

# Inflammation 6

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FRCPath

# Outcome of acute inflammation

- 1. resolution: regeneration and repair.
- 2. chronic inflammation.
- 3. scarring

# Resolution - definition

- Restoration of normal structure and function.

# resolution

Happens when:

1. Injury is limited and short lived.
2. No or minimal tissue damage.
3. Injured tissue can regenerate.

# Chronic inflammation

Happens if:

- Offending agent not removed.

# scarring

- If there is extensive tissue destruction
- Or in tissues that cannot regenerate.

# Serous inflammation

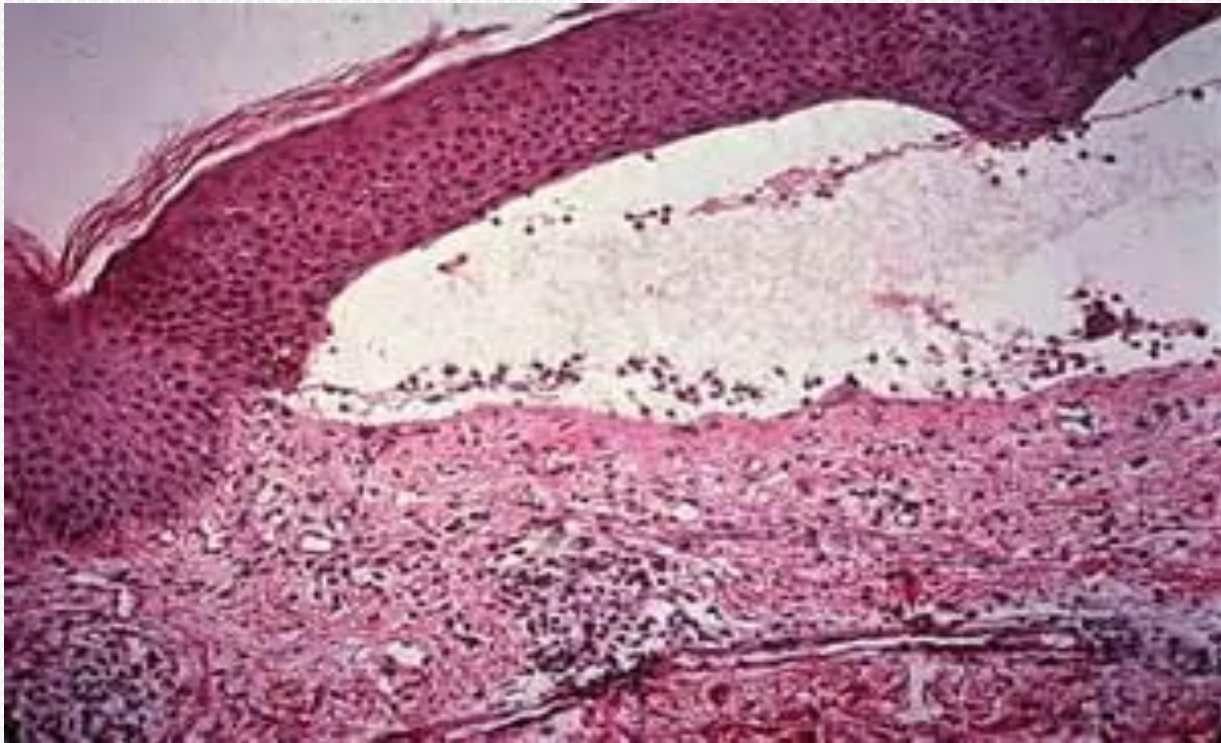
- Occurs if injury is mild.
- Watery protein-poor fluid.
- Happens in skin blisters due to burns or viral infections
- Occurs also in body cavities: serous effusions.

# Serous inflammation

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# Serous inflammation



# Fibrinous inflammation

- More severe injury.
- Greater vascular permeability.
- So: edema fluid contains fibrin.
- In body cavity lining... pericarditis.

# fibrin

- Protein.
- Eosinophilic meshwork of threads.

# Fibrinous inflammation

Outcome:

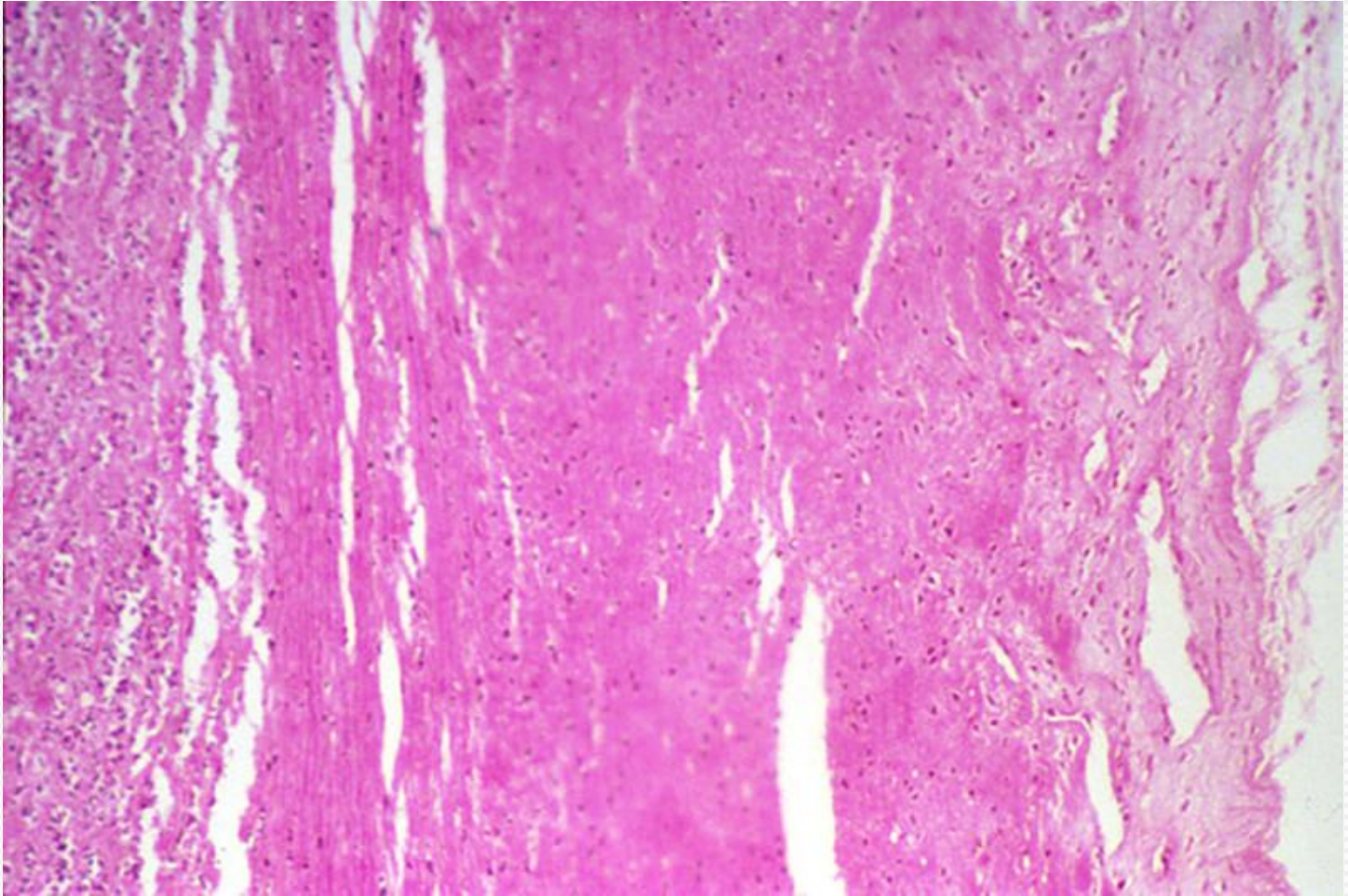
- Resolution if mild
- Scarring if more severe.

# Fibrinous pericarditis



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# Fibrinous pericarditis



# Suppurative inflammation

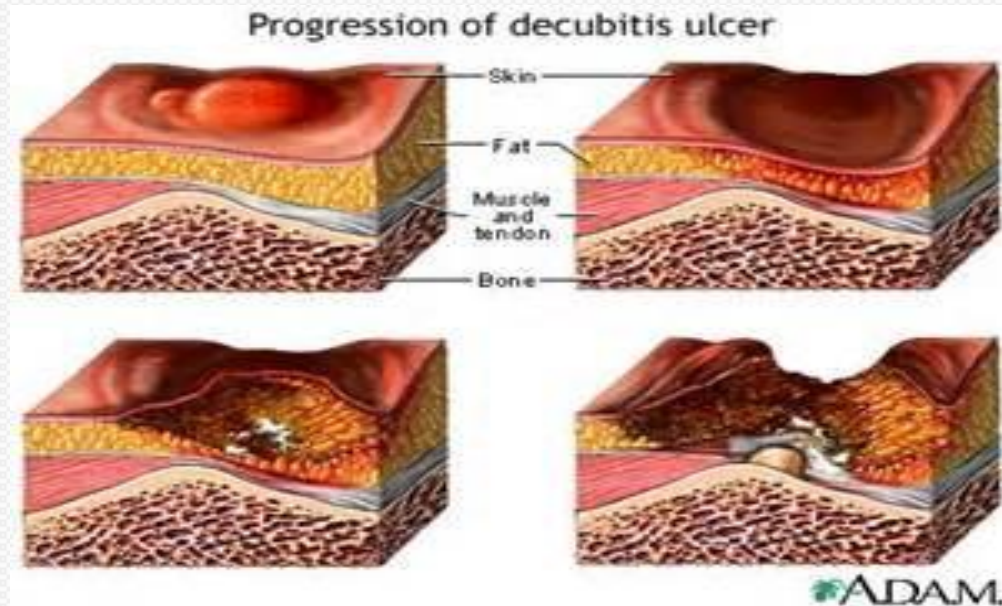
- In bacterial infections.
- Pus formation... composed of neutrophils.
- Abscess: focal collections of pus.
- Usual outcome: scarring

# ulcer

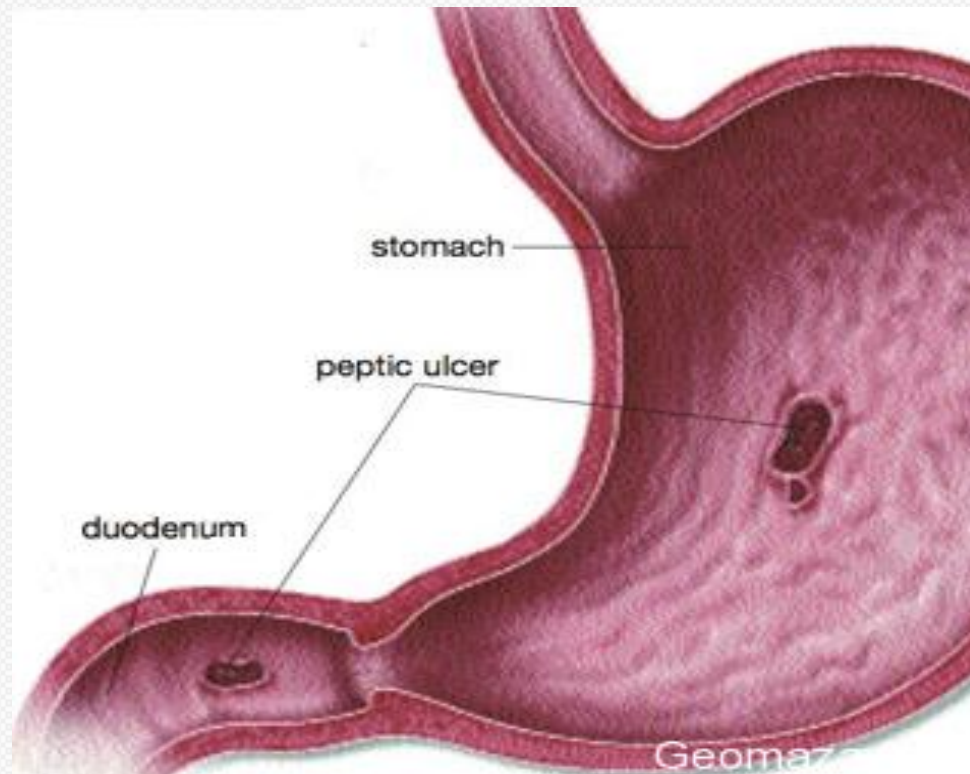
- Local defect of the surface of an organ or tissue produced by cellular necrosis and shedding of necrotic and inflammatory tissue.



# ulcer



# ulcer



ulcer



# Chronic inflammation



# Chronic Inflammation

Characterized by:

1. Infiltration with mononuclear cells.
2. Tissue destruction
3. Repair ( new angiogenesis & Fibrosis)



# Causes of chronic inflammation

- 1- Unresolving acute inflammation**
- 2- Persistent infections:** (TB, Syphilis, Fungi, viruses)
- 3- Prolonged exposure to toxic agents.** (Silica, plasma lipids like in atherosclerosis)
- 4- Autoimmune disease** (Rheumatoid arthritis, Inflammatory bowel disease)



# Cells of chronic inflammation

- Macrophages.
- Lymphocytes
- Plasma cells
- Eosinophils
- Mast cells

# Macrophages

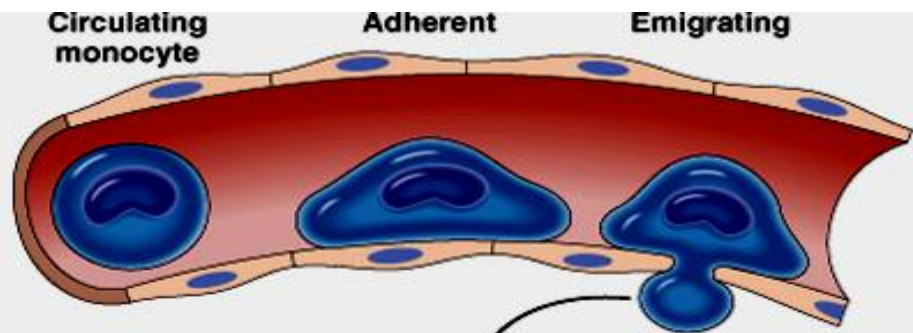
- Are the **dominant** cells of chronic inflammation
- derived from blood **monocytes**
- normally diffusely scattered in most connective tissues, and also in organs such as the liver (*Kupffer cells*), spleen and lymph nodes (called *sinus histiocytes*), central nervous system (*microglial cells*), and lungs (*alveolar macrophages*).





# M1 and M2 macrophages

- M1 .. Classical pathway... inflammation
- M2... alternative pathway... anti-inflammation and tissue repair and necrosis.



Tissue macrophage

IMMUNE  
RESPONSE:  
Activated T cell

ACTIVATION  
BY MICROBES,  
DEAD CELLS,  
ETC.

Cytokine  
(IFN- $\gamma$ )

Activated macrophage

**TISSUE INJURY AND  
INFLAMMATION**

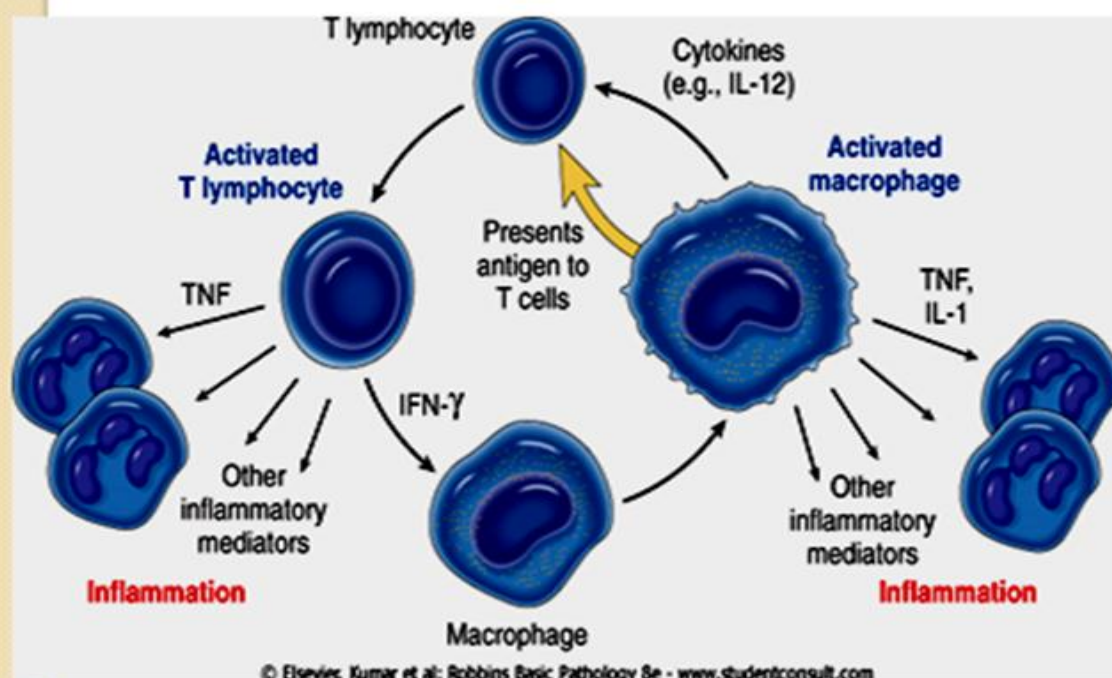
- Reactive oxygen and nitrogen species
- Proteases
- Cytokines, including chemokines
- Coagulation factors
- AA metabolites

**FIBROSIS**

- Growth factors (PDGF, FGF, TGF $\beta$ )
- Fibrogenic cytokines
- Angiogenesis factors (FGF)

# Lymphocytes and Plasma Cells

- Lymphocytes and macrophages interact in a **bidirectional** way, and these interactions play an important role in chronic inflammation
- **Activated T lymphocytes** produce cytokines, including **IFN- $\gamma$** , a powerful activator of macrophages,



# Plasma cells

- develop from **activated B lymphocytes**.
- produce antibodies against persistent antigens or against altered tissue components.

# *Eosinophils*

- characteristically found in inflammatory sites around:
  1. **parasitic infections or**
  2. **as part of immune reactions mediated by IgE, typically associated with *allergies*.**

# eosinophils



# Granulomatous Inflammation

- Aggregates of epithelioid histiocytes

- Mechanism:

1. Persistent T-cell response to certain microbes as M. tuberculosis,
2. Foreign bodies. E.g. suture, splinter.



**Diseases associated with granulomatous inflammation include:**

1. Tuberculosis
2. Leprosy
3. sarcoidosis

# Granuloma morphology

- **Macrophages in granulomas have pink, granular cytoplasm with indistinct cell boundaries (epithelioid macrophages)**
- **These are surrounded by a collar of lymphocytes**
- **Older granulomas may have a rim of fibroblasts and connective tissue, and frequently, multinucleated giant cells**

# Caseating granulomas

- hypoxia and free-radical injury leads to a central zone of **necrosis**.
- Grossly, this has a granular, cheesy appearance (**caseous necrosis**)
- Microscopically, amorphous, structure-less, granular debris, with complete loss of cellular details.
- Seen mainly in TB.

# Caseating granulomas

- a combination of **hypoxia** and **free-radical injury** leads to a central zone of necrosis. Grossly, this has a granular, cheesy appearance and is therefore called **caseous necrosis**.

Microscopically, this necrotic material appears as amorphous, structureless, granular debris, with complete loss of cellular details.

- Healing of granulomas is accompanied by fibrosis that may be quite extensive.

# Granuloma: microscopic morphology

