeability, and that it is during these transients, while the wall is adapting to its new fluid dynamic environment, that most of the uptake of fats by the wall takes place. The hypothesis is tested in animal experiments using swine.

A related study seeks a better understanding of the influence of fluid dynamic stresses on the vessel wall in the normal adapted state. Of particular interest are the effects of shear stress gradients and the high frequency components of the shear stress profile at a site. The effect of these variables are examined in the swine experiments noted above, and in custom designed flow systems that expose cultured endothelial cells to defined shear stress environments. Research opportunities in these two studies, in addition to the experiments themselves, include calculation of the flow field and wall shear stress distributions generated experimentally, and measurement of the response of the vascular endothelium at the tissue (permeability - in vivo experiments only), cellular (junctions, cytoskeleton) and molecular (gene expression, RNA, protein) levels.

A second focus of Dr. Friedman's laboratory explores the relationship between blood vessel geometry and dynamics and the natural history of atherosclerotic disease in the coronary arteries. Our interest in this area arises because of increasing evidence that wall mechanical forces play a role in the progression of atherosclerosis and its fatal consequences. The coronary arteries are tethered to the surface of the heart and deform as the heart beats. Since vascular geometry, such as the angle at an arterial branch, and vascular deformation, such as the flexing of an arterial bend, influence the wall stress distribution in the vessel, it follows that certain geometric or dynamic features of an individual's arteries might affect that person's susceptibility to disease. This hypothesis is being tested using both clinical data and studies in genetically modified mice that develop humanoid atherosclerosis in a matter of weeks. Research opportunities include the analysis of coronary artery images obtained using cineangiography (human), stereomicroscopy (mouse) and intravascular ultrasound (human), mechanical modeling of the vascular wall to elucidate the relation between vessel dynamics and plague rupture, and proteomic and genomic studies on the mouse tissues.

Note: the most current description of Dr. Friedman's research and listing of recent publications with links can be found at the Cardiovascular Simulations Laboratory website, whose URL is given below.

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| | http://biotrans.pratt.duke.edu |
| | Book: Principles and Models of Biological Transport |

Education:

PhD, University of Michigan, 1961 MSE, University of Michigan, 1958 BSE, Cornell University, 1957

Specialties:

Atherosclerosis Hemodynamics Biological Transport Mathematical Modeling Angiographic and Ultrasound Image Processing Computational Fluid Dynamics

Research Interests:

Dr. Friedman's research interests focus on the role of mechanical forces in the localization and evolution of vascular disease, particularly atherosclerosis. Specific areas of research are the influence of adaptation on vascular biology and physiology, the influence of shear stress gradients and harmonic content on the vascular endothelium, and the influence of coronary artery geometry and dynamics on plaque formation and rupture.

Awards, Honors, and Distinctions

Dedicated Service Award, American Society of Mechanical Engineers Distinguished Service Award, AIMBE Elected Member, World Council of Biomechanics Fellow, American Association for the Advancement of Science Fellow, American Heart Association, Council on Arteriosclerosis, Thrombosis & Vascular Biology Fellow, American Society of Mechanical Engineers Fellow, American Institute of Chemists Founding Fellow, American Institute of Medical and Biological Engineering H.R. Lissner Medal, American Society of Mechanical Engineers Inaugural Fellow, Biomedical Engineering Society National Capital Award, District of Columbia Council of Engineering and Architectural Societies Richard Skalak Best Paper Award, American Society of Mechanical Engineers, Journal of Biomedical Engineering, 2006

Recent Publications (More Publications)

- G. A. Ateshian and M. H. Friedman, *Integrative biomechanics: A paradigm for clinical applications of fundamental mechanics*, Journal Of Biomechanics, vol. 42 no. 10 (July, 2009), pp. 1444 --1451 [abs].
- 2. H. Zhu and Z. H. Ding and R. N. Piana and T. R. Gehrig and M. H. Friedman, *Cataloguing the geometry of the human coronary arteries: A potential tool for predicting risk of coronary artery disease*, International Journal Of Cardiology, vol. 135 no. 1 (June, 2009), pp. 43 -- 52 [abs].
- 3. M. H. Friedman, Variability of arterial wall shear stress, its dependence on vessel diameter and implications for Murray's Law, Atherosclerosis, vol. 204 no. 1 (May, 2009), pp. 18 -- 19.
- 4. M. H. Friedman, Variability of arterial wall shear stress, its dependence on vessel diameter and implications for Murray's Law, Atherosclerosis, vol. 203 no. 1 (March, 2009), pp. 47 -- 48.
- Y. Liang and H. Zhu and M. H. Friedman, *Estimation of the transverse strain tensor in the arterial wall using ivus image registration*, Ultrasound In Medicine And Biology, vol. 34 no. 11 (November, 2008), pp. 1832 -- 1845 [abs].