

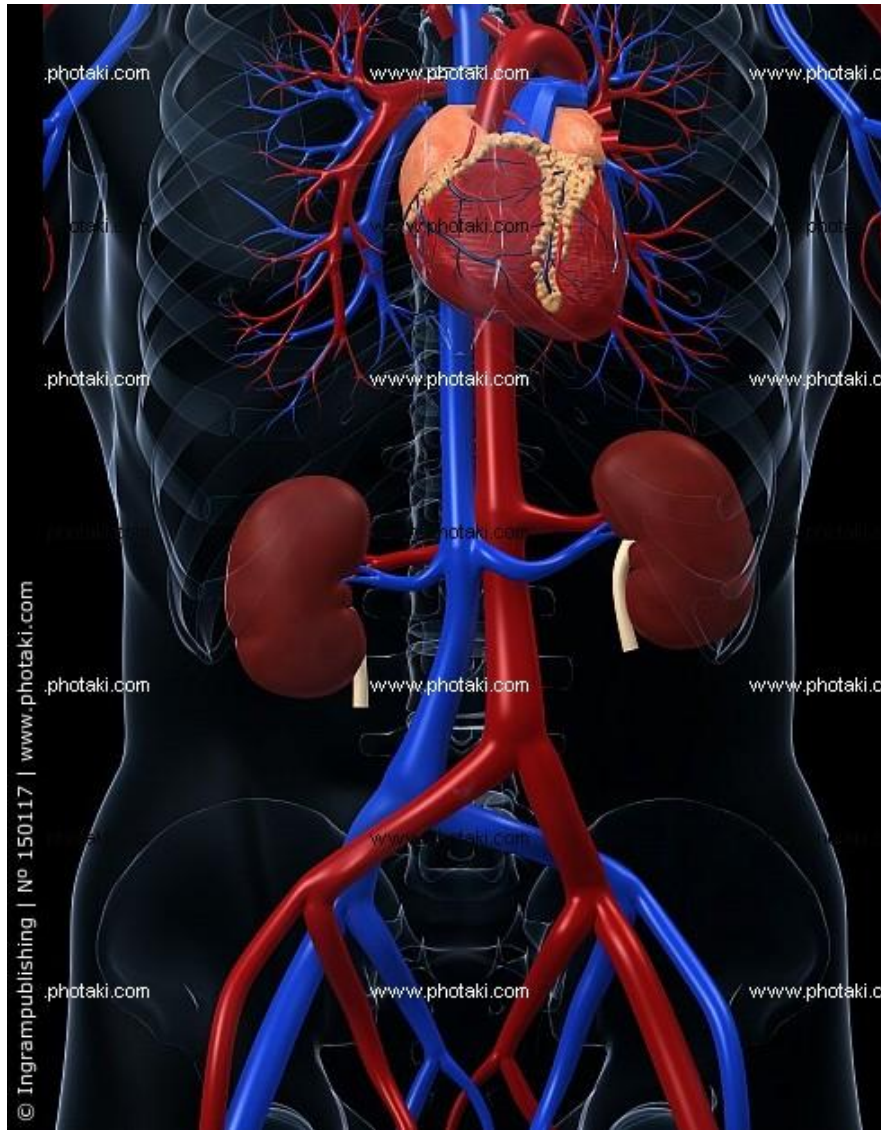
Blood Vessel Mechanics

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Department of Bioengineering

BIOEN 326

10/31/2014

Vasculature in our body



A. structure:

- 100,000 km of pipes!
- total surface area 800–1000 m²
- 60,000 miles of capillaries
- diameters from 10 μm to 2 cm
- double network connected at smallest scale (anastomosed)

B. function:

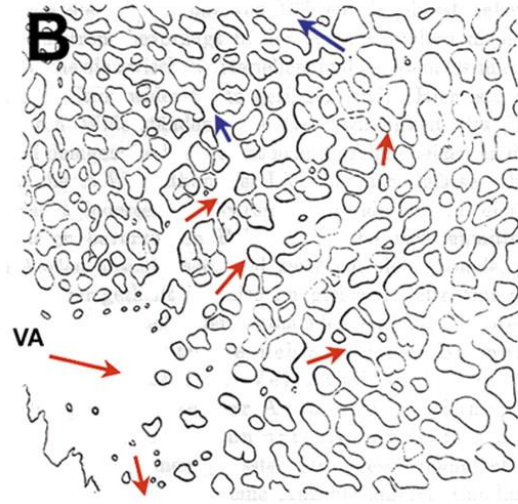
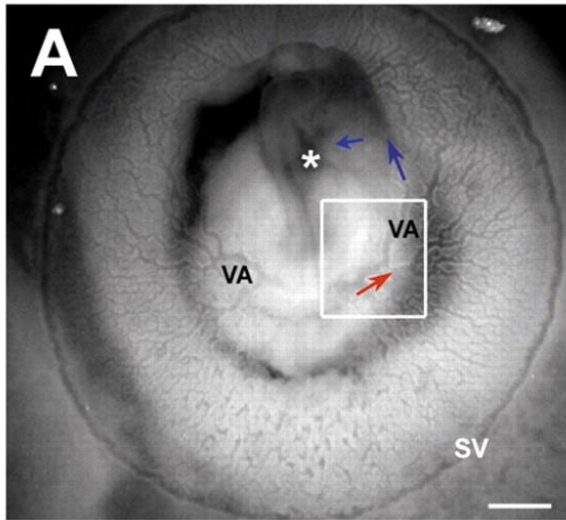
- provide nutrients, oxygen to tissues and remove waste
- self-regulation/homeostasis, tissue remodeling and healing
- cellular, molecular trafficking

C. mechanics:

- Pressure: 5 – 120 mmHg
- Flow: 0.03 – 40 cm/s

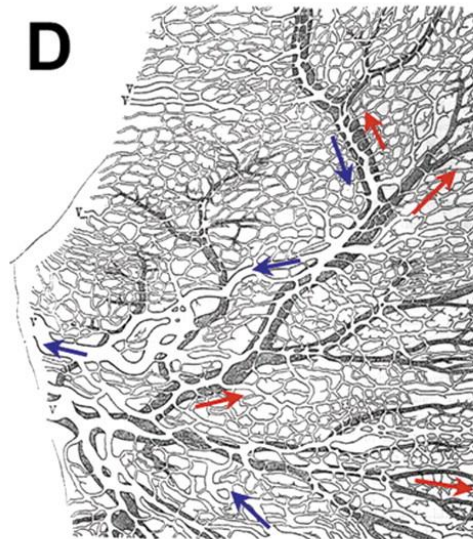
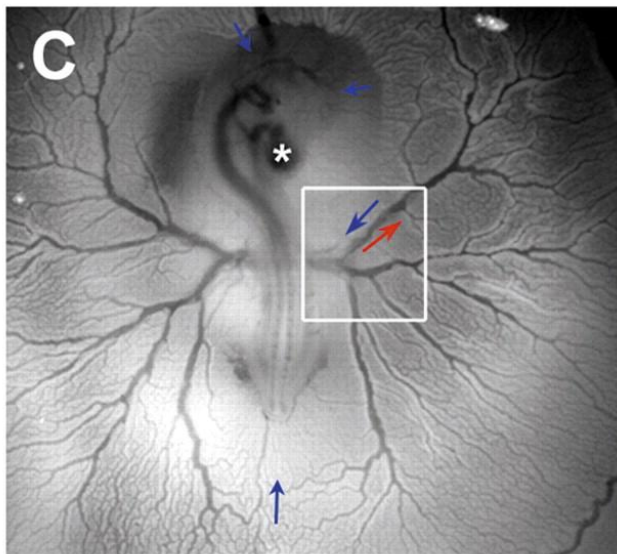
Development of the Vasculature

Chicken embryos



Yolk sac vessels just after the onset of perfusion.

Connected tube formed.



Embryo 26 hours later than in A

Hierarchical structure formed.

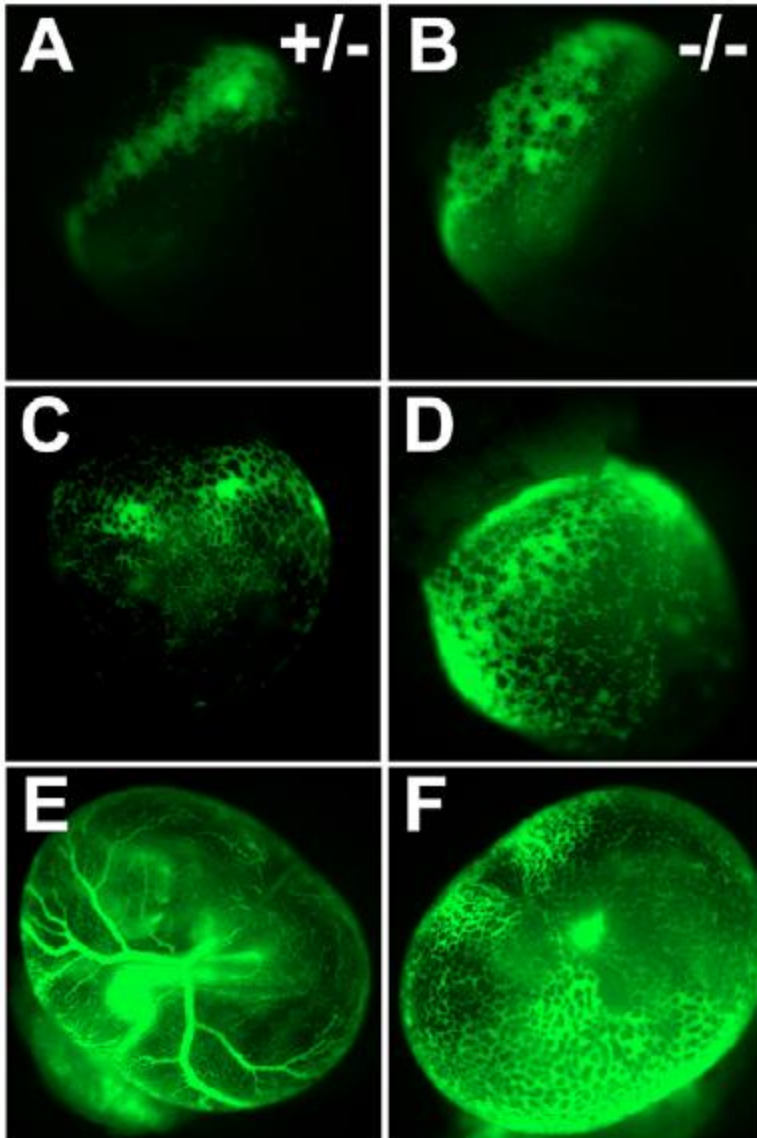
(le Noble, Development 2004)

Development of the Vasculature

Mouse embryos:

Normal

impaired heart function (impaired contractility *Mlc2a*^{-/-}).

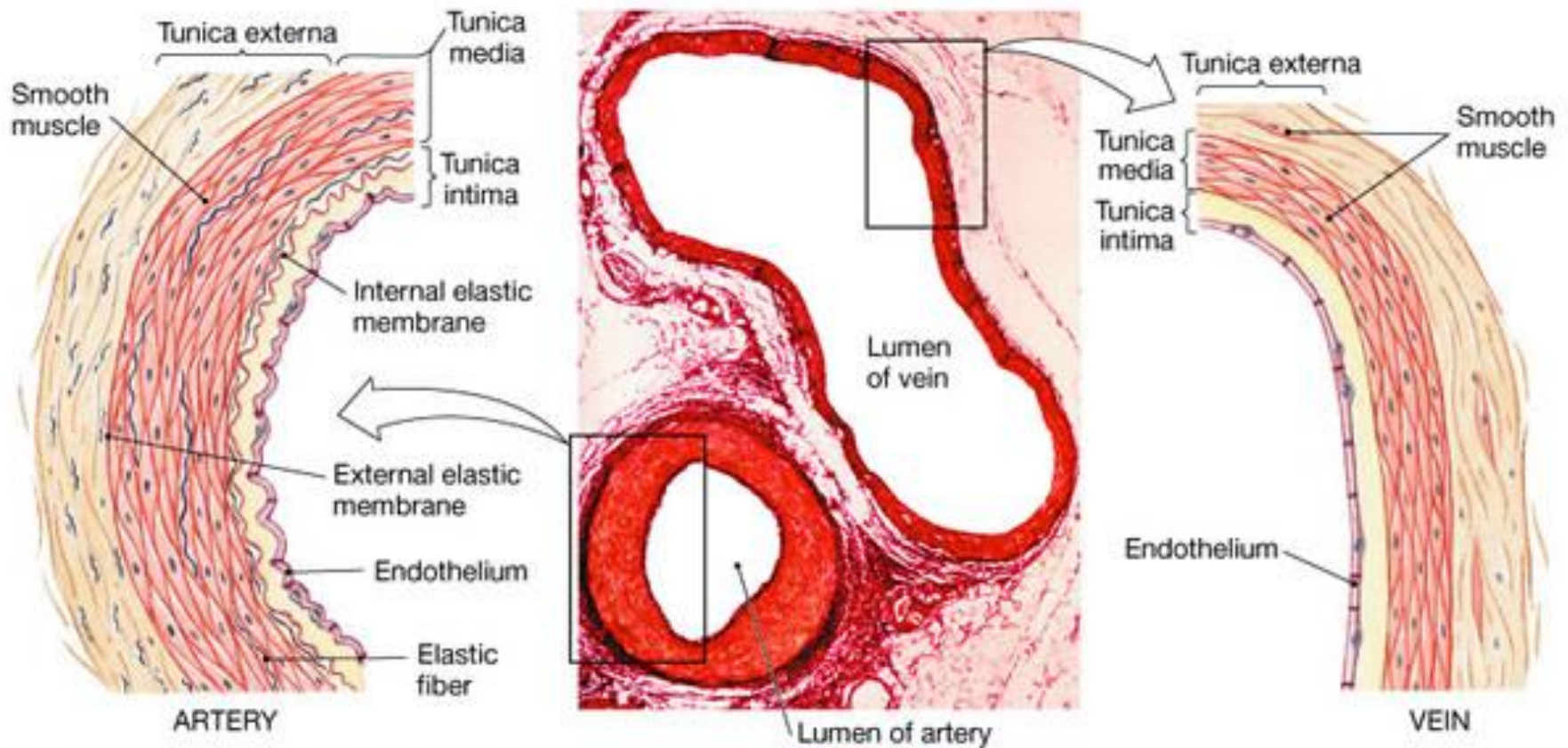


When heart function is impaired, hierarchical branching does not develop

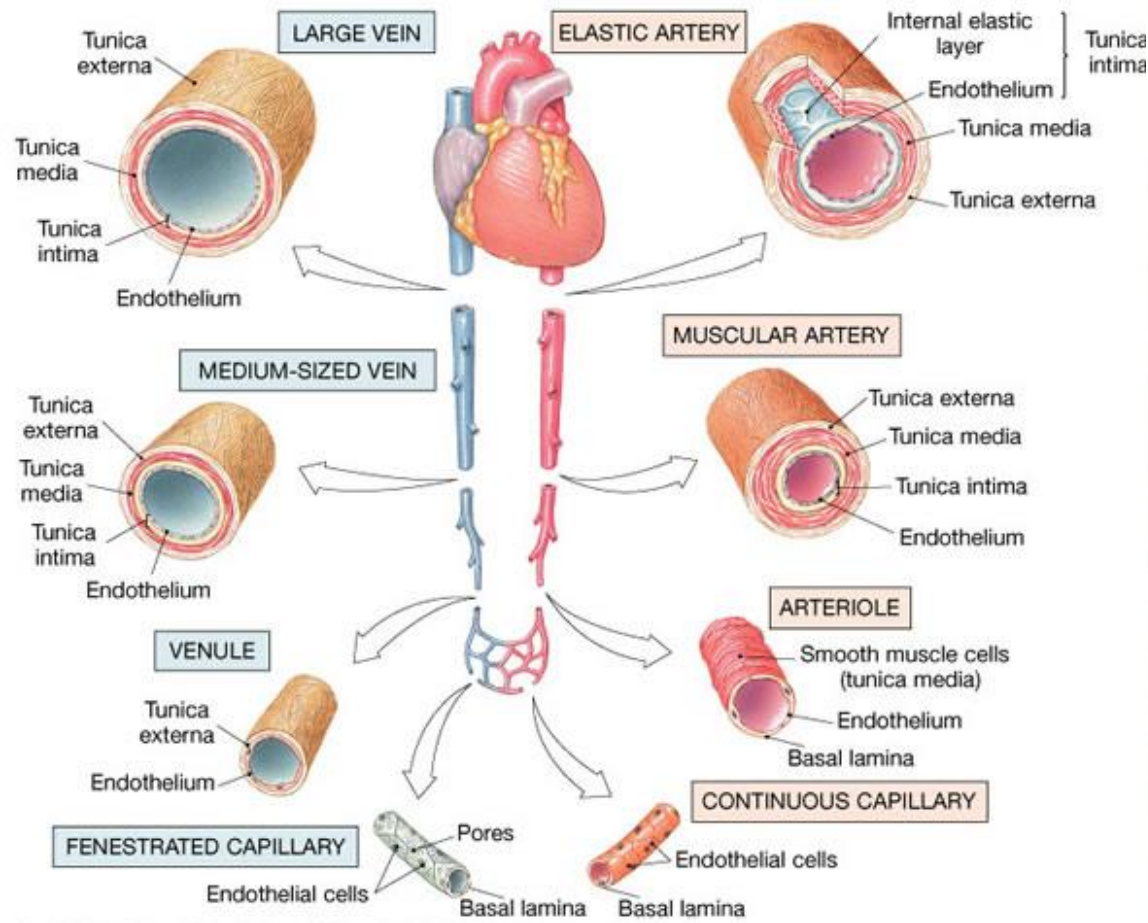
Flow and hydraulic pressure control vascular structure

(Lucitti, *Development* 2007)

A Typical Artery and a Typical Vein

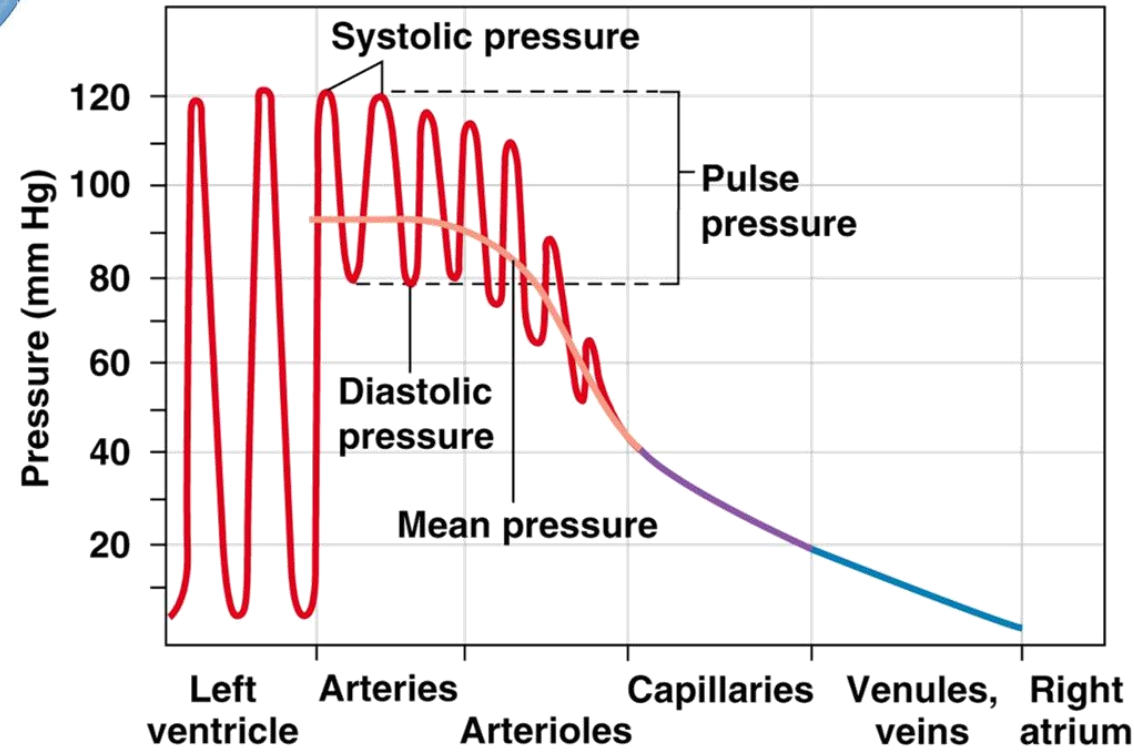
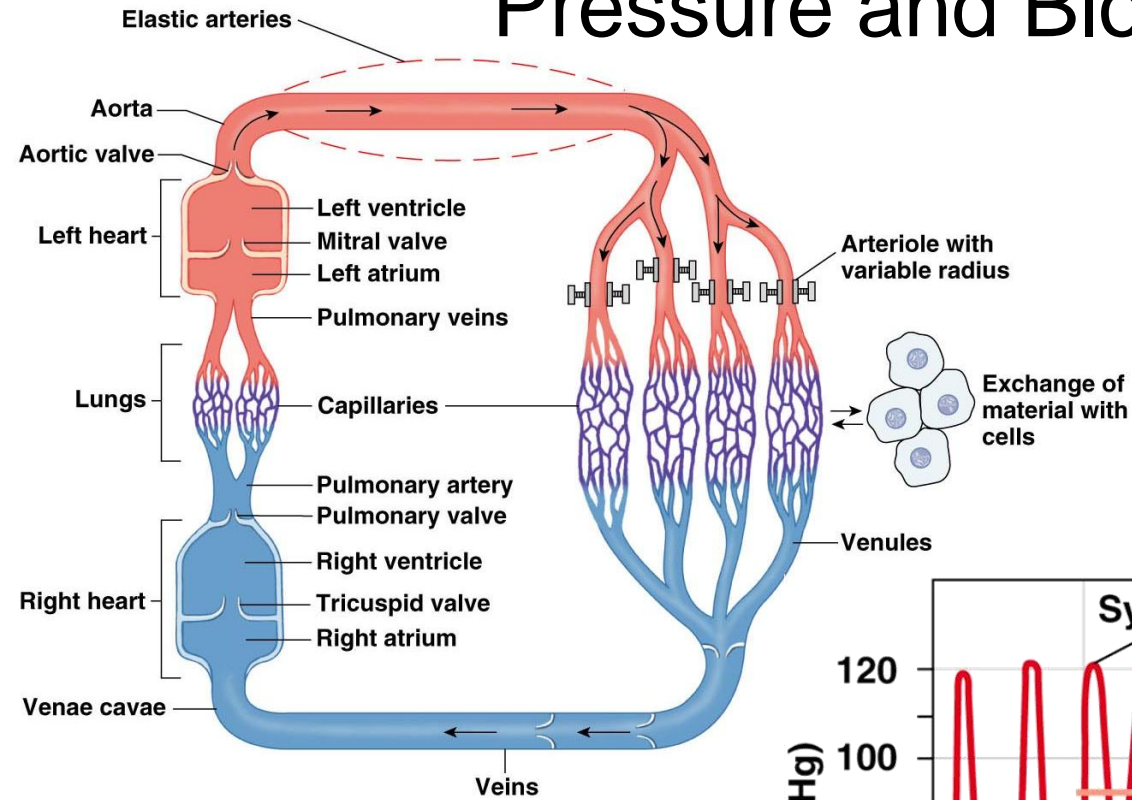


Blood Vessel Structure



| | Mean diameter | Mean wall thickness | Endothelium | Elastic tissue | Smooth muscle | Fibrous tissue | |
|-----------|---------------|---------------------|-------------|----------------|---------------|----------------|--|
| Artery | 4.0 mm | 1.0 mm | Low | High | High | Low | |
| Arteriole | 30.0 μm | 6.0 μm | Low | Low | High | Low | |
| Capillary | 8.0 μm | 0.5 μm | High | Low | Low | Low | |
| Venule | 20.0 μm | 1.0 μm | Low | Low | Low | High | |
| Vein | 5.0 mm | 0.5 mm | Low | Low | Low | High | |

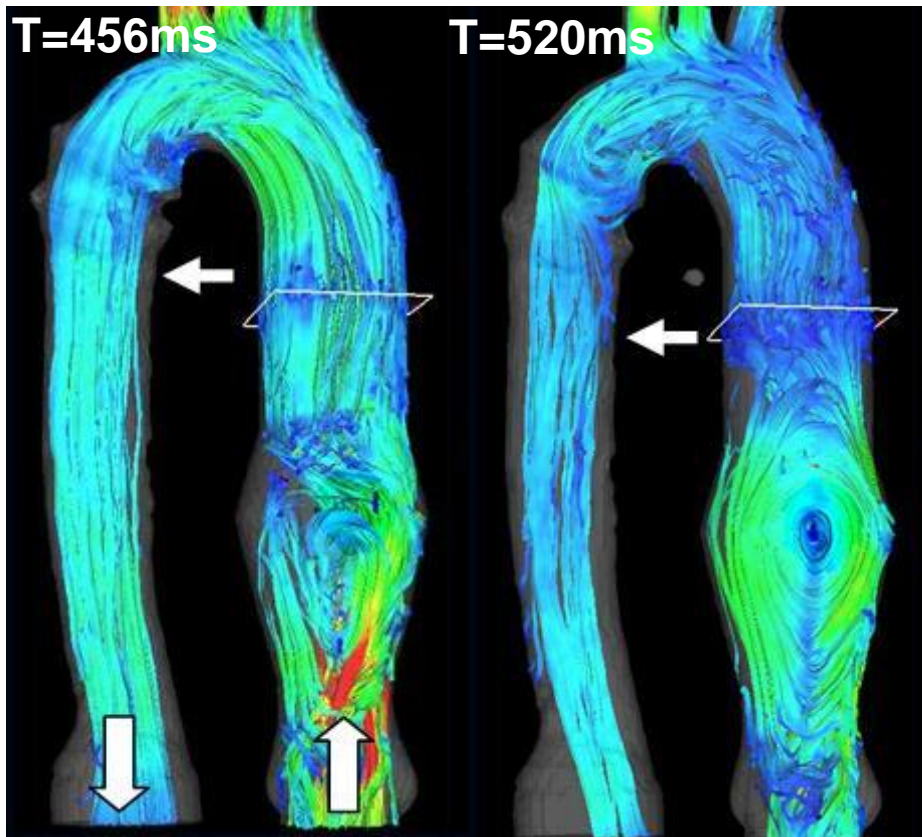
Pressure and Blood Flow



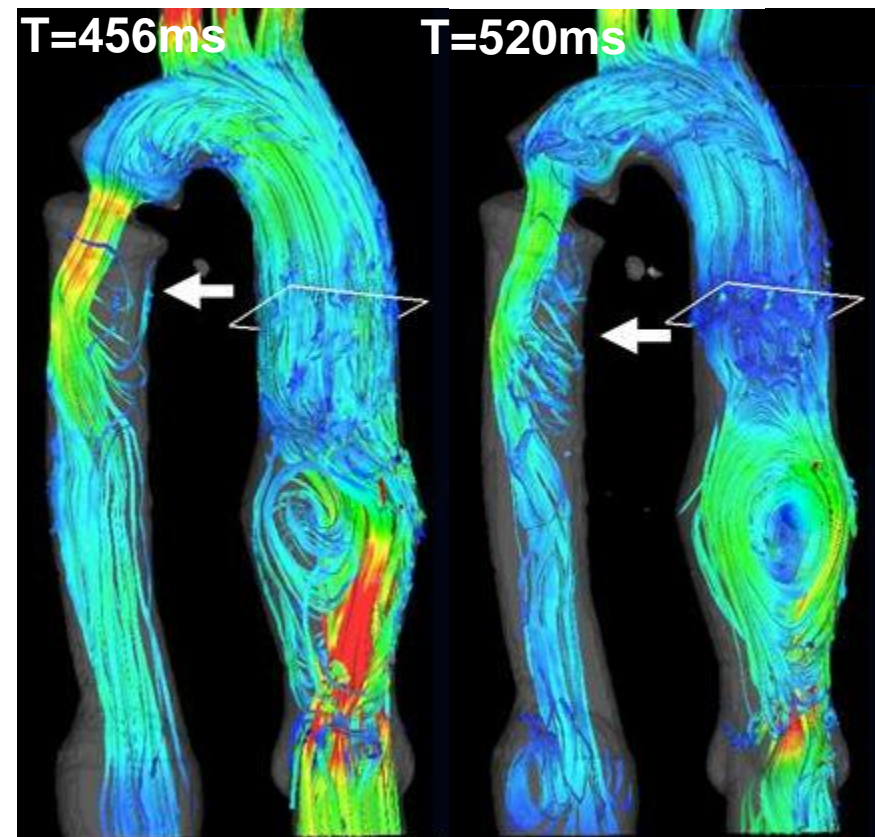
Pressure and blood flow

Poiseuille's relationship: $\Delta P = \frac{8\mu l Q}{\pi r^4}$ (steady, laminar, pipe flow)

MRI, velocity mapping in thoracic aorta



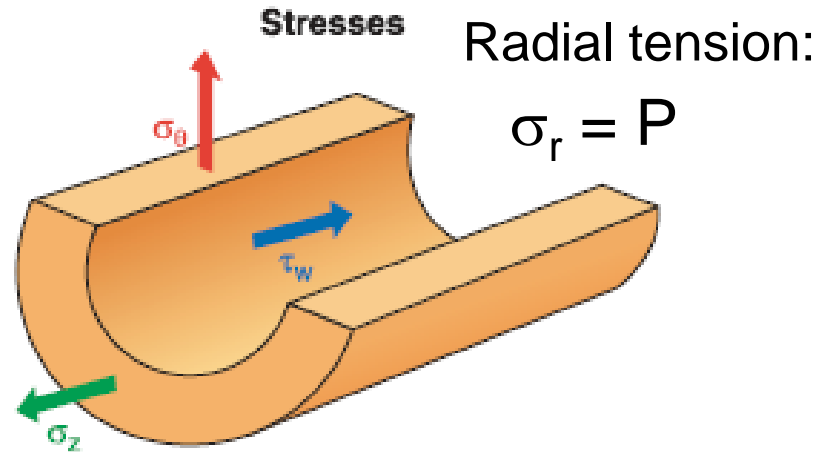
Stenosis, w/o



Stenosis, 54%

(Canstein, MRM, 2006, 2007)

Mechanics: vessel wall



Longitudinal stress:

$$\begin{aligned}\sigma_z &= F/A \\ &= Pd^2 / ((d+2t)^2 - d^2)\end{aligned}$$

Hoop stress:

$$\sigma_\theta = PD_m / 2t$$

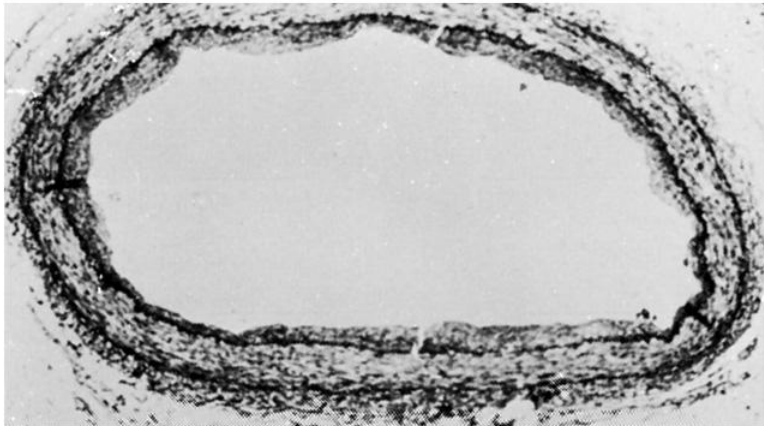
Vessel Wall Associated Pathologies

1. Atherosclerosis

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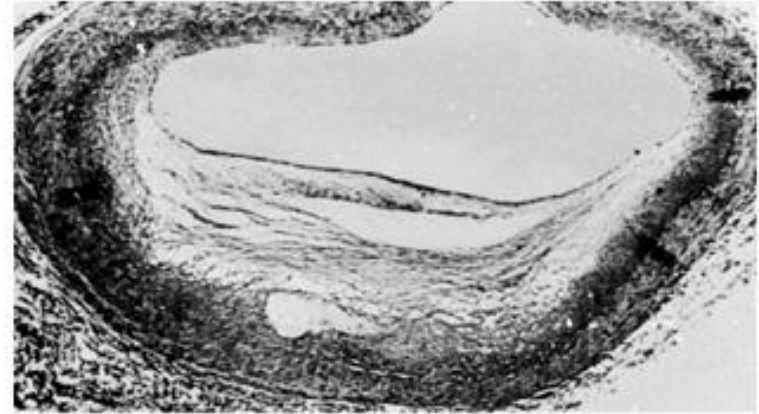
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Normal Arteriole

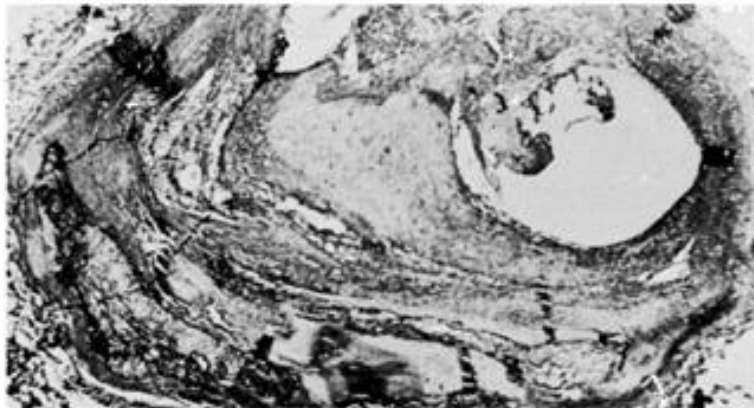


(a)

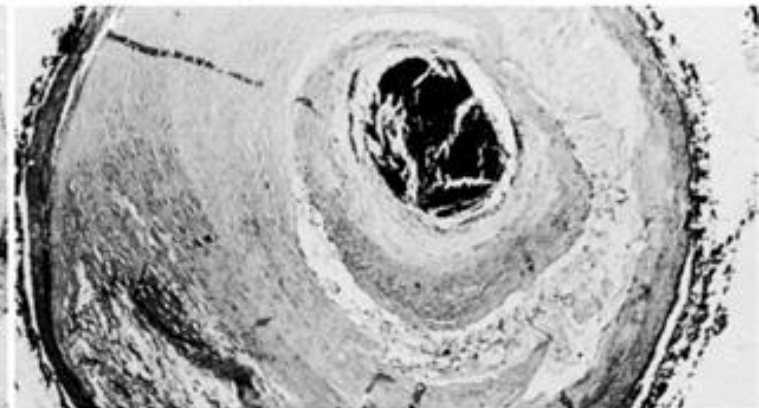
Courtesy of the American Heart Association



(b)



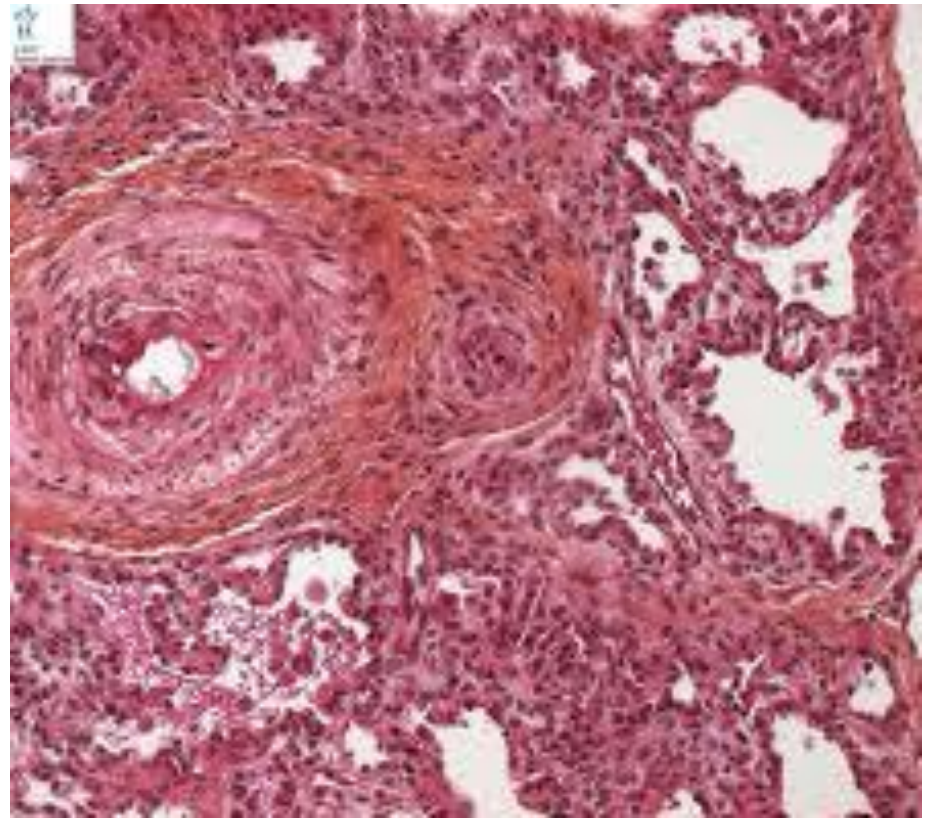
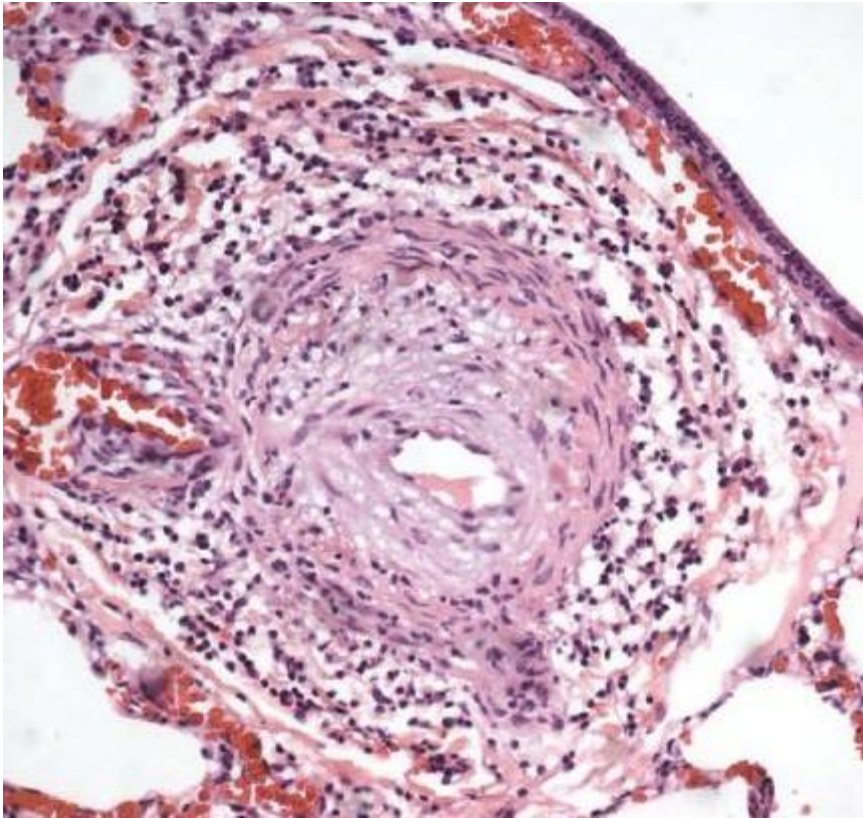
(c)



(d)

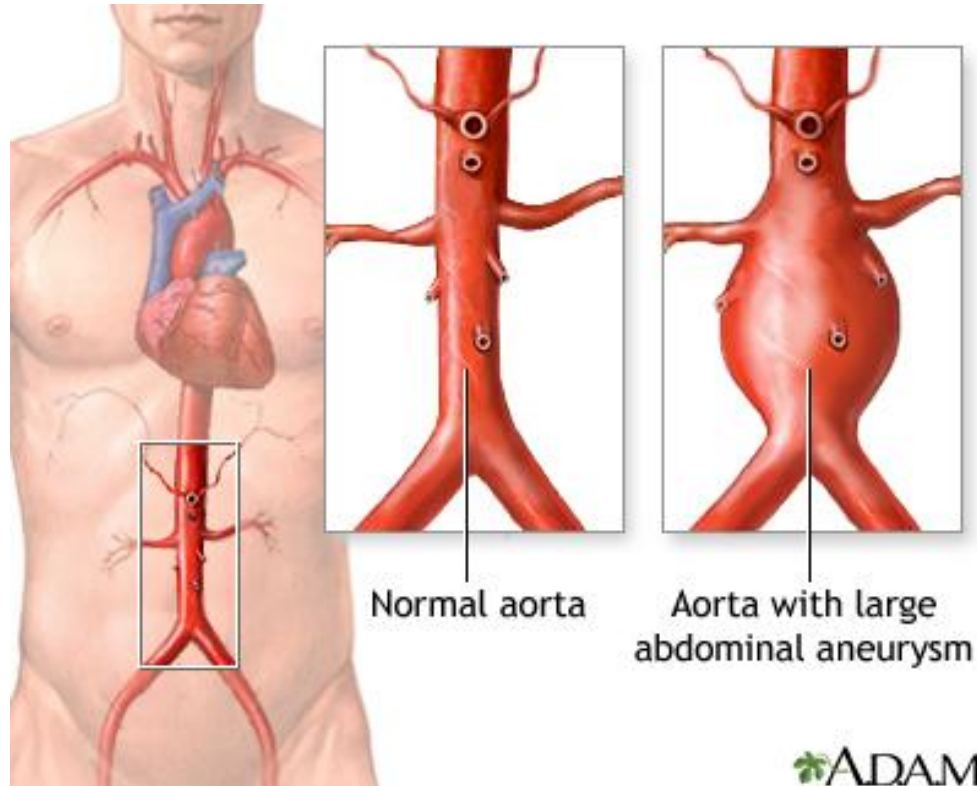
Vessel Wall Induced Pathologies

2. Hypertensive Vascular Disease



Vessel Wall Induced Pathologies

3. Aneurysms



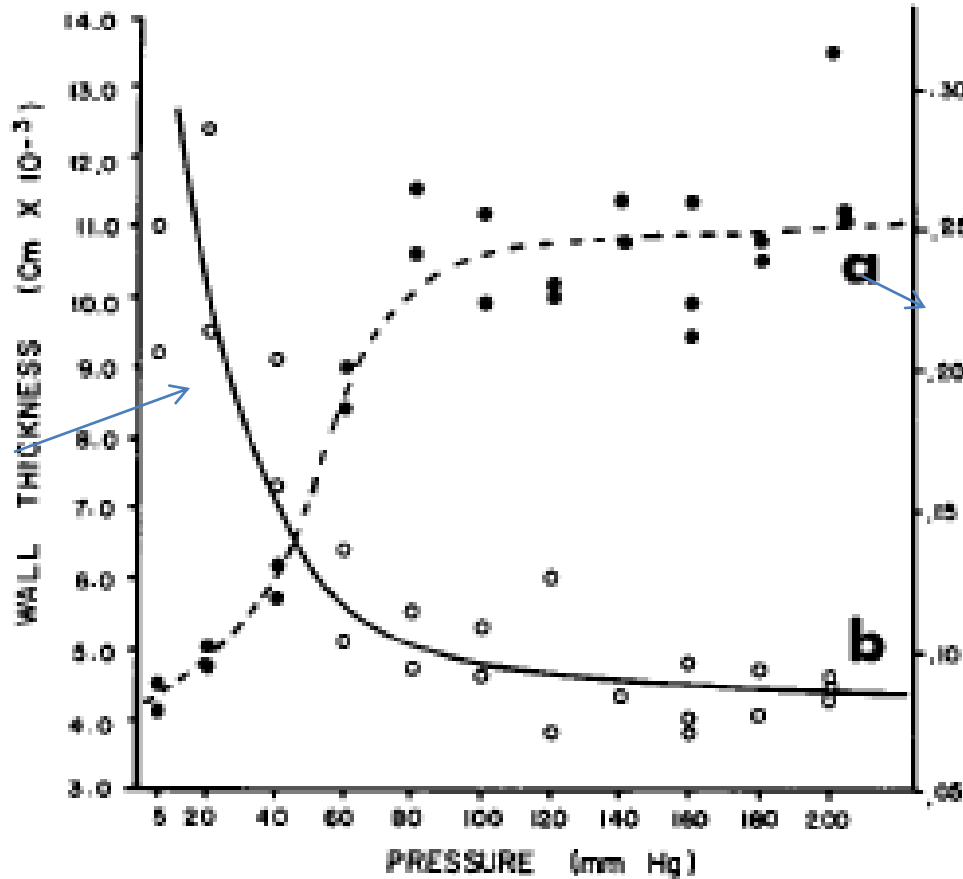
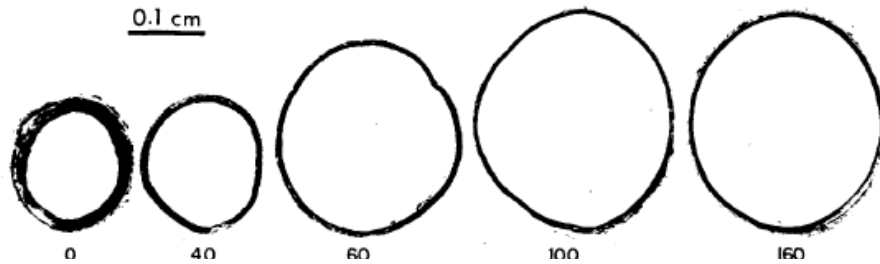
High risk of rupture and bleeding



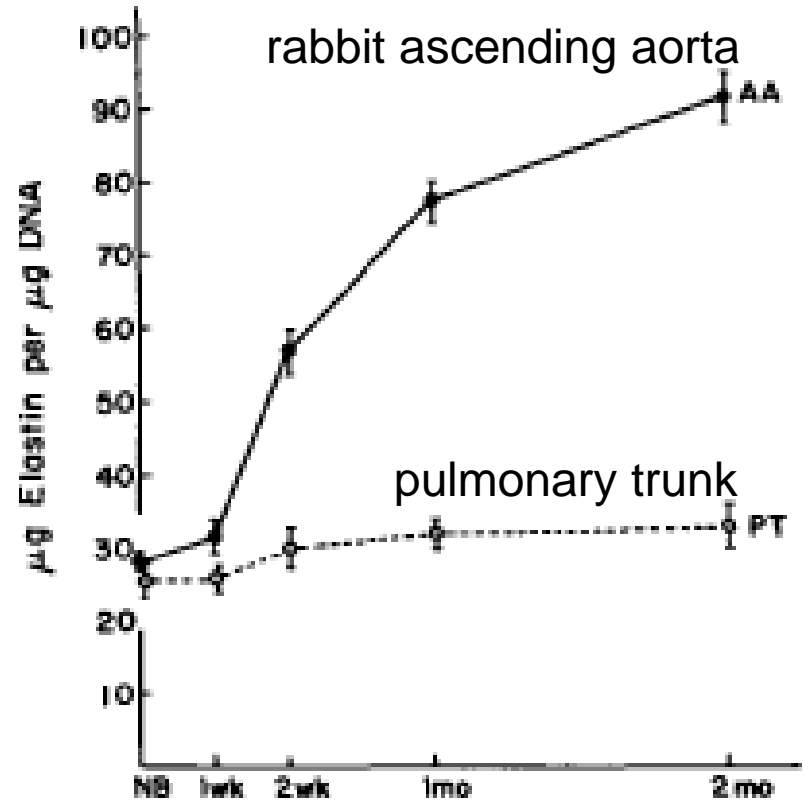
What are the structural components?

| ARTERY/VEIN | % H ₂ O | % COLLAGEN | % ELASTIN | C:E RATIO |
|--------------|--------------------|---------------|------------|-------------|
| Aorta | 70.4 ± 0.4 | 45.5 ± 1.7 | 30.1 ± 1.7 | 1.58 ± 0.15 |
| Carotid | 71.1 ± 0.1 | 50.7 ± 2.1 | 20.1 ± 1.0 | 2.55 ± 0.13 |
| Coronary | 63.2 ± 1.0 | 47.9 ± 2.6 | 15.6 ± 0.7 | 3.12 ± 0.12 |
| Femoral | 68.0 ± 0.3 | 44.5 ± 1.4 | 24.5 ± 1.6 | 1.89 ± 0.14 |
| Mesentary | 70.8 ± 0.5 | 38.1 ± 1.7 | 26.5 ± 1.7 | 1.51 ± 0.15 |
| Renal | 70.4 ± 0.7 | 42.6 ± 1.6 | 18.7 ± 1.8 | 2.46 ± 0.27 |
| Vena cava | | 35.07 ± 2.1 | 21.0 ± 3.7 | 1.67 ± 0.18 |
| Jugular vein | | 41.8 ± 2.8 | 47.1 ± 3.1 | 0.89 ± 0.09 |
| Femoral vein | | 47.0 ± 4.7 | 45.3 ± 2.6 | 1.04 ± 0.11 |

Structure of aorta

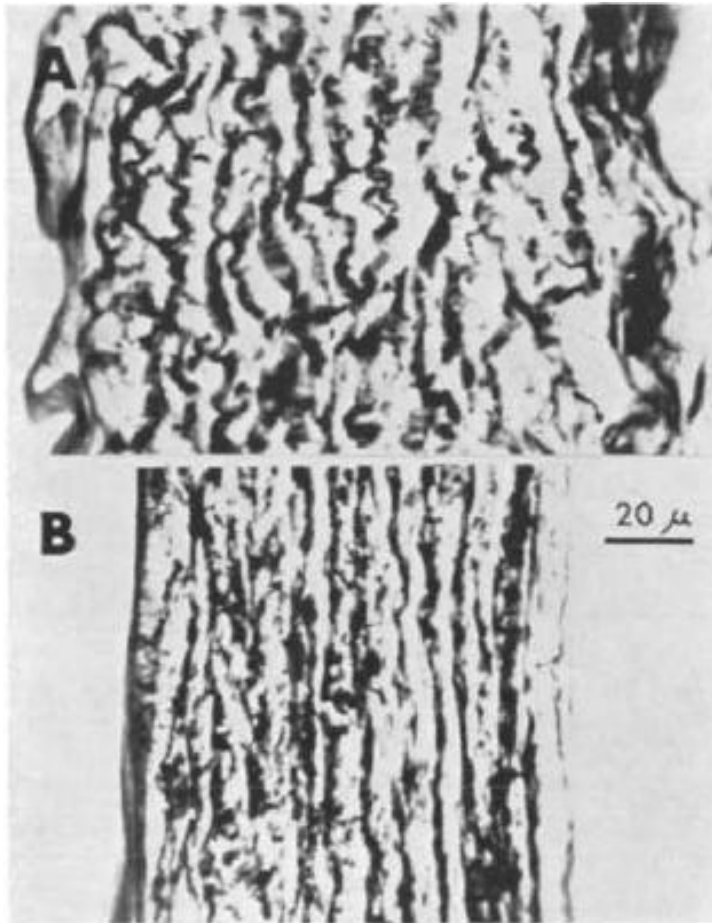


(Wolinsky, *Cir Res*, 1964)

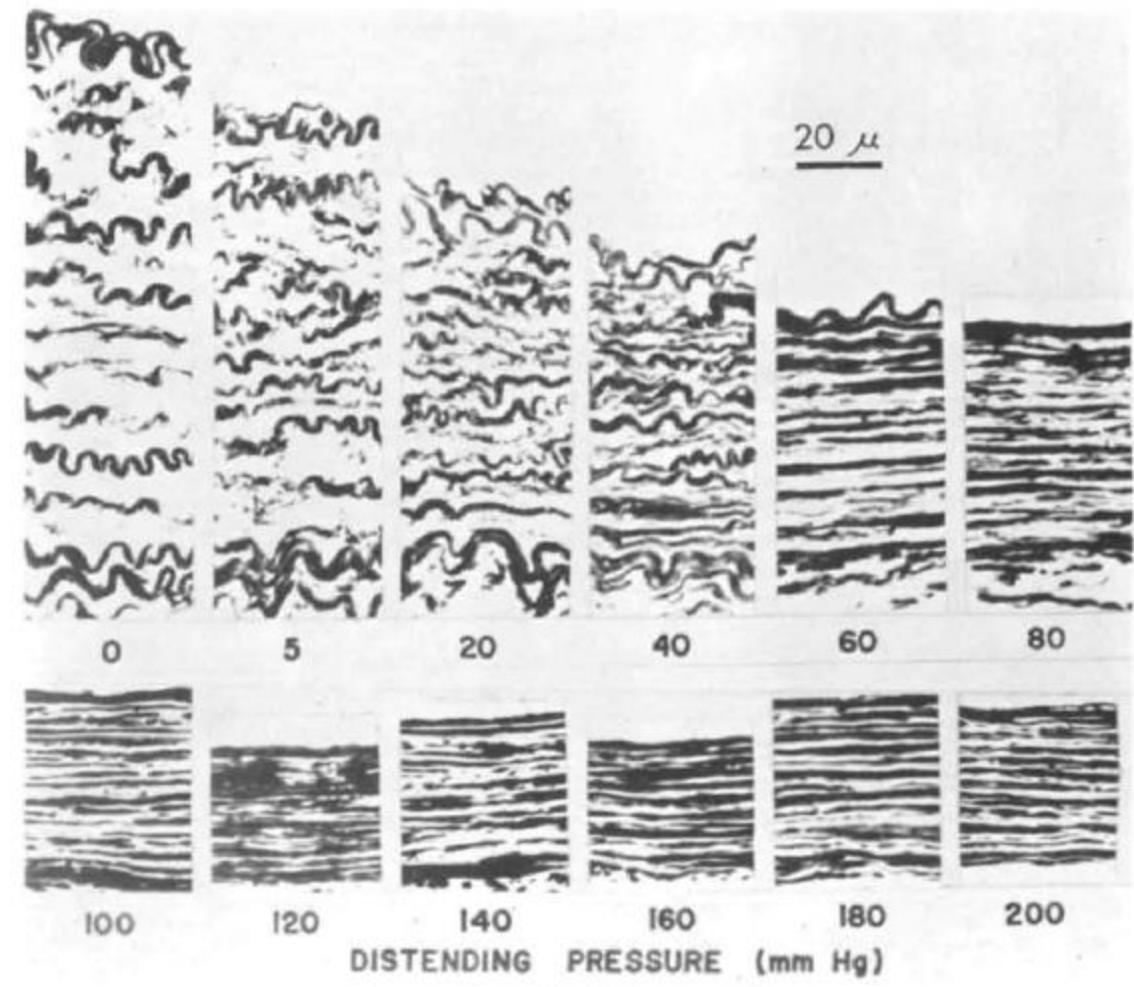


$T_{AA} = T_{PT}$ $T_{AA} = 6T_{PT}$
 Medial tension (Leung, *Circ Res*, 1977)

Vessel wall composition – aortic elastin



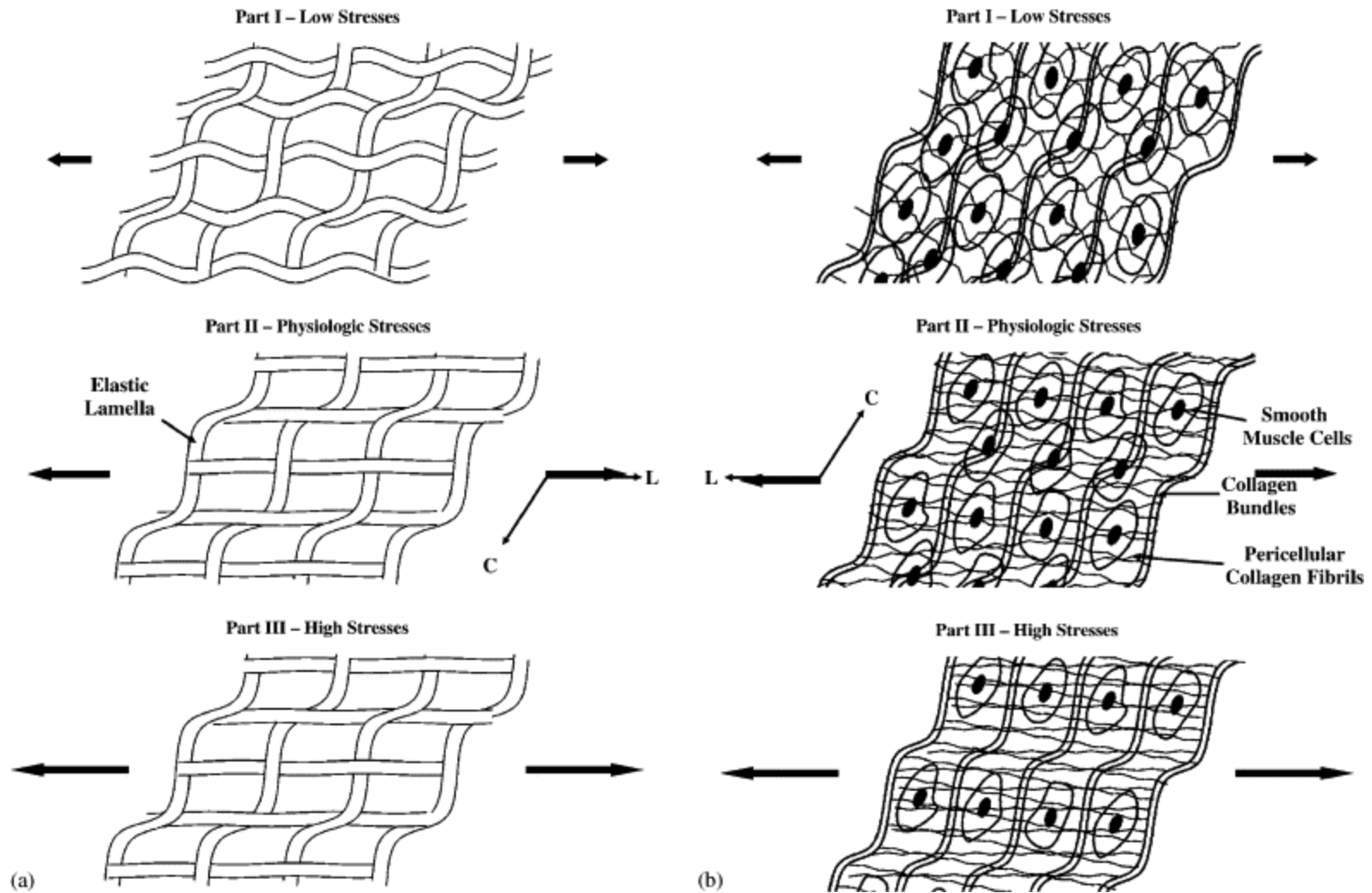
Longitudinal section



Cross-sectional section

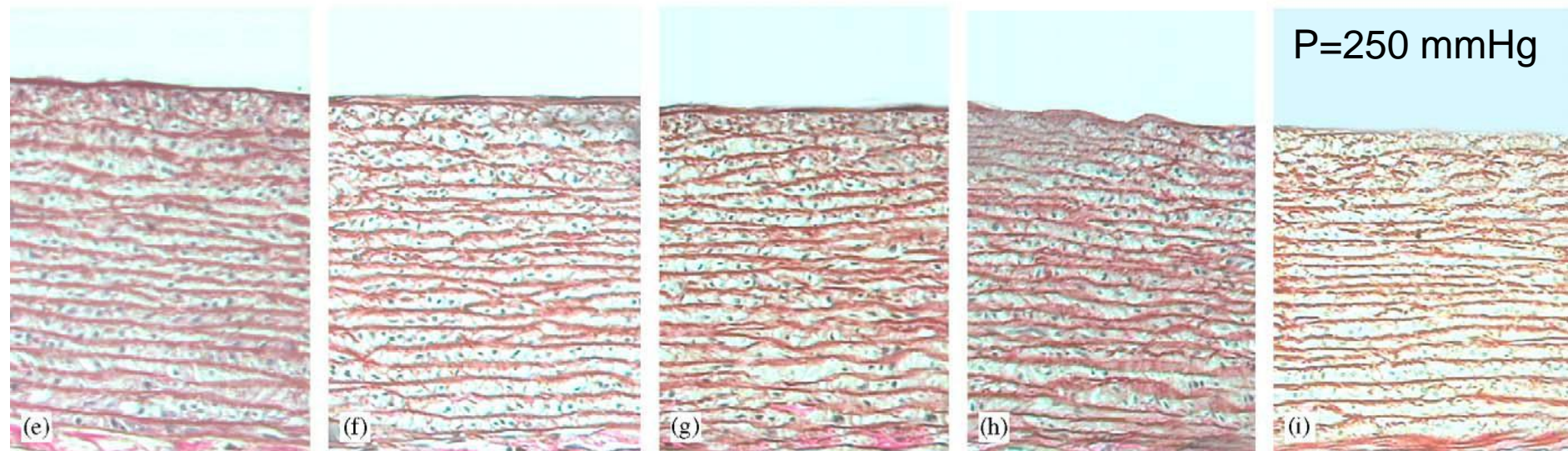
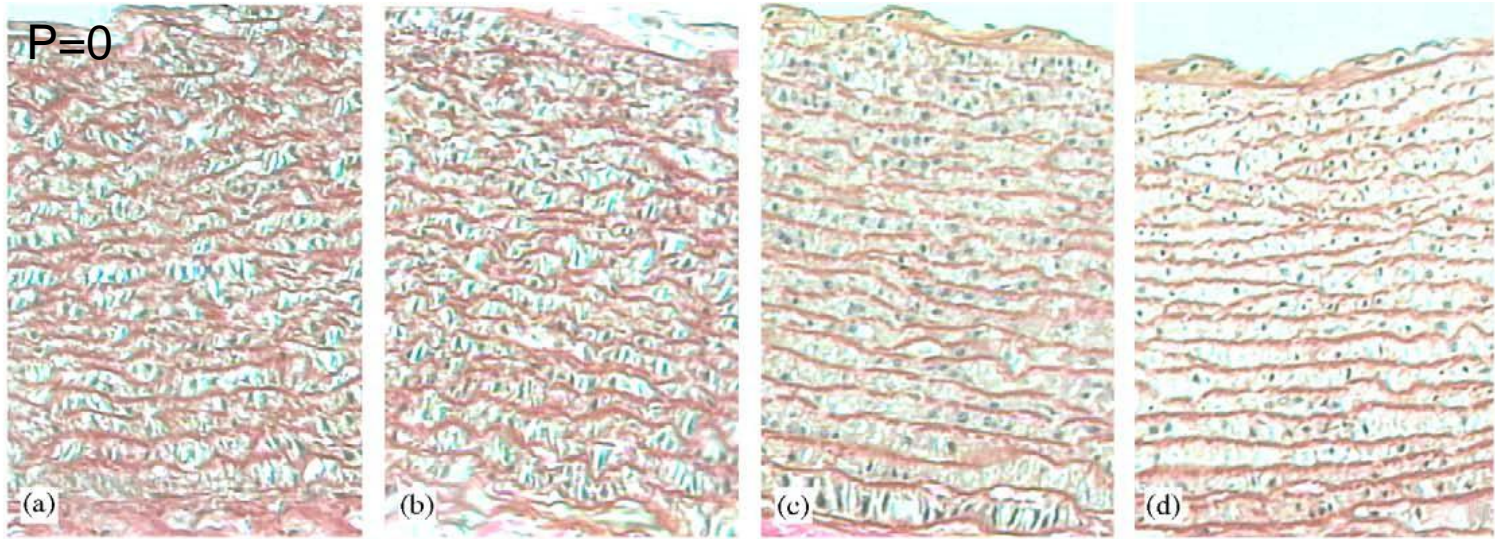
(Wolinsky, *Cir Res*, 1964)

Vessel wall composition – aortic elastin



Aortic wall composition – elastin

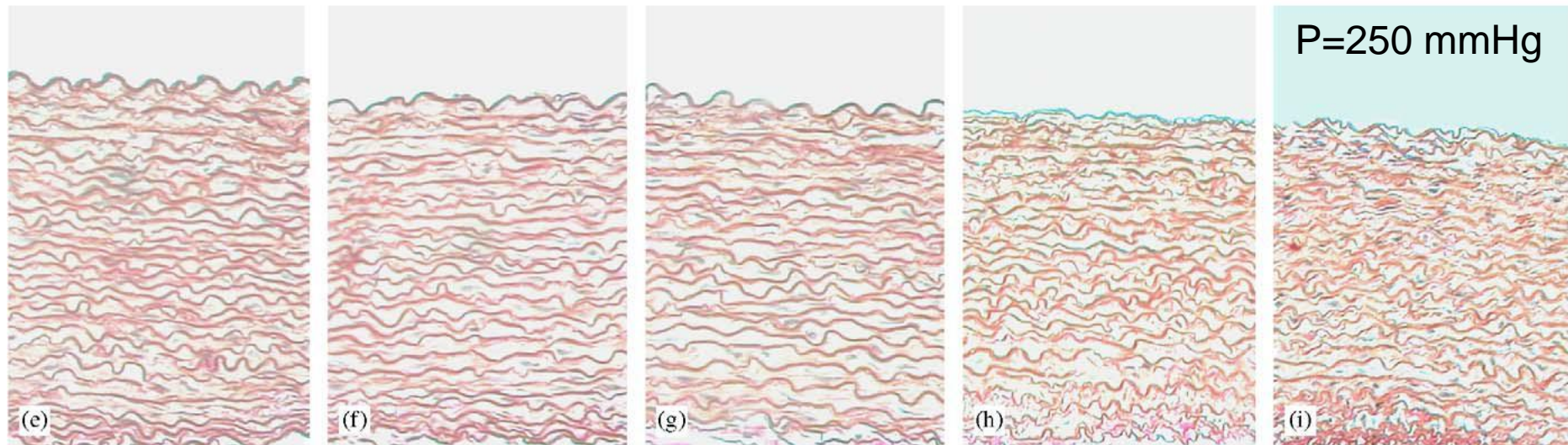
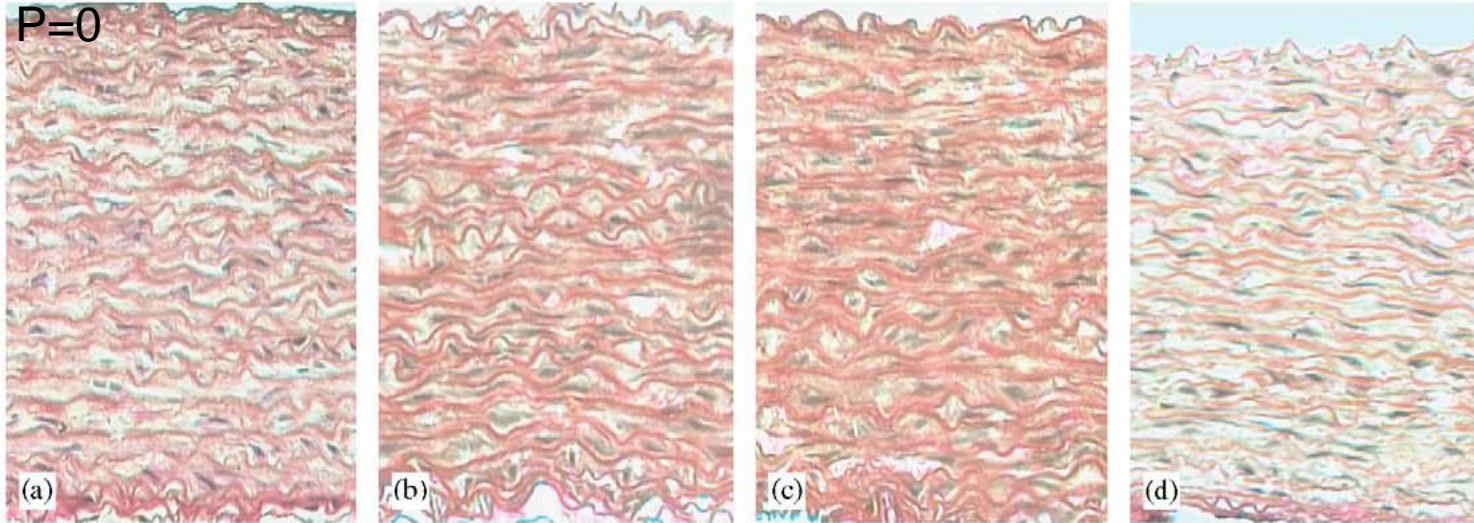
Longitudinal section



(Sokolis, *J Biomechanics*, 2006)

Aortic wall composition – elastin

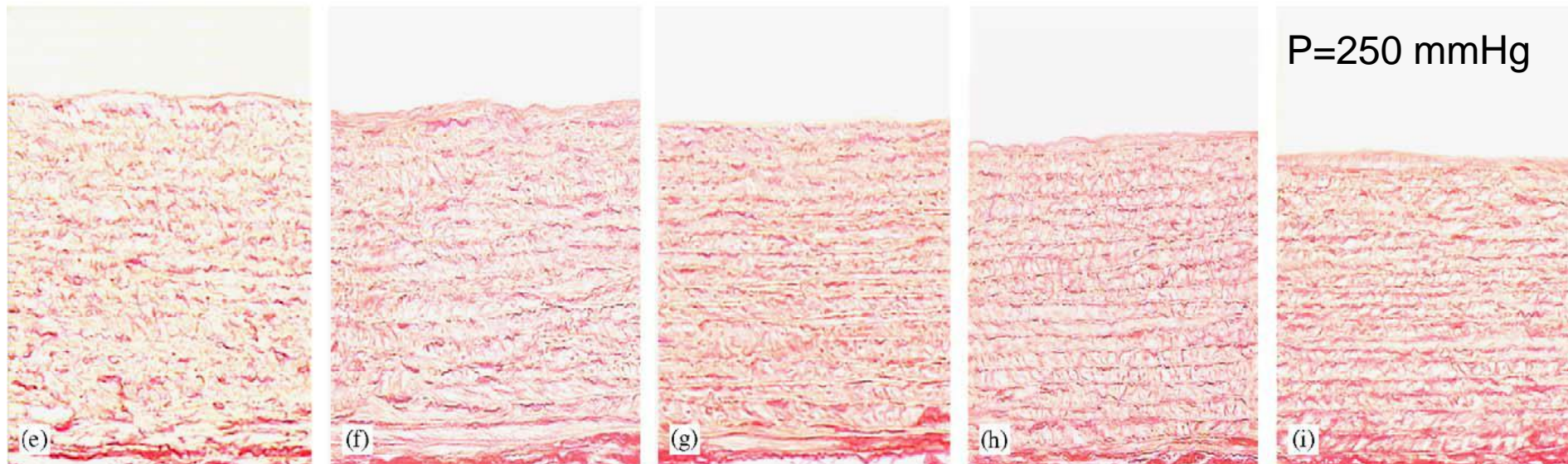
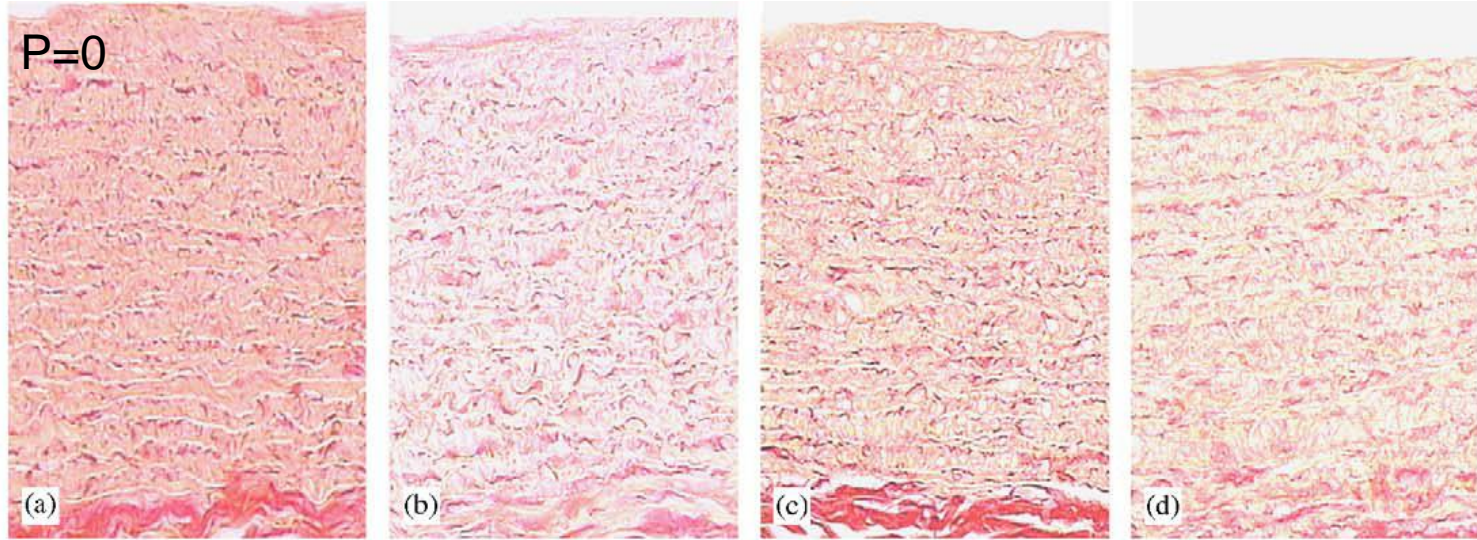
circumferential section



(Sokolis, *J Biomechanics*, 2006)

Aortic wall composition – collagen

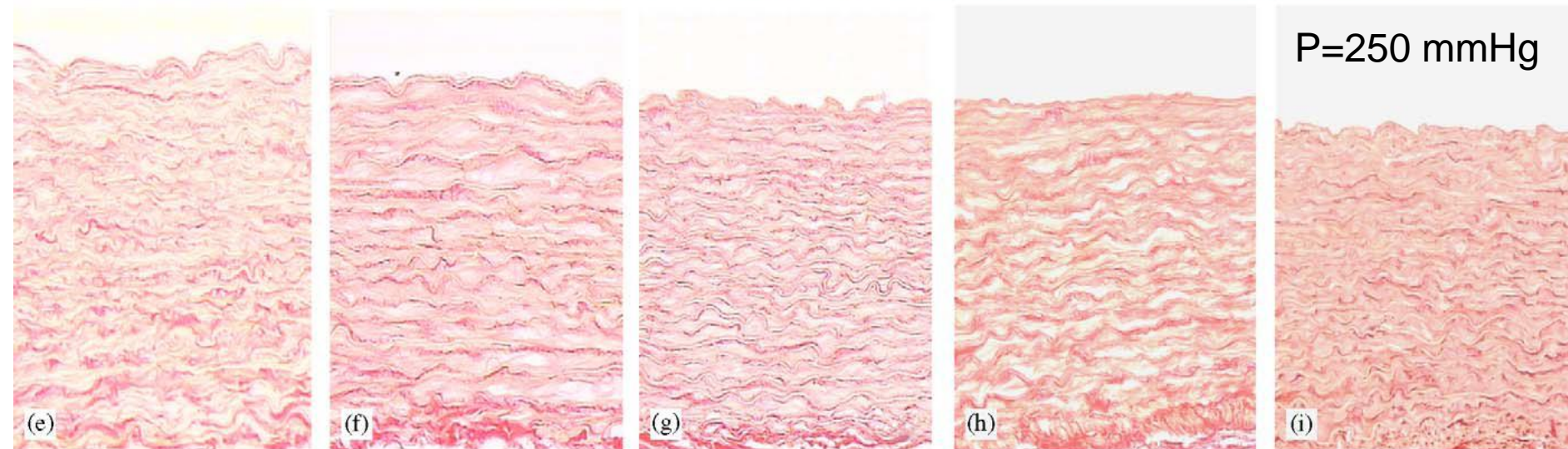
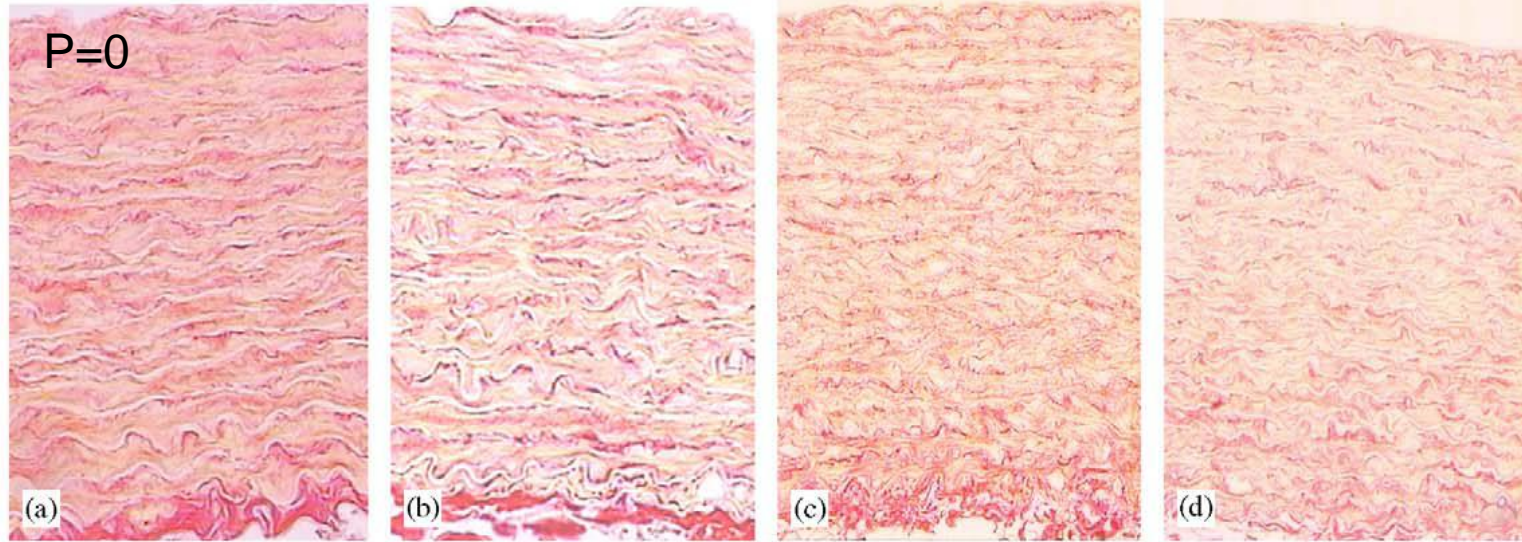
Longitudinal section



(Sokolis, *J Biomechanics*, 2006)

Aortic wall composition –collagen

circumferential section



(Sokolis, *J Biomechanics*, 2006)

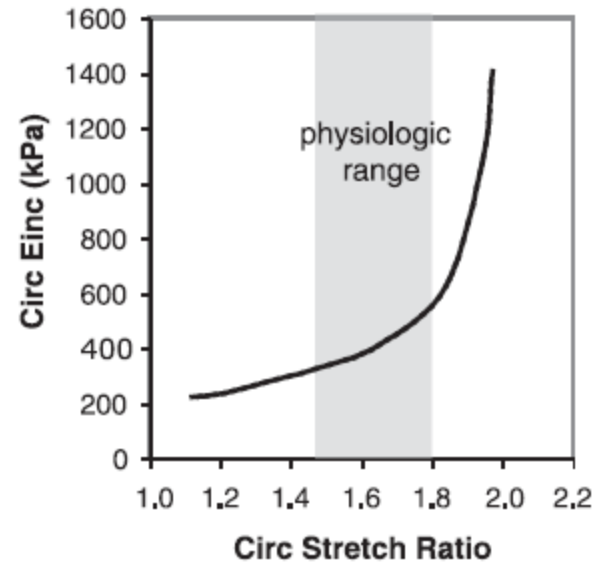
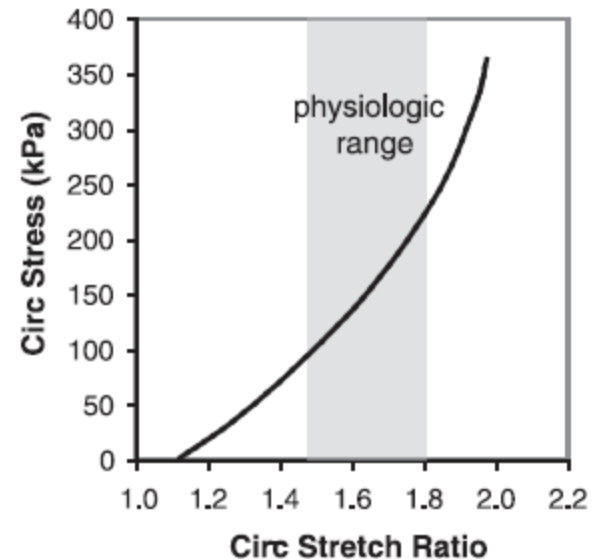
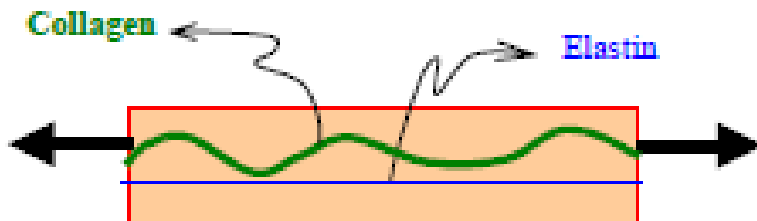
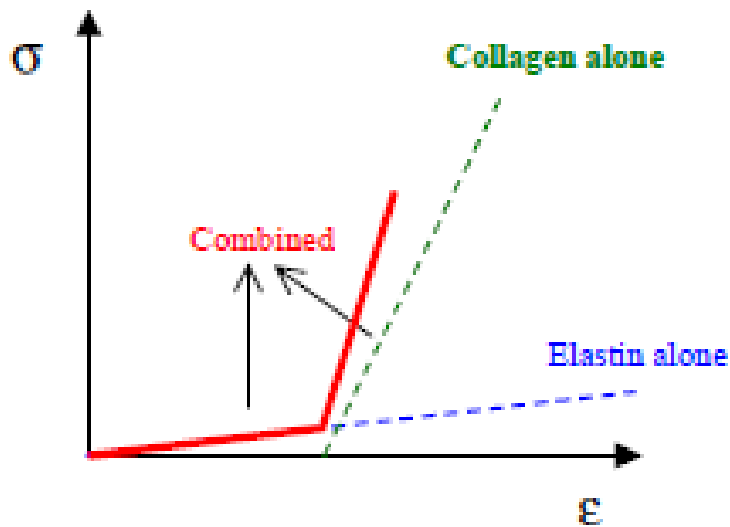
Vessel wall – Non-linear elasticity

Heterogeneity: Two-phase materials

Collagen: $E = 10^9$ dynes/cm²

Elastin: $E = 3 \times 10^6$ dynes/cm²

$$\sigma = E\varepsilon$$

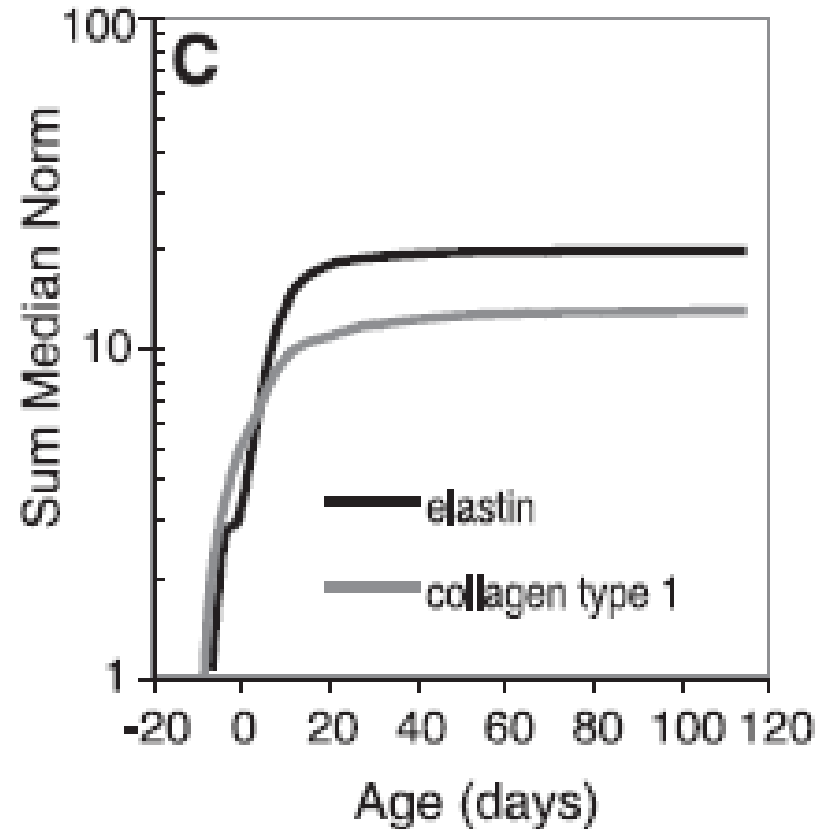
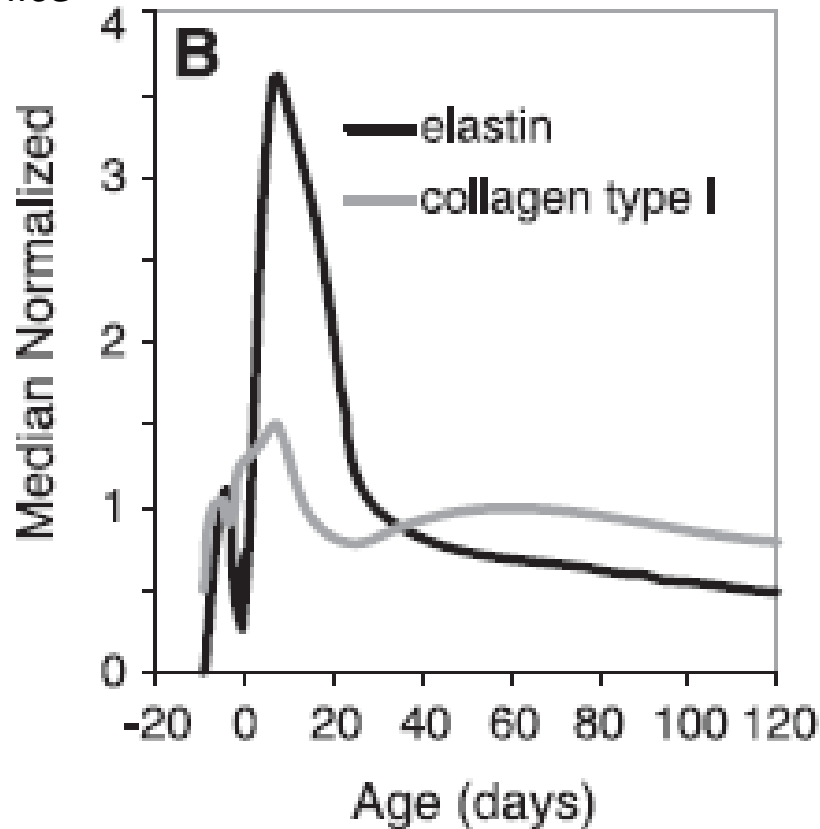


(Wagenseil, Mecham, 2005)

Vessel wall composition - Aging

Collagen and Elastin Expression

mice

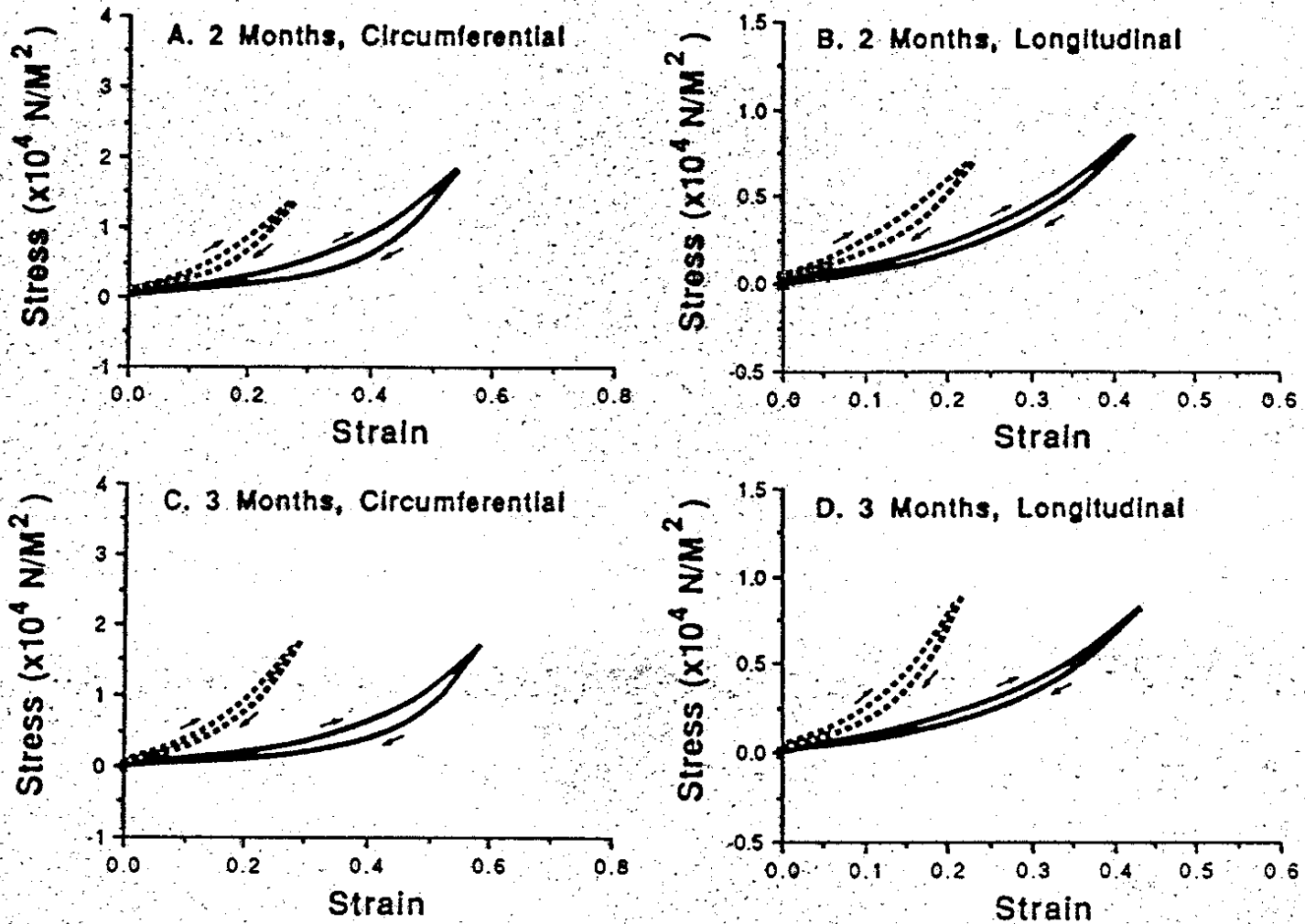


Diminished windkessel effect, hardening of the artery (fragmentation and loss of elastin)

(Wagenseil and Mecham, *Physiol Rev* 2009)

Vessel wall function - disease

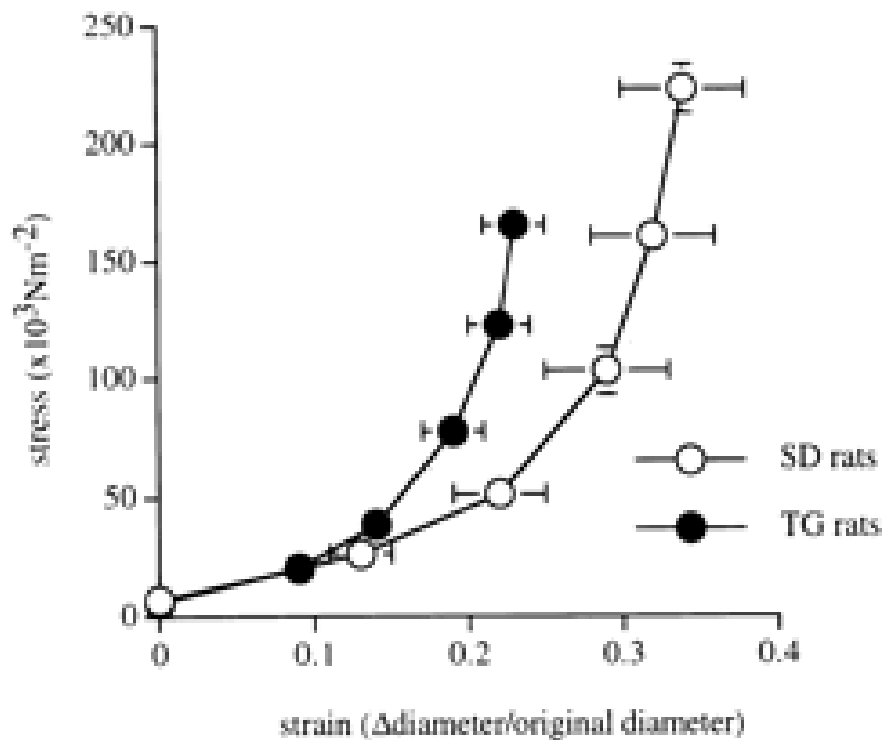
pulmonary arteries – rat smoking.



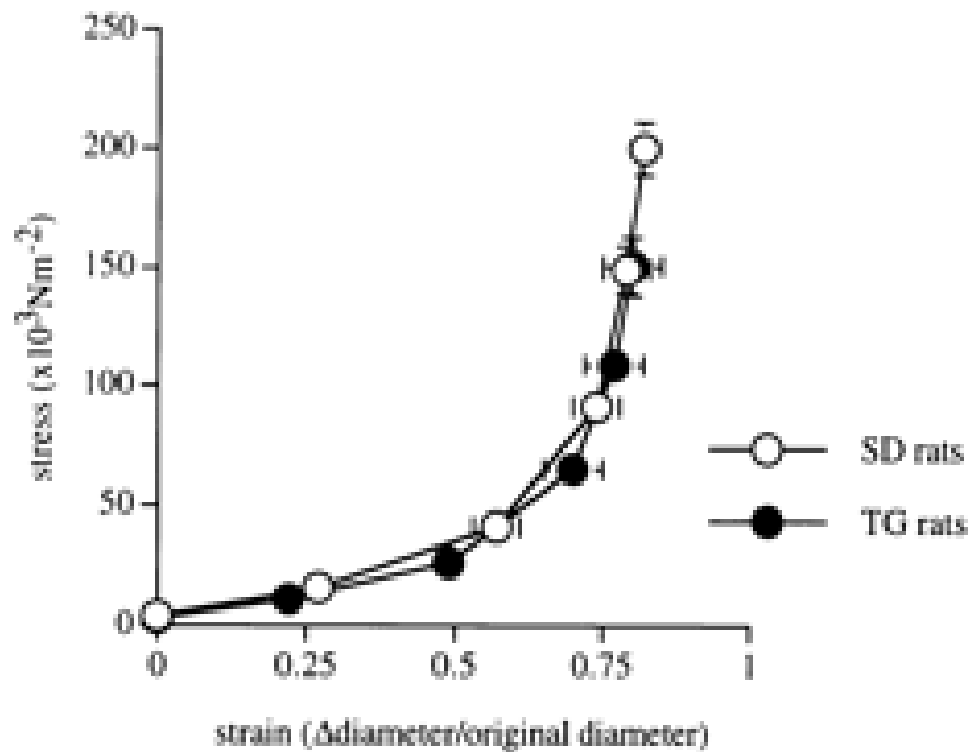
(Liu and Fung, *J Biomechanics*, 1992)

Vessel wall function - hypertension

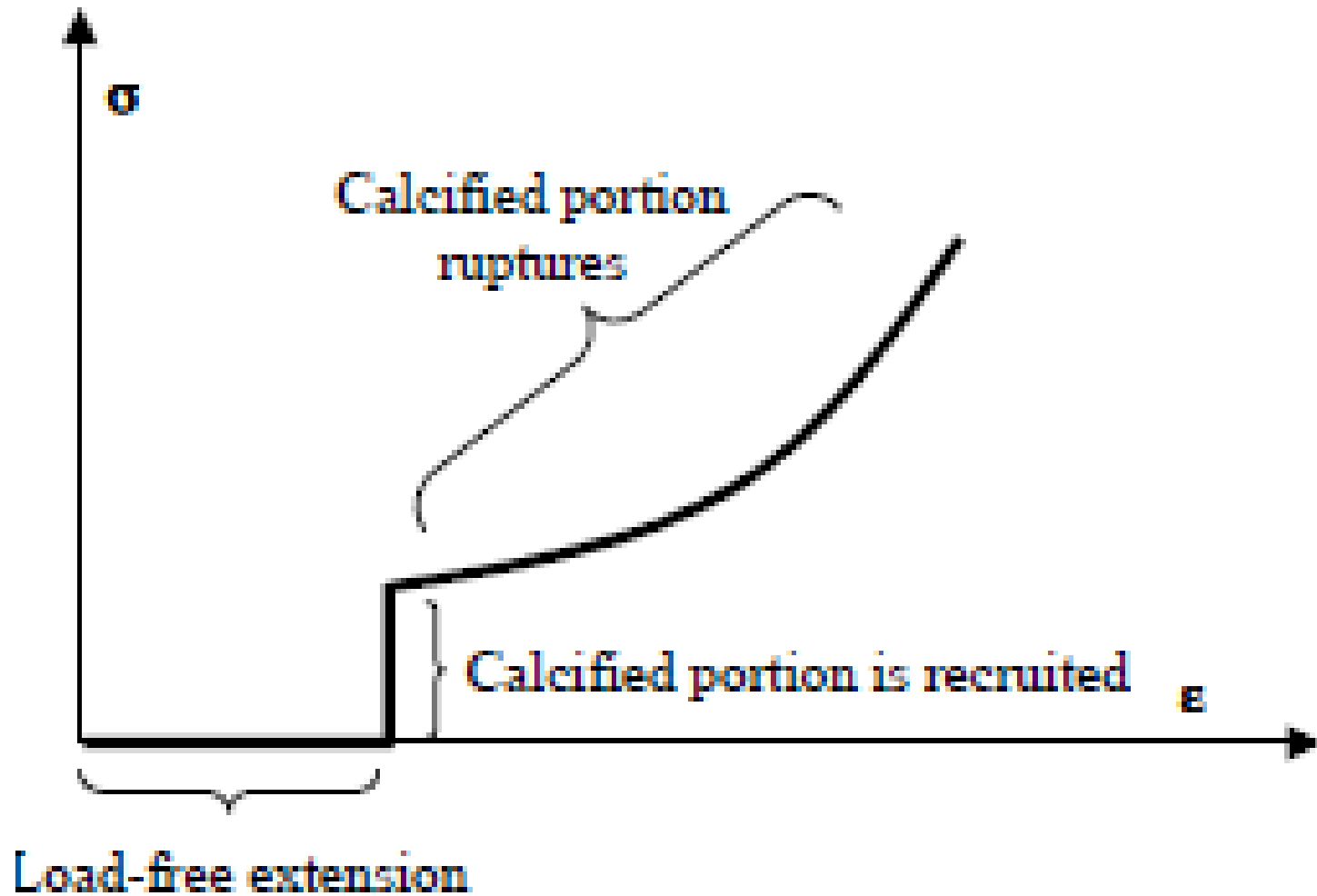
Rat cerebral artery



3-order resistive vessel



Vessel wall function - disease



What causes the change of vessel structure and function?

Learn from the development:

In 1893, Thomas :

Vessel lumen size depends on blood flow

Vessel length depends on longitudinal force on connective tissues

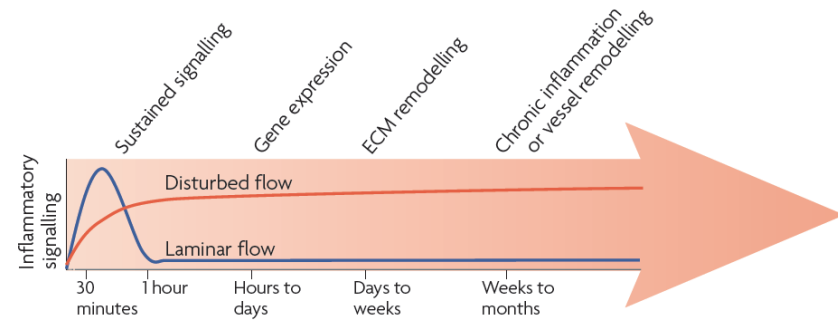
Vessel wall thickness depend on pressure

Changes by the cells

ECs:

short term -> secrete vasoconstrictor or vasodilator to constrict or relax the smooth muscle cells

long term -> generate basement membrane



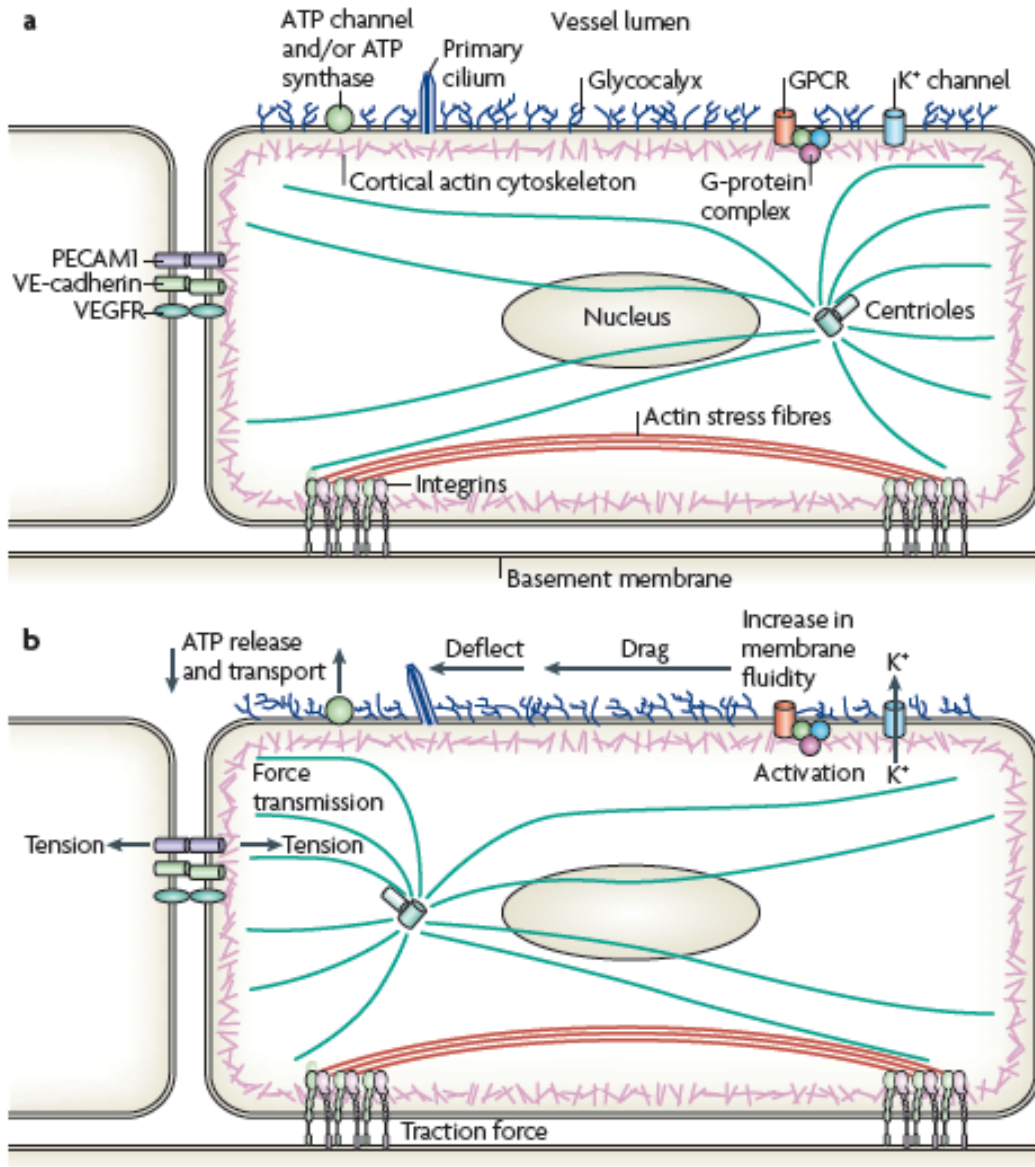
SMCs:

short term -> change diameter of artery wall in response to flow change.

long term -> change of elastin/collagen content (aneurysm), SMC replication

Collagen: I, III, V -> fibril-forming, responsible for vessel strength

Mechanotransduction of ECs



Ion channels, integrins, receptor Tyr kinases, apical glycocalyx, primary cilia, heterotrimeric G proteins, PECAM1, VE cadherin

(Hahn and Schwartz, *Nat Rev Mol Cell Biology*, 2009)

Mechanotransduction of SMCs

What do they sense:

Transmural pressure (120/80mmHg in arteries, 30-40mmHg in capillaries)

Vascular wall strain by pulsative pressure (coronary artery, carotid artery)

Circumferential, axial wall tension; radial compression

Passive or active mechanics, myogenic tone

Shear stress from luminal flow

Results: thicken, stiffen, lengthen the vessel wall.



Mechanotransduction of SMCs

How do they sense:

Increased transmural pressure



VSM membrane depolarization



Activating calcium entry



Vessel constriction

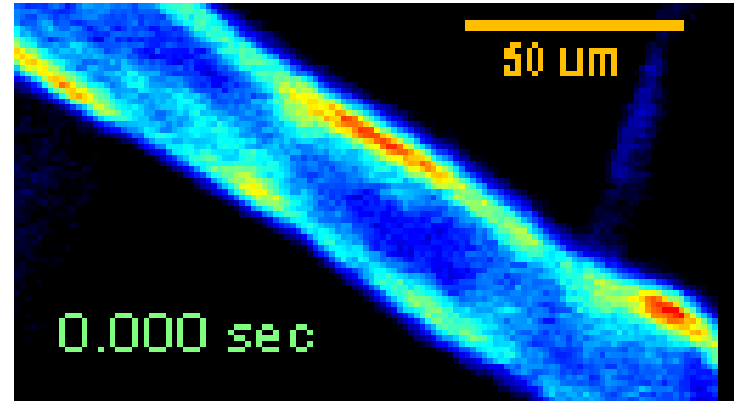


Hyperpolarization,



Activation of K_{ca} channels

GCaMP2 Transgenic Mice, Ach stimulation



(Tallini, *Circ Res*, 2007)

Application and Vascular Engineering

Acute hypertension

Atherosclerosis – SMC proliferation, matrix calcification

Coronary bypass vein grafts – when veins becomes artery – VSM induced fibrosis (collagen deposition)

Engineered vessel grafts always lack of elastin

