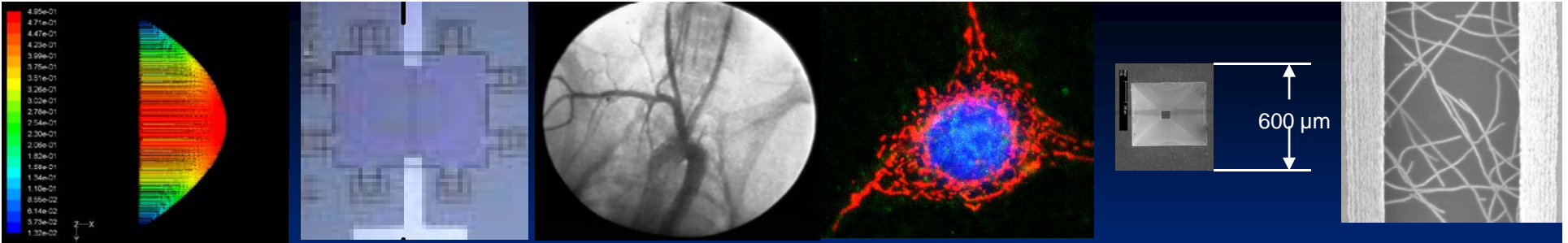


Cardiovasculature: A Dynamic Sensing and Actuating System

Tzung Hsiai, MD, PhD

Cardiovascular Engineering Research Core
School of Engineering &
School of Medicine
University of Southern California

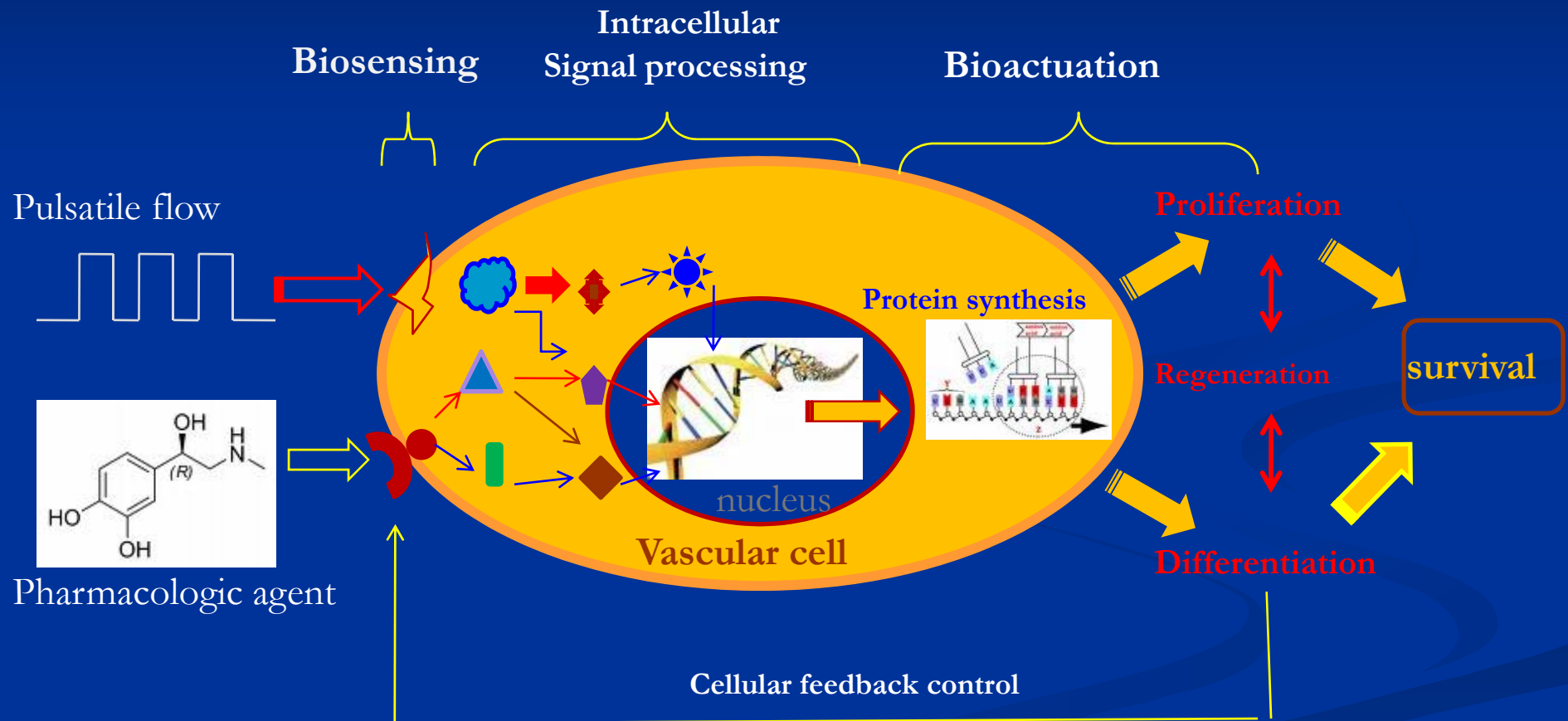


Objectives

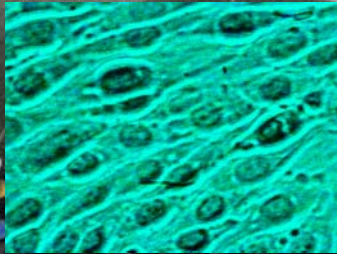
- Introduce vascular systems
- Interface vascular dynamics with biology
- Translate vascular dynamics to in vivo models
- Bridge bioengineering, industry, and medicine



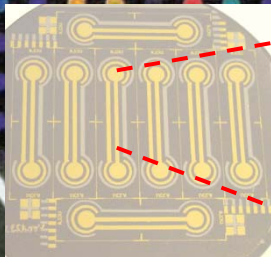
Cardiovasculature: A Dynamic Sensing and Actuating System



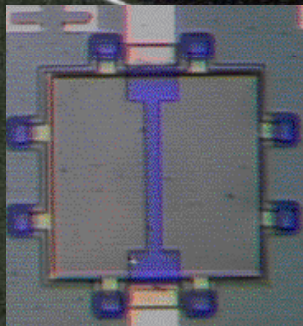
Biomedical Research in the Post-Genomic Era



Shear stress slew rates
-remodeling
-monocyte binding



DNA/RNA sensors
-gene expression



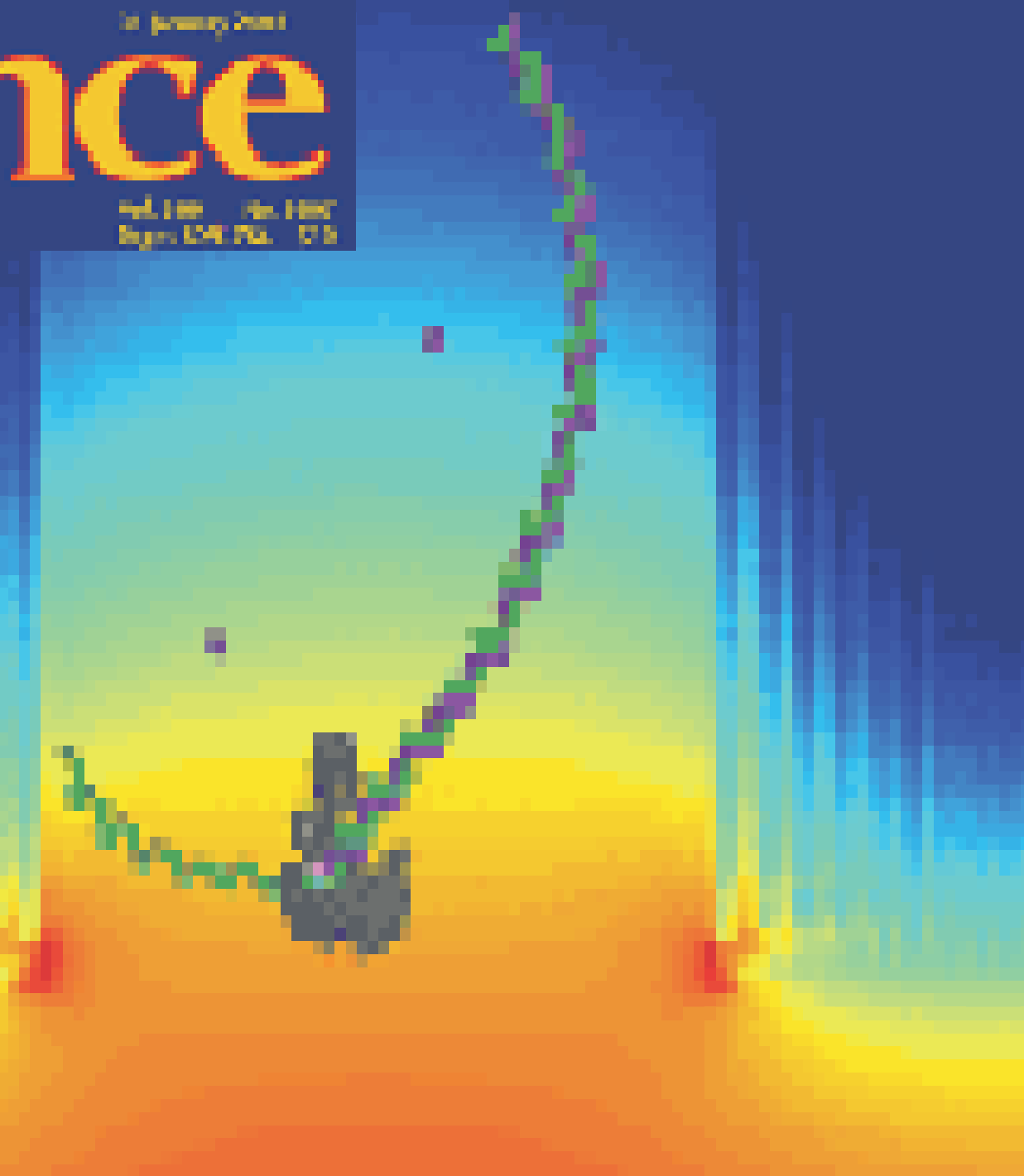
Shear stress sensors
-direct and precise measurement

EC Dynamics

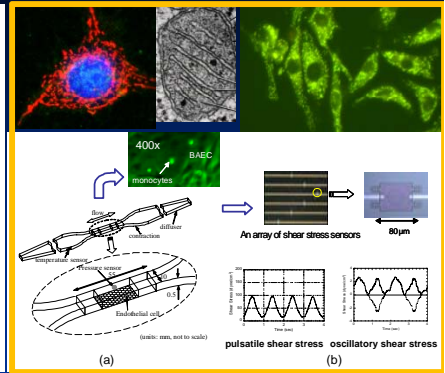
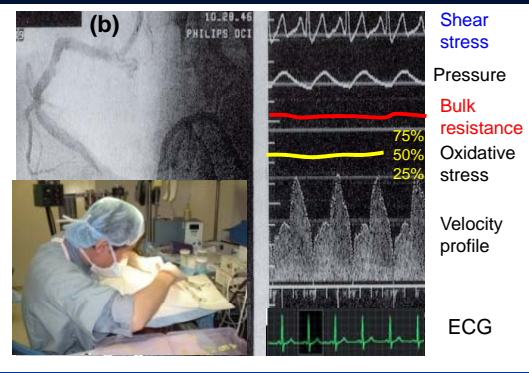
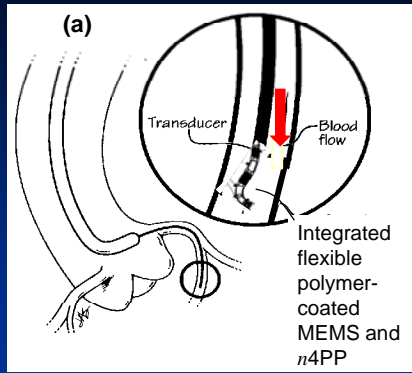
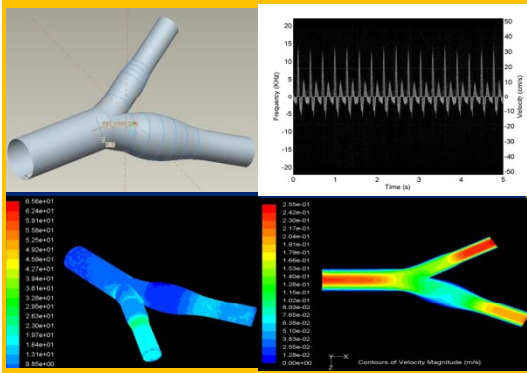
Science

1st January 2008

Vol. 318 No. 5857
Pages 1241-1312



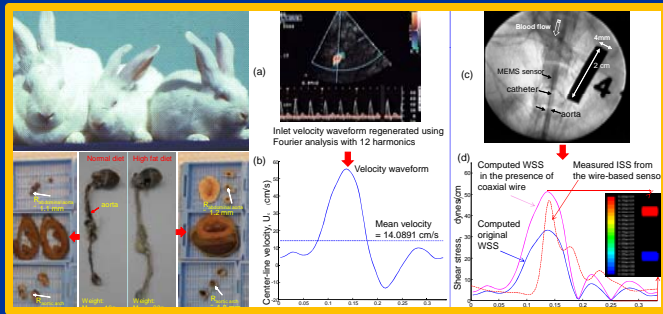
Cardiovascular Engineering Research Core



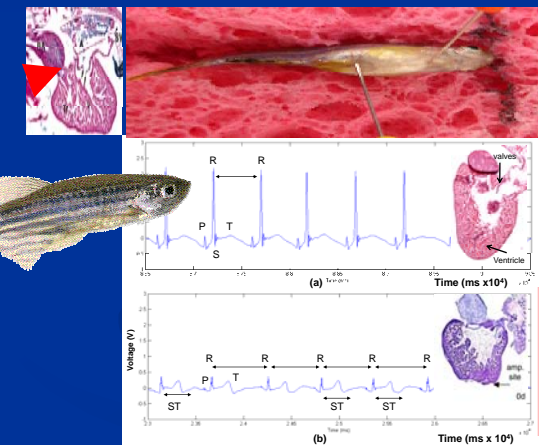
(1) Computational Fluid Dynamics

(2) Nanosensing of vascular oxidative stress and mechanically unstable plaque to predict acute coronary syndromes

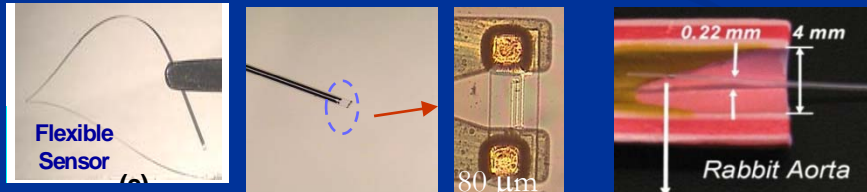
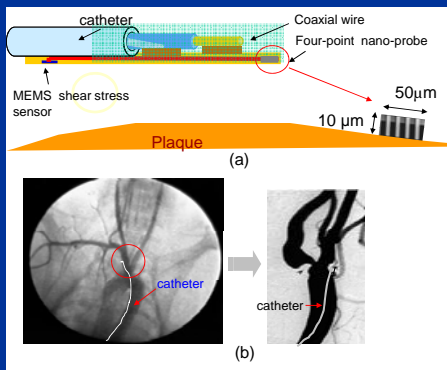
(3) Mechano-transduction and vascular biology



Collaborators:
 Fengzhu Sun, PhD, College LAS, USC
 Ellen Lien, PhD, Children's Hospital Los Angeles, USC
 Michael Kahn, PhD, Biochem & Molecular Bio, KSoM, USC
 Mark Barr, MD, Cardiothoracic Surgery, USC
 Fred A. Weaver, MD, Vascular Surgery, USC
 Robert Kloner, MD, PhD, Good Samaritan Hospital, USC
 Q. Zhou, PhD, E. S. Kim, PhD, E. Meng, PhD, C.Sioutas, PhD, VS
 Joe Wu, MD, PhD, Stanford University School of Medicine
 Julie Freishlag, MD, Johns Hopkins School of Medicine
 Linda L. Demer, MD, PhD, UCLA School of Medicine
 Judith Berliner, PhD, UCLA School of Medicine



(4) Rabbit model of atherosclerosis



(6) MEMS shear stress sensors and flexible polymer sensors

(8) HAEC and tube formation

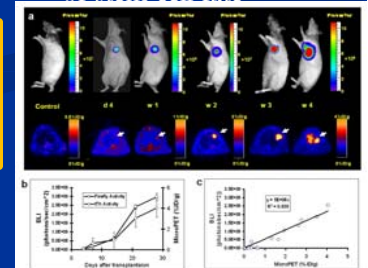


(5) Deployment of intravascular sensors in rabbit aortas

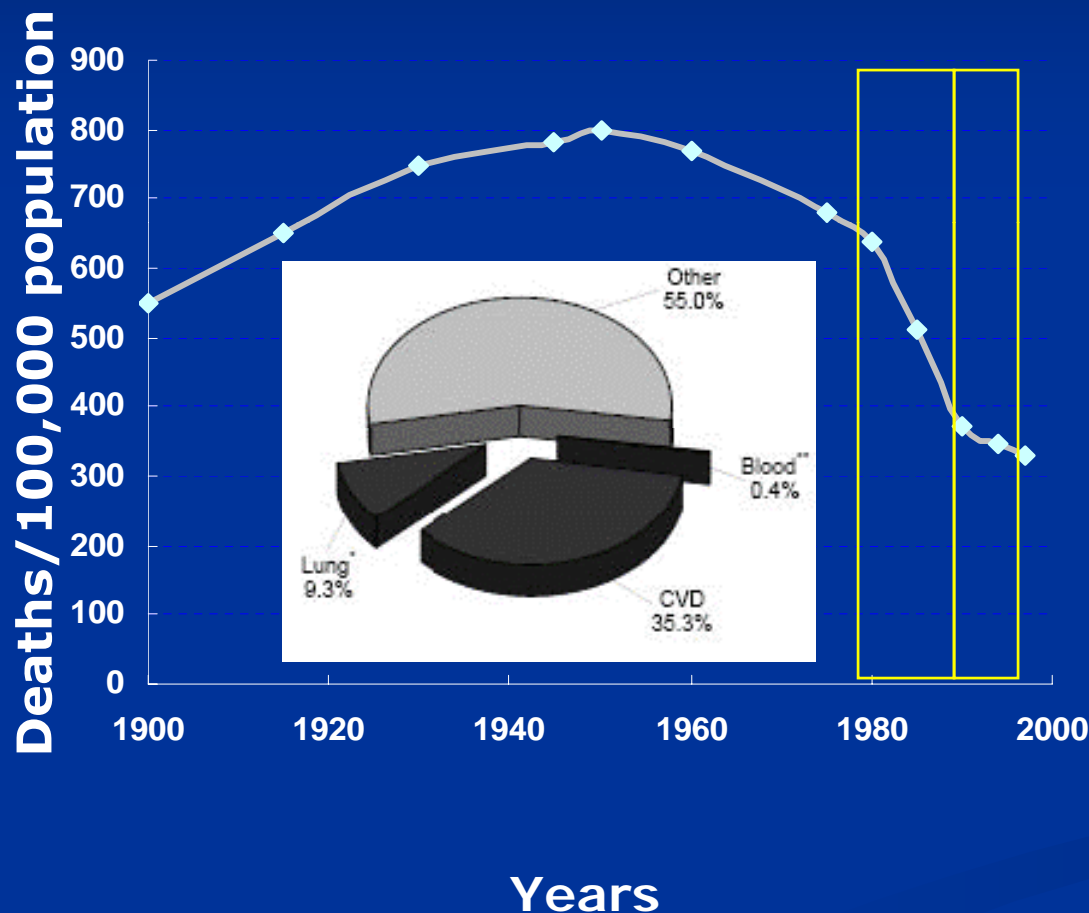
(9) Nano particles and environmental health

(7) Micro-ECG to monitor Zebrafish heart regeneration

(10) Molecular imaging of stem cell fate



Death Rates in Cardiovascular Disease: 35.3 % of all deaths in USA



Number of deaths for leading causes of death:

Heart disease: 631,636

Cancer: 559,888

Stroke (cerebrovascular diseases): 137,119

Chronic lower respiratory diseases: 124,583

Accidents (unintentional injuries): 121,599

Diabetes: 72,449

Alzheimer's disease: 72,432

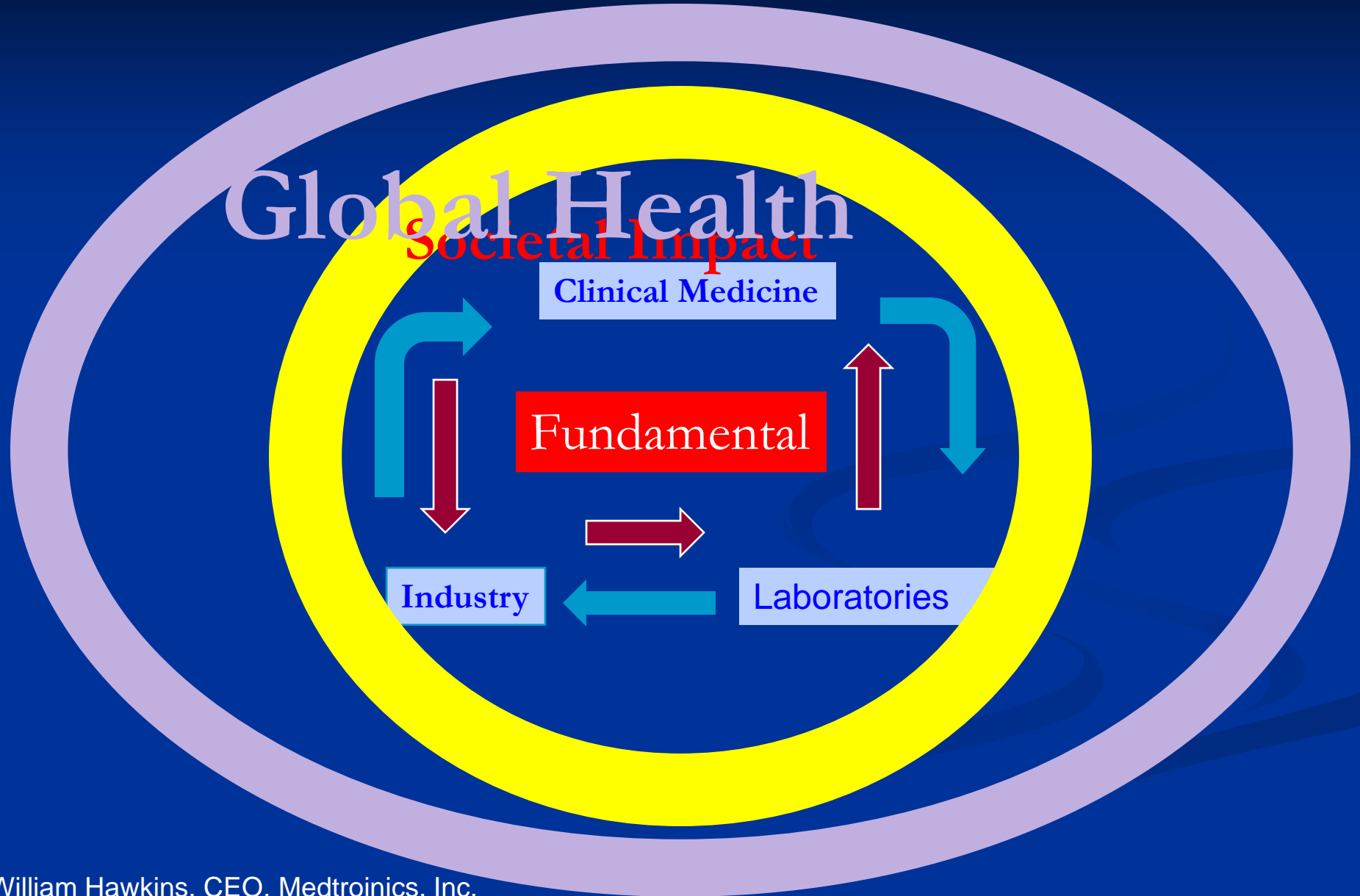
Influenza and Pneumonia: 56,326

Nephritis, nephrotic syndrome, and nephrosis: 45,344

Septicemia: 34,234

BMES2009

Three rivers of biology, engineering, and medicine

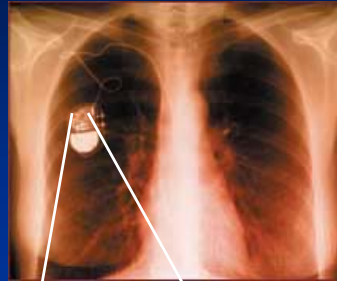
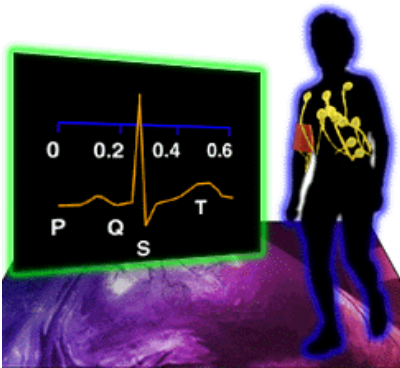


Impact of Biomedical Research on Medicine

Macro



ECG/EKG
(electrocardiogram)



Chest X-Ray

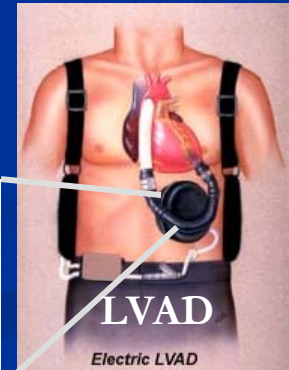
Echocardiogram



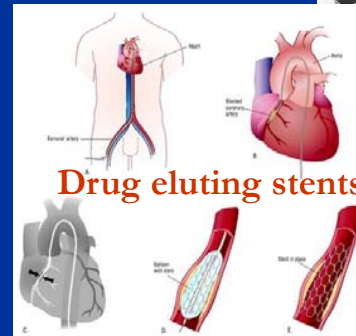
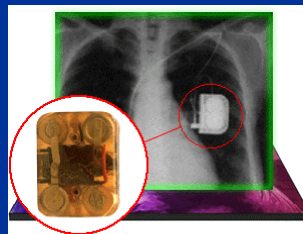
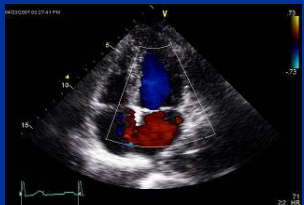
Imaging: MRI



Defibrillator



LVAD
Electric LVAD



Drug eluting stents



A wide-open territory

Micro



Molecular and nano approach to stem cell and regenerative medicine, tissue engineering, materials science, chemistry. Molecular biology, and alike.

Nano



Biology, nanotechnology and nanophotonics

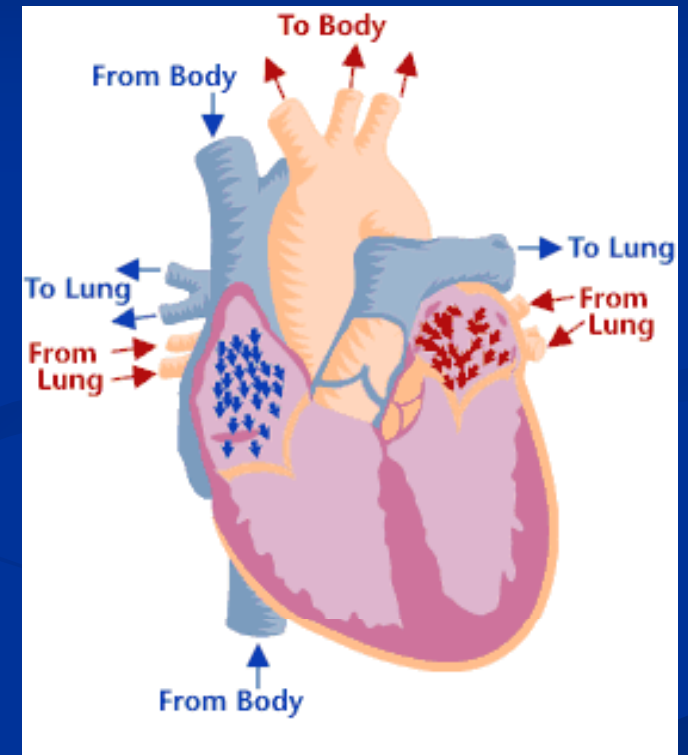
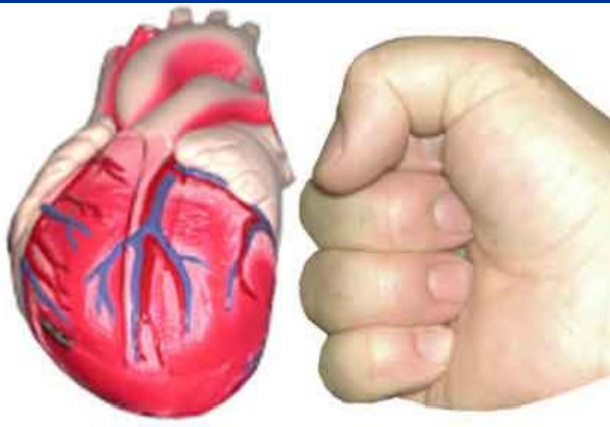
Dr. Hsiai's group at Edwards Lifesciences,
Irvine, Ca.



Objectives

- **Introduce vascular systems**
- **Interface vascular dynamics with biology**
- **Translate vascular dynamics to in vivo models**
- **Bridge bioengineering, industry, and medicine**

Vascular Dynamics



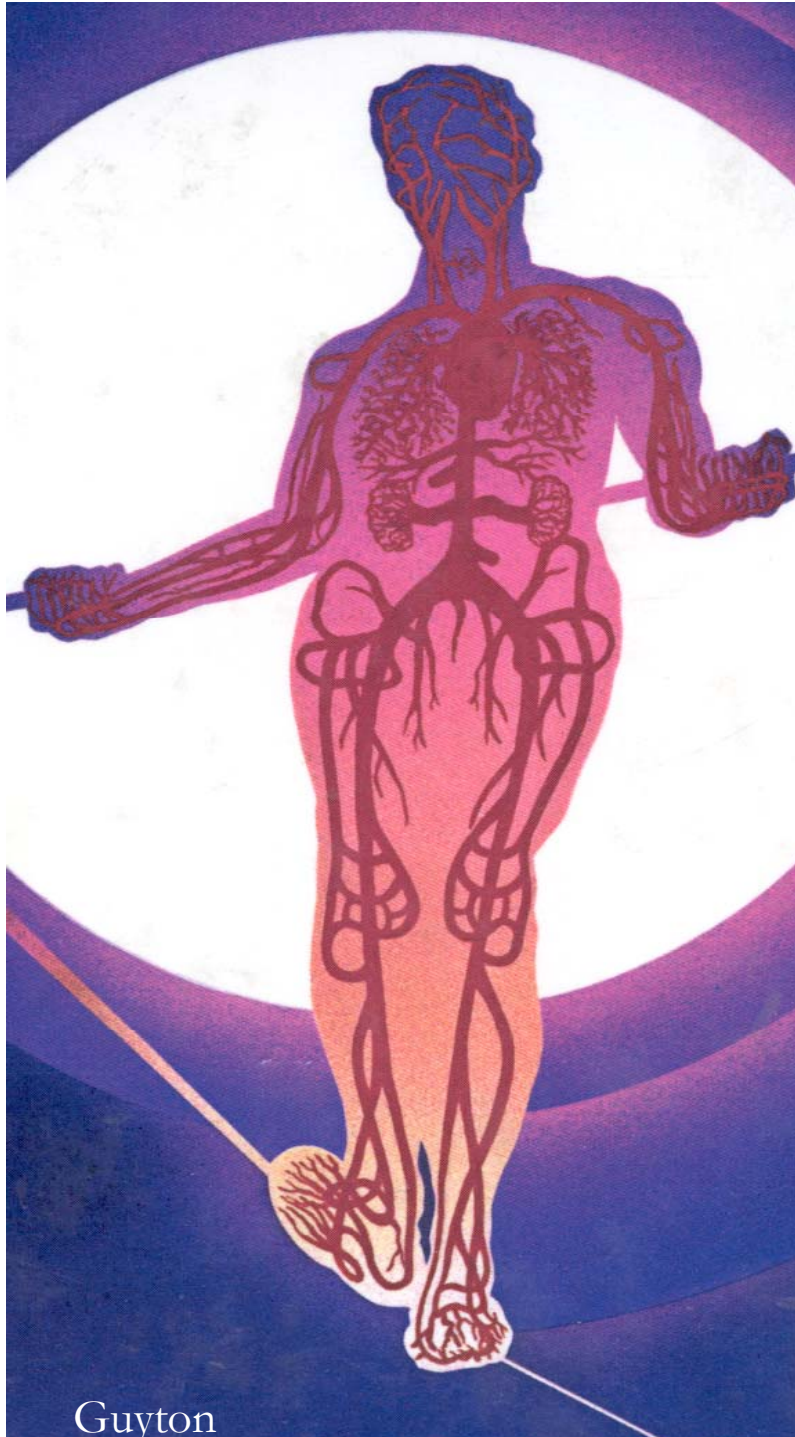
The Circuitry

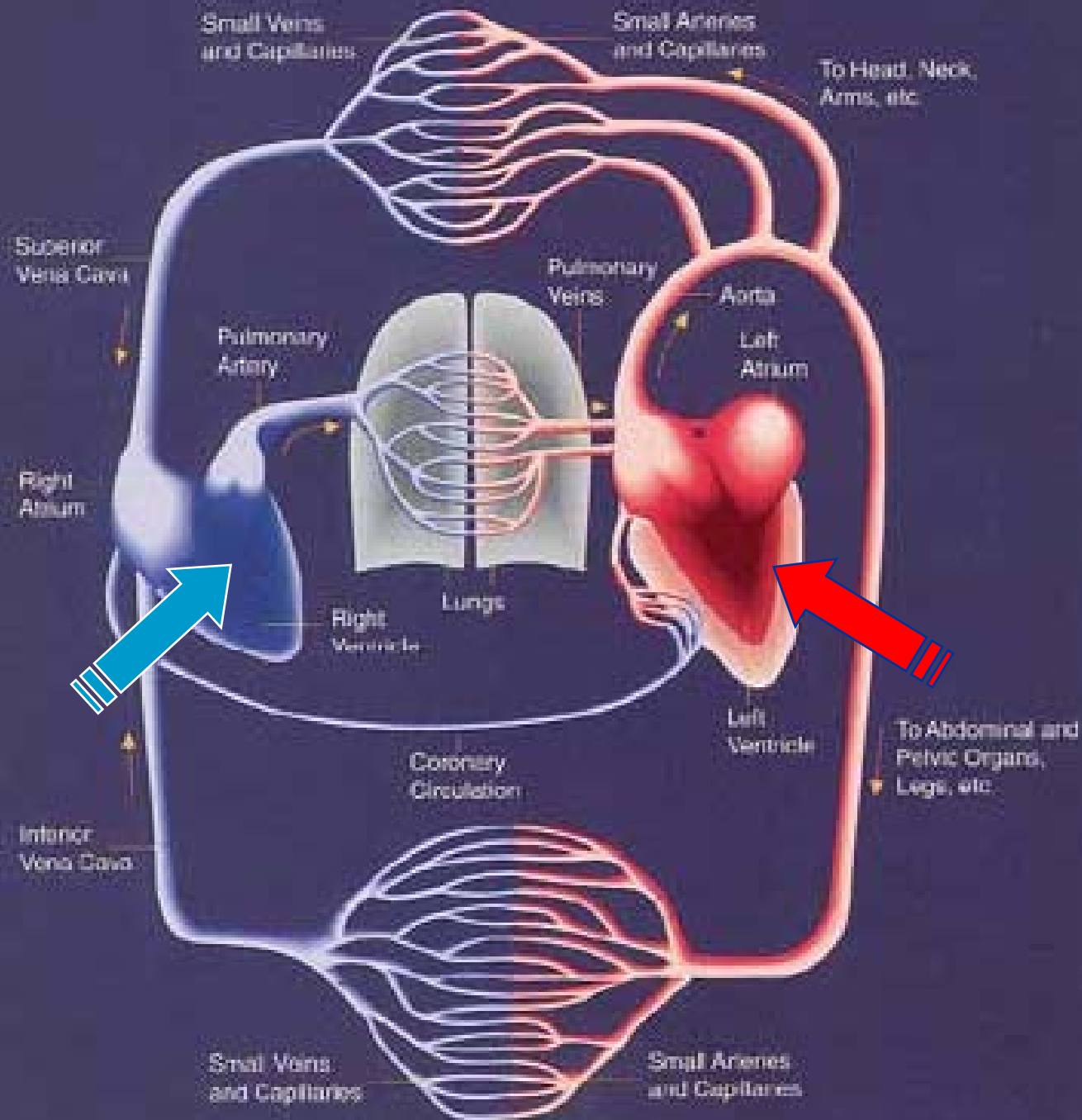
Cardiovascular system

-a pump

-a series of distributing and collecting tubes

-an extensive system of thin vessels that permits rapid exchange of substances between the tissues and the vascular channels





-Pulsatile flow

-Laminar flow

-Viscous/frictional resistance

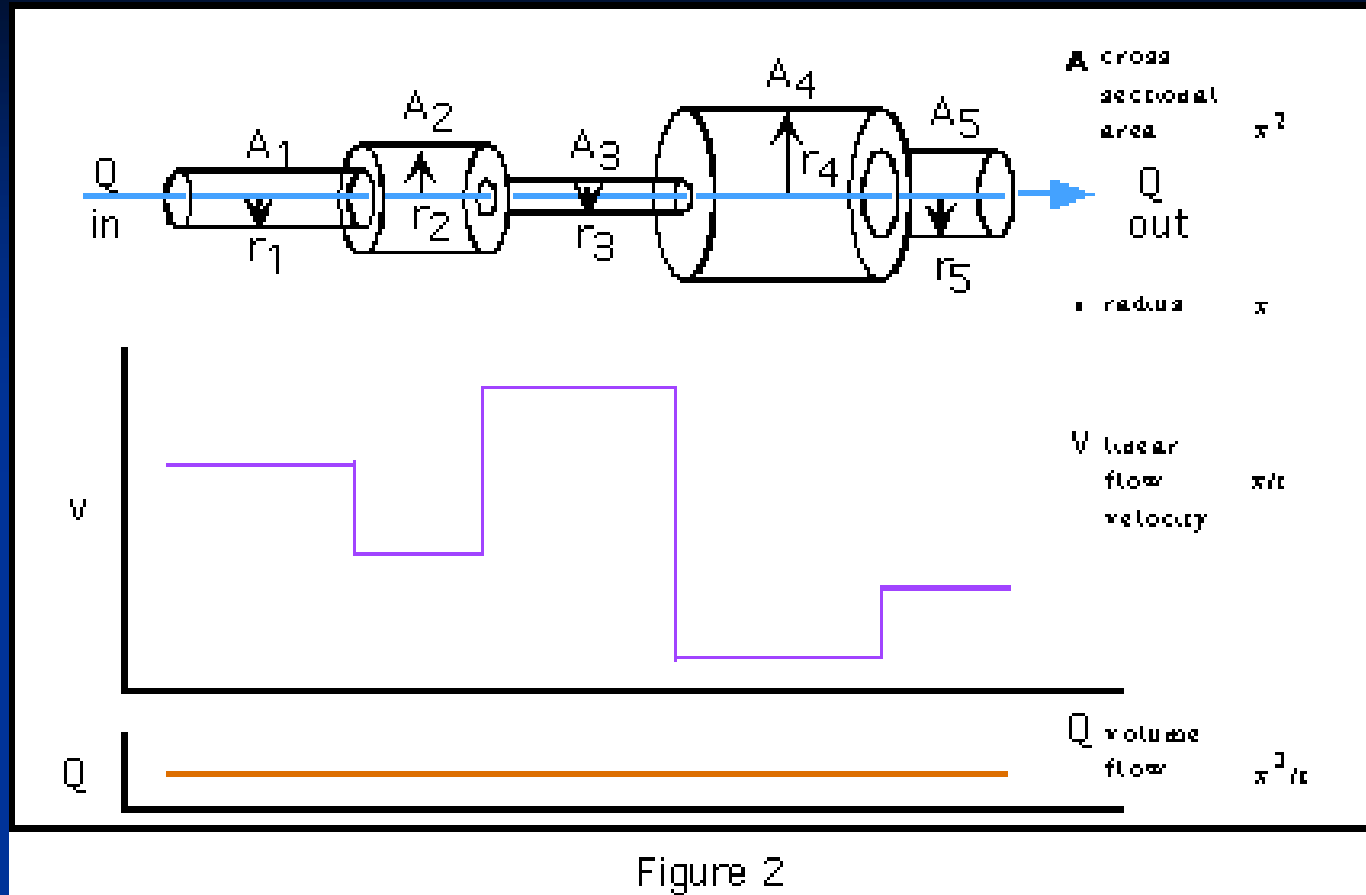
-Right vs left ventricular wall thickness

-Arterial resistance: Aorta vs. capillaries

-Capillaries: Resistance in parallel

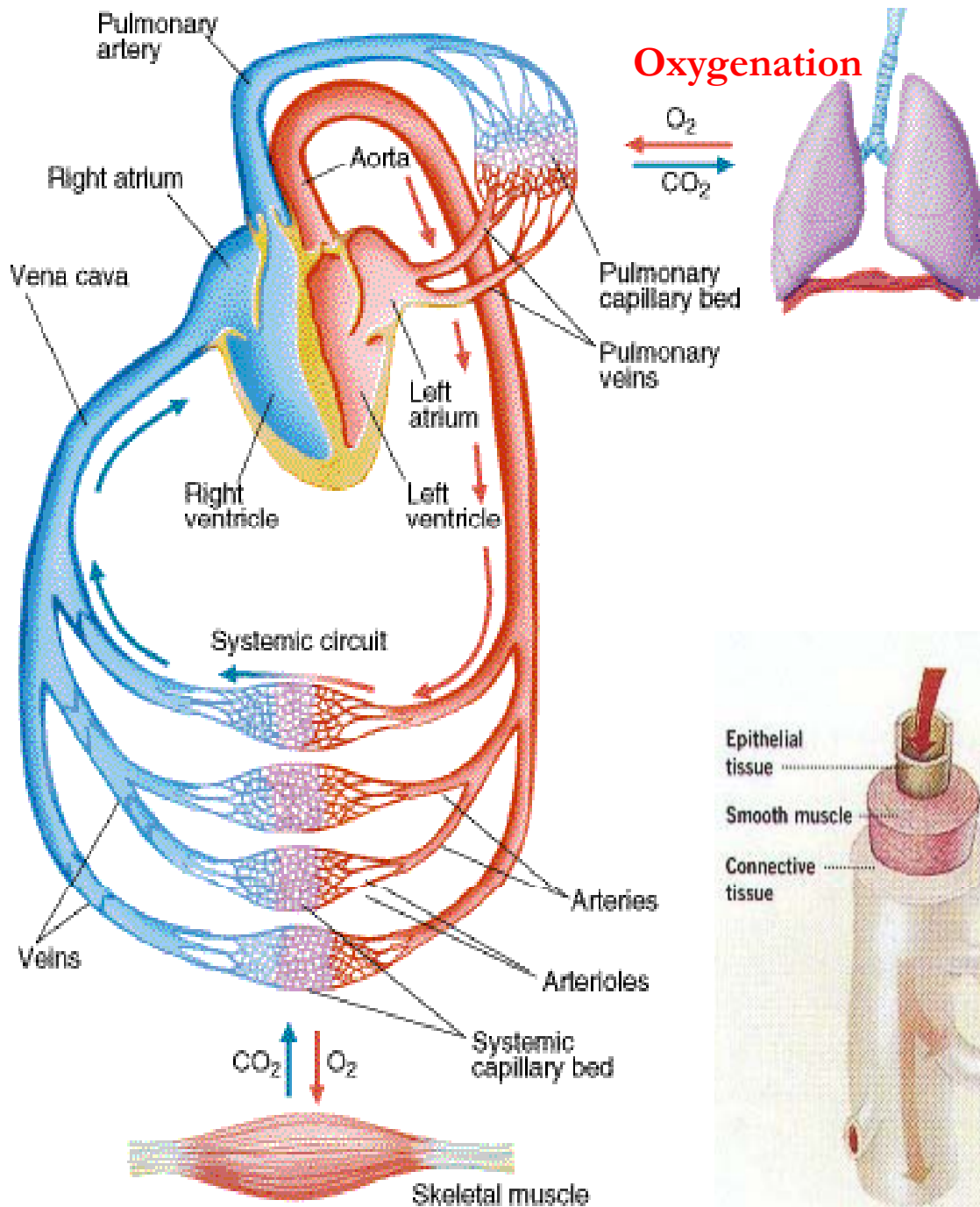
-surface area: Aorta vs. capillaries

Law of Continuity of Flow



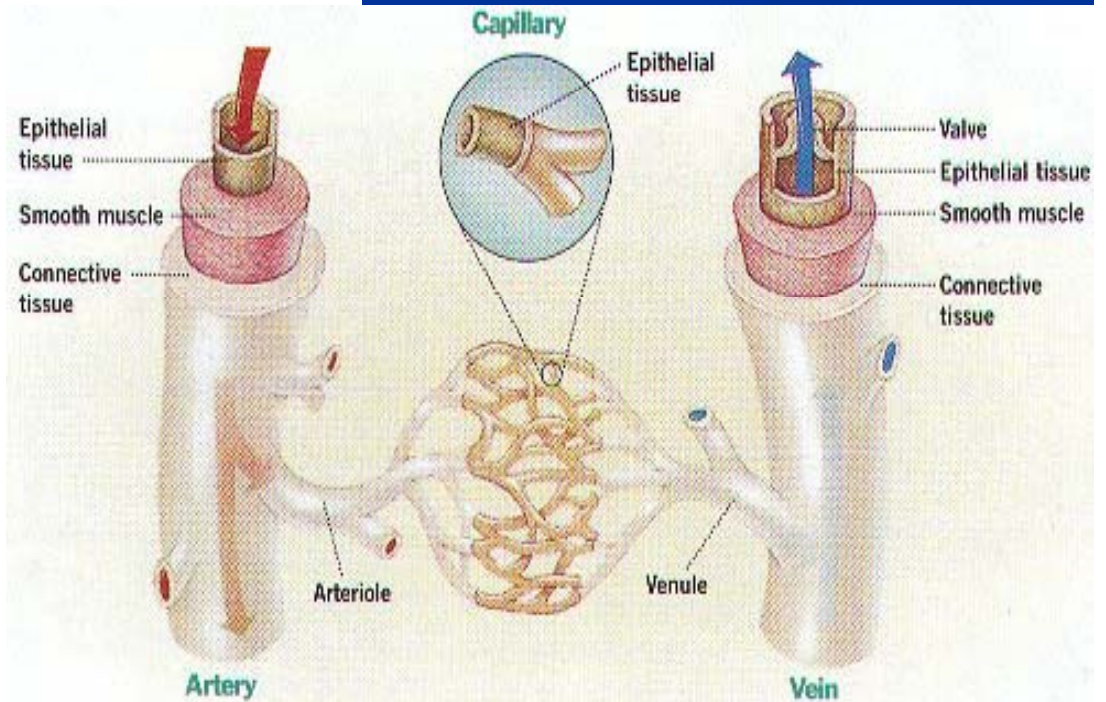
Blood flow in vessels of different sizes is constant in each segment of a given cross-sectional area regardless of the magnitude of that area. If vessels run side by side (in parallel), the flow will be additive. When one vessels lead one directly into another (in series), what flows in is what flows out:

$$Q_{\text{total}} = Q_1 + Q_2 = (V_1 A_1) + (V_2 A_2) = Q_{\text{in}} = Q_{\text{out}}$$



Compositions of vessel wall

- Endothelial cells
- Intima (collagen fibers)
- Elastic lamina (arteries)
- smooth muscle cells (arteries vs. veins)
- tunica media
- adventia
- vaso vasorum



Navier-Stokes Equations

3-D unsteady flow

Coordinates: (x,y,z)

Time : t Pressure: p

Heat Flux: q

Density: ρ Stress: τ

Reynolds Number: Re

Velocity Components: (u,v,w)

Total Energy: Et

Prandtl Number: Pr

Continuity:
$$\frac{\partial \rho}{\partial t} + \frac{\partial(\rho u)}{\partial x} + \frac{\partial(\rho v)}{\partial y} + \frac{\partial(\rho w)}{\partial z} = 0$$

X – Momentum:
$$\frac{\partial(\rho u)}{\partial t} + \frac{\partial(\rho u^2)}{\partial x} + \frac{\partial(\rho uv)}{\partial y} + \frac{\partial(\rho uw)}{\partial z} = -\frac{\partial p}{\partial x} + \frac{1}{Re_r} \left[\frac{\partial \tau_{xx}}{\partial x} + \frac{\partial \tau_{xy}}{\partial y} + \frac{\partial \tau_{xz}}{\partial z} \right]$$

Y – Momentum:
$$\frac{\partial(\rho v)}{\partial t} + \frac{\partial(\rho uv)}{\partial x} + \frac{\partial(\rho v^2)}{\partial y} + \frac{\partial(\rho vw)}{\partial z} = -\frac{\partial p}{\partial y} + \frac{1}{Re_r} \left[\frac{\partial \tau_{xy}}{\partial x} + \frac{\partial \tau_{yy}}{\partial y} + \frac{\partial \tau_{yz}}{\partial z} \right]$$

Z – Momentum:
$$\frac{\partial(\rho w)}{\partial t} + \frac{\partial(\rho uw)}{\partial x} + \frac{\partial(\rho vw)}{\partial y} + \frac{\partial(\rho w^2)}{\partial z} = -\frac{\partial p}{\partial z} + \frac{1}{Re_r} \left[\frac{\partial \tau_{xz}}{\partial x} + \frac{\partial \tau_{yz}}{\partial y} + \frac{\partial \tau_{zz}}{\partial z} \right]$$

Energy:

$$\frac{\partial(E_T)}{\partial t} + \frac{\partial(uE_T)}{\partial x} + \frac{\partial(vE_T)}{\partial y} + \frac{\partial(wE_T)}{\partial z} = -\frac{\partial(up)}{\partial x} - \frac{\partial(vp)}{\partial y} - \frac{\partial(wp)}{\partial z} - \frac{1}{Re_r Pr_r} \left[\frac{\partial q_x}{\partial x} + \frac{\partial q_y}{\partial y} + \frac{\partial q_z}{\partial z} \right] + \frac{1}{Re_r} \left[\frac{\partial}{\partial x} (u \tau_{xx} + v \tau_{xy} + w \tau_{xz}) + \frac{\partial}{\partial y} (u \tau_{xy} + v \tau_{yy} + w \tau_{yz}) + \frac{\partial}{\partial z} (u \tau_{xz} + v \tau_{yz} + w \tau_{zz}) \right]$$

Navier-Stokes Equation:

Newtonian fluids of constant density and viscosity

At steady flow

Hydrostatic equilibrium

An ideal fluid

$$\rho \left(\frac{\partial \mathbf{v}}{\partial t} + \mathbf{v} \cdot \nabla \mathbf{v} \right) = -\nabla \rho + \mu \nabla^2 \mathbf{v} + \mathbf{f}$$

Transient
Inertial
force

+

Convective inertial
force

=

Pressure
force

+

Viscous
force

+

Body
Force
= $\rho \mathbf{g}$

For pulsatile flow, compare the transient inertial force term with the viscous force term:

$$\frac{\textit{inertia}}{\textit{viscous}} = \frac{\rho \omega U}{\mu U L^{-2}} = \frac{\rho \omega L^2}{\mu} = \frac{\omega L^2}{\nu} = \text{a frequency parameter called the Stokes' number}$$

$$\text{The square root} = \alpha = N_w = \frac{D}{2} \sqrt{\frac{\omega}{\nu}} = \text{Womersley's number}$$

In aorta: α in a man ~ 20 , in a dog ~ 14 , a cat ~ 8 and a rat ~ 3

Reynolds number

Compare convective inertial force with the viscous force terms:

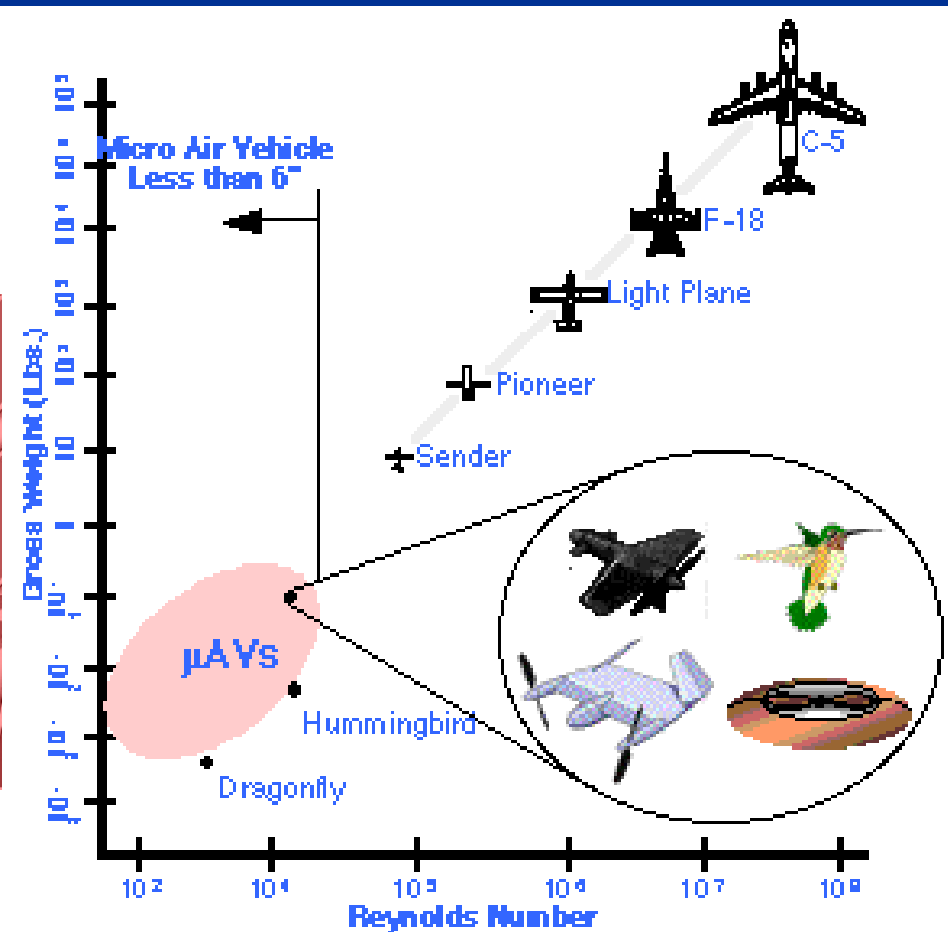
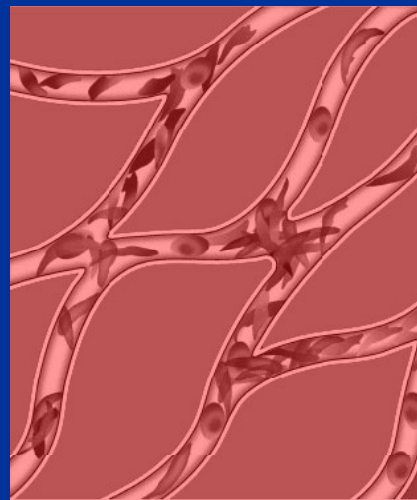
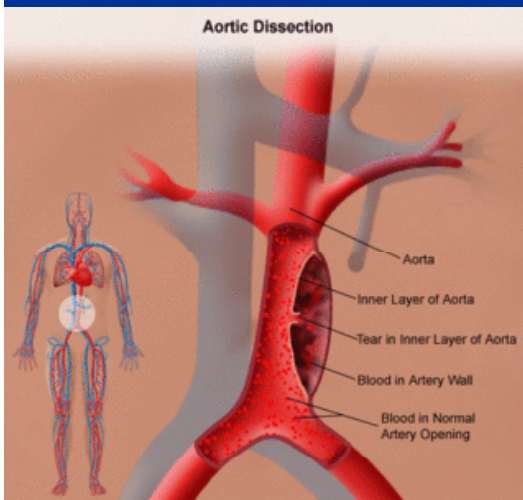
$$\frac{\textit{inertia}}{\textit{viscous}} = \frac{\rho U^2}{\mu U / L} = \frac{\rho U L}{\mu} = \textit{Reynolds}$$

A large Re #: a preponderant inertia effect.

A small Re #: a predominant viscous force effect.

Re # in aorta: 2,000-3,000

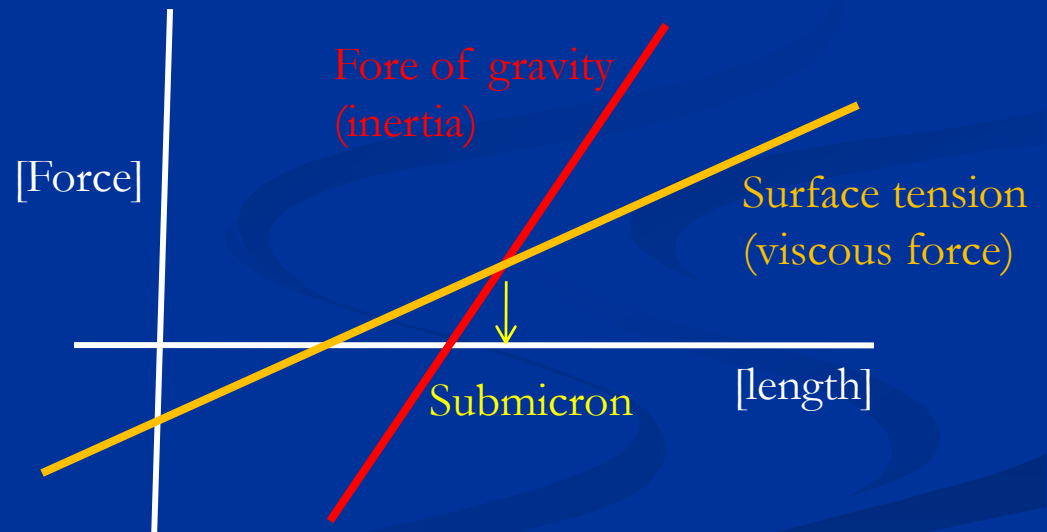
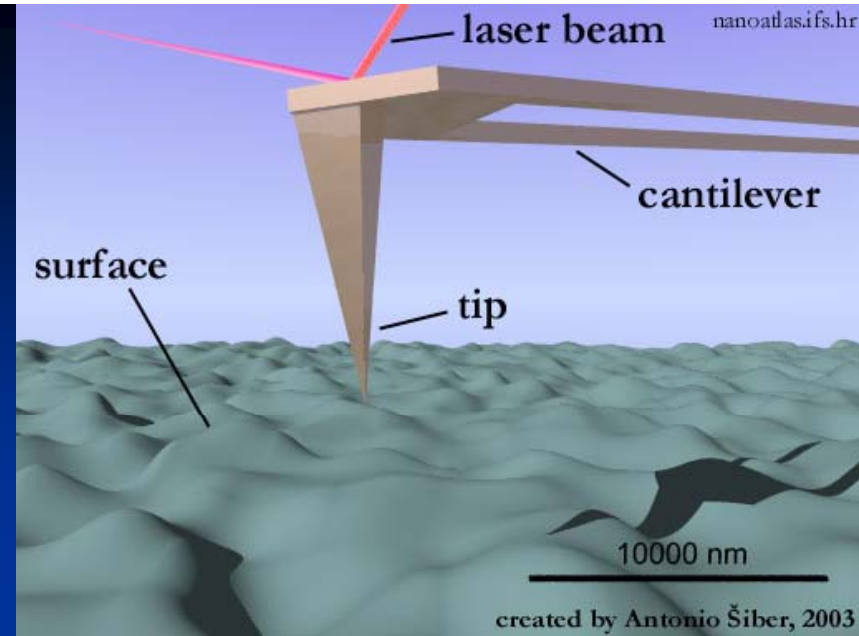
Re # in capillary: 0.001 to 0.01



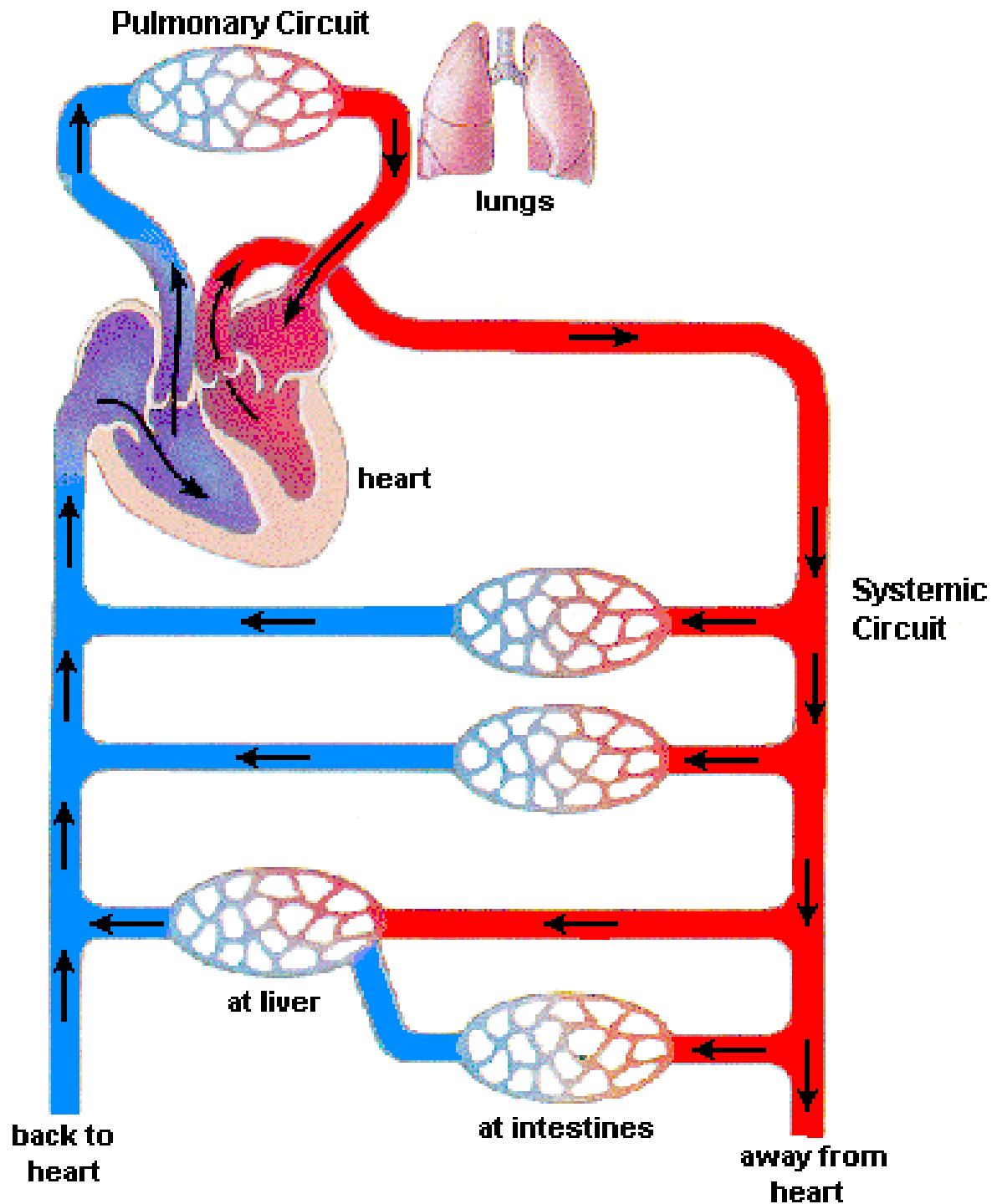
Paradigm shift in the micro- and nano-scale dynamics

Key points

- Semiconductors
- MEMS
- Microfluidics
- Gene Chips
- MEMS cardiovascular sensors
- Nanotechnology
- Nanotubes
- Quantum dots
- Green fluorescent protein
- Optics/molecular imaging
- Biodefense
- Nanotoxicity



$$\frac{\textit{inertia}}{\textit{viscous}} = \frac{\rho U^2}{\mu U / L} = \frac{\rho U L}{\mu} = \textit{Reynolds}$$



Reynolds numbers:

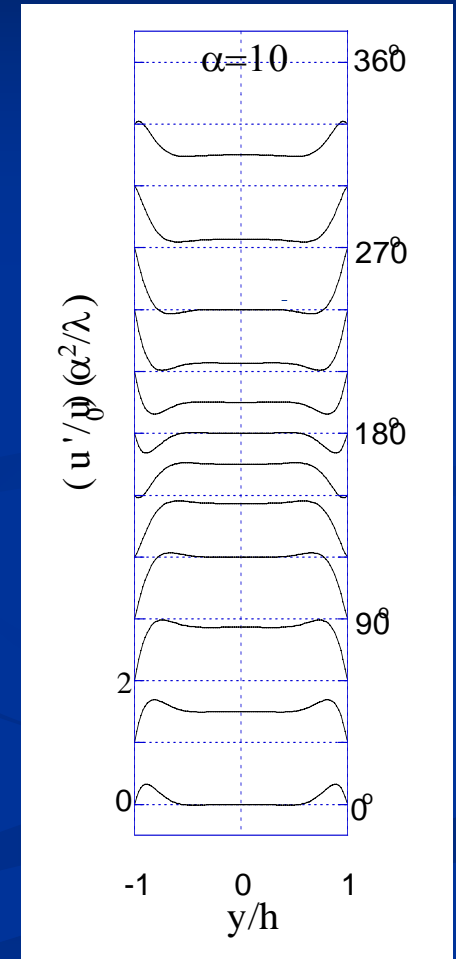
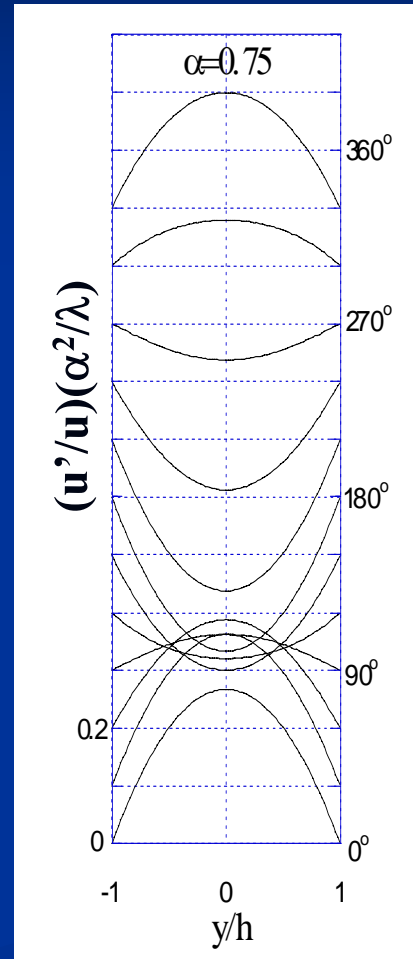
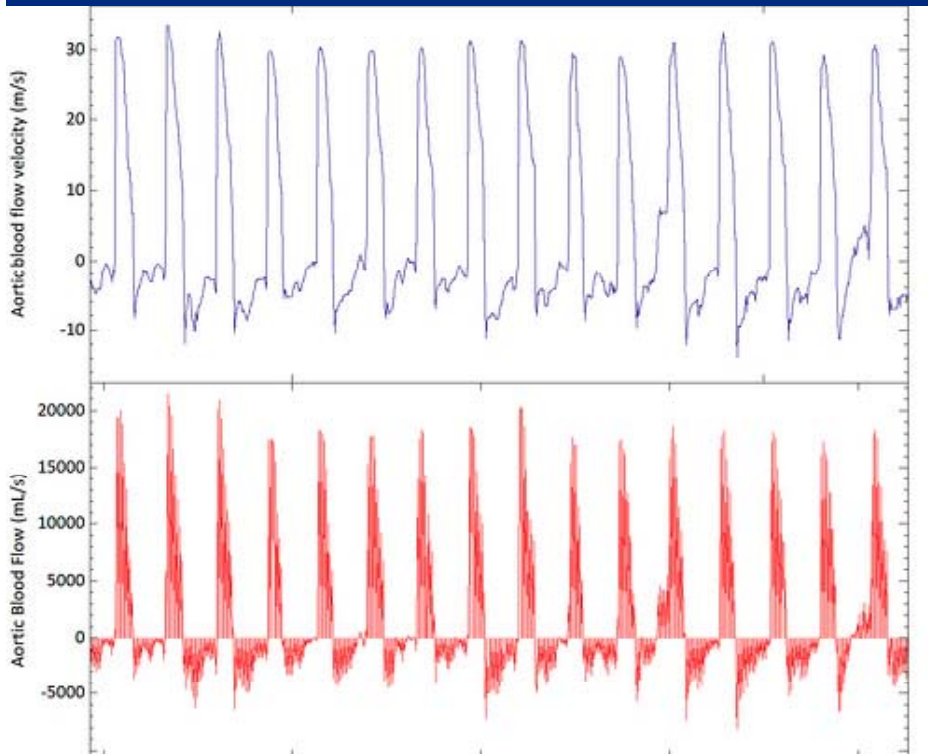
- aorta
- arterioles
- capillaries
- veins

Womerley's number:

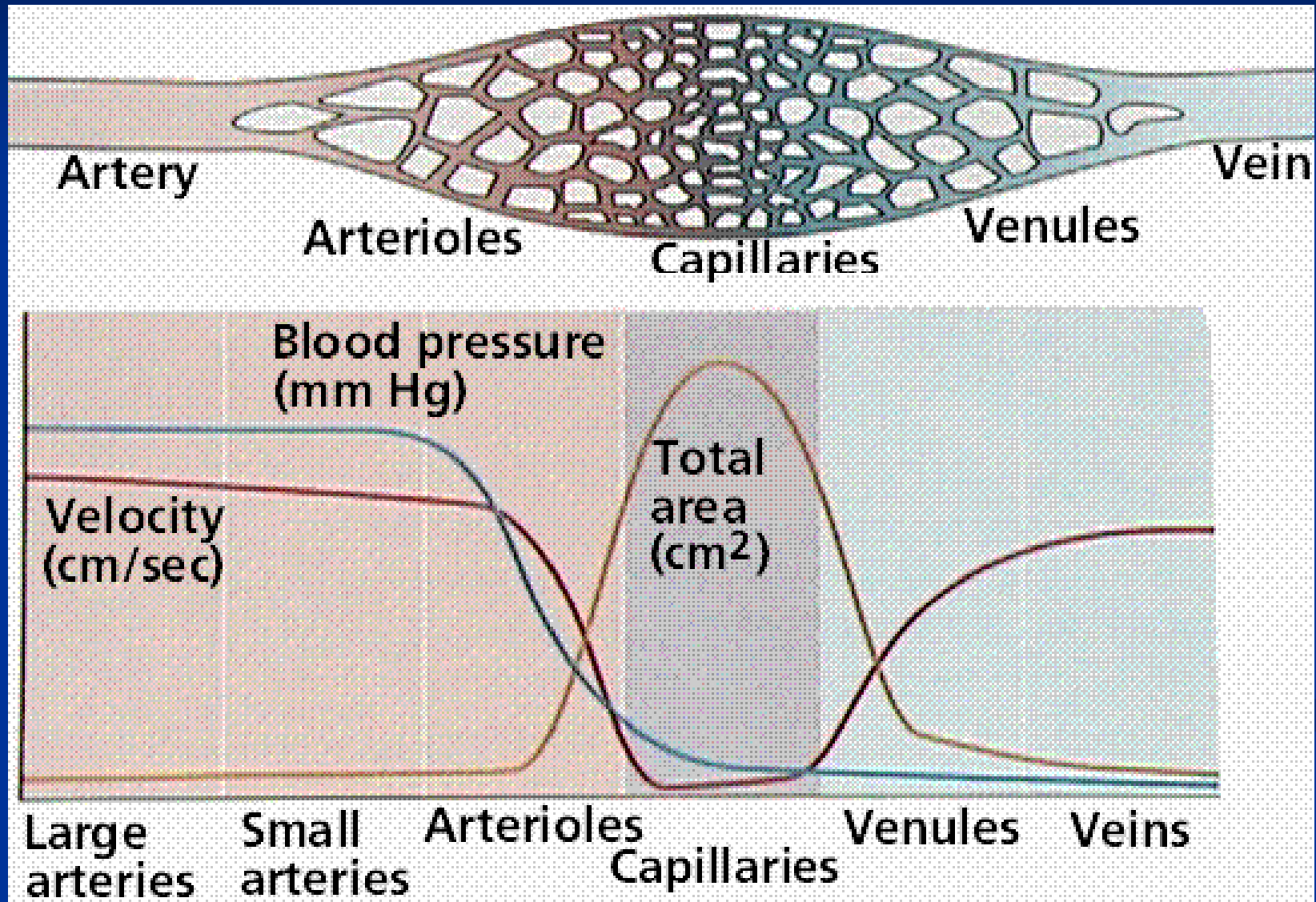
- aorta
- arterioles
- capillaries
- veins

Velocity Profiles of Pulsatile Flow:

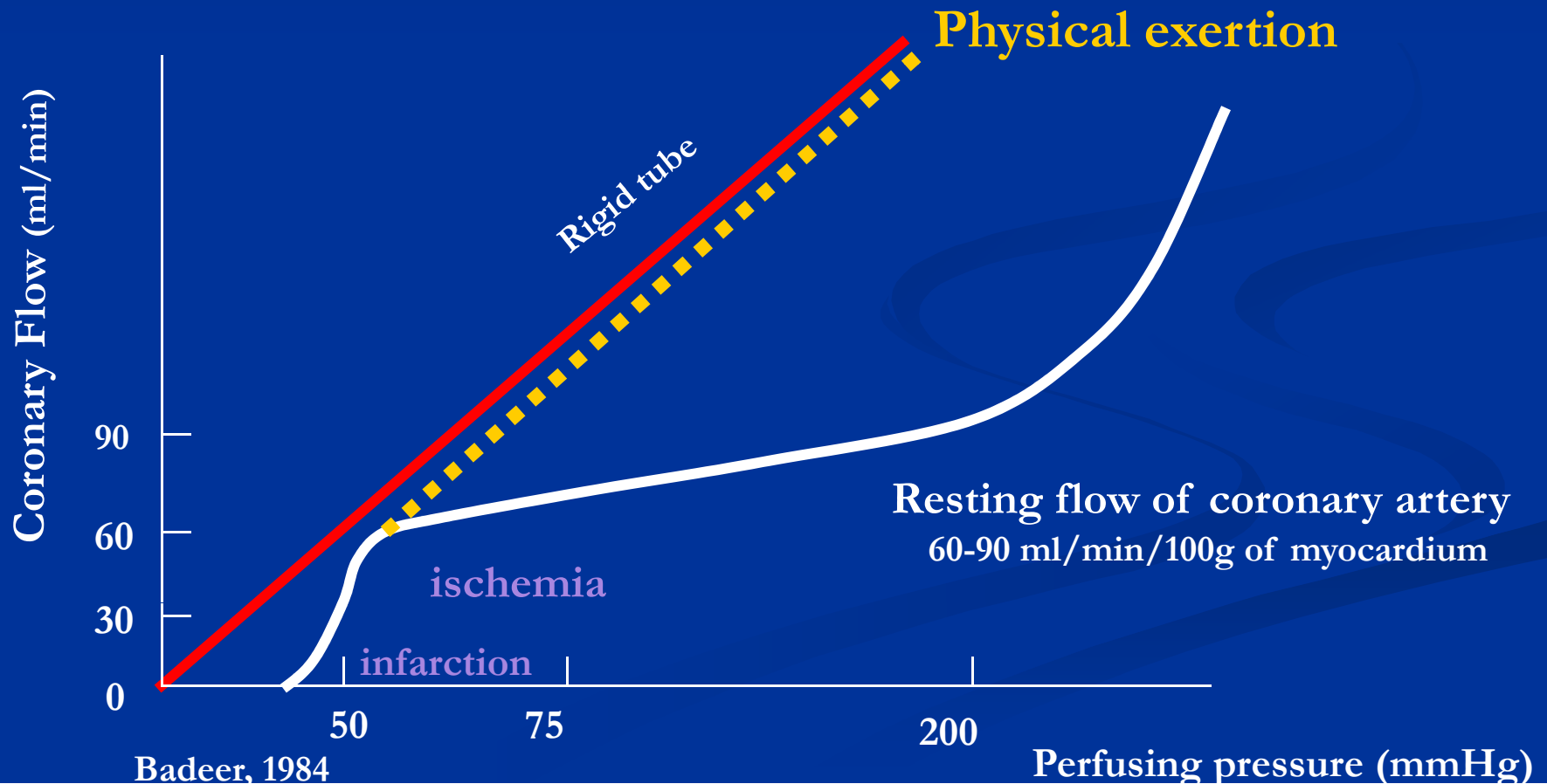
$\alpha = 0.75$ vs. 10



Blood Pressure



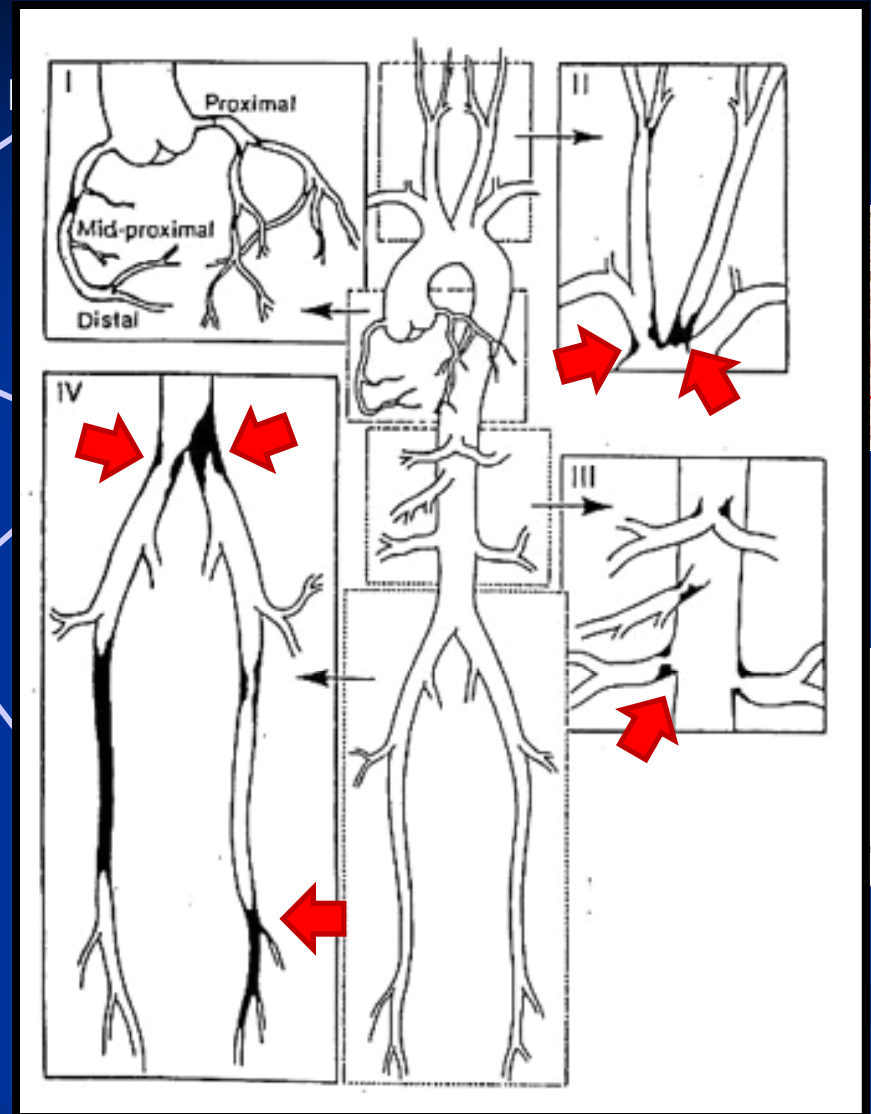
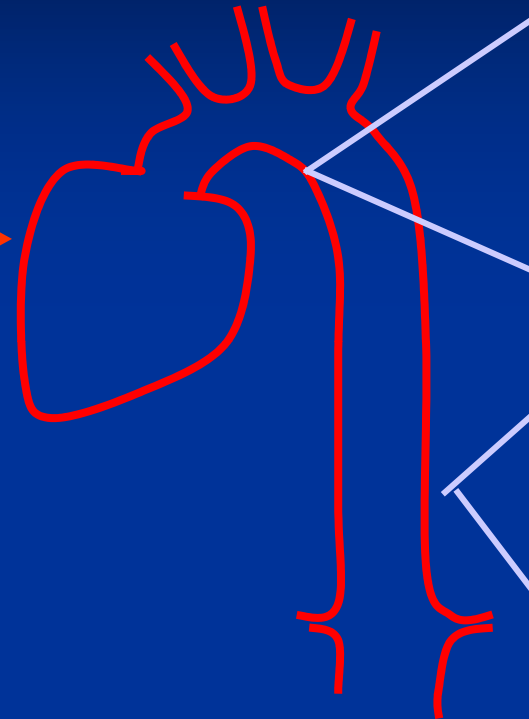
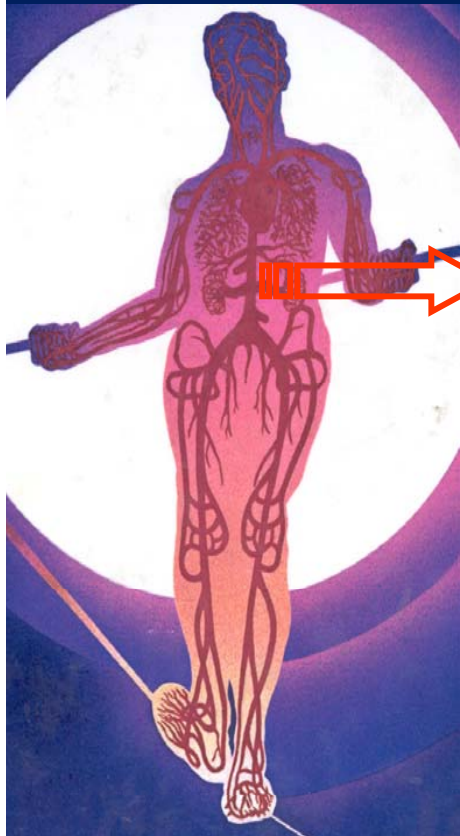
Rigid tube vs. Coronary vascular bed



Objectives

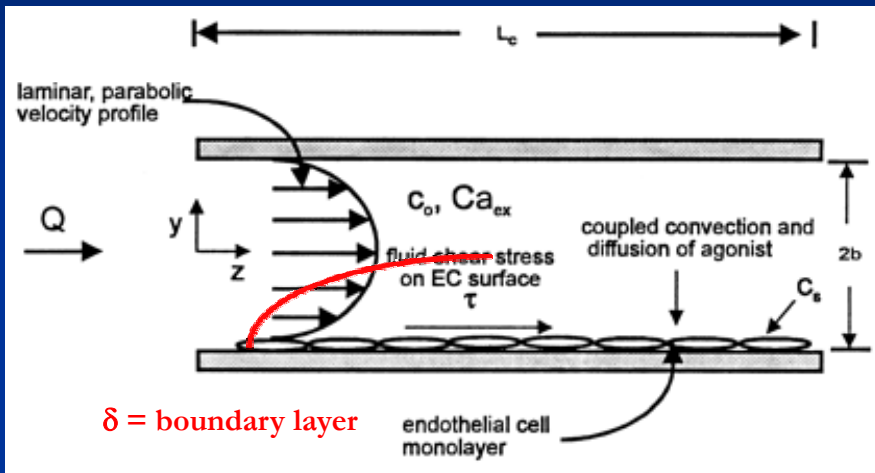
- **Introduce vascular systems**
 - Inertia vs. viscous force
- Interface vascular dynamics with vascular biology
- Translate vascular dynamics to an in vivo model
- Bridge bioengineering, industry, and medicine

Interplay between Hemodynamics and Vascular Biology



Shu Chien, John Cook, John Frango, Scott Simon, R. Narem, M. Gimbrone, P. Davies, L. McIntyre, Don Giddens, David Ku, C. Taylor, C. Zarins

Atherosclerosis and Hemodynamics



Navier-Stokes Equation

$$\rho \left(\frac{\partial \mathbf{v}}{\partial t} + \mathbf{v} \cdot \nabla \mathbf{v} \right) = -\nabla \rho + \mu \nabla^2 \mathbf{v} + \mathbf{f}$$

Transient inertial force

Convective initial force

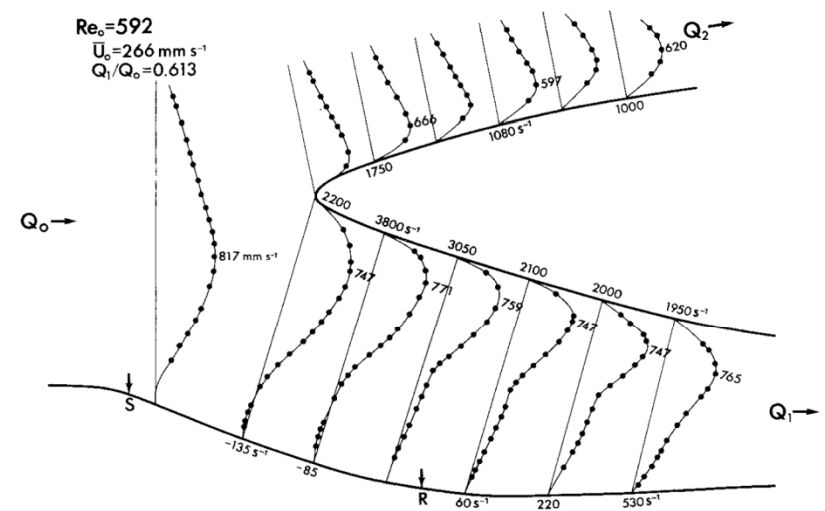
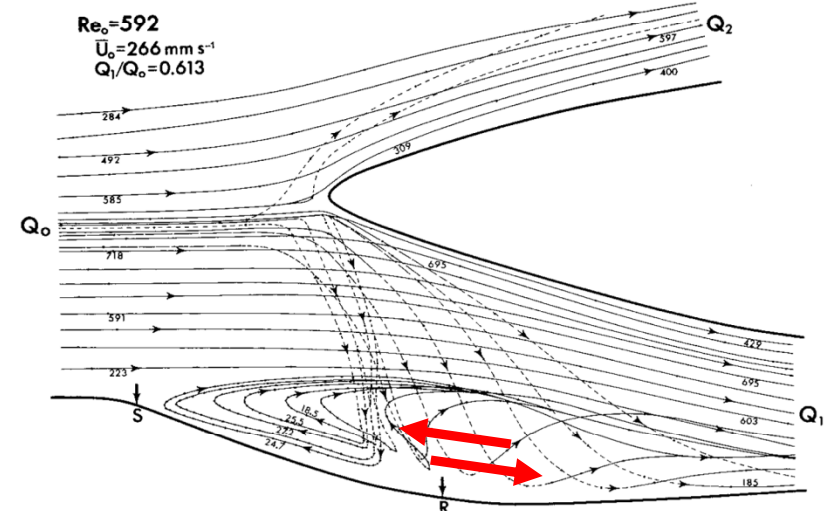
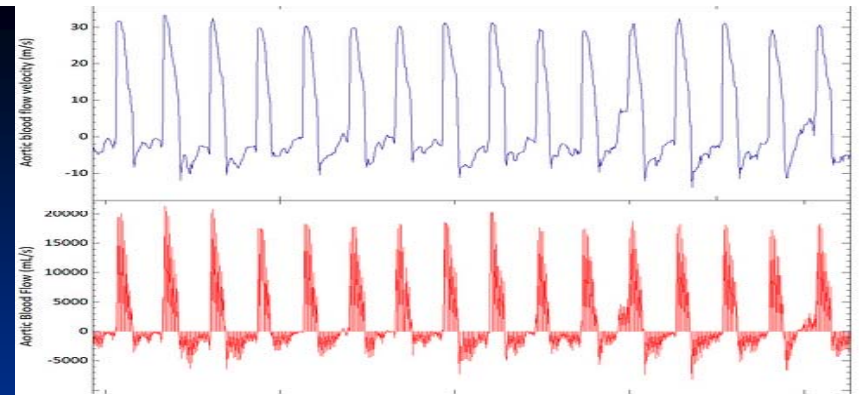
Pressure force

Viscous force

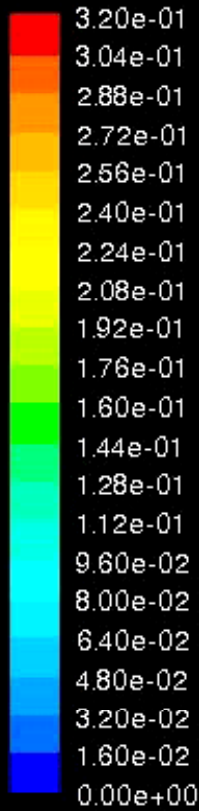
Body force

$$\frac{\partial \tau}{\partial t} = 0, \alpha \leq 1, Re < 1000$$

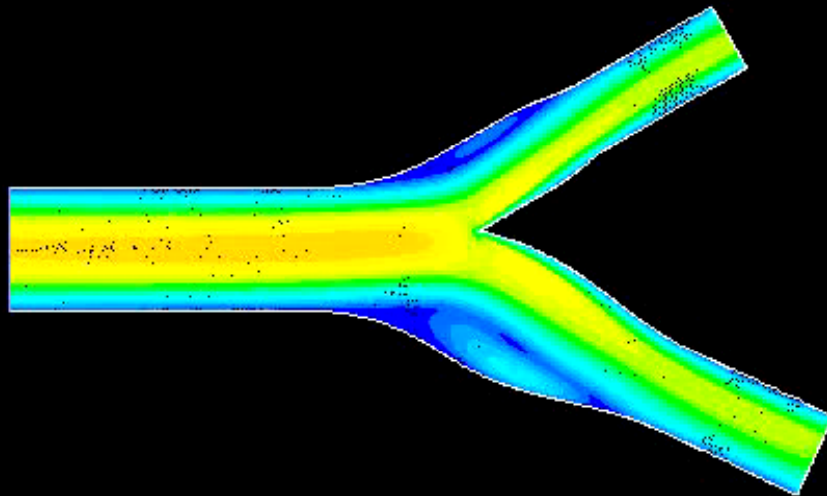
$$\tau_w = \mu \left. \frac{\partial u}{\partial y} \right|_{y=0}$$



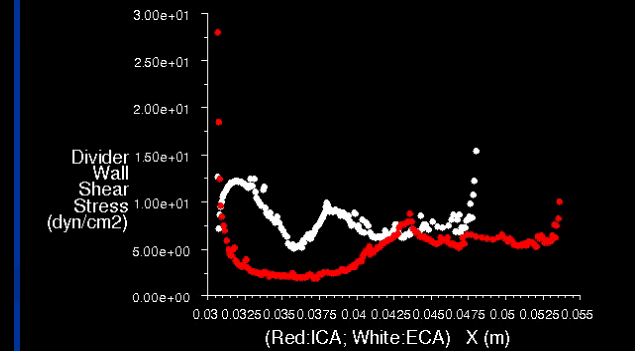
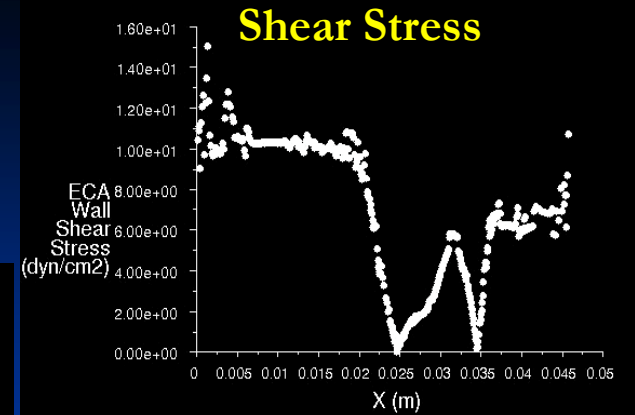
Unsteady State CFD of Human Carotid Artery



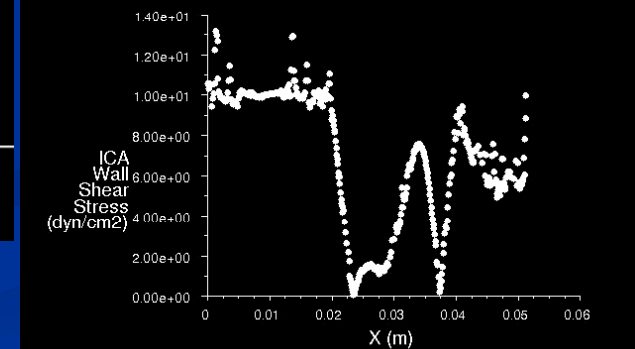
Velocity Profiles



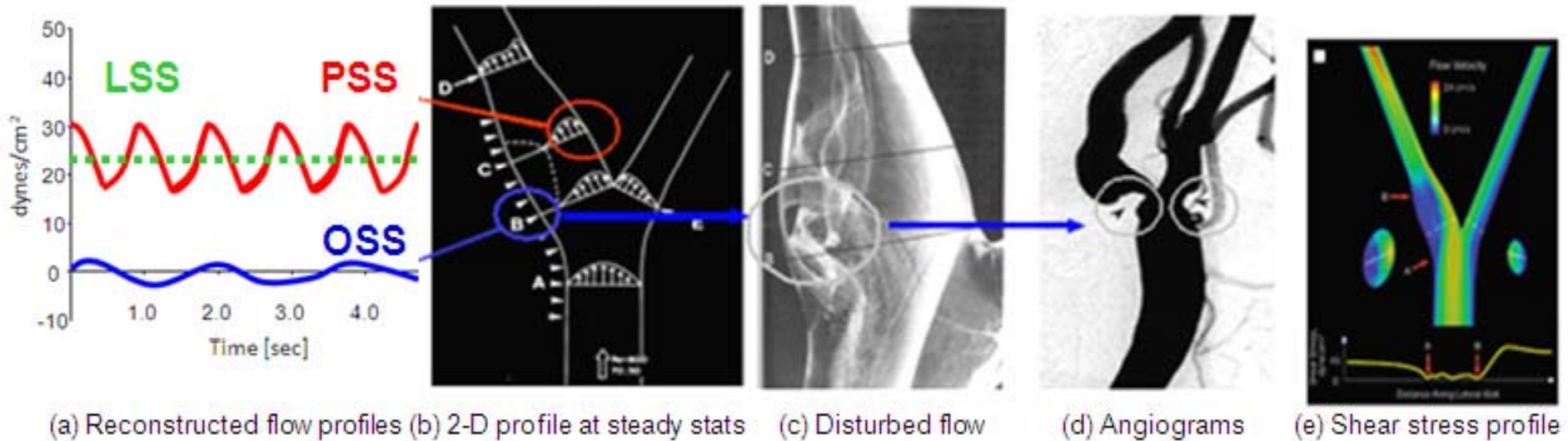
Contours of Velocity Magnitude (m/s) (Time=1.0100e+00) Feb 03, 2007
FLUENT 6.2 (2d, segregated, lam, unsteady)



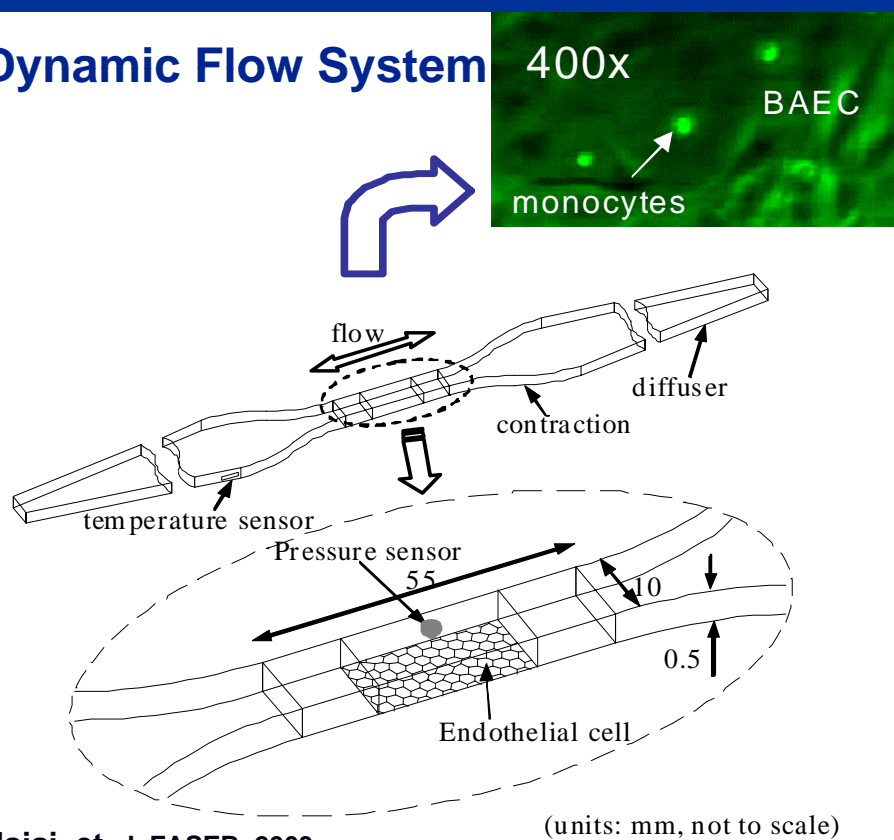
Wall Shear Stress (Time=1.0100e+00) Feb 03, 2007
FLUENT 6.2 (2d, segregated, lam, unsteady)



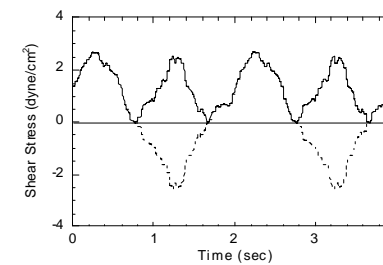
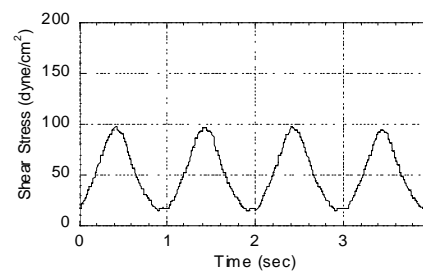
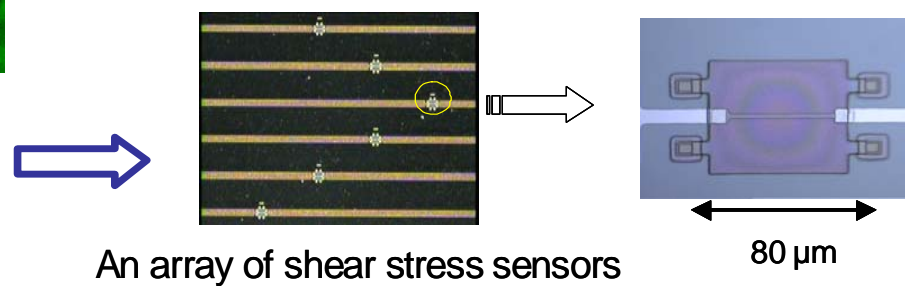
Wall Shear Stress (Time=1.0100e+00) Feb 03, 2007
FLUENT 6.2 (2d, segregated, lam, unsteady)



Dynamic Flow System



Hsiai, et al, FASEB, 2003



pulsatile shear stress oscillatory shear stress

(1)

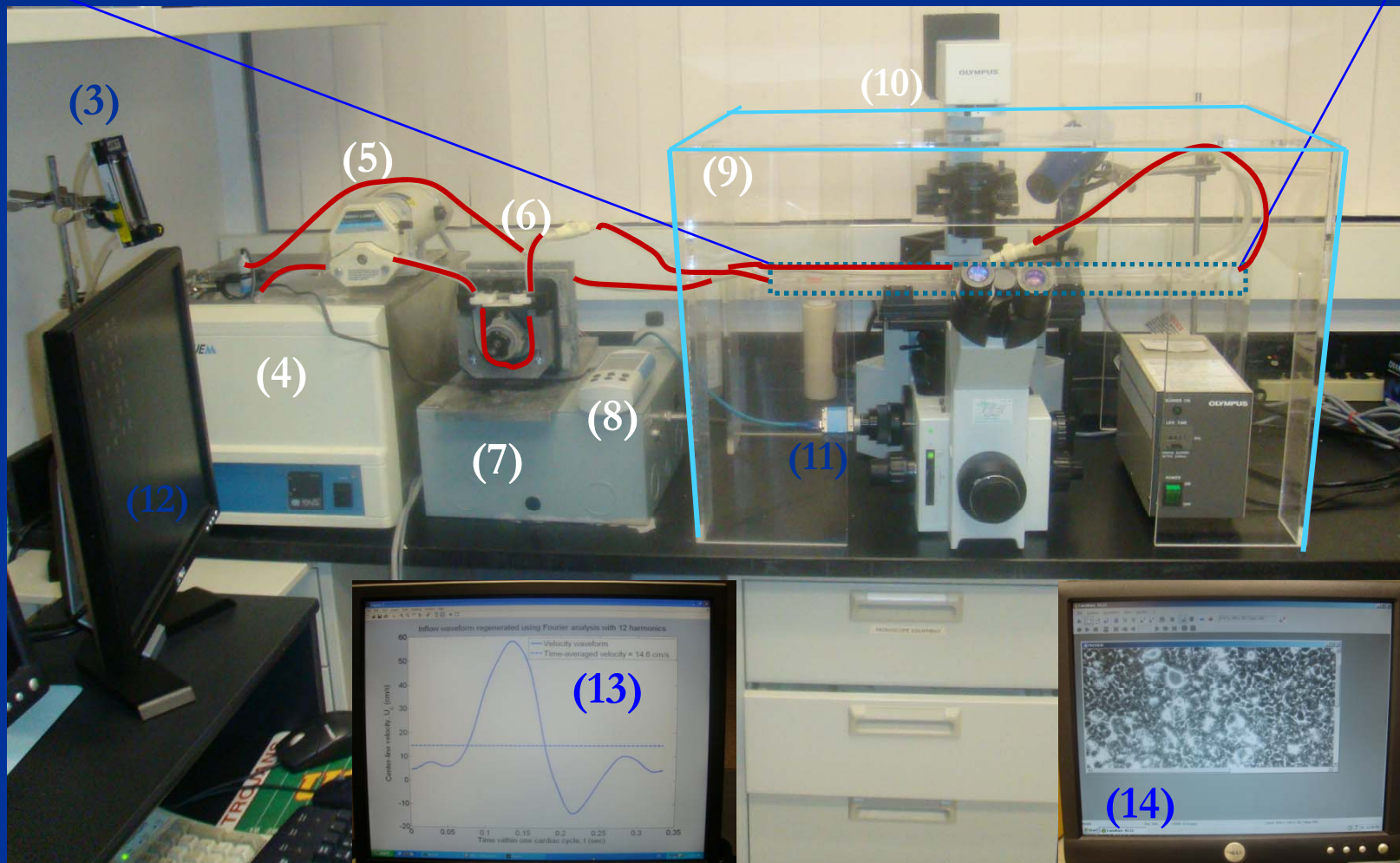


Introduction of MEMS sensors

(2)



(3)



(4)

(5)

(6)

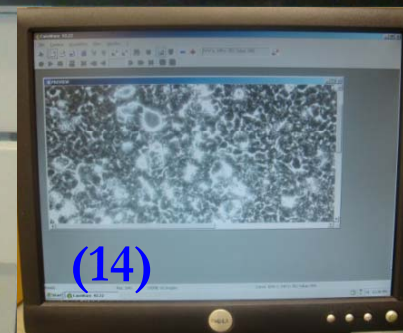
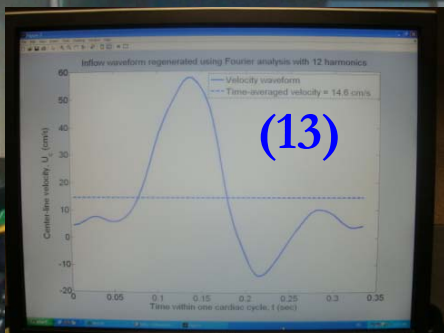
(7)

(8)

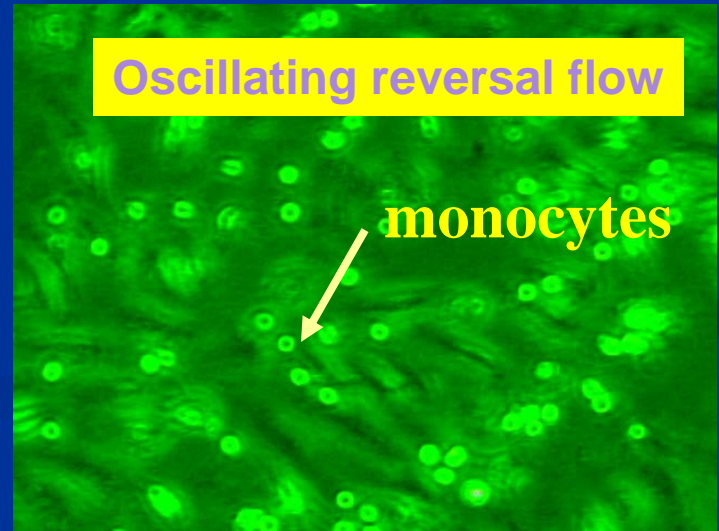
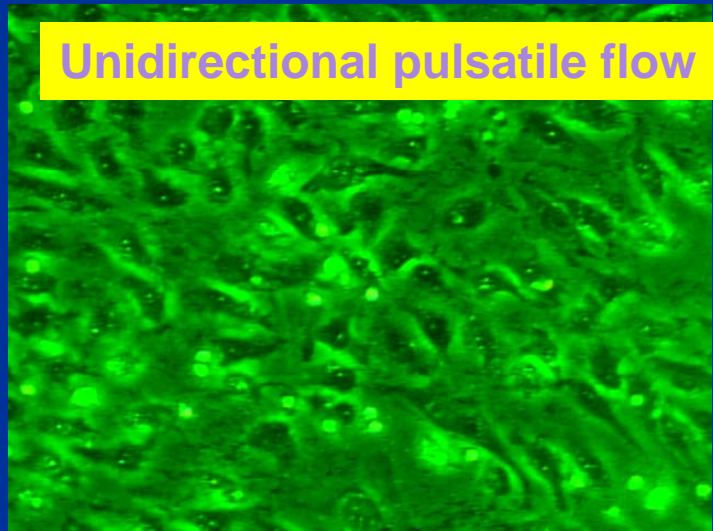
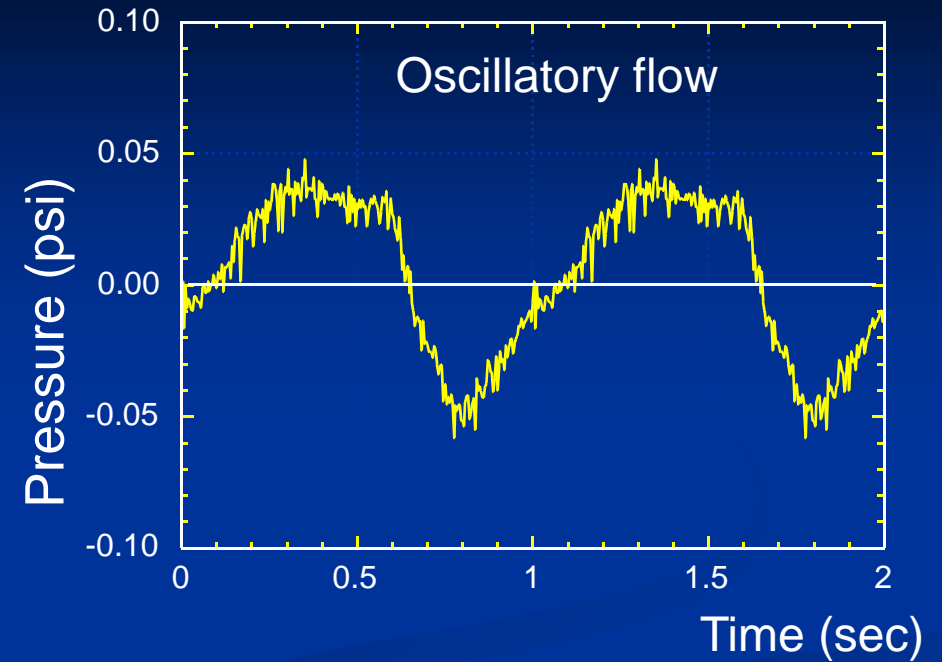
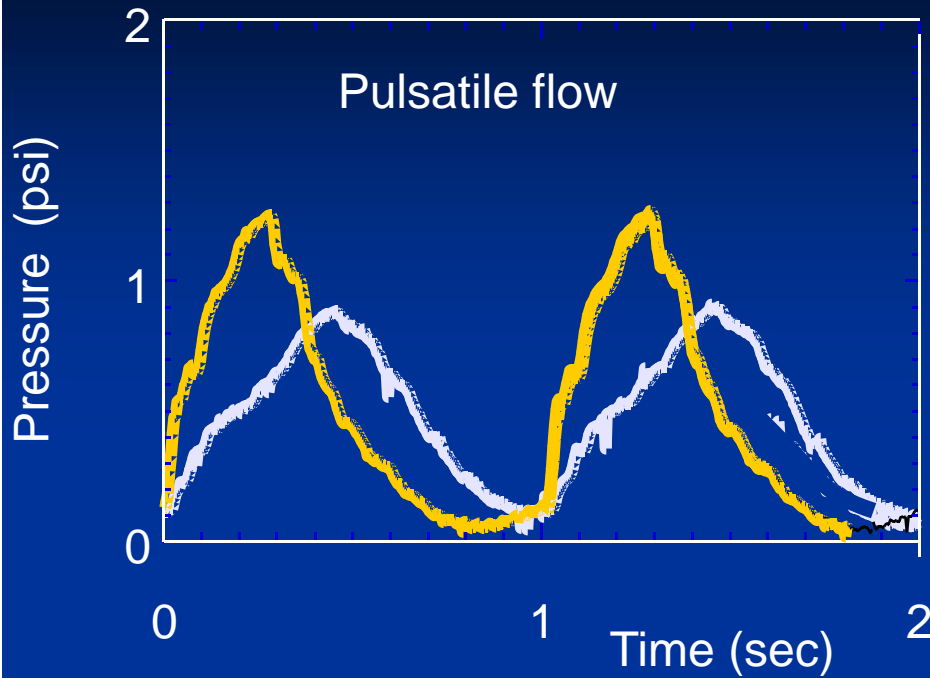
(9)

(10)

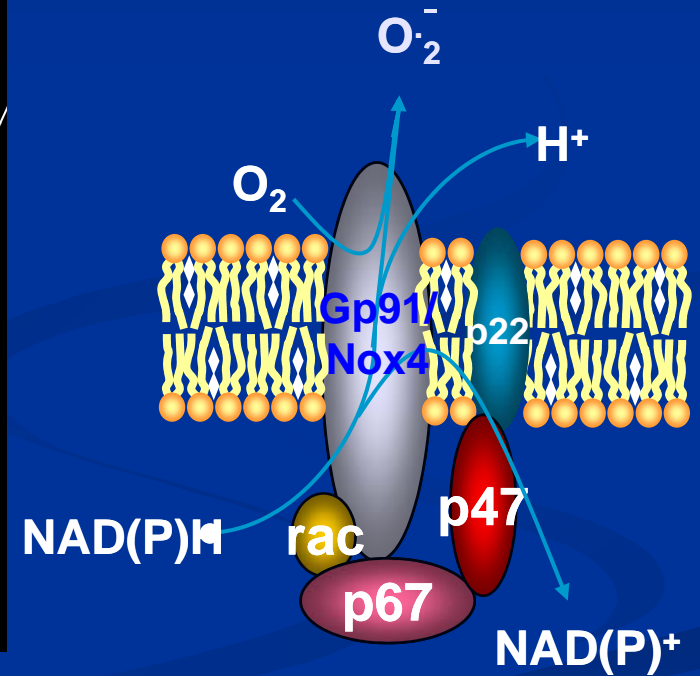
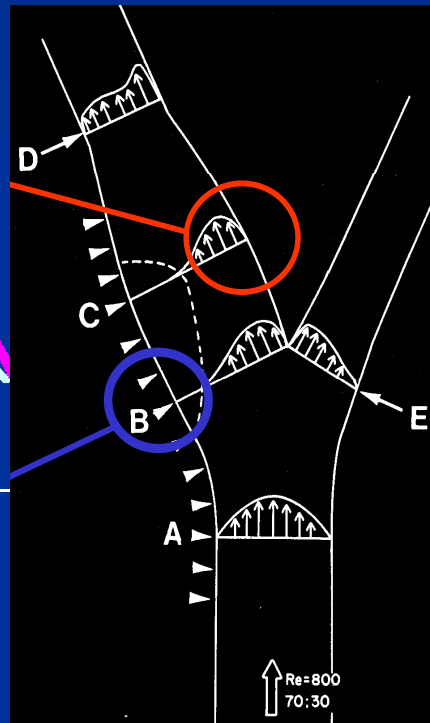
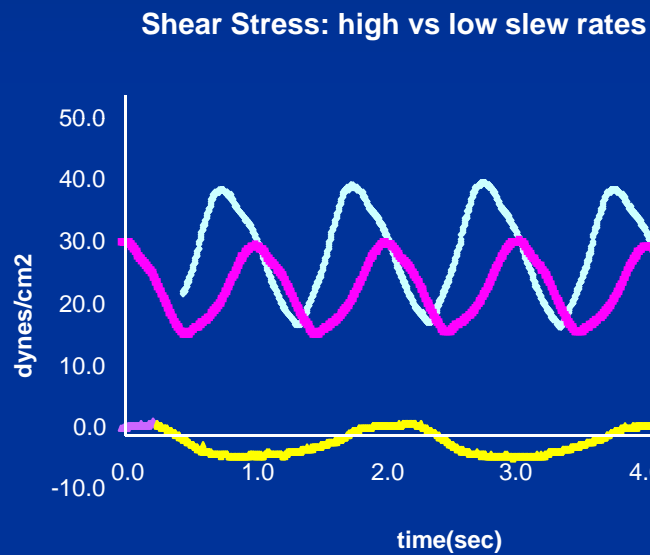
(12)



Monocyte Attachment to Endothelial Cells

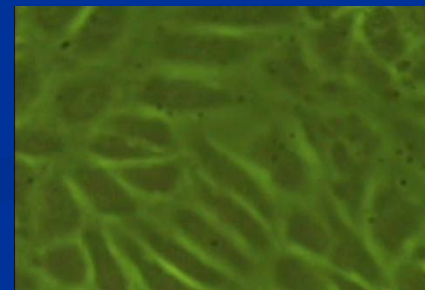
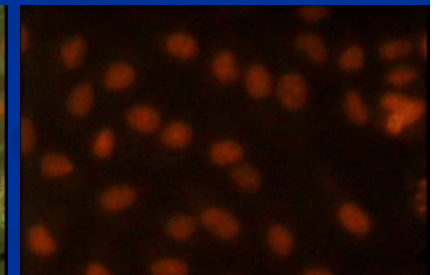
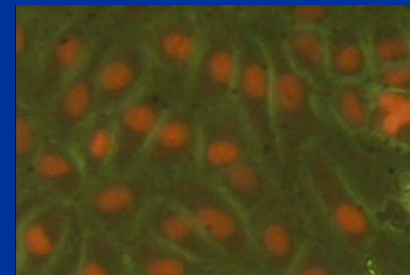
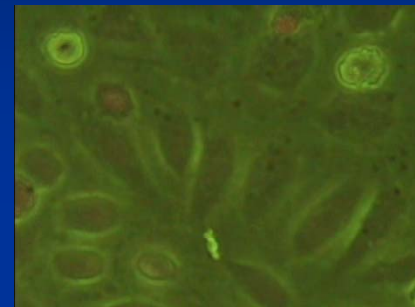
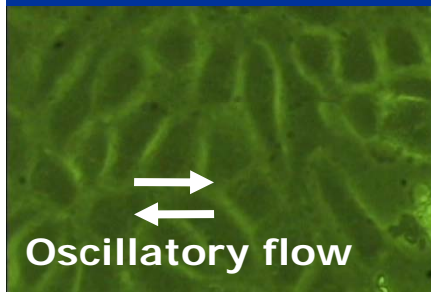
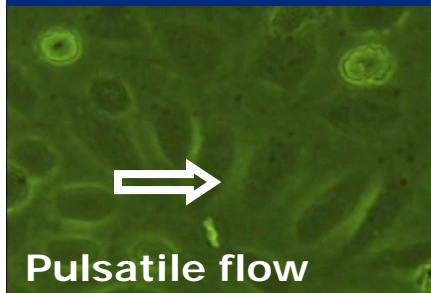


Shear stress induces NADPH oxidase system



Shear Stress and Vascular Oxidative Stress

Dihydroethidium Bromide for $O_2^{\cdot-}$

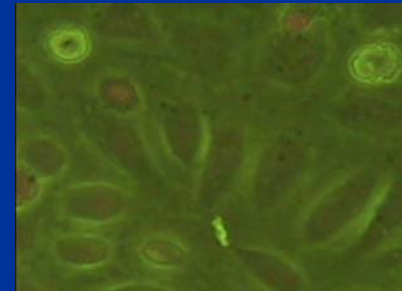
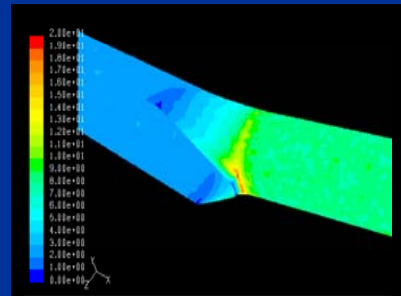
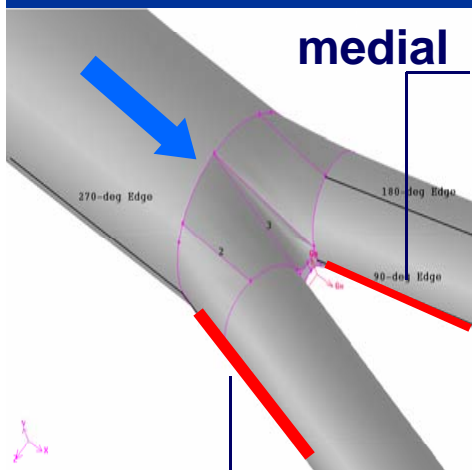


Ku, Giddens, Zarins

Hsiai et al, Circ Res. 2003

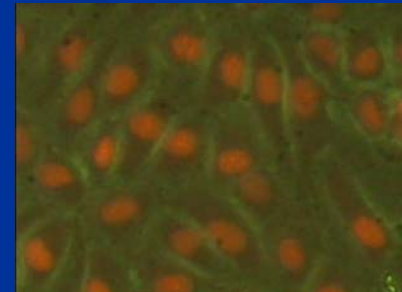
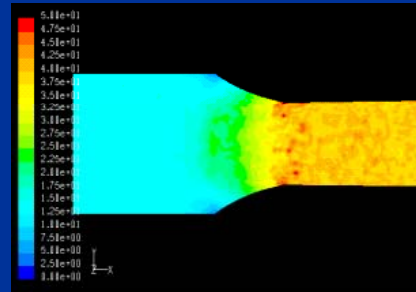
Spatial variations in shear stress and vascular oxidative stress

LDL oxidation and nitration



\uparrow eNOS
 \downarrow Nox4
 \downarrow O₂⁻

} \uparrow RNS/ROS

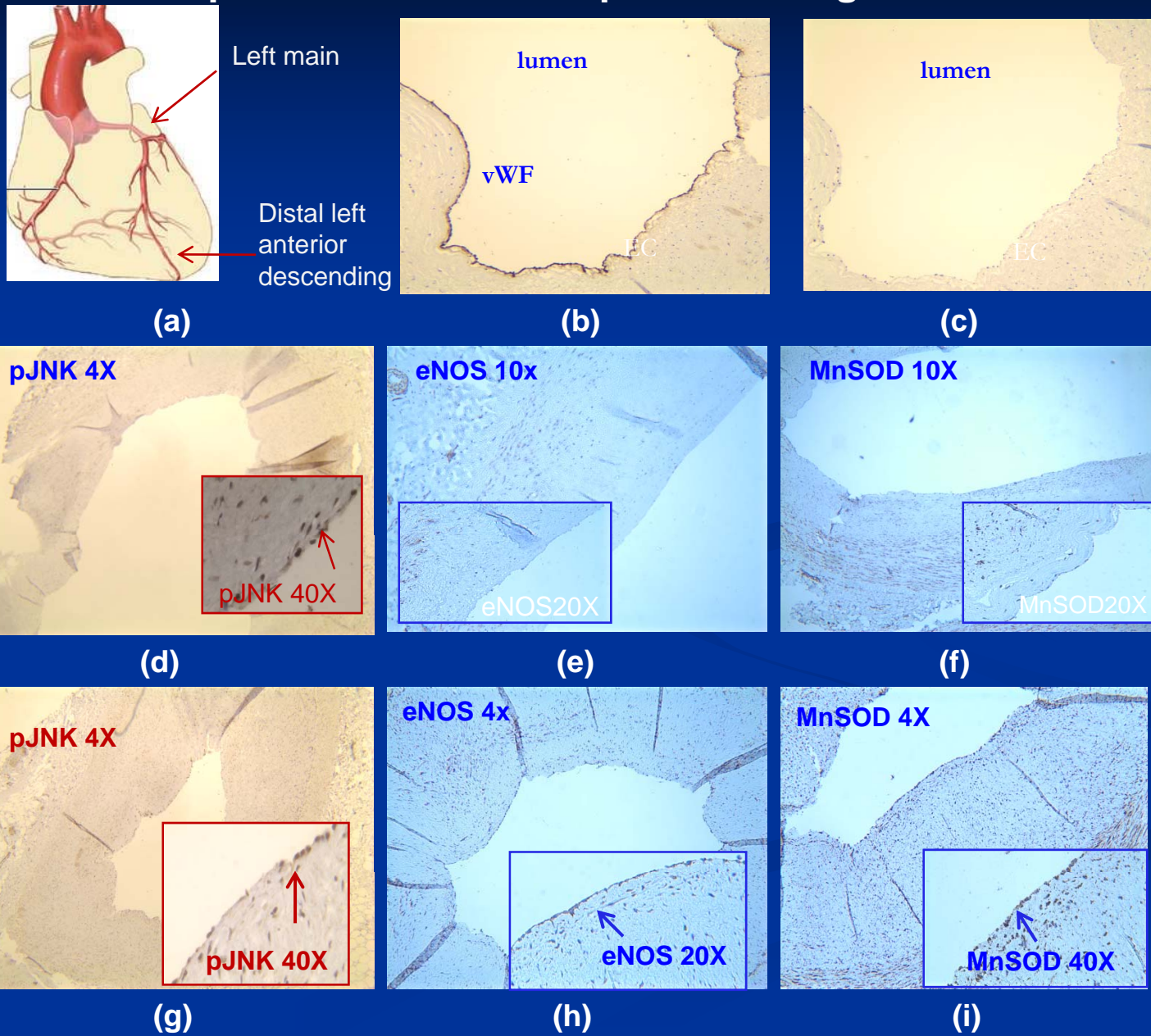


\downarrow eNOS
 \uparrow Nox4
 \uparrow O₂⁻

} \downarrow RNS/ROS

lateral

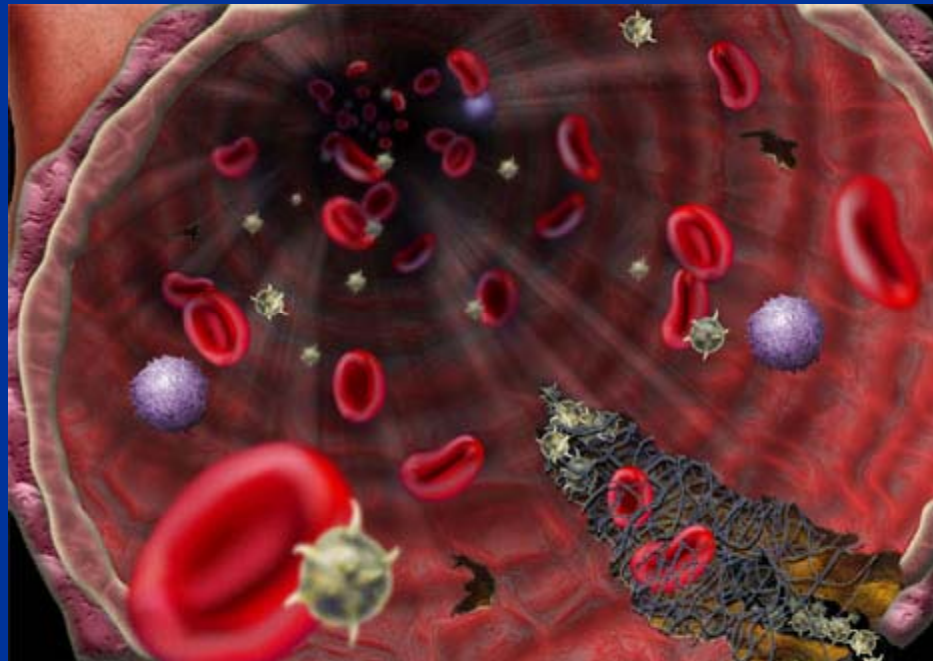
JNK activation and antioxidant expression in the athero-prone versus athero-protective regions



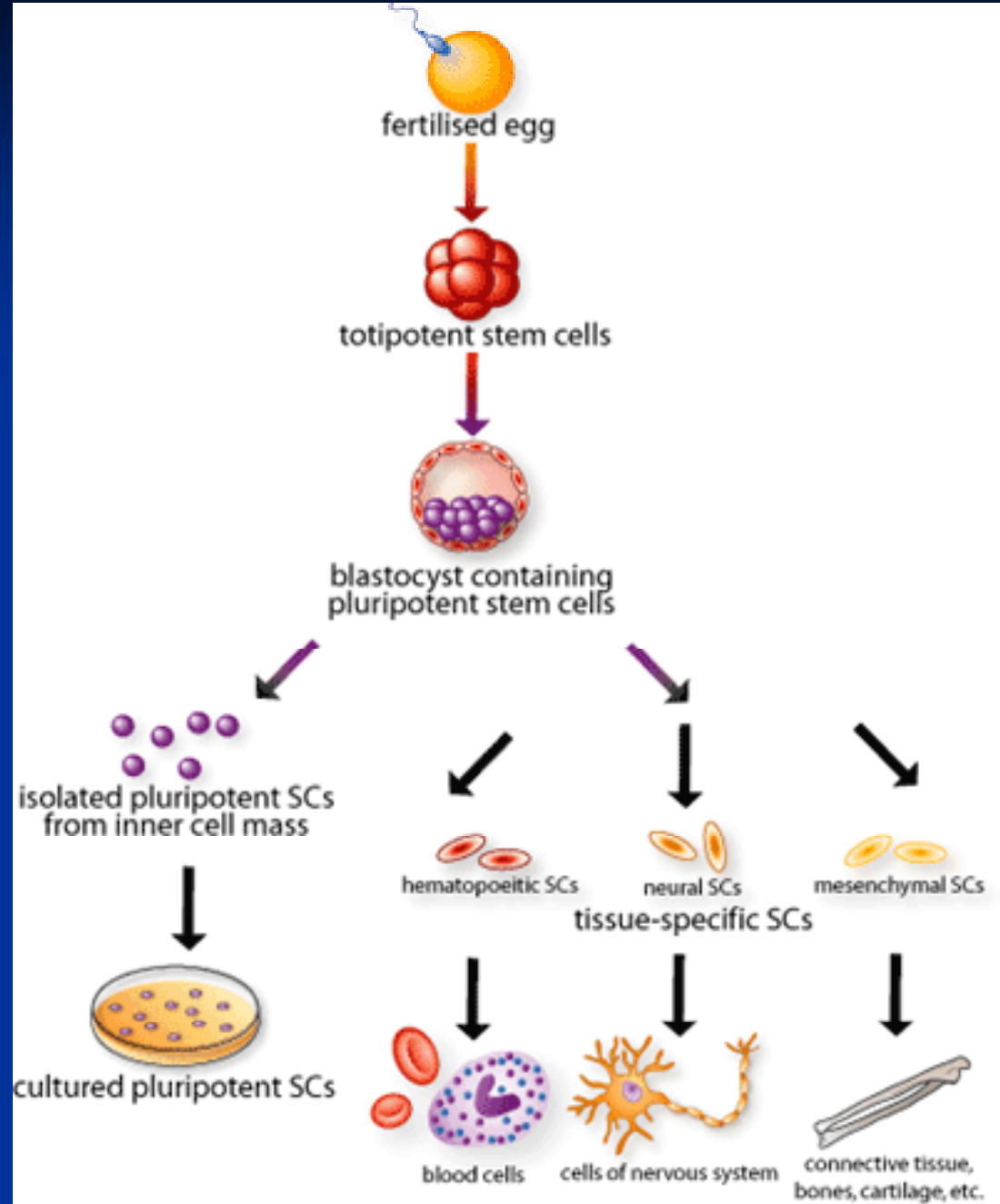
Hemostasis

In response to blood vessel injuries, the blood under three process to stem the flow of blood:

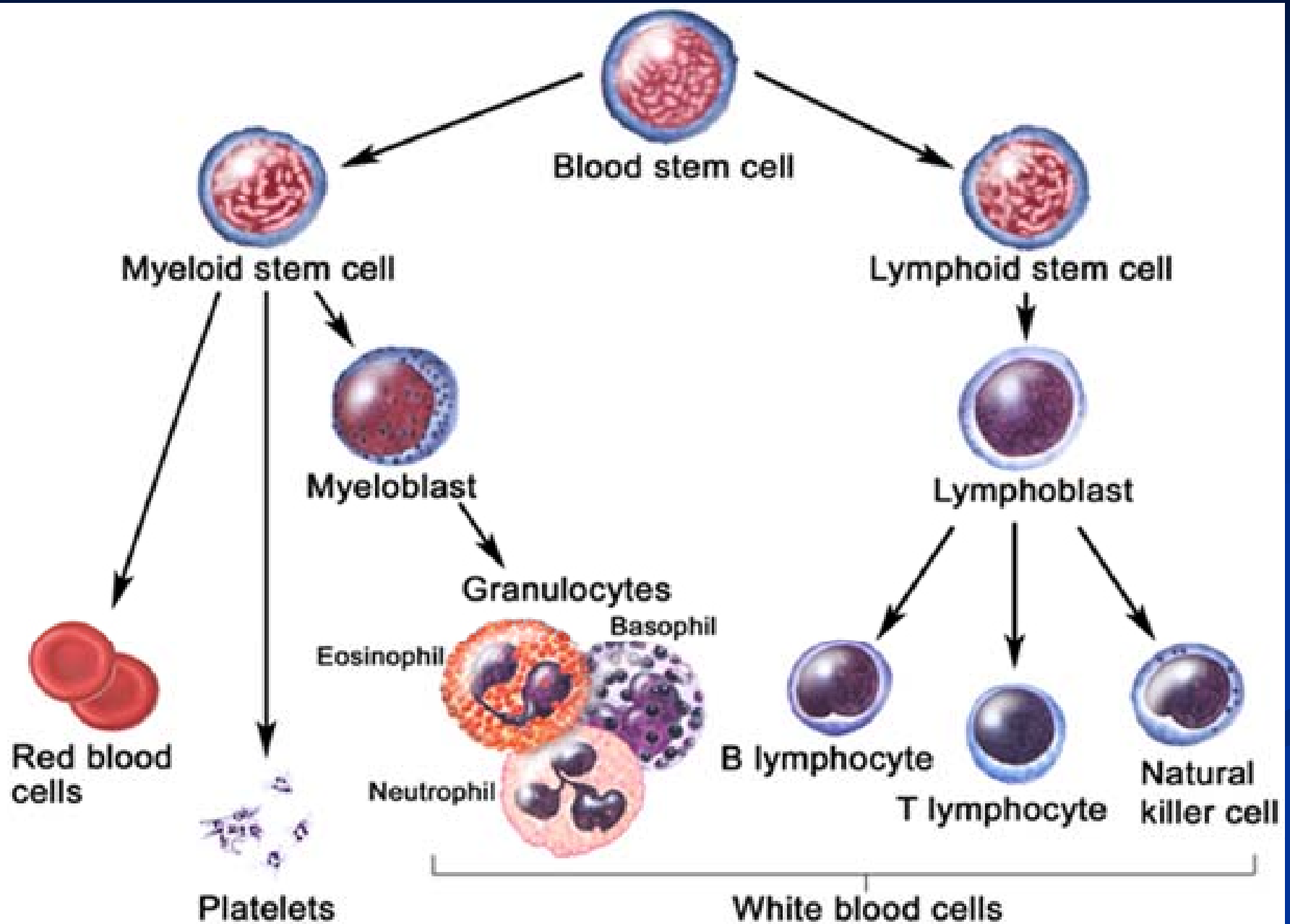
1. Vasoconstriction (endothelium, smooth muscle cells)
2. Platelet aggregation, and
3. Blood coagulation (coagulation cascade)



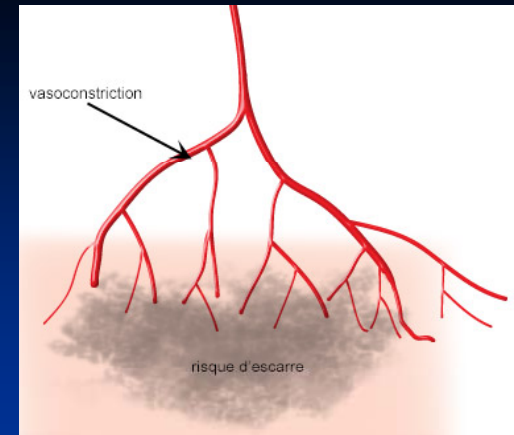
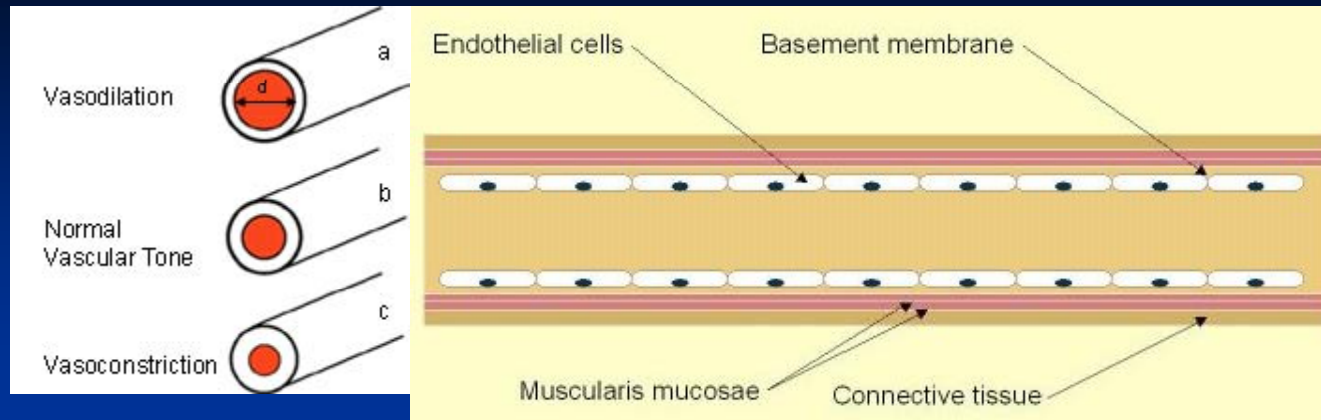
Stem Cells



Part I: Blood cells



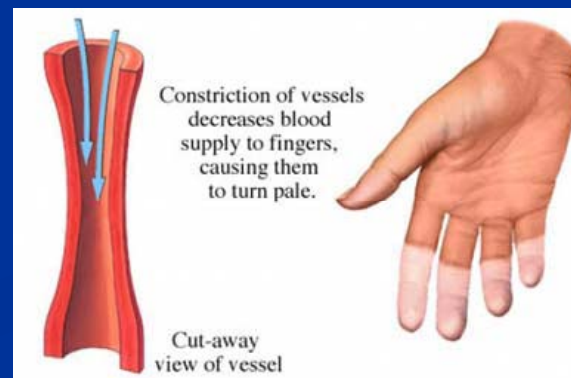
Vasoconstriction



-Injury to a blood vessel elicits a contractile response by the vascular smooth muscle, resulting in a narrowing of the vessel.

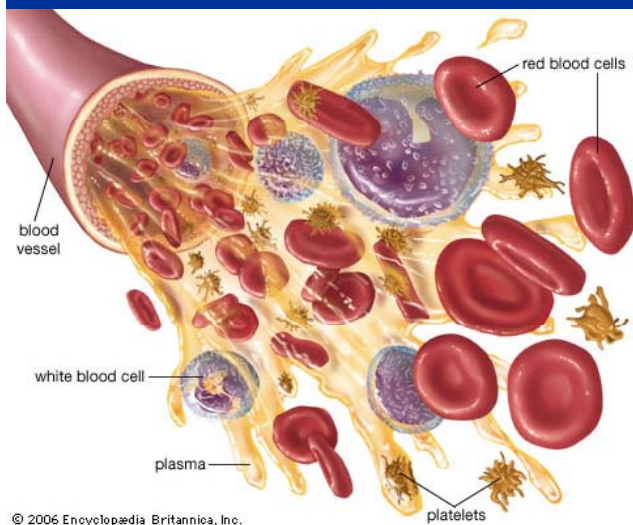
-Vasoconstriction in severed arterioles or small arteries can completely obliterate the lumen of the vessel to stop blood flow.

-The contraction of the vascular smooth muscle is induced by direct mechanical stimulation (GTP, increased in intracellular calcium, etc.) as well as by mechanical stimulation of perivascular nerves.

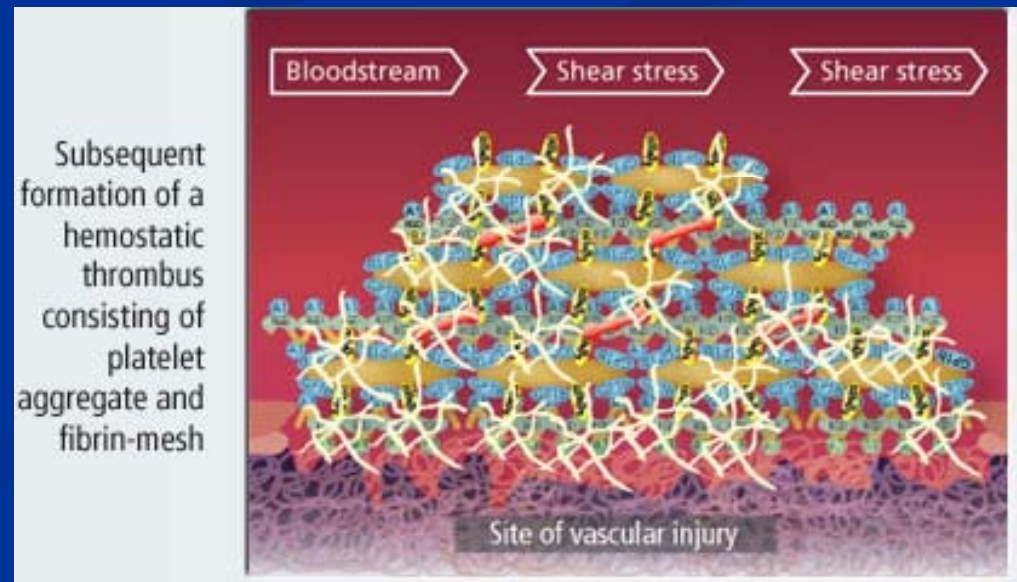


Hemostasis

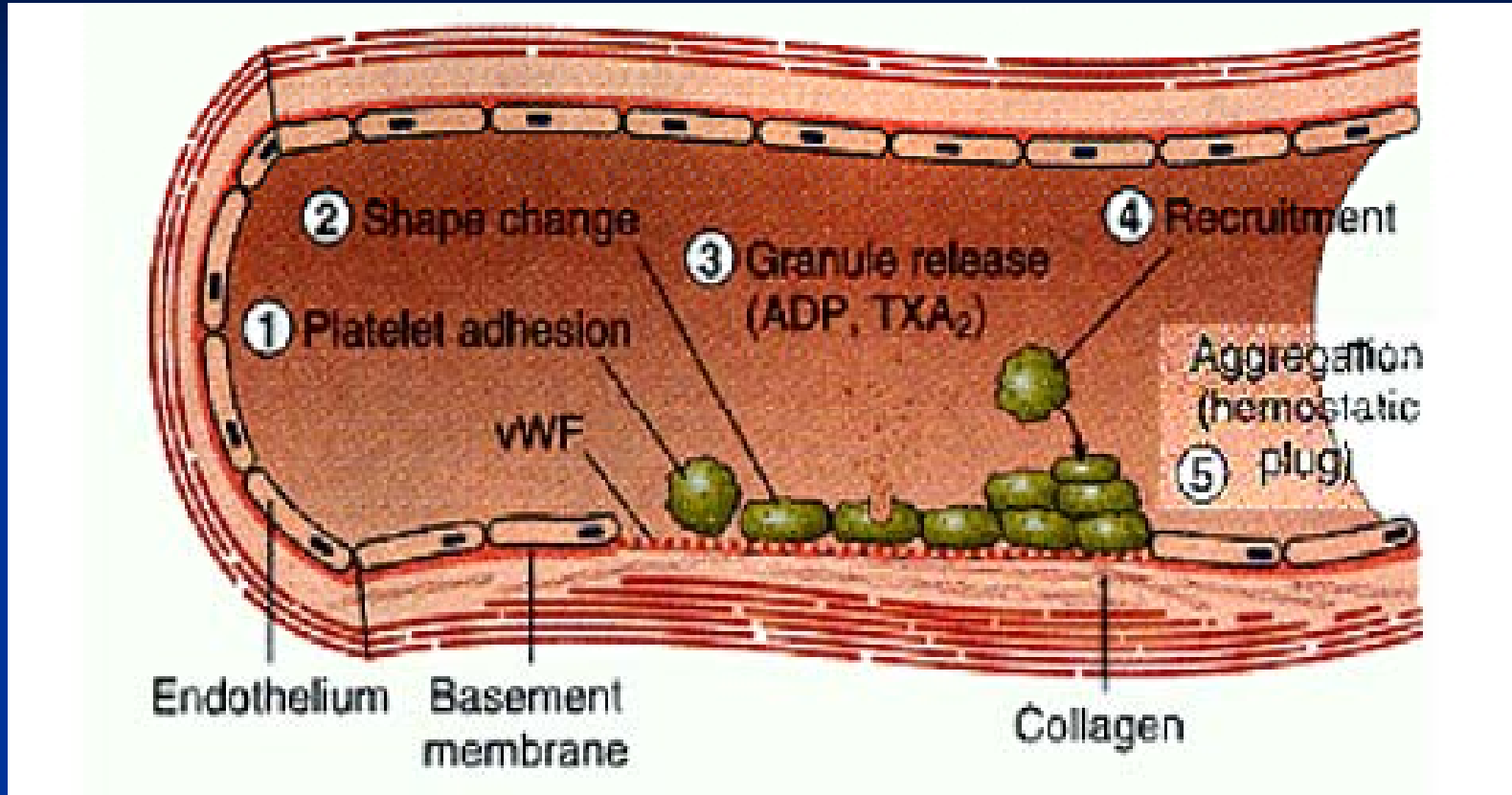
- Following injury to a blood vessel, all of the systems are activated.
- The hemostatic process is divided into 2 components;
 - primary hemostasis
 - Primary hemostasis depends upon the response of the platelet and blood vessel wall to the injury. When the small blood vessels are injured, blood platelets adhere and aggregate at the site of injury, reducing and finally arresting bleeding.
 - secondary hemostasis
 - Secondary hemostasis starts when the cascade system of coagulation is activated by substances released at the time of blood vessel injury.



© 2006 Encyclopædia Britannica, Inc.



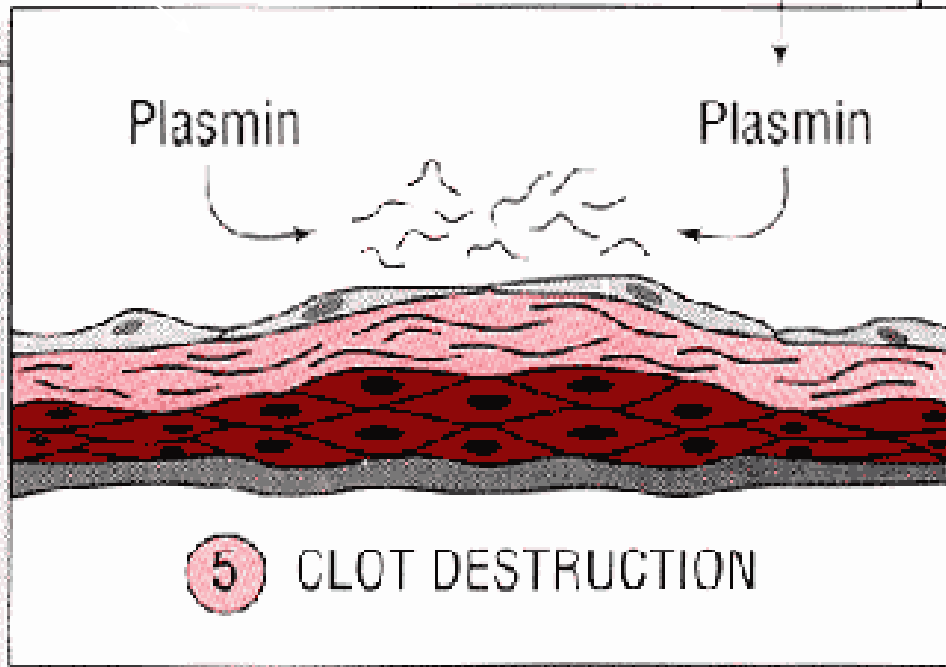
Primary Hemostasis



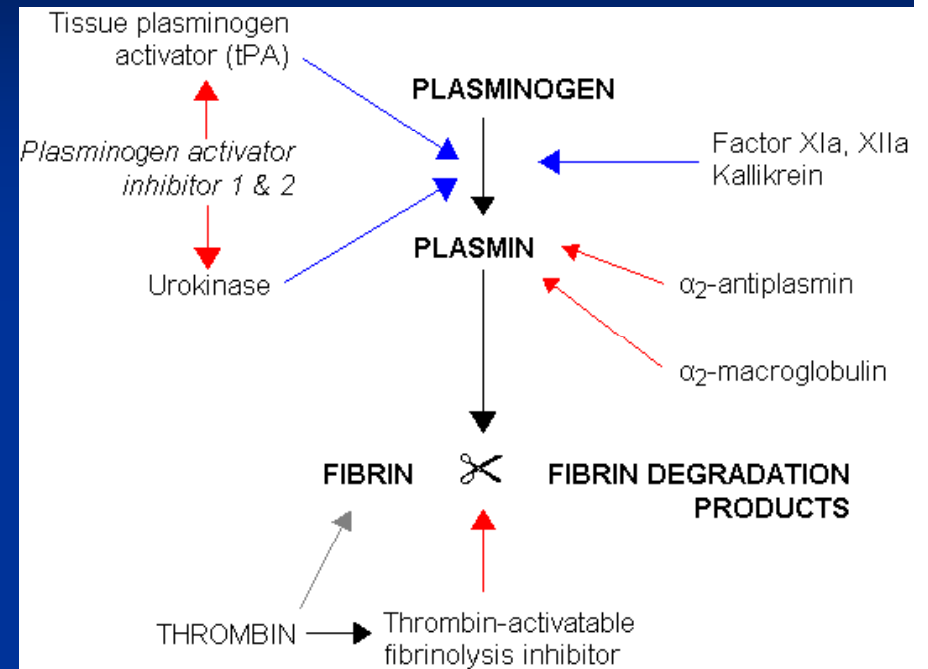
Primary hemostasis depends upon the response of the platelet and blood vessel wall to the injury. When the small blood vessels are injured, blood platelets adhere and aggregate at the site of injury, reducing and finally arresting bleeding.

Plasminogen

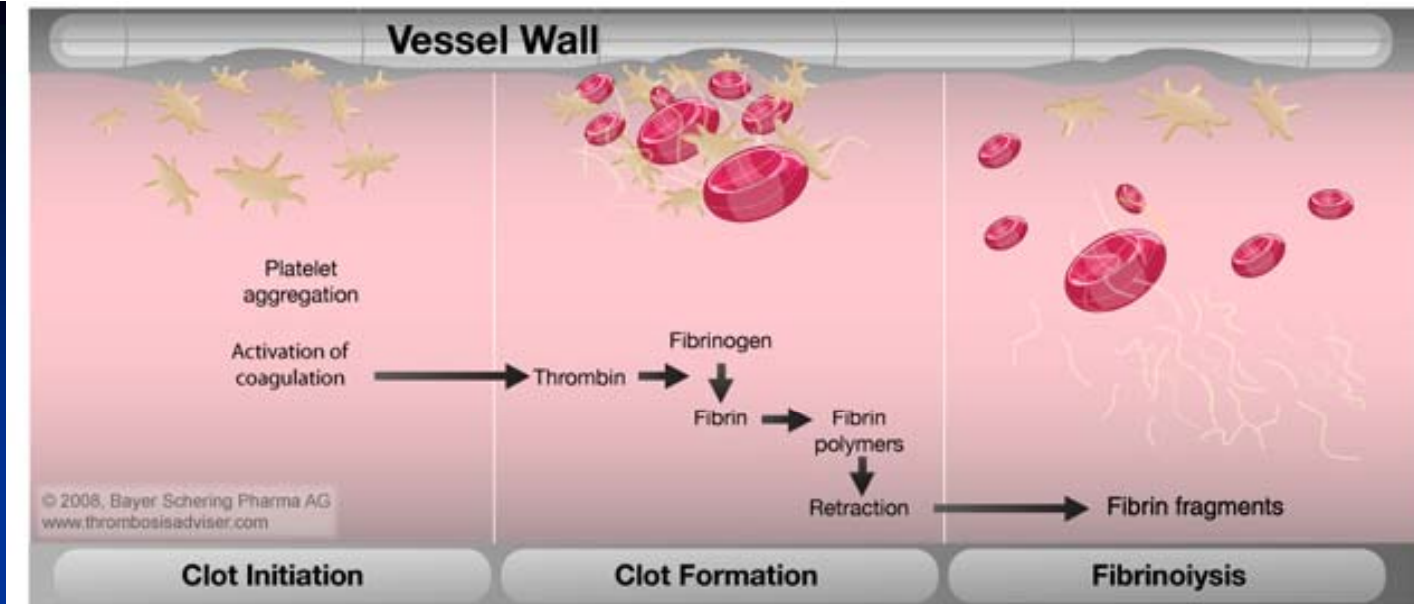
Clot Lysis: fibrinolysis



Enzymatic destruction of clot



The last stage of coagulation is fibrinolysis, which is the dissolution and localization of a fibrin clot. These functions are carried out by enzymes and their inhibitors. A disruption or breach of the fine balance of this fibrinolytic system can result in bleeding or thrombosis.



Fibrinolysis is mediated by activation of plasminogen to plasmin.

This is accomplished by:

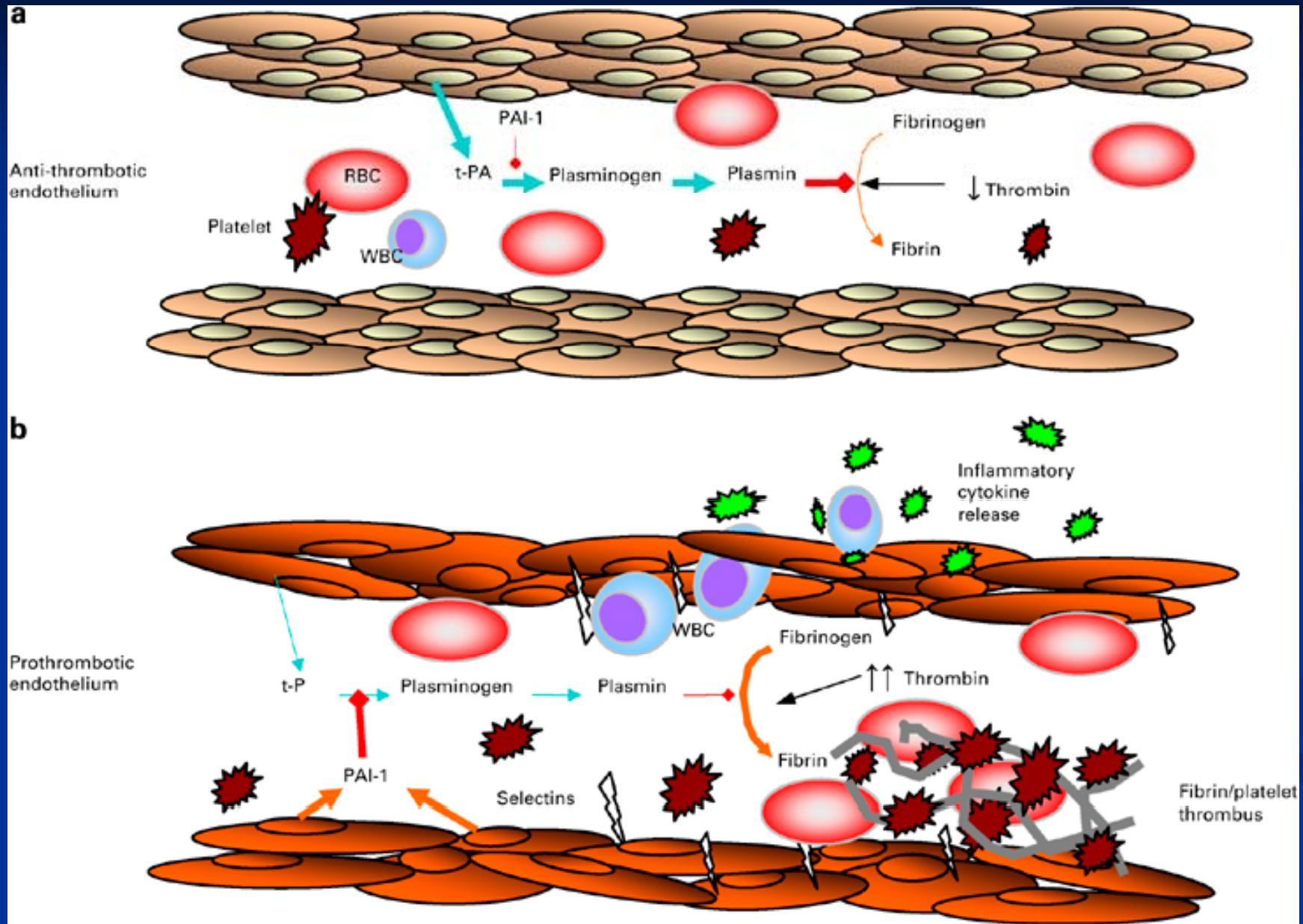
Intrinsic activation (plasma based) initiated through Factor XIIa and allikrien. Thus, the contact system of coagulation serves as an intrinsic activator.

Extrinsic activation (cellular based) initiated by way of stimuli such as vascular injury, ischemia, exercise, stress and pyrogens.

Exogenous

(Therapeutic) activation (drug based) includes streptokinase, urokinase and tPA tissue plasminogen activator).

Summary



At equilibrium

Key Concepts of Part II

Blood coagulation

Clot lysis

Blood coagulation

Bleeding

Clot lysis

Blood clots

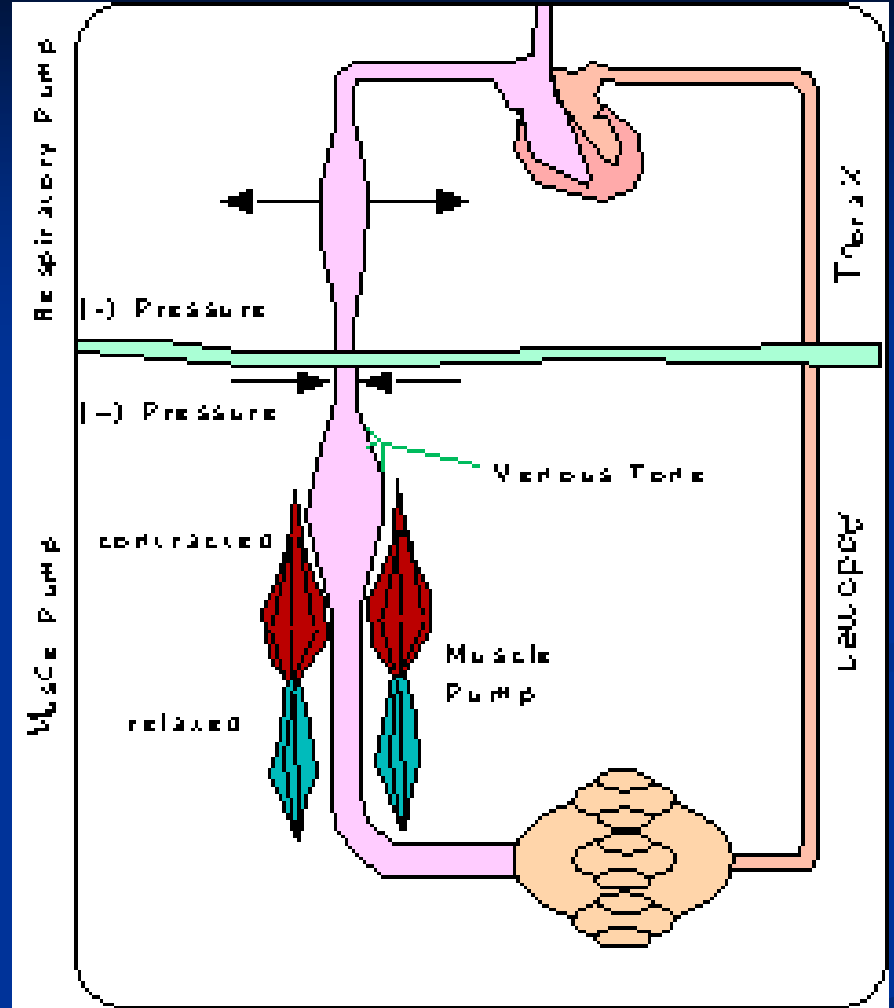
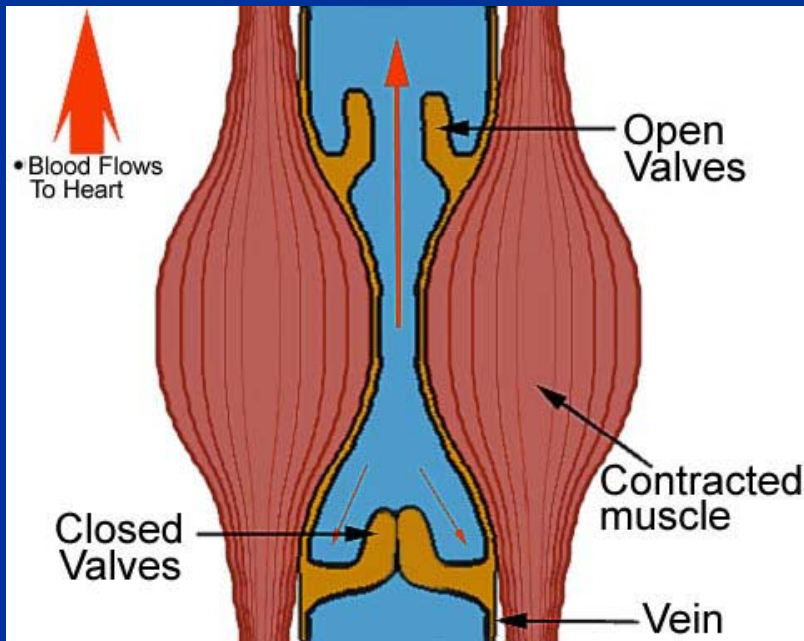
Clot lysis

Blood coagulation

Deep Venous Thrombosis



Flickr/Corbis/Bett Co. | Photo: Corbis/Bett Co. | Photo: Corbis/Bett Co.



Contraction of muscles helps the blood to be pushed up the vein.

1998 Nobel Prize to cardiovascular system: NO (nitric oxide)

ROBERT F. FURCHGOTT (State University of New York, Downstate)

LOUIS J. IGNARRO (UCLA)

FERID MURAD (U Texas, Huston)

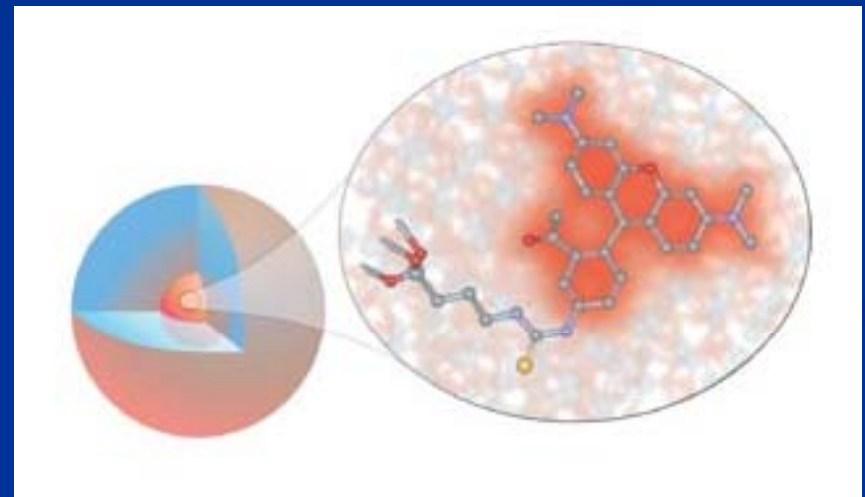
for their discoveries concerning nitric oxide as a signaling molecule in the cardiovascular system.

Objectives

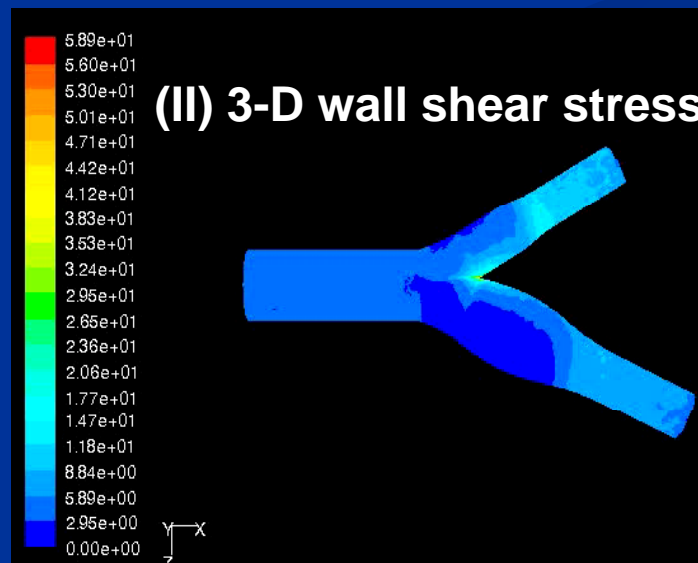
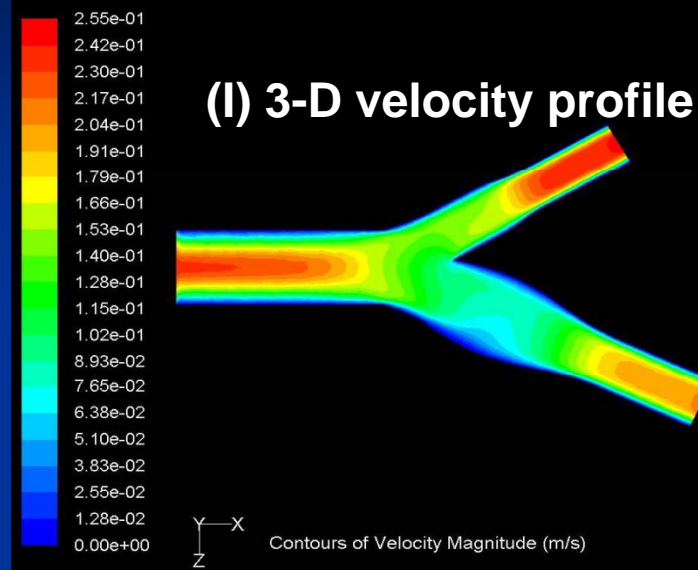
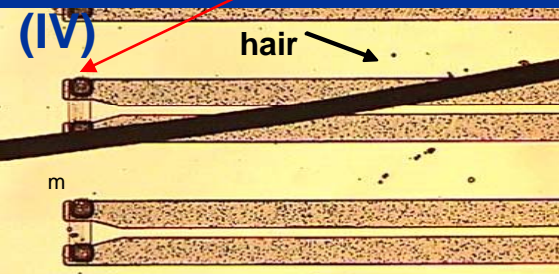
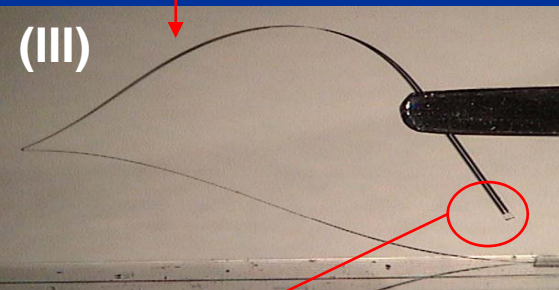
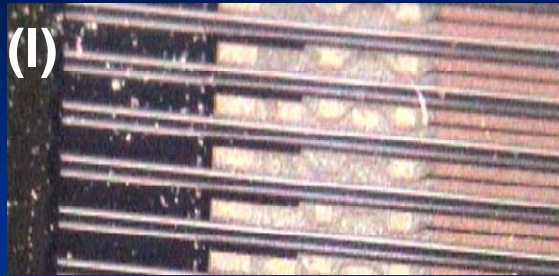
- **Introduce vascular systems**
 - Inertia vs. viscous force
- **Interface vascular dynamics with vascular biology**
 - Hemodynamics and vascular oxidative stress
 - Balance between thrombosis and thrombolysis
- **Translate vascular dynamics to an *in vivo* model**
- **Bridge bioengineering, industry and medicine**

Cardiovasculature: A Dynamic Sensing and Actuating System

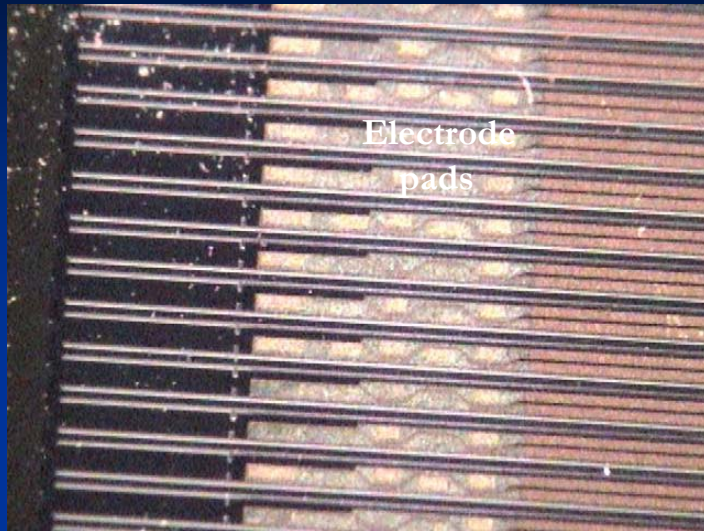
- Too monotonous
- Too easy
- Too biological
- Too clinical
- Too little MEMS
- Too little class interactions
- Too little engineering
- Too much basic sciences
- Too little depth
- Too little breadth



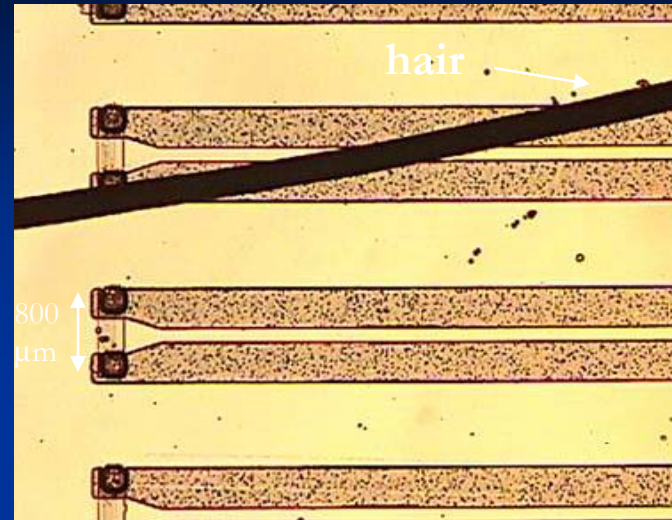
Translating MEMS to Assess Pathophysiology of Atherosclerosis



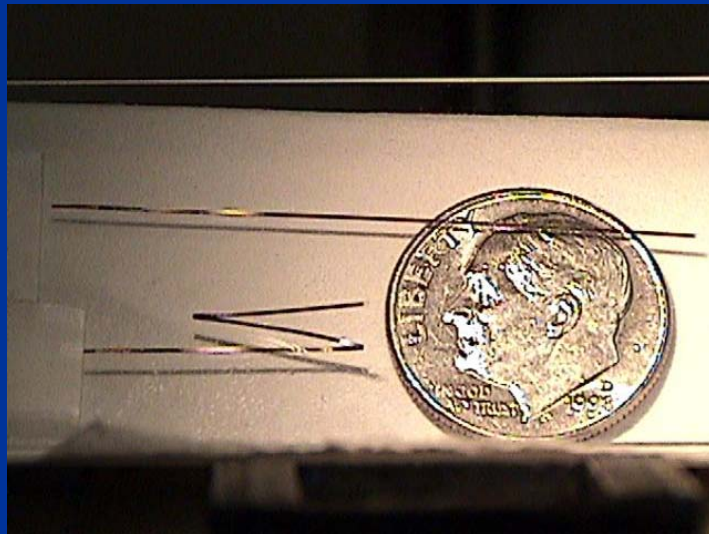
Flexible Polymer MEMS Sensors



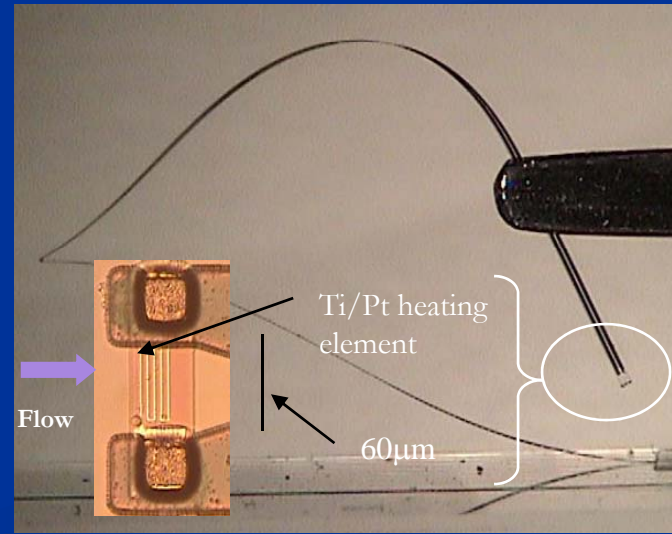
(a)



(b)

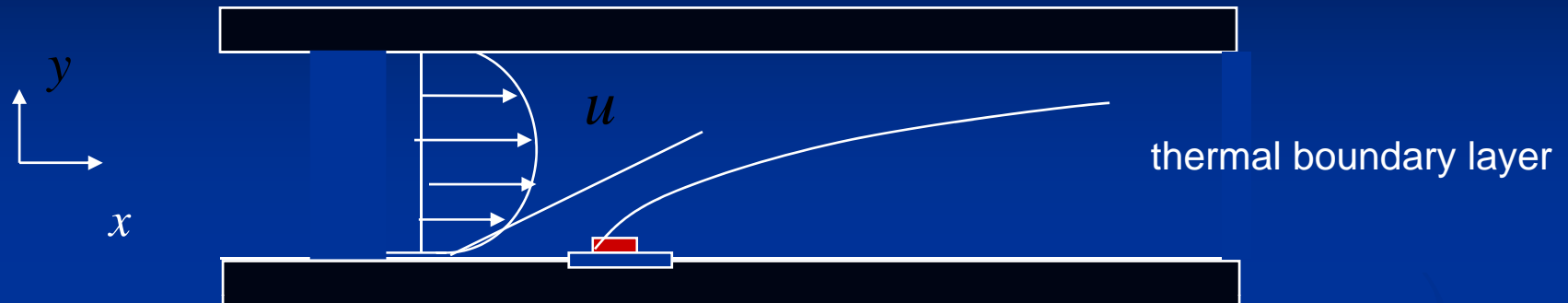


(c)



(d)

Operating Principle of Shear Stress Sensors: Relation between convective heat transfer and shear stress



$$\tau_w = \mu \left. \frac{\partial u}{\partial y} \right|_{y=0}$$

Indirectly measured by heat convection:

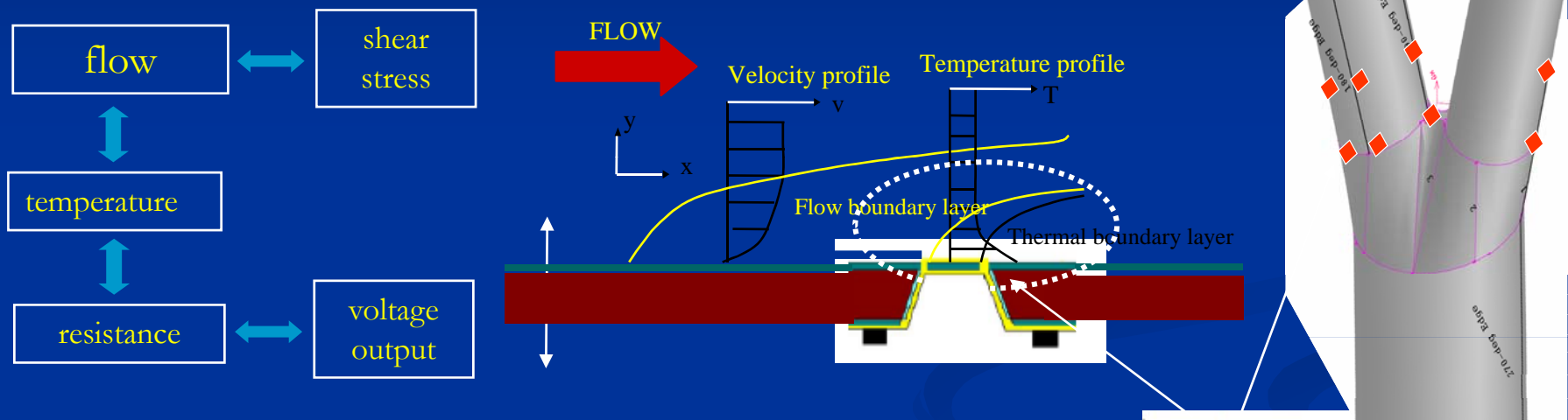
$$V_o^2 / R_s \approx Q_{conv} \propto \tau_w^{1/3}$$

The heat convection from a resistively heated element to the flowing fluid is measured, from which a value for shear stress is inferred

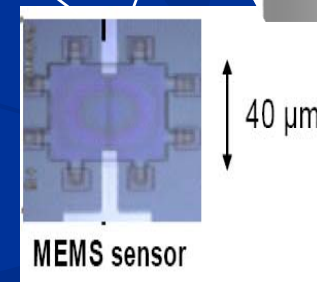
Liu et al, J MEMS 2000
Huang et al, 1995

MEMS Shear Stress Sensors

- MicroElectroMechanical Systems (MEMS) for high temporal and spatial resolution



- The operational principle: Convective heat transfer
 - Sensing element: an electrical resistor
 - Resistance varies with temperature
 - An electrical current is heating up the sensing element
 - Flow past the sensing element changes the temperature and resistance



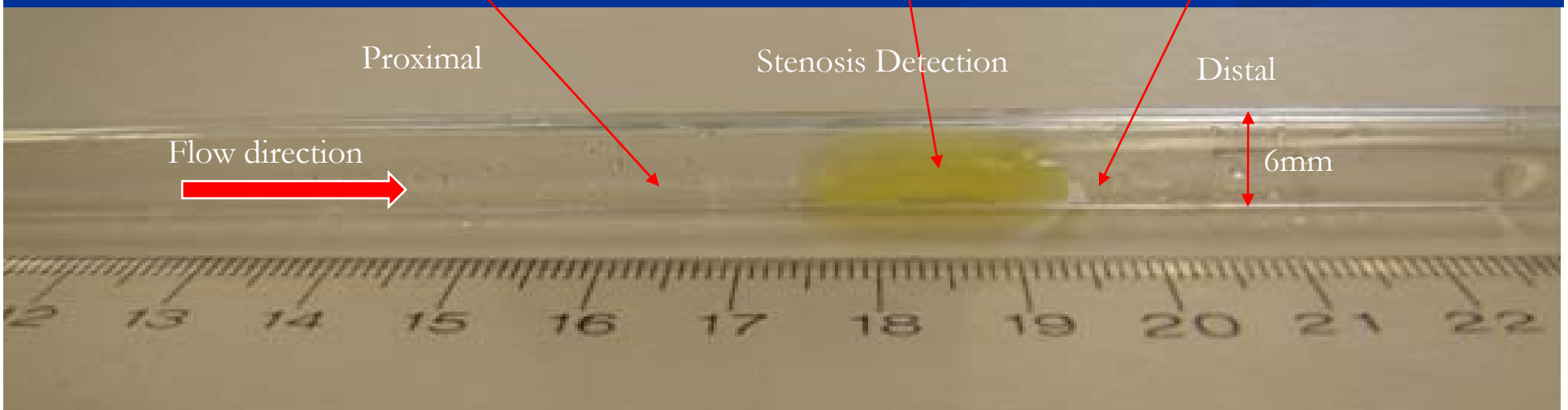
In Vitro Stenotic model: Combining Doppler, CFD, and MEMS sensors



(a)

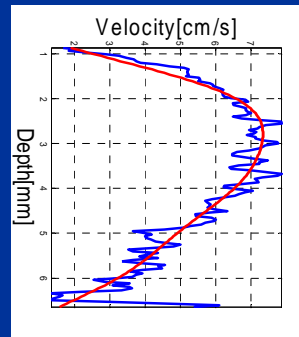
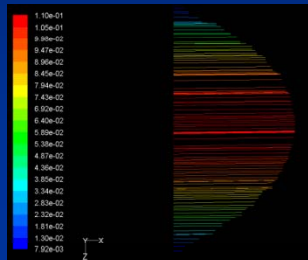
(b)

(c)

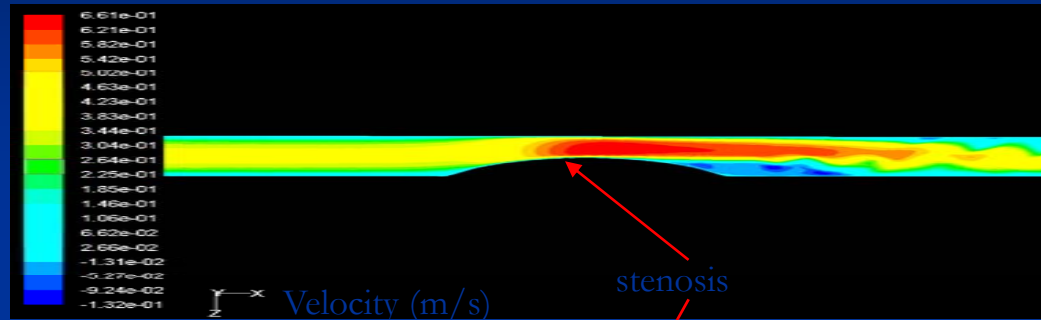


(d)

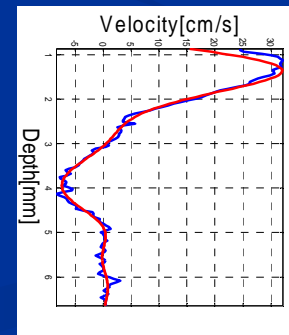
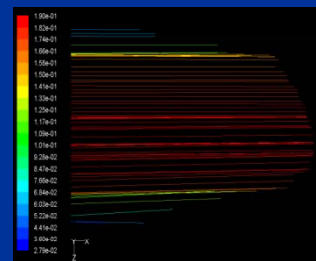
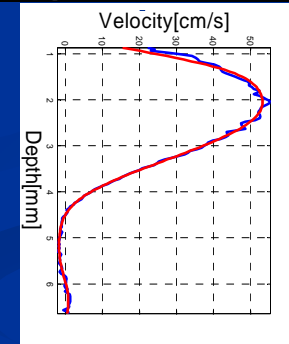
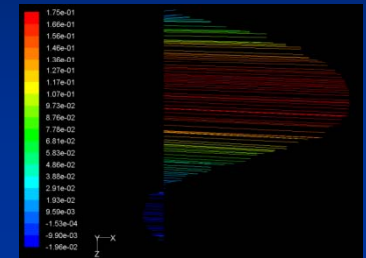
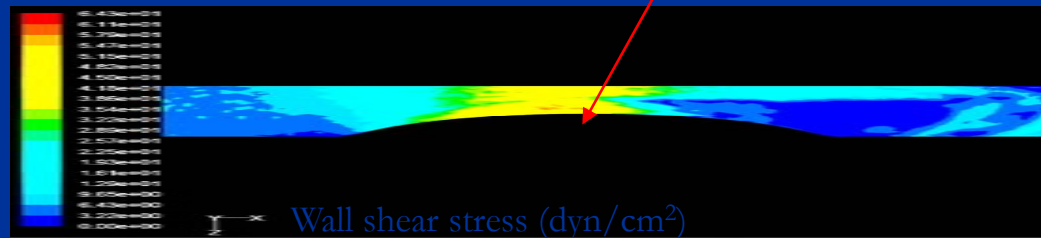
Flow Separation and Flow Reversal post stenotic region



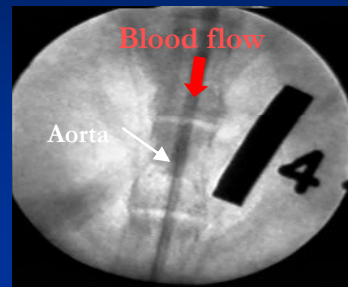
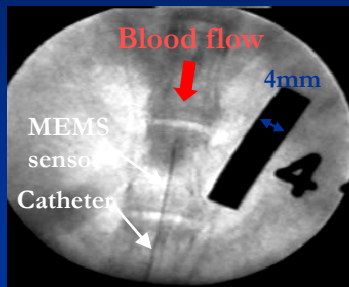
(a)



(b)



Translating *In Vitro* to *In Vivo* Models



MEMS sensor in the rabbit arterial system

The angiogram delineated the diameter of the rabbit aorta after contrast dye injection.



The angiogram revealed the aortic arch, descending aorta, and left carotid arteries.



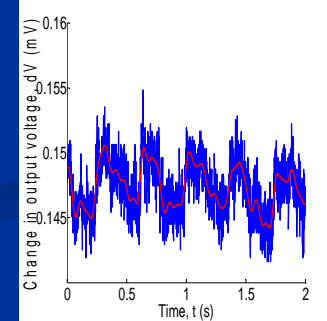
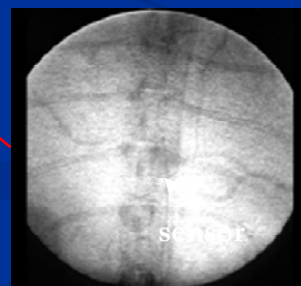
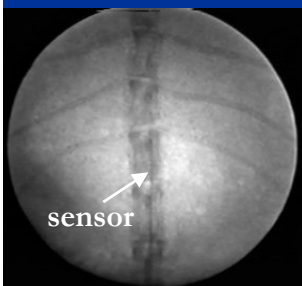
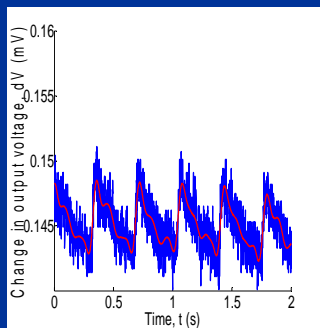
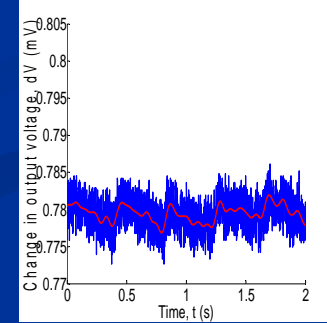
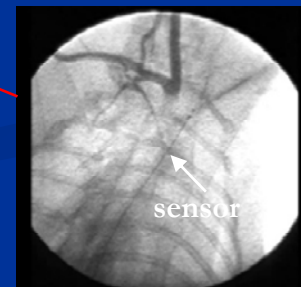
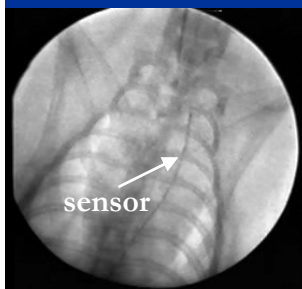
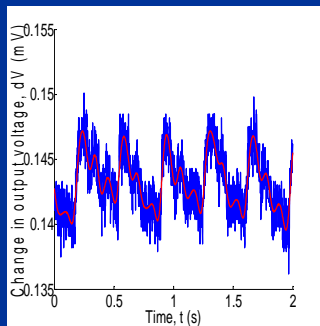
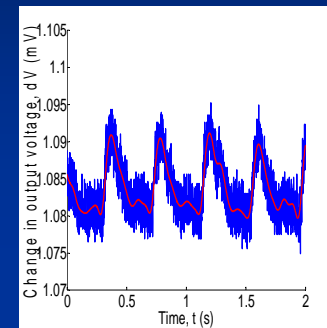
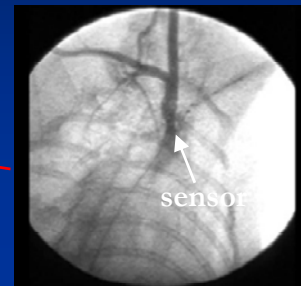
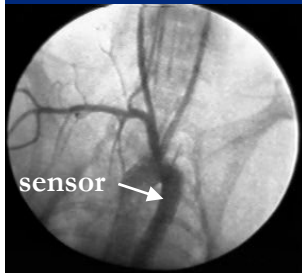
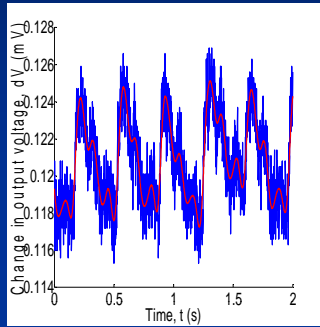
A physician was deploying the sensor via the femoral-cut-down approach.

Yu, H., Ai, L., Hsiai, T. K., et al. Flexible Polymer Sensors for In Vivo Intravascular Shear Stress Analysis, *IEEE/ASME J MEMS*, 2008

Descending aorta

Normal chow diet

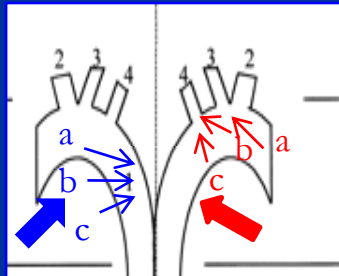
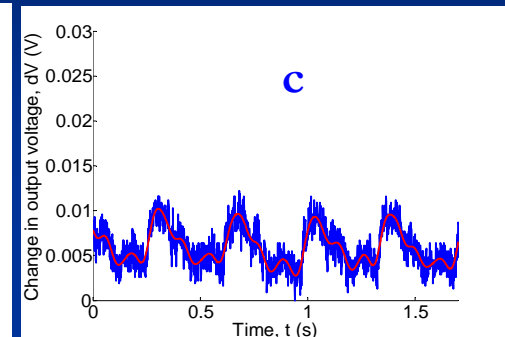
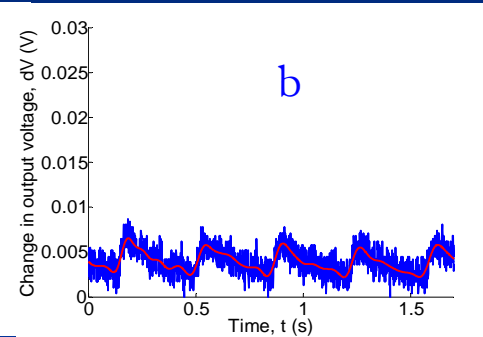
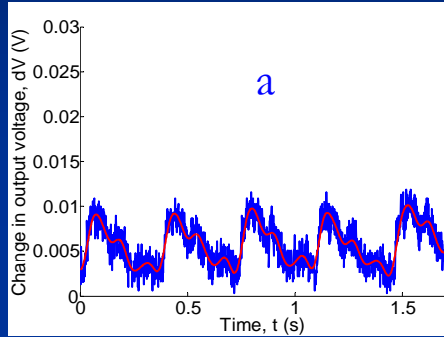
High fat/cholesterol diet



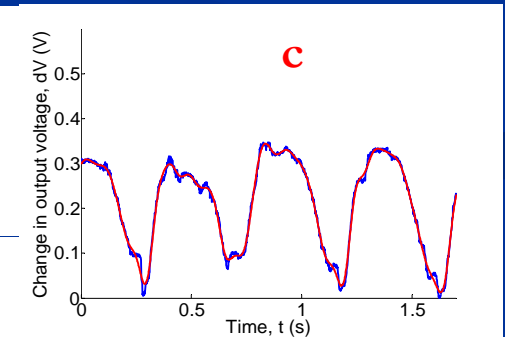
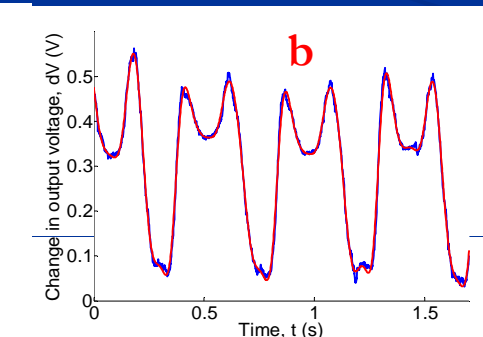
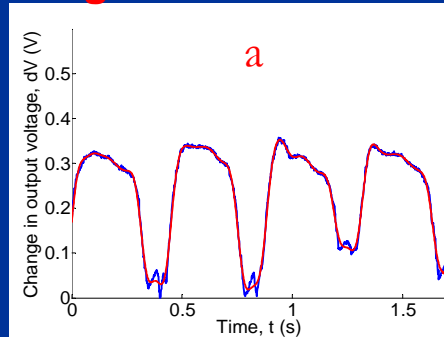
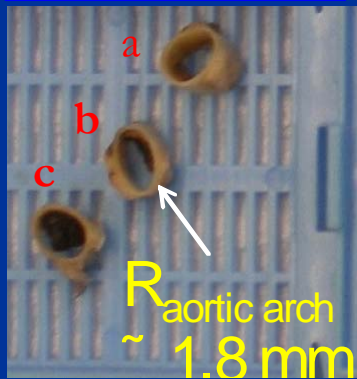
Aortic arch



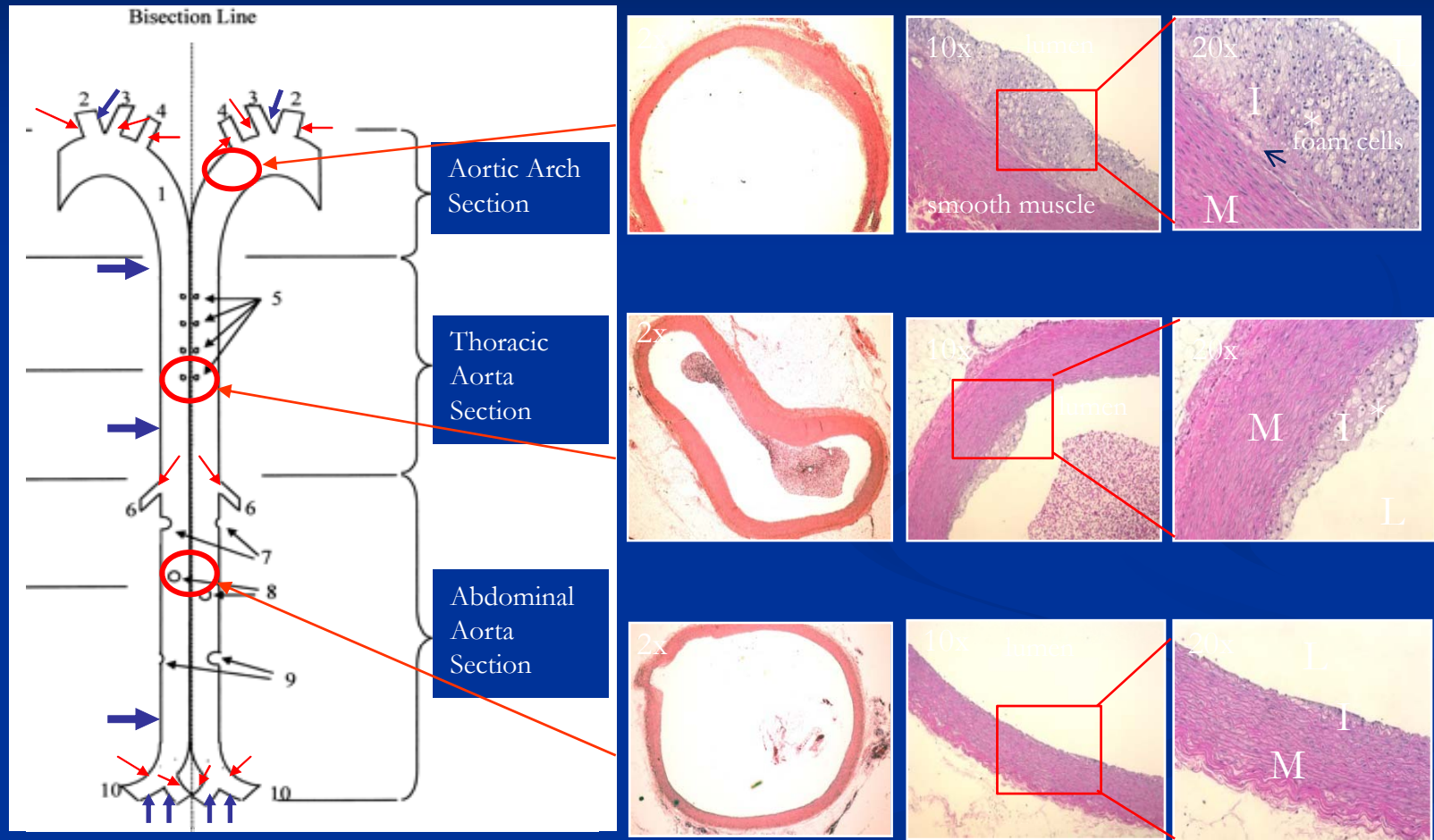
Normal chow diet



High fat/cholesterol diet

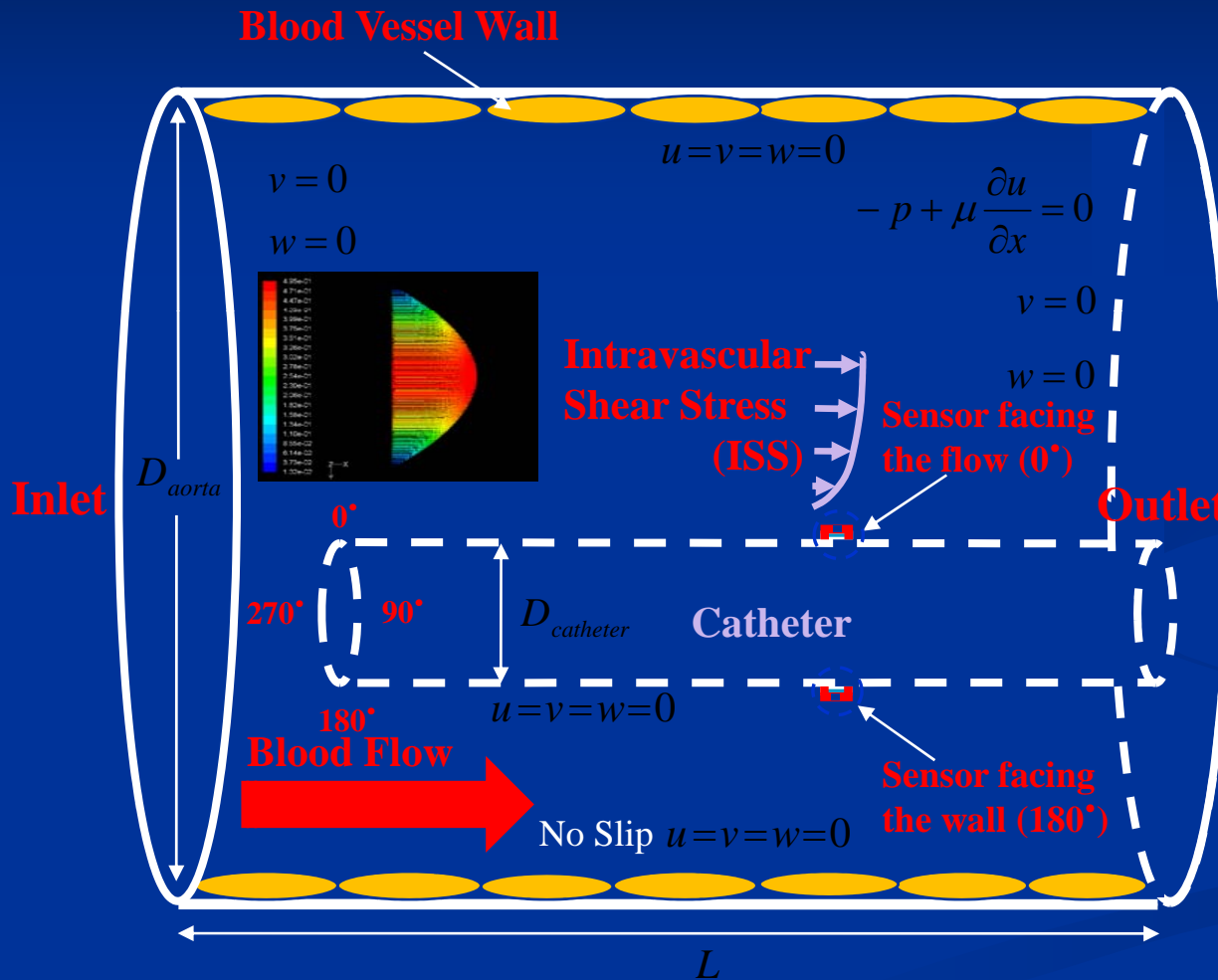


Immunohistochemistry

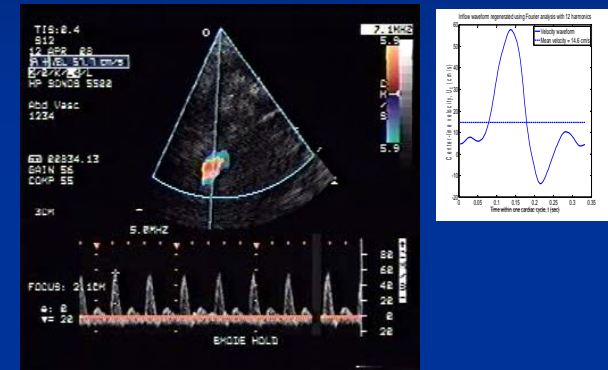


Heart rate at 200 beats/minute; Respiratory rate at 30 times/minute

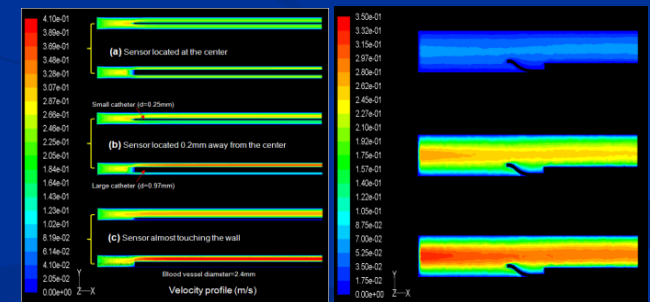
Geometry and Boundary Conditions



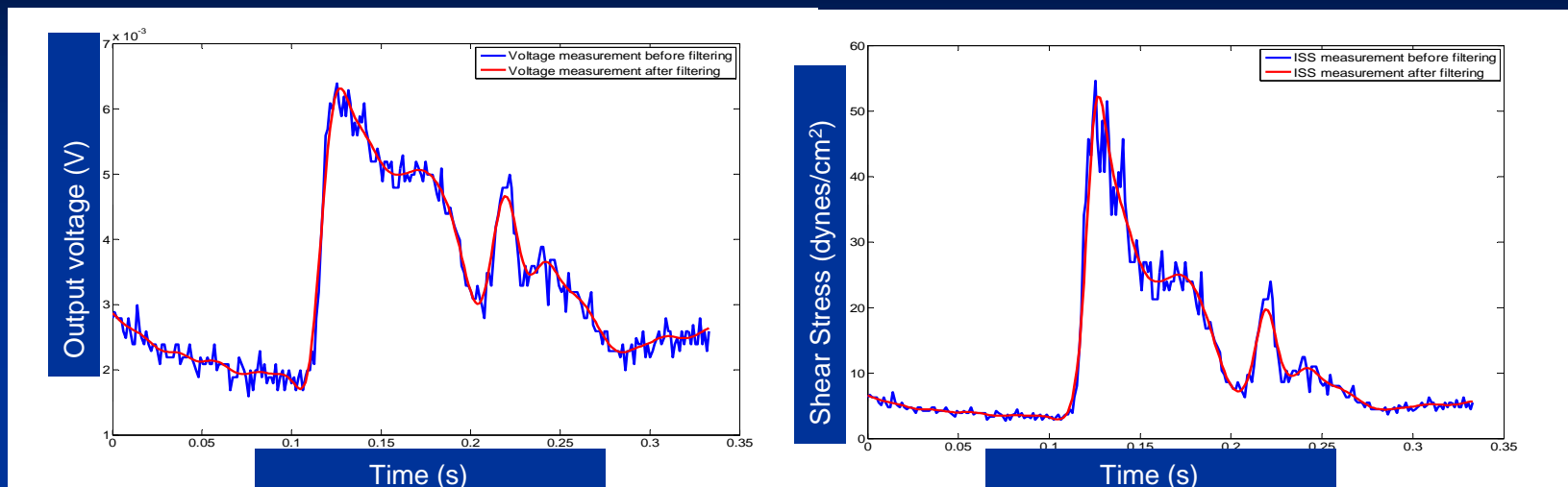
Measured and regenerated waveforms



Effects of catheter and flow rates

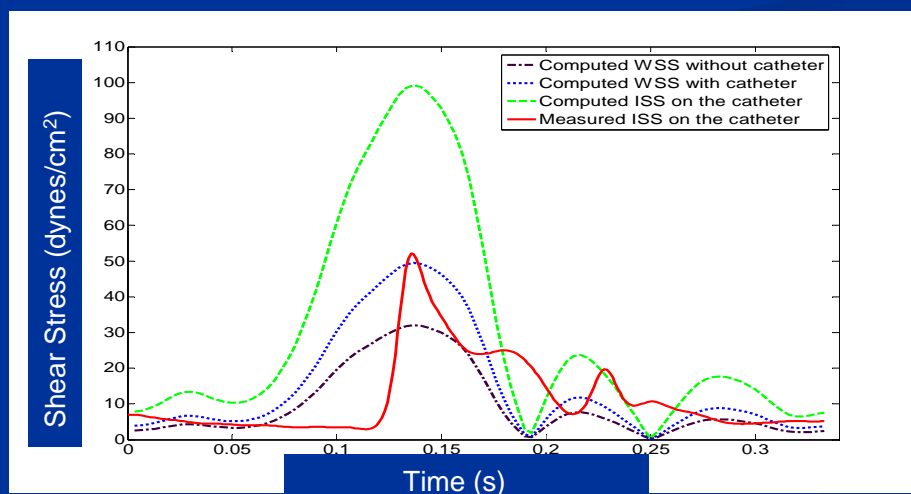


Conversion of output voltage to shear stress

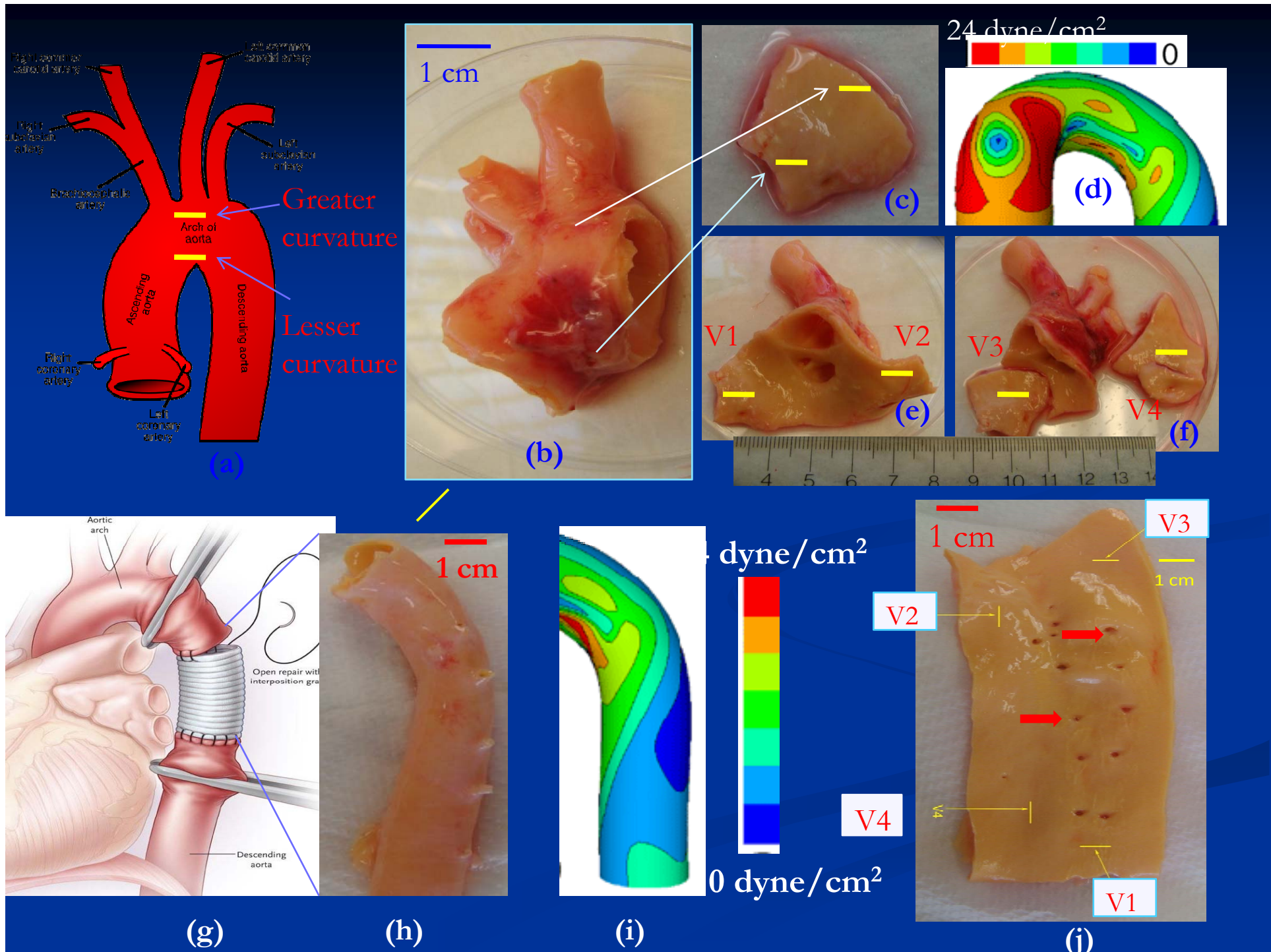


(a) Real-time voltage signals

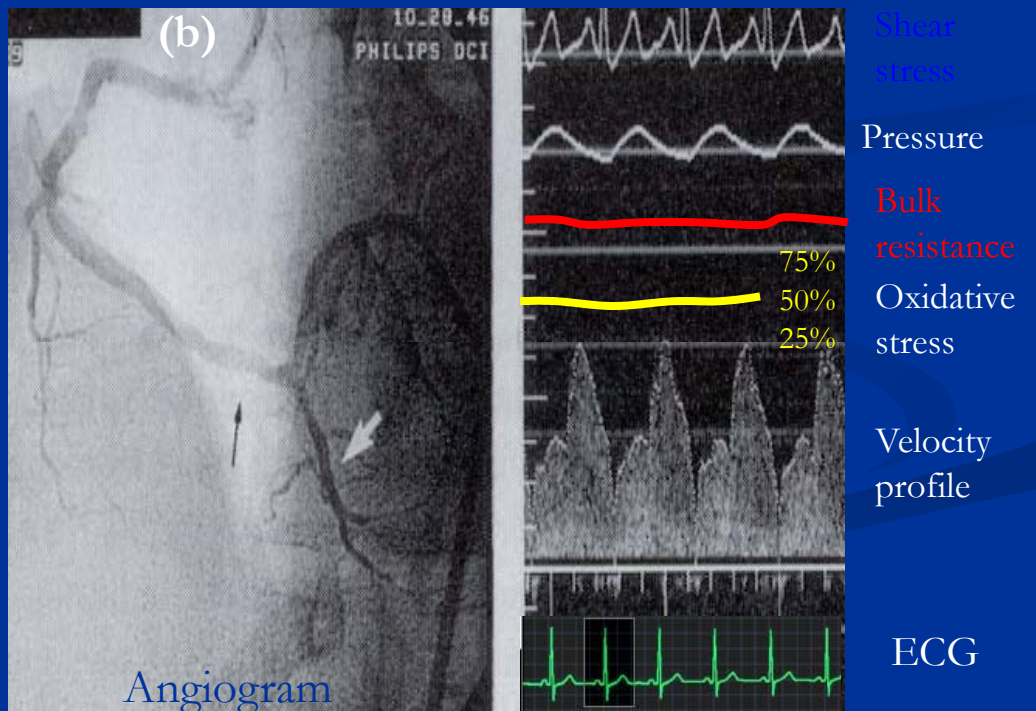
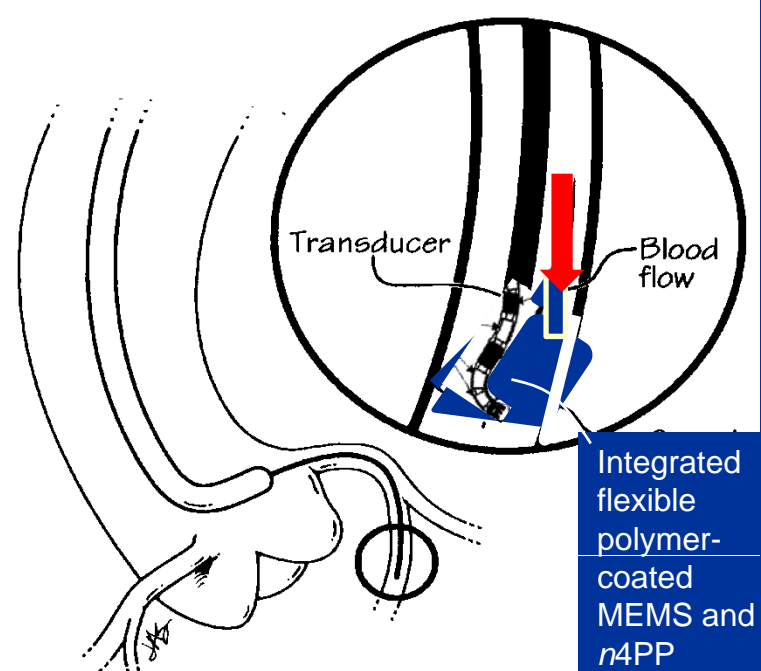
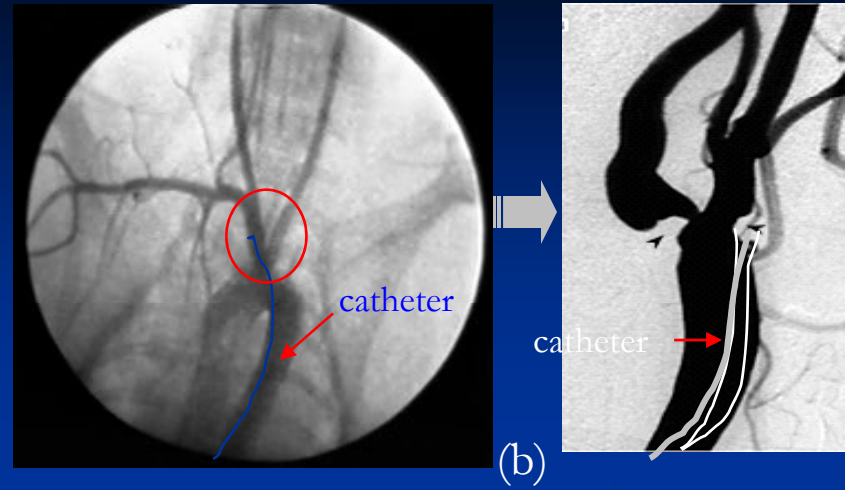
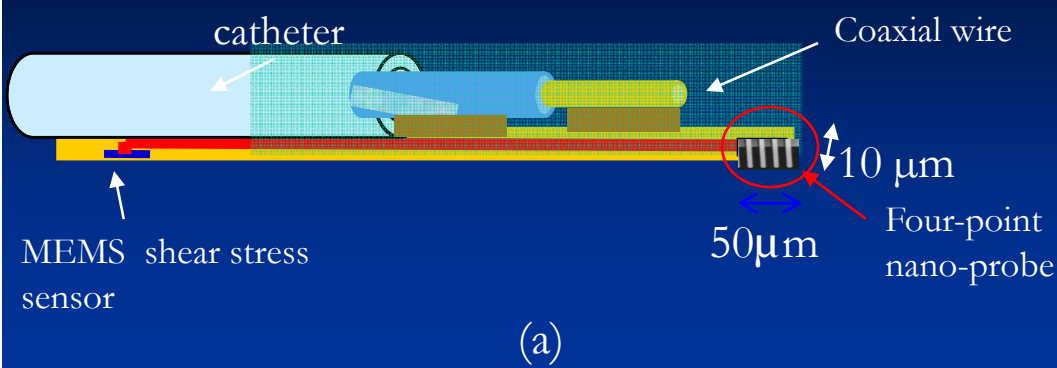
(b) Shear stress converted from voltage signals.



(c) Experimental shear stress curve (in red) compared with computational results



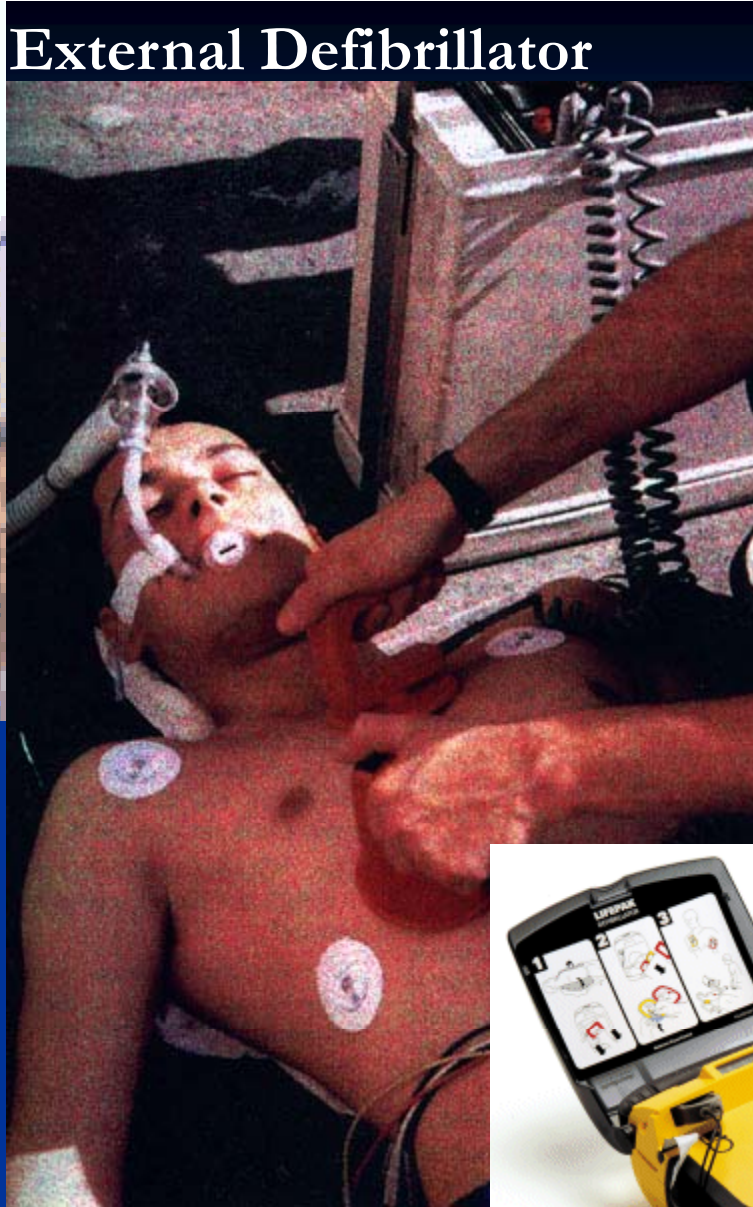
Early detection of Mechanically Unstable Plaque for Selective Intervention



Objectives

- **Introduce vascular systems**
 - Inertia vs. viscous force
- **Interface vascular dynamics with vascular biology**
 - Hemodynamics and vascular oxidative stress
 - Balance between thrombosis and thrombolysis
- **Translate vascular dynamics to an *in vivo* model**
 - Heat transfer strategy to sense secondary flow
 - Bench to pre-clinical study for safety and efficacy
- **Bridge bioengineering, industry and medicine**

External Defibrillator

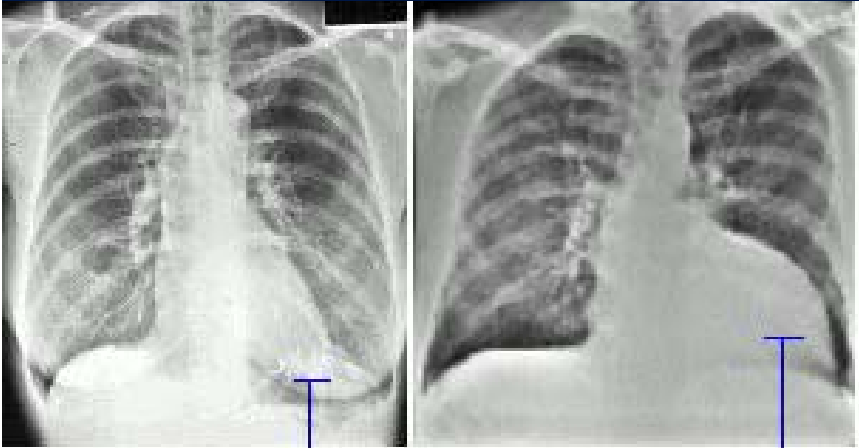


Emergency Room



Diagnostic modalities

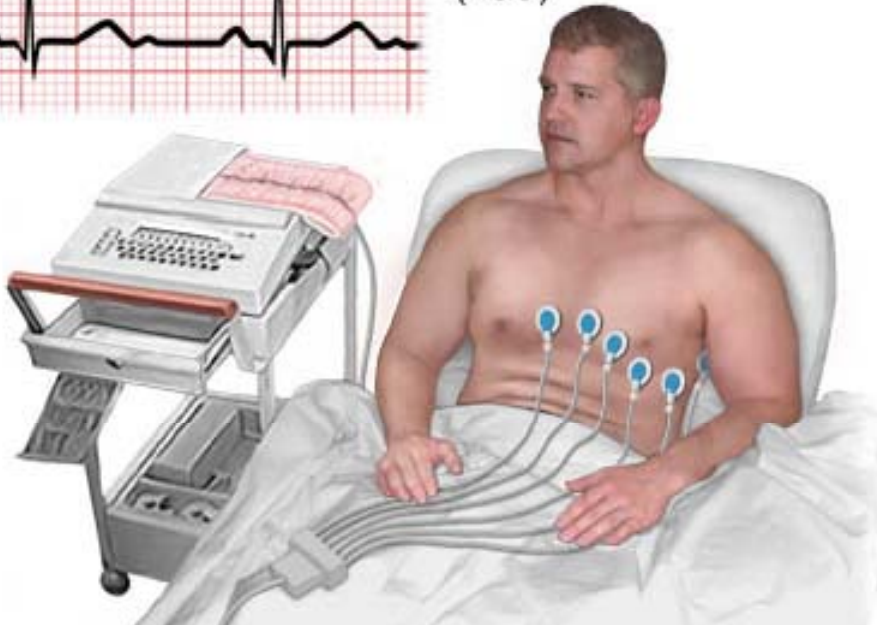
Chest X-Ray



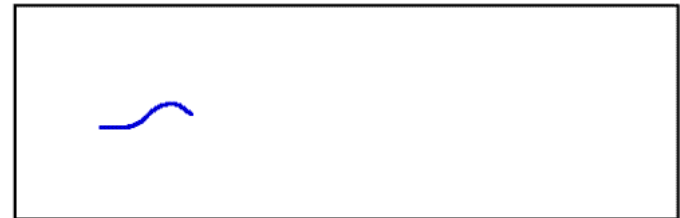
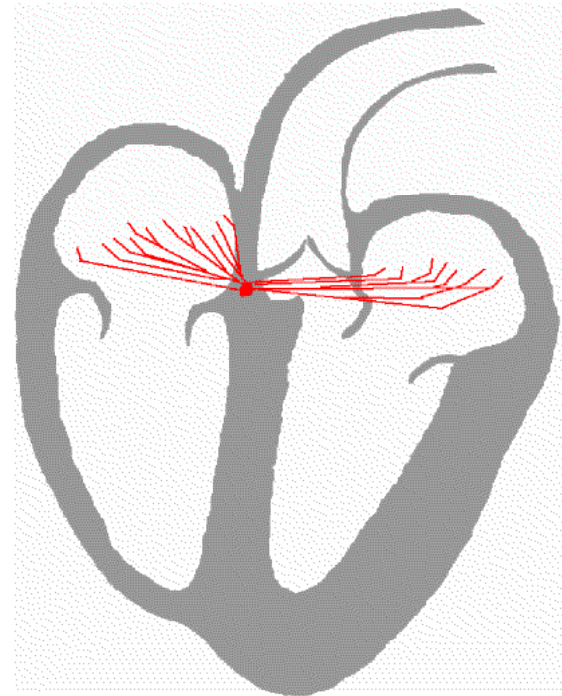
The X-Ray on the left shows a normal heart.
On the right, the heart is enlarged.



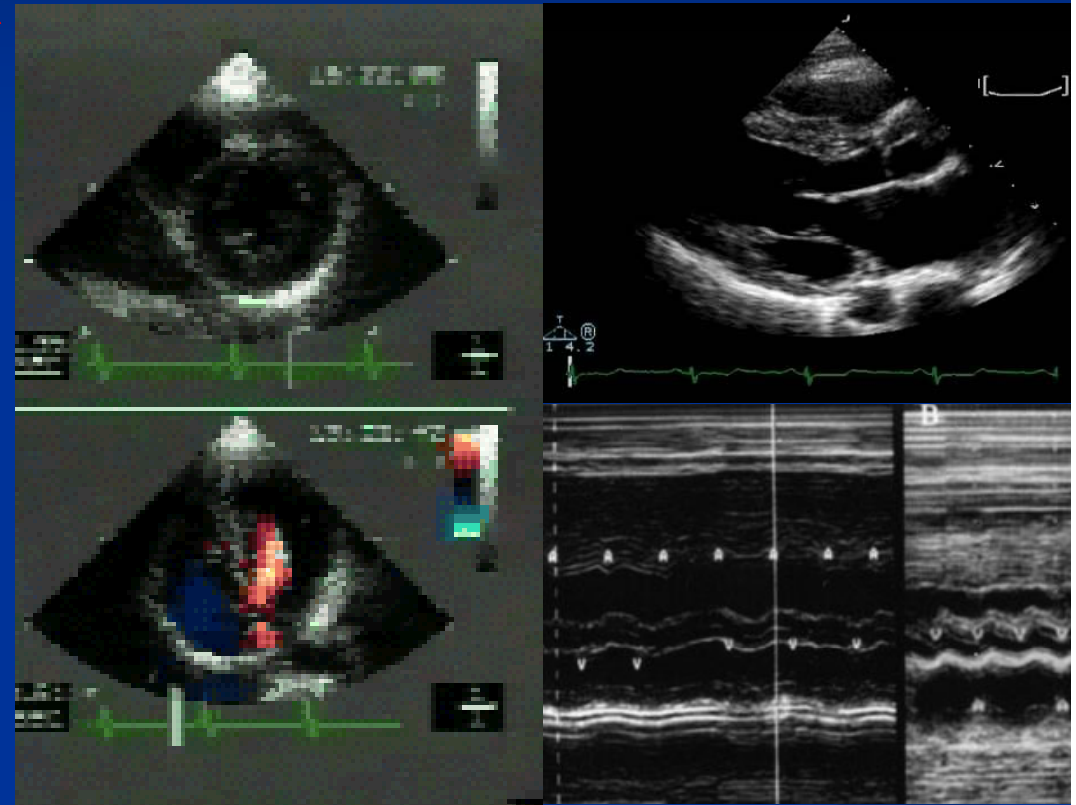
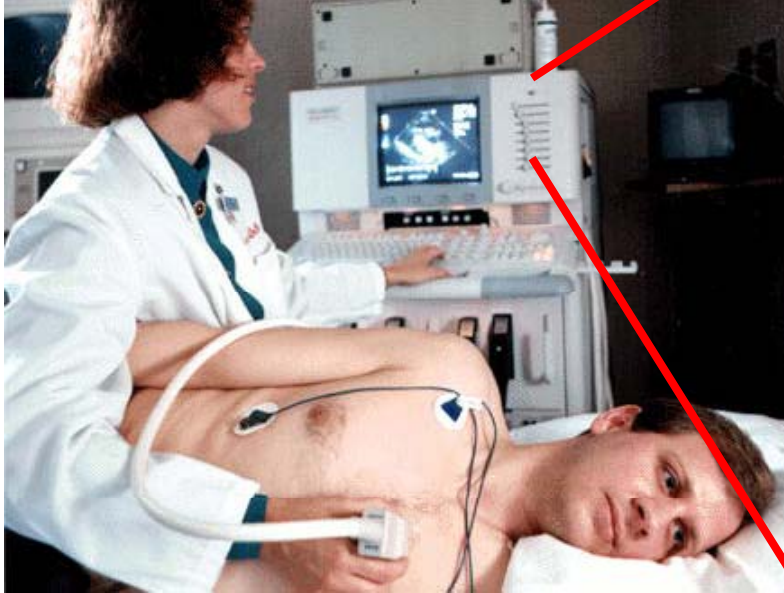
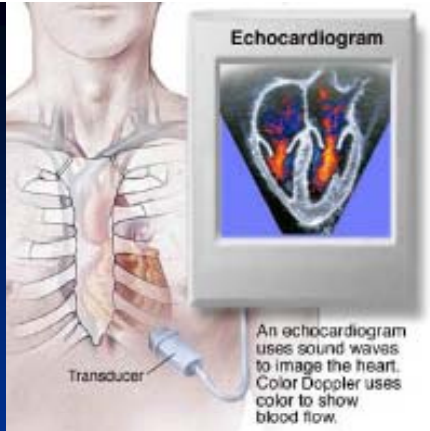
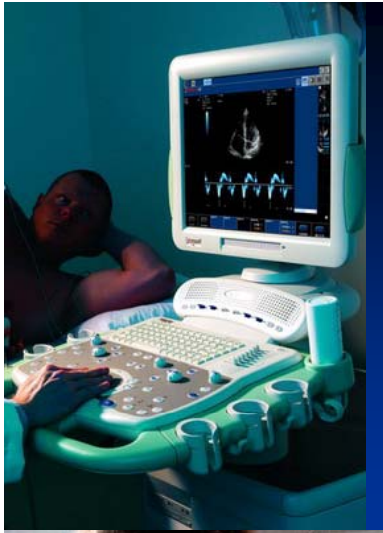
Electrocardiogram
(ECG)



ECG/EKG (electrocardiogram)



Echocardiogram



Echocardiography is the process of mapping the heart through echoes. The pulses are sent into the chest and the high-frequency sound waves bounce off of the heart's walls and valves. The returning echoes are electronically plotted to produce a picture of the heart called an echocardiogram.

Ultra Fast Computed Tomography (CT)



Computed tomography (CT or CAT scan) is a diagnostic imaging procedure that uses a combination of x-rays and computer technology to produce cross-sectional images (often called slices), both horizontally and vertically, of the body.

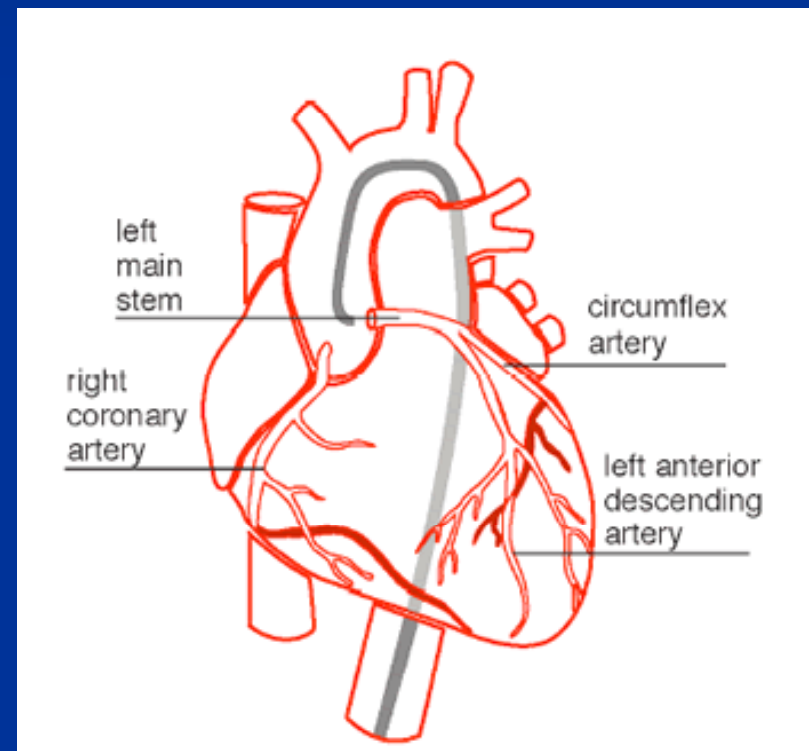
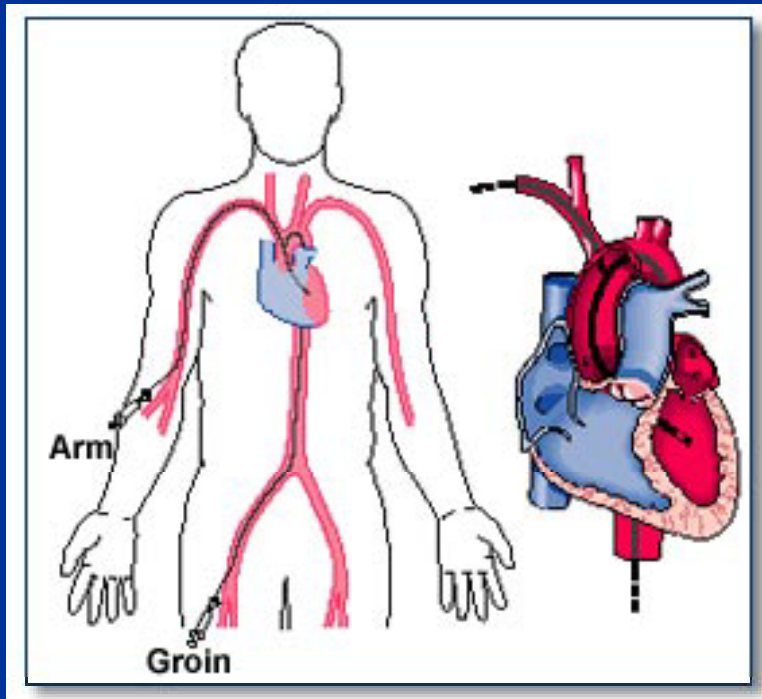
In standard x-rays, a beam of energy is aimed at the part of the body being studied. A plate behind the body part captures the variations of the energy beam after it passes through skin, bone, muscle, and other tissue.

Ultrafast CT scans can take multiple images of the heart within the time of a single heartbeat, thus providing much more detail about the heart's function and structures, and also greatly decreasing the amount of time required for a study.

Cardiac Cath Lab

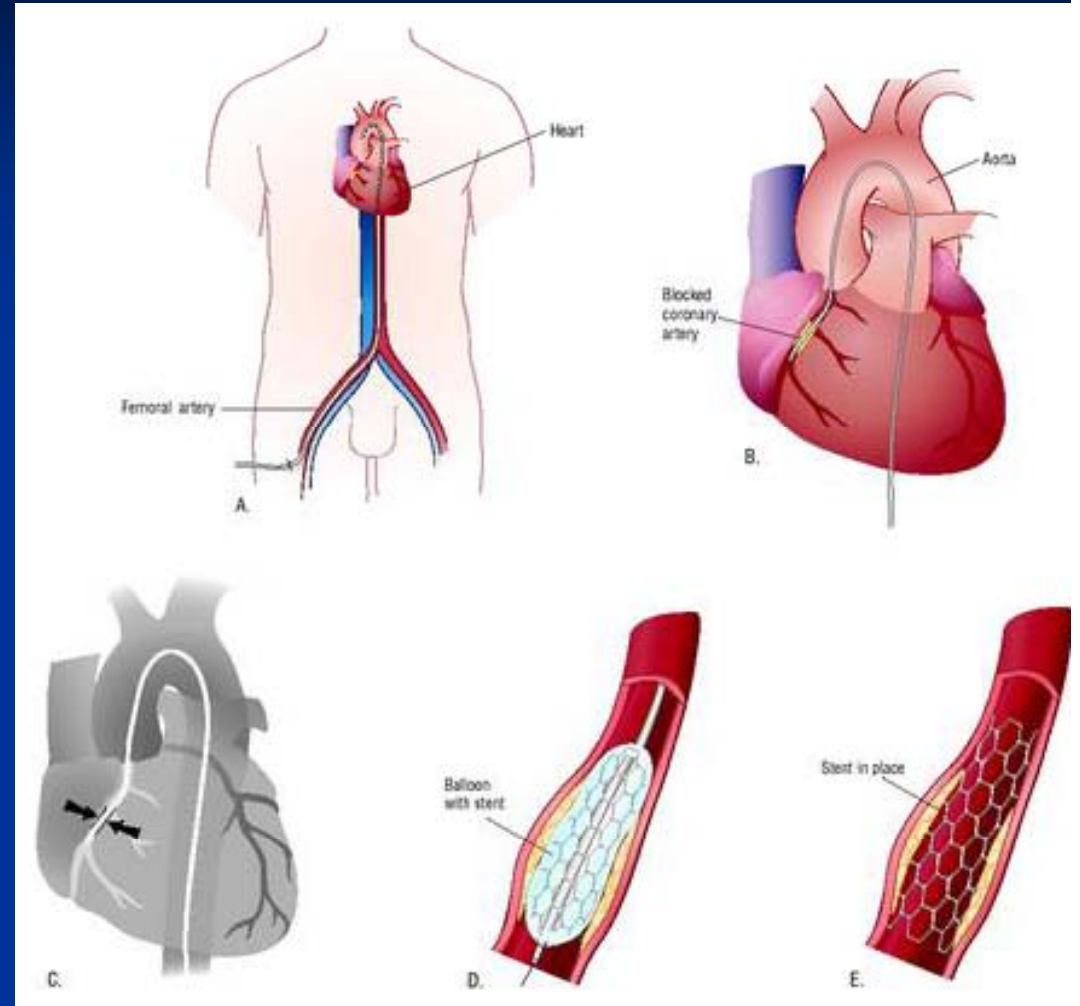
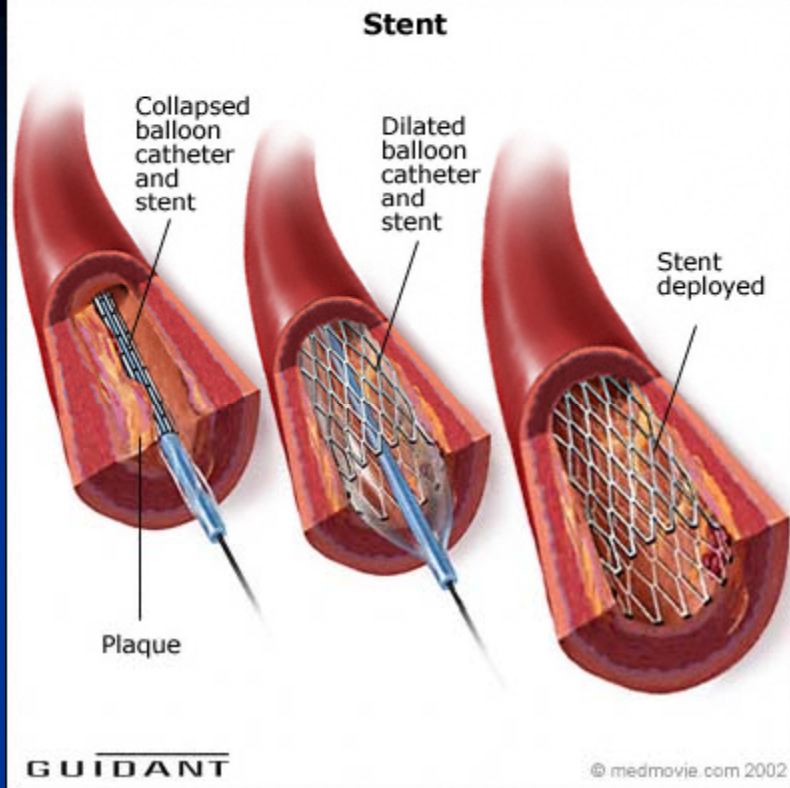


Coronary arteries



Angioplasty

Stents



During angioplasty, a catheter is fed into the femoral artery of the upper leg (A). The catheter is fed up to coronary arteries to an area of blockage (B). A dye is released, allowing visualization of the blockage (C).

A stent is placed on the balloon-tipped catheter. The balloon is inflated, opening the artery (D). The stent holds the artery open after the catheter is removed (E).

Left Ventricular Assist Device (LVAD)

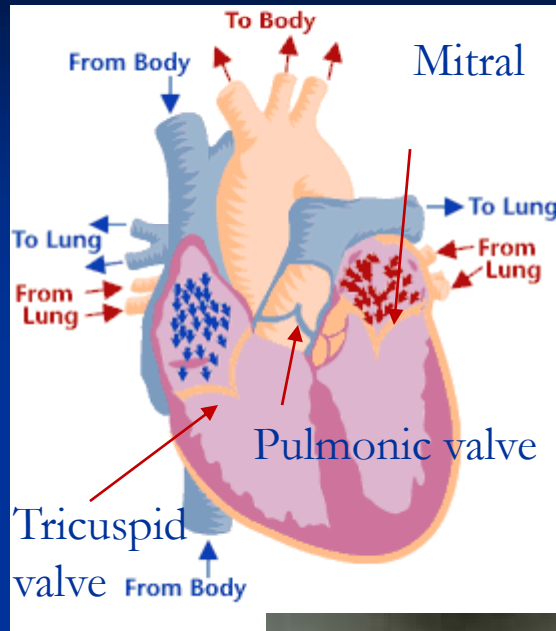


Because of the shortage of donor hearts, heart assist devices are often used to keep patients alive while they await heart transplant. Assist devices, which take over the majority of the heart's pumping function, allow the heart to rest, heal, and grow stronger.

As a result, patients often become healthier and stronger before they undergo transplant surgery.



Prosthetic Heart Valves



Heart valves have key roles in regulating blood flow the heart, opening and closing in sequence with each heartbeat.

These valves act like one-way doors, allowing blood to flow either forward into the next chamber, or out of the heart via one of two main blood vessels that carry blood away from the heart. The valves close to prevent back flow.



The two main prosthetic valve designs include mechanical and bioprosthetic (tissue) heart valves, some of which are shown below.

Objectives

- **Introduce vascular systems**
 - Inertia vs. viscous force
- **Interface vascular dynamics with vascular biology**
 - Hemodynamics and vascular oxidative stress
 - Balance between thrombosis and thrombolysis
- **Translate vascular dynamics to an *in vivo* model**
 - Heat transfer strategy to sense secondary flow
 - Bench to pre-clinical study for safety and efficacy
- **Bridge bioengineering, industry and medicine**



Althea Lyman (Provost Scholar)
Jeff Cui (Rose Hills Scholar)
Farhad Darbandi (Provost Scholar)
Raj Kalsa (Rose Hills Scholar)
Elizabeth Park (Trustee Scholar)
Tyler Bebee (Provost Scholar)
Mahsa Rouhanizadeh, PhD,
Fei Yu (Singapore National U)
Wakako Takabe, PhD (U of Tokyo)
Lisong Ai (UCR Dean's Fellowship)
Hongyu Yu, PhD, AHA PDG (Tsinghua U, Beijing)
Rongsong Li, PhD (Tuft Univ)

Collaborators/Consultants

Donald Heistad
Robert Kloner
Ellen Lien
Fengzhu Sun
E. S. Kim
Joe Wu
Randall Lee
Judith Berliner
Shu Chien
Kathy Griendling
Hanjoong Jo
Stan Hazen

University of Iowa
Heart Institute, Good Samaritan Hospital, Los Angeles
Children's Hospital Los Angeles, USC
Computational Biology, USC
Electrical Engineering, USC
Stanford University
UCSF
UCLA
UCSD
Emory U
Georgia Tech/Emory U
Cleveland Clinic

NIH NHLBI KO8 (HL068689) AHA GIA (0655051Y)
 NIH NHLBI RO1 (HL083015) AHA PDF (0425053Y)
 NIH R21 (HL091302) AHA PDF (0615063Y)



**The search for truth is one way hard another easy,
for no one can master it fully nor miss it fully, each
adds a little knowledge to our nature, and from all
things assembled there arises a certain grandeur.**

Aristotle

Thank You!