

## **OBJECTIVES**

#### At the end of this lecture you should be able to:

- Describe different types and formation of WBC
- Recognize the general functions of WBC
- Describe genesis and site of formation of WBC
- Describe stages of neutrophil formation
- Describe role of neutrophils in defending
- against infection
- Describe the process of phagocytosis.



Note: Macrophages are key components of the innate immunity and activate adaptive immunity by transforming into Antigen Presenting Cells



## WBC TYPES (CLASSIFICATION)

## Granulocytes

- Polymorphonuclear leukocytes (PMNs)
  - Neutrophils (10-16um, 2-5lobes of Nucleus)
  - Eosinophils (12-18um, Bilobed, coarse red granules)
  - Basophils (10-14um, rarely segmented Nucleus hidden by large round bluish granules)

#### Agranulocytes

- Lymphocytes (Round Nucleus, Small [5-8um]& large [9-15um])
  - T lymphocyte
  - B lymphocyte

■ Monocytes (15-20um, Kidney shaped Nucleus)→ make macrophage system

#### **Concentration (Normal Counts)**

Cells	Approximate Normal range (/μL)	Percentage of Total WBC	Life Span
Total WBC	4000-11000		
Granulocytes •Neutrophils • Eosinophils • Basophils	3000-6000 150-300 0-100	50-70 1-4 0.4	4-8 hours in blood and 4-5 days in tissues
Lymphocytes	1500-4000	20-40	Weeks-months
Monocytes (macrophages)	300-600	2-8	10-20 hours (months)
Polymorphonuclear neutrophils Polymorphonuclear eosinophils Polymorphonuclear basophils Monocytes Lymphocytes		62.0% 2.3% 0.4% 5.3% 30.0%	









Fig. 8 - Neutrophil



Fig. 9 - Eosinophil



fig. 10 - Basophil











Fig. 11 - Lymphocyte

#### Macrophage and Neutrophil Responses During Inflammation

- 1<sup>st</sup> line of defense Tissue macrophages & Physical Barriers
- 2<sup>nd</sup> line of defense Neutrophil Invasion of the inflamed area
- <sup>3rd</sup> line of defense Monocytes –macrophage invasion of inflamed area
- 4<sup>th</sup> line of defense Increased production of granulocytes and Monocytes by Bone marrow



## **Genesis of WBC**

- **Two major lineage of WBC are formed:**
- 1. Myelocytic: granular, monocytes
- 2. Lymphocytic: lymphocytes

#### **Sites of WBC Formation**

- Granulocytes (neutrophil, basophil, eosinophil) in bone marrow
- Monocytes- bone marrow
- lymphocytes- bone marrow, thymus, lymphoid tissues







Genesis of white blood cells. The different cells of the myelocyte series are 1, myeloblast; 2, promyelocyte; 3, megakaryocyte; 4, neutrophil myelocyte; 5, young neutrophil metamyelocyte; 6. "band" neutrophil metamyelocyte; 7, polymorphonuclear neutrophil; 8, eosinophil myelocyte; 9, eosinophil metamyelocyte; 10, polymorphonuclear eosinophil; 11, basophil myelocyte; 12, polymorphonuclear basophil; 13–16, stages of monocyte formation.

#### Genesis of white blood cells



## NEUTROPHILS

Most Abundant WBCs 60-70 %
Size: 15-20 µm
Nucleus: Multilobed 2-5 lobes
Life span: 6-8 hours

#### DEFENSIVE PROPERTIES OF MACROPHAGES & NEUTROPHILS

- **1**. Margination
- 2. Diapedesis
- **3.** Chemotaxis
- 4. Opsonization
- **5.** Degranulation
- 6. Phagocytosis & Digestion



### Chemotaxis

- The attraction of the neutrophils to inflamed area following chemotactic substances release from infected site:
- Chemotactic substances:
  - Bacterial toxin
  - Degenerative products of inflamed tissue
  - Complement system
  - Reaction product of plasma
  - clotting



## **Margination & Diapedesis**

WBC marginate along the wall of blood capillaries

WBC squeezes itself through endothelial holes leaving blood capillaries (diapedesis)
WBC move by amoeboid motion towards inflammation area following chemotactic substance released from site of infection
Upon reaching the site of infection neutrophils start to engulf infecting organism



## Phagocytosis

**Selective process: foreign substance recognize by: 1.**Rough surface 2.No protective protein coat, which prevents phagocytosis **3.**Marked by certain substance e.g **Complement 3 or antibodies making them ready for** killing a process known as opsonization. Neutrophils encircled the bacteria with pseudopodia and engulf it inside into a vacuole (phagosome), takes 3-20 bacteria

#### **Opsonization & Phagocytosis**



## PMNs Digestive System (Antimicrobial system)

## **ENZYMATIC** Granules

- Heparin
- Histamine
- Bradykinin
- Serotinin
- Defensins
- Lysosomal enzymes
- Slow reacting substance of anaphylaxis



PMNs Digestive System (Antimicrobial system)

#### **NON ENZYMATIC respiratory burst**

- O2 Free Radicals (O-2, H2O2, -OH)
- NADPH-oxidase
- Myeloperoxidase
- $Cl \rightarrow HoCl$
- Hypochlorous acid "very toxic"

#### **Microbial killing**

- Digestion of organism inside the phagosom
- Fusion of intracellular lysosomes with phagosome vacuole
- Lysosomes discharge its proteolytic enzymes such as yeloperoxidase, catalase into the vacuole, killing and digesting the engulfed bacteria. and or Release of bactericidal such as superoxide, hydrogen peroxide to kill the bacteria





#### Figure 15.2

Phagocytosis by a neutrophil or macrophage. A phagocytic cell extends its pseudopods around the object to be engulfed (such as a bacterium). (Blue dots represent lysosomal enzymes.) (1) If the pseudopods fuse to form a complete food vacuole, lysosomal enzymes are restricted to the organelle formed by the lysosome and food vacuole. (2) If the lysosome fuses with the vacuole before fusion of the pseudopods is complete, lysosomal enzymes are released into the infected area of tissue.

477

# MONOCYTES



**No Granules but Vacoules** -Size: 15-20 μm (active cells 60-80 μm) More Efficient than Neutrophils (100 bacteria vs 3-20 by Neutr, larger particles like RBCs & malarial parasites) Life span: 10-20 hours in blood Two types: Mobile & Fixed Lysosomes contain lipases unlike Neut.

#### **RETICULOENDOTHELIAL SYSTEM**

•Monocytes transform themselves into macrophages in tissue these macrophages are mononuclear cells, & this system of phagocytes is called as Monocye-Macrophage Cell System

 This system of cells was known as reticuloendothelial system although neither they are reticular in appearance nor they have endothelial origin

•Therefore, the term reticuloendothelial system is obsolete.