Inflammation II
Outcomes of inflammation with clinical examples

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Acute Inflammation: Sequelae

**RESOLUTION**
(see Acute Inflammation 1 lecture)

**SUPPURATION**
+/- abscess formation
+/- discharge of pus

**CHRONIC INFLAMMATION**

**ORGANISATION AND REPAIR**

**FIBROSIS**
SUPPURATION

Definitions

**Suppuration:** the formation of pus.

**Pus:** an accumulation of dead and living neutrophils, dead and living bacteria (when inflammation caused by **pyogenic bacteria**), protein (especially fibrin) and other particulate matter (eg cell fragments etc).

**Abcess:** a pus-filled cavity

**Empyema:** an accumulation of pus in a naturally occurring body cavity

This type of exudate if referred to as ‘**suppurative**’ or ‘**purulent**’
Suppurative meningitis

Note creamy yellow exudate around vessels in subarachnoid space

Micro: Exudate predominantly neutrophils, fibrin, bacteria
Evolution of an abscess

Starts as an acute inflammatory exudate with many neutrophils. Proteins (mainly fibrin), bacteria and polymorphs aggregate to form a mass

**Tissue death (necrosis)** ensues

New capillaries and fibroblasts develop at edge of accumulated material = ‘granulation tissue’ (process of organisation)

Fibroblasts start to lay down scar tissue (collagen)

Pus resorbed (if small amount) or can burst onto (‘point’) to external surface (sinus) or adjacent body cavity (fistula) and be discharged in this way

Collagen deposition proceeds to formation of mature scar
Brain abscess
Pus filled cavity with peripheral organisation
Wall of abscess
Note: suppurative exudate and surrounding organisation
EXAMPLES of ACUTE INFLAMMATORY REACTION IN THE LUNG caused by PYOGENIC BACTERIA

(Bacterial Pneumonia)

• Lobar pneumonia
• Bronchopneumonia
LOBAR PNEUMONIA
S. Pneumoniae
Note consolidation (hepatisation) of lower lobe
Consolidation of entire lobe
Lobar pneumonia - often caused by *Streptococcus pneumoniae*. Alveolar spaces filled with exudate - neutrophils, fibrin, dead bacteria. (Referred to as consolidation or hepatisation)
Outcomes of pneumonia

- Resolution
- Abscess formation
- Empyema
- Fibrosis and scarring
- Septicaemia
- Death
**Bronchopneumonia**

Patchy distribution of consolidation, related to bronchi
Abscess formation complicating pneumonia
Description:

- Inflammation enduring longer than acute inflammation
- May be primary but often results from acute inflammation when causative agent cannot be removed
- Polymorphs (neutrophils) largely replaced by lymphocytes, plasma cells (and macrophages)
- Macrophages often fuse to form giant cells
- Often proliferation of vascular endothelium and fibroblasts esp at periphery (= organisation)
- Fibrosis
Lung – chronic inflammation
Note lymphocytic aggregate (*), interstitial fibrosis (long arrows), Type 2 pneumocytes (blunt arrows)
Example of chronic inflammatory reaction

CHRONIC PEPTIC ULCER
Chronic ulcer occurring in an area of acid pepsin digestion
Commonly stomach duodenum oesophagus
Often associated with *Helicobacter pylori* infection
Chronic peptic ulcer of stomach (*)
CHRONIC PEPTIC ULCER
OUTCOMES OF CHRONIC PEPTIC ULCER

• Resolution - rare without appropriate therapy
• Haemorrhage
• Fibrosis (± stenosis)
• Perforation
• Penetration (± fistula formation)
• Malignant transformation very rare
Chronic peptic ulcer complications

Perforation (left)
Haemorrhage (right)
Example of chronic inflammatory disease

Tuberculosis

*Mycobacterium tuberculosis*
Tuberculosis
Tuberculous meningitis - above
Tuberculosis of spine (Pott’s disease) - right
Tuberculosis
Often associated with ‘caseous’ necrosis

LUNG
Caseous tuberculosis and tuberculous bronchopneumonia
Tuberculosis:
Often associated with *granulomatous inflammation*
Note epithelioid macrophages and giant cells
Outcomes of inflammation

Healing and repair
Healing

Resolution
- Removal of exudate
- Regeneration of tissue if possible
- Complete return to normal

Repair
- Occurs when resolution impossible (severe, ongoing damage, or tissue cannot regenerate)
- Involves formation of granulation tissue (organisation)
- Maturation of granulation tissue to scar tissue (fibrosis)
Organisation

Definition

• The growth of new capillaries and fibroblasts into the damaged tissue together with migration of macrophages. Macrophages remove debris, fibroblasts lay down collagen.
• New capillaries and fibroblasts = ‘granulation tissue’ which matures to form fibrous tissue (collagen).
• Often occurs when exudation or damage is excessive and cannot be removed
• Is the process involved in repair (healing) of tissue (when resolution is not possible)
Example of healing
Healing of a skin wound
Healing of a skin wound

Healing by primary intention
• Occurs in clean incised wounds with apposed edges (e.g., surgical wounds)
• Results in minimal scarring
• Occurs in shorter time (mainly healed in a week or two – stitches can be removed)
• Strengthening, devascularisation continues longer

Healing by secondary intention
• Occurs in open wounds (loss of tissue, necrosis or infection)
• Often results in significant scarring (fibrosis)
• Process may continue for months or years
Healing by primary intention

**Immediate**: small cavity fills with blood and fibrin

**2-3 hours**: minor inflammation

**2-3 days**: macrophage, fibroblast activity, new vessel formation (ie minimal granulation tissue)

**10-14 days**: re-epithelialisation complete, weak fibrous union

**Weeks**: good fibrous union continues strengthening for months to years.

Devascularisation. Minimal scarring

Healing by secondary intention

**Immediate**: large cavity fills with blood and fibrin. Acute inflammation begins.

**Days**: epithelium begins to regenerate to cover lesion. Overlying exudate = scab.

Contraction of wound

New capillary loops form, bring macrophages, neutrophils (prominent granulation tissue). Fibroblasts proliferate

**Weeks-months**: epithelium restored, collagen bundles thickened. Often extensive scarring
Granulation tissue
(New blood vessels and fibroblastic proliferation)
Granulation tissue (top) maturing to Fibroblastic tissue (below)
Mature scar tissue (collagen)
FIBROSIS

End result of organisation in wound healing and chronic inflammation

The process:

• Fibrocytes stimulated by polypeptides from surrounding damaged cells
• Become active fibroblasts. Commence protein synthesis
• Secretion of ground substance including fibrinonectins
• Secretion of procollagen
• Condensation to fine reticulin fibres
• Further condensation to mature collagen fibres
• Binding and weaving to form scar tissue
• Fibroblasts revert to fibrocytes
Factors adversely affecting wound healing

Local
Poor blood supply
Infection
Excessive movement or irritation
Foreign material

General
Deficiency of Vitamin C, essential amino acids, zinc
Excess adrenal corticosteroids
Intercurrent debilitating chronic disease
COMPLICATIONS OF WOUND HEALING

- Infection
- Failure to heal
- Breakdown of wound
- Scarring/Stricture
- Keloid formation
- Pseudoepitheliomatous hyperplasia
- Malignancy
Excessive epithelial proliferation
(pseudoepitheliomatosus hyperplasia)
Complication of wound healing: excess collagen formation (keloid)