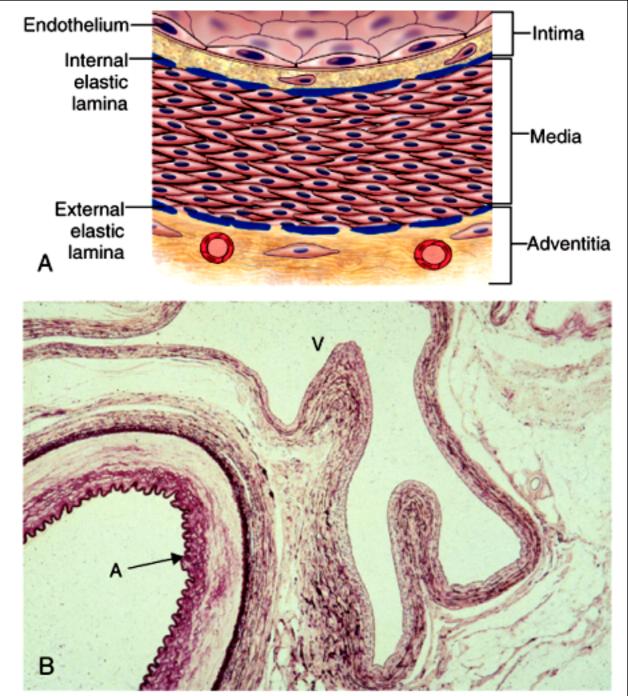
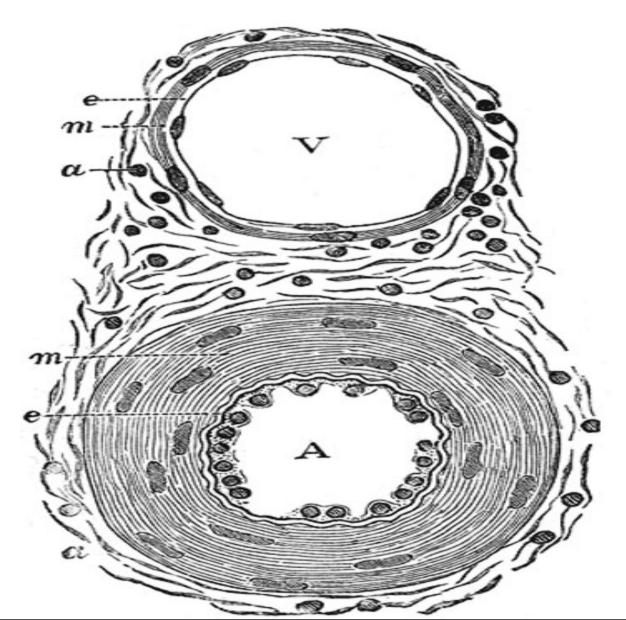
Normal blood vessels
A= artery
V= vein



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Artery (A) versus vein (V)

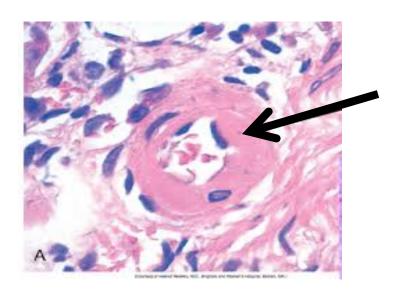


ARTERIOSCLEROSIS

- Arteriosclerosis ="hardening of the arteries"
- <u>arterial</u> wall thickening and loss of elasticity.
- Three patterns are recognized, with different clinical and pathologic consequences:

1-Arteriolosclerosis

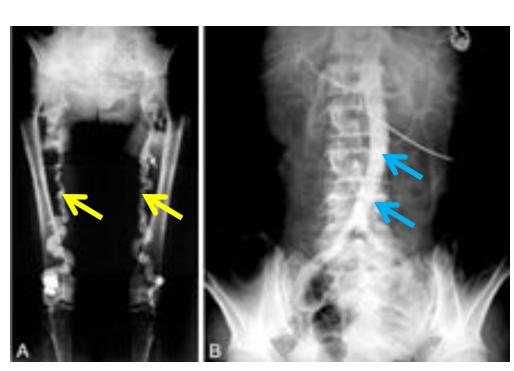
- affects small arteries and arterioles
- associated with hypertension and/or diabetes mellitus

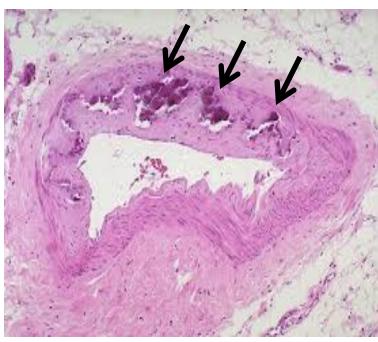


2- Mönckeberg medial calcific sclerosis

- calcific deposits in muscular arteries
- typically in persons > age 50
- radiographically visible (x-rays, etc...)
- palpable vessels
- do not encroach on vessel lumen and are usually not clinically significant

2-Mönckeberg medial calcific sclerosis





3-Atherosclerosis

- Greek word="gruel", "hardening,"
- most frequent and clinically important pattern of arteriosclerosis
- characterized by intimal lesions = atheromas (a.k.a. atherosclerotic plaques)

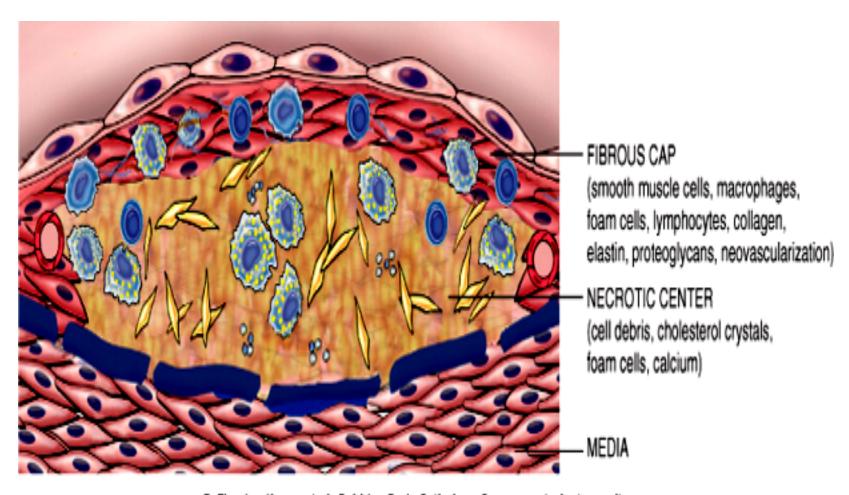
• atheromatous plaque = raised lesion with a core of lipid (cholesterol and cholesterol esters) covered by a firm, white fibrous cap

Pathogenesis

- not fully understood
- ? inflammatory process in endothelial cells of vessel wall associated with retained <u>low-density lipoprotein</u> (LDL) particles → ? a cause, an effect, or both, of underlying inflammatory process

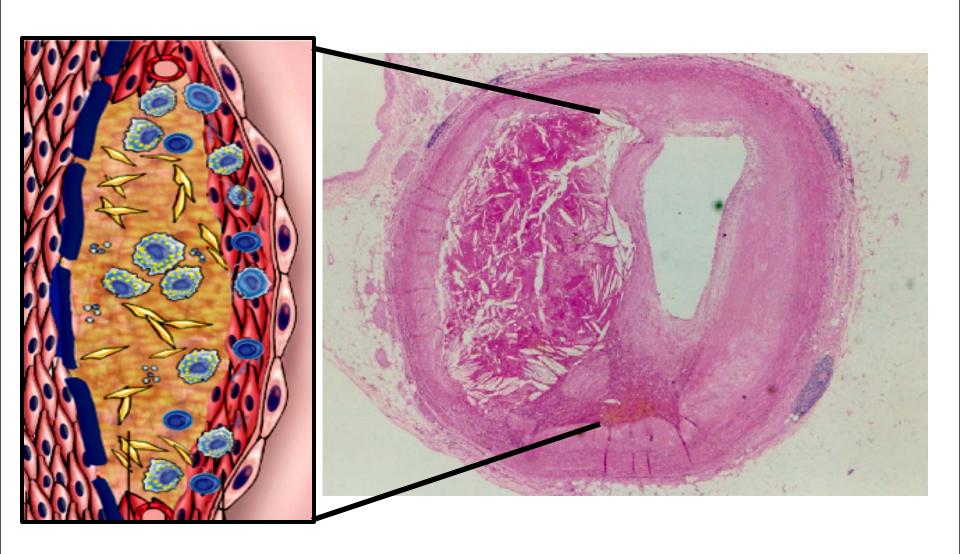
• <u>initiation of inflammatory process</u> LDL particles and their content are susceptible to oxidation by <u>free radicals</u> → <u>endothelial</u> activation

The major components of a well-developed intimal atheromatous plaque

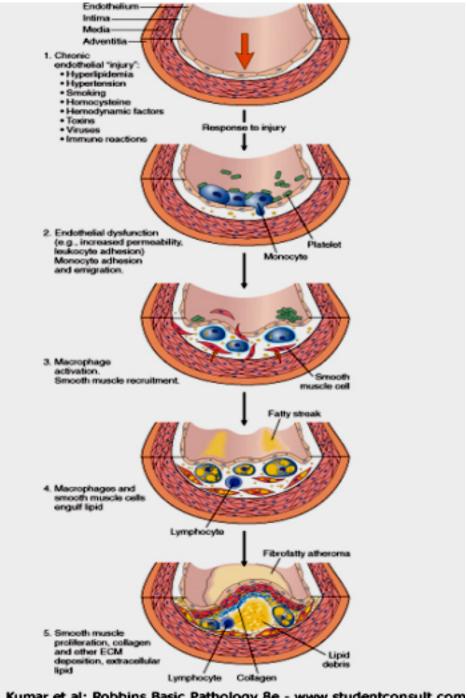


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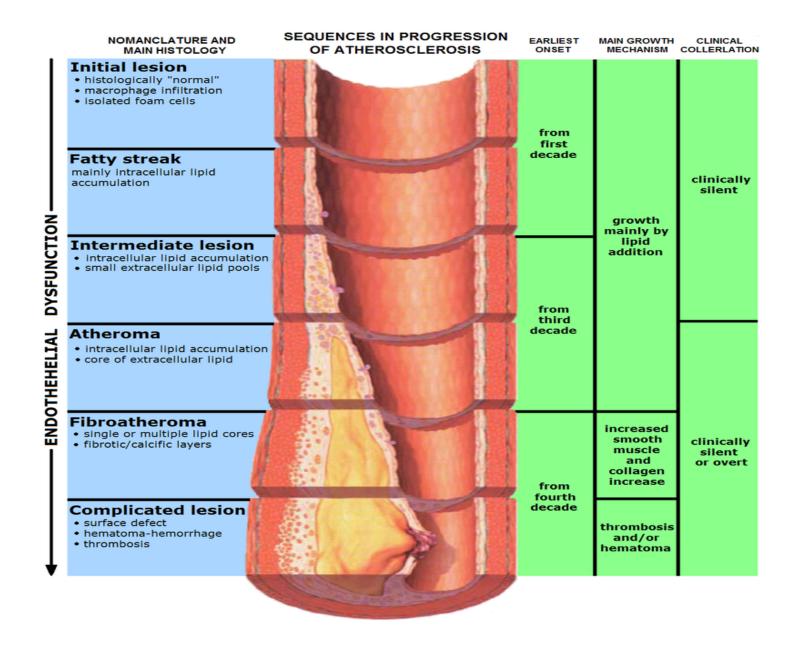
Atheromatous plaque



Formation of atheromatous plaque

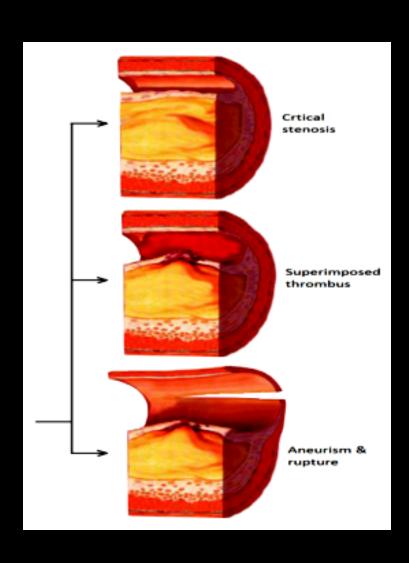


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Atherosclerosis progression



- Epidemiology
- Multiple risk factors have a multiplicative effect: 2 risk factors increase the risk 4X.
- E.g. if 3 risk factors are present (e.g., hyperlipidemia, hypertension, and smoking), the rate of myocardial infarction is increased 7X.

Risk Factors for Atherosclerosis

	Lesser, Uncertain, or Non-
Major Risks	quantitated Risks
Non-modifiable	Obsesity
Increasing age	Physical inactivity
Male gender	Stress ("type A personality)
Family history	Postmenopausal estrogen deficiency
Genetic abnormalities	High carbohydrate intake
	Lipoprotein(a)
Potentially Controllable	Hardened (trans)unsaturated fat intake
Hyperlipidemia	
Hypertension	Chlamydia pneumoniae infection
Cigarette smoking	
Diabetes	
C-reactive protein	

Major Constitutional Risk Factors for atherosclerosis

• Major Risks (Nonmodifiable):

- *Increasing age
- *Male gender
- *Family history
- *Genetic abnormalities

• Potentially Controllable/modifiable:

- Hyperlipidemia
- Hypertension
- Cigarette smoking
- Diabetes
- C-reactive protein

1-age

- ages 40 to 60, incidence of MI in men increases 5 x
- Death rates from IHD rise with each decade

2-Gender

- Premenopausal* → protected against atherosclerosis compared with age-matched men.
- * = unless they are otherwise predisposed by diabetes, hyperlipidemia, or severe hypertension.
- After menopause → incidence of atherosclerosisrelated diseases increases

3-Genetics

- familial predisposition is multifactorial.
- Either:
- 1- familial clustering of other risk factors
- e.g. HTN or DM

or:

- 2- well-defined genetic derangements in lipoprotein metabolism
- e.g. familial hypercholesterolemia

Additional Risk Factors for atherosclerosis

- 20% of all cardiovascular events occur without any major risk factor
- 1-Inflammation as marked by C-reactive protein
- 2-Hyperhomocystinemia
- 3-Lipoprotein a
- **4-Factors Affecting Hemostasis**
- Other Risk Factors
- 1-lack of exercise
- 2-competitive, stressful lifestyle ("type A" personality)
- **3-obesity**
- 4-Postmenopausal estrogen deficiency
- 5-High carbohydrate intake