# Cells and Tissues of the Immune System

#### Aims

 To become familiar with the cells and tissues involved in the immune response

#### Learning outcomes

- List all the cells involved in immunity and identify their key characteristics
- Explain how to distinguish between the different cells
- Describe the structure and function of lymphoid tissue

### Lymphatic System

- The lymphatic system consists of lymphatic vessels and the lymphoid organs.
- This system, which is closely associated with the cardiovascular system
- , has three main functions that contribute to homeostasis
- : (1) lymphatic capillaries take up excess tissue fluid and return it to the blood stream;
- (2) lacteals receive lipoproteins1 at the intestinal villi and transport them to the blood stream and
- (3) the lymphatic system helps defend the
- body against disease.

### The lymphatic system

 Blood components from capillaries enter tissues making extracellular fluid that provides food & gas exchange

• Extracellular fluid drains into vessels called LYMPHATICS VESSELS

• Lymphocytes and other leucocytes use this system to migrate around the body

### Lymph flows one way from a capillary to ever larger

 lymphatic vessels and finally to a lymphatic duct, which enters a sub clavian vein.

## Lymphoid tissues

 Lymphocytes found in blood, but majority are either in discrete clusters or organised in specific tissues

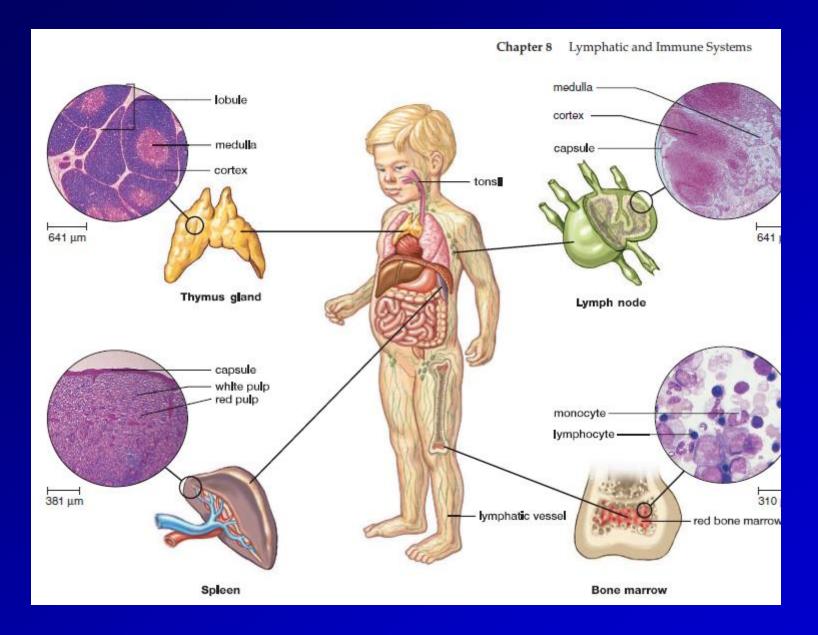
 Components are either primary, secondary or tertiary lymphoid tissues

#### **Primary lymphoid tissues**

 Involved in development & differentiation of lymphocytes & include:

'Bursa equivalent' tissues: foetal liver & adult bone marrow in man produced blood cell & (B cells) maturation

Thymus gland: (T cells) maturation



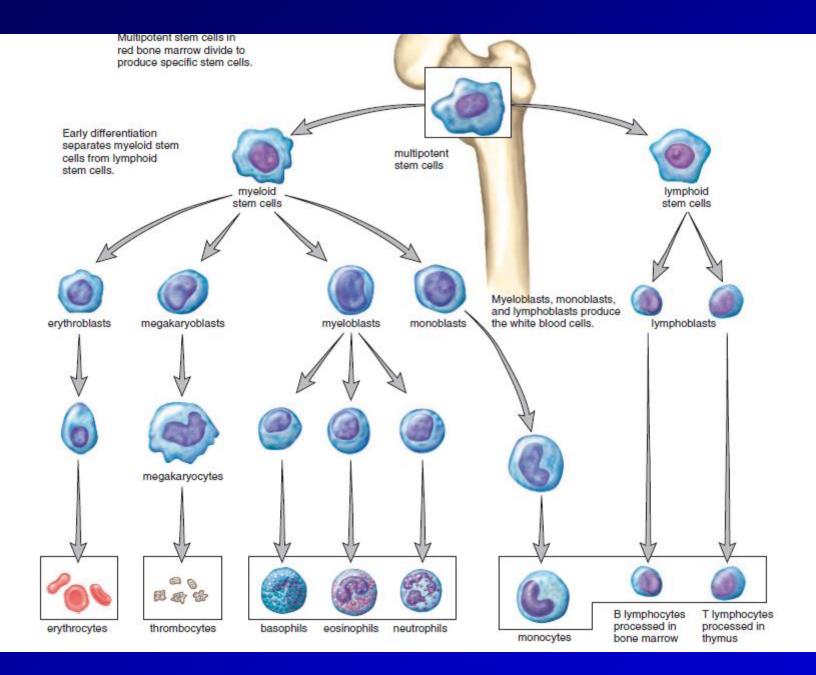
### Bone marrow

- foetal liver & adult bone marrow in man.
- They have stem cells which produce blood cells

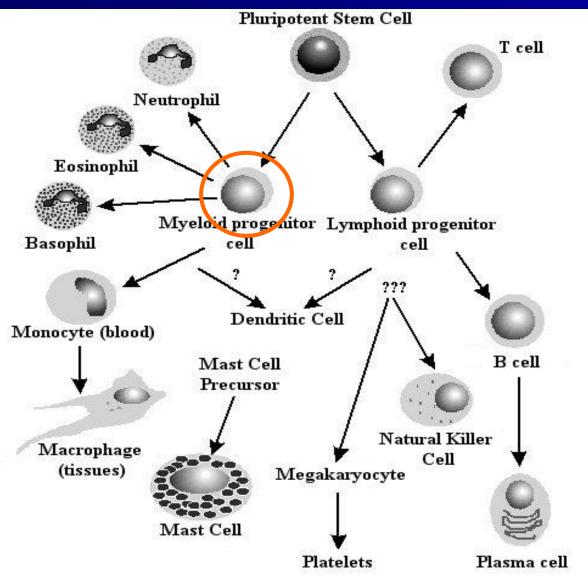
#### **The Red Blood Cells**

- Blood is composed of cells and a fluid containing many inorganic and organic molecules. Red blood cells are very abundant in blood and function in oxygen transport.
- The White Blood Cells There are several types of white blood
- cells, and each type has a specific function in defending the body against disease.
- Platelets are fragments of larger cells that function in blood clotting **Plasma**
- Plasma is over 90% water and contains a variety
- of proteins and other molecules.

FORMED ELEMENTS	Function and Description	Source		PLASMA	Function	Source
Red Blood Cells (erythrocytes)	Transport $O_2$ and help transport $CO_2$	Red bone marrow		Vater 90–92% of plasma)	Maintains blood volume; transports molecules	Absorbed from intestine
4 million–6 million per mm <sup>3</sup> blood	7–8μm in diameter Bright-red to dark-purple biconcave disks without nuclei			Plasma proteins 7–8% of plasma) Albumin	Maintain blood osmotic pressure and pH Maintain blood volume and pressure	Liver
White Blood Cells (leukocytes) 4,000–11,000 per mm <sup>3</sup> blood	Fight infection	Red bone marrow		Globulins Fibrinogen	Transport; fight infection Clotting	
Granular leukocytes • Basophils	10–12 $\mu$ m in diameter Spherical cells with lobed	Æ		Salts (less than 1% of plasma)	Maintain blood osmotic pressure and pH; aid metabolism	Absorbed from intestine
20–50 per mm <sup>3</sup> blood	nuclei; large, irregularly shaped, deep-blue granules in cytoplasm; release histamine which promotes blood flow to injured tissues	Plasma 55% Formed elements 45%		Gases Oxygen Carbon dioxide	Cellular respiration End product of metabolism	Lungs Tissues
• Eosinophils	10–14 μm in diameter Spherical cells with bilobed nuclei; coarse, deep-red, uniformly sized granules in cytoplasm; phagocytize antigen-antibody complexes and allergens		76	Nutrients Lipids Glucose Amino acids	Food for cells	Absorbed from intestine
• Neutrophils	10–14μm in diameter Spherical cells with multilobed nuclei; fine, pink granules in cytoplasm; phagocytize pathogens		ents	Nitrogenous wastes Urea Uric acid	Excretion by kidneys	Liver
er mm <sup>3</sup> blood Agranular leukocytes • Lymphocytes	$5-17\mu$ m in diameter (average 9-10 $\mu$ m)			Other Hormones, vitamins, etc.	Aid metabolism	Varied
1,500–3,000 per mm <sup>3</sup> blood	Spherical cells with large round nuclei; responsible for specific immunity		• with	Wright's stain		
• Monocytes	10–24 $\mu$ m in diameter Large spherical cells with kidney-shaped, round, or lobed nuclei; become macrophages which phagocytize pathogens and cellular debris					
• Platelets (thrombocytes) (thrombocytes) (thrombocytes) (thrombocytes) (thrombocytes) (thrombocytes)	Aid clotting 2–4 $\mu$ m in diameter Disk-shaped cell fragments with no nuclei; purple granules in cytoplasm	Red bone marrow				



### **Cells involved in immunity**



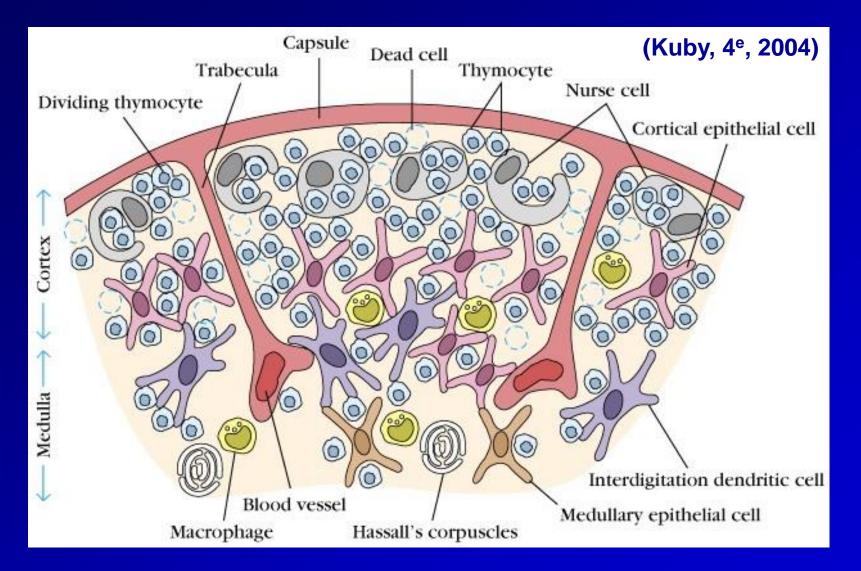
### **The Thymus**

 First organ to received from bone marrow & provides environmen for T cell maturation and education

• 2 lobes divided into lobules by trabeculae (or connective tissue walls)

- Each lobe has outer CORTEX & inner MEDULLA
- Thymocytes are surrounded by EPITHELIAL cells
- In cortex, THYMIC NURSE cells affect thymocyte development
- In medulla & corticomedullary junction where most blood vessels are found there is a special type of dendritic cell: interdigitating cells
- Cells pass from cortex to medulla while differentiating into T cells: Enter expressing neither CD4 or CD8), pass through expressing either CD4 (helper) or CD8 (cytotoxic)

### THE THYMUS



Secondary Lymphoid Tissues 1-Lymph nodes 2-The Spleen 3-Mucosa-associated lymphoid tissue Payer's patches

Secondary lymphoid tissues function

- Allow accumulation of Antigen

 Present Antigen to both naïve & memory lymphocytes

### Secondary Lymphoid Tissues Lymph nodes

 Small, bean-shaped, tissue aggregates at junction of major lymph vessels, 1-25mm diameter, becoming much larger in infection

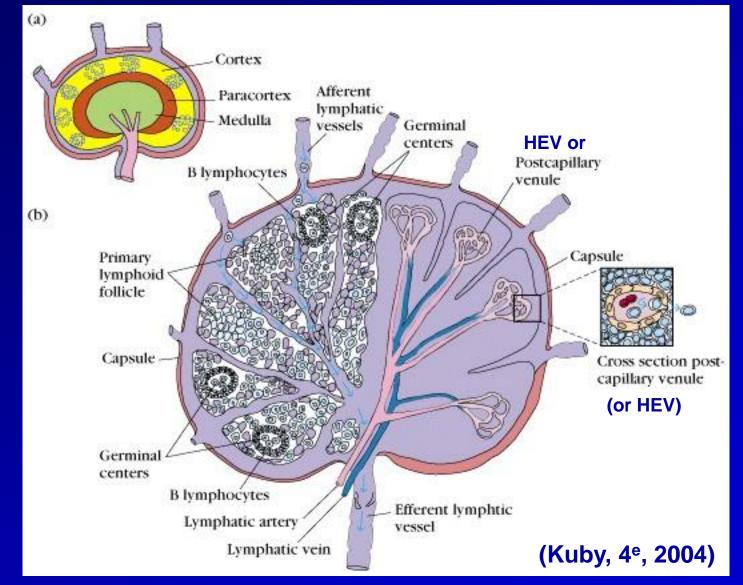
 Surrounded by a capsule & supported by a reticulum, 3 main areas: cortex, para cortical area, and medulla

• The Lymph carries Ag from tissues to lymph nodes allowing cells of the immune system to get together with antigen to start specific immune responses

 Cells flow to medulla & drain into efferent lymphatic at the hilus

 Blood lymphocytes enter lymph nodes at High Endothelial Veinules (HEV) & return via thoracic duct

## Lymph Node



## Lymph nodes

- T cells collect in paracortical areas, B cells in outer cortex
- Paracortical region contains large lymphocytes & activated cells –BLAST cells
- Medulla contains plasma cells secreting Ab
- B cells form dense aggregates –FOLLICLES
- Primary follicles are via dense & uniform centre contains larger cells associated with macrophages – GERMINAL CENTRE
- After Ag exposure, lymph node shows increased turnover of lymphocytes

# **The Spleen**

 lymphoid organ with other non-immunological functions

Capsule with fibrous partitions (SEPTAE)

Lymphocytes enter and leave mainly via blood stream

• 2 types of tissue: **RED & WHITE PULP** 

• Red pulp: filters damaged or aged red cells

## **The Spleen**

#### White pulp:

 cells & tissues surrounding major arterial branches & associated clusters of lymphocytes –

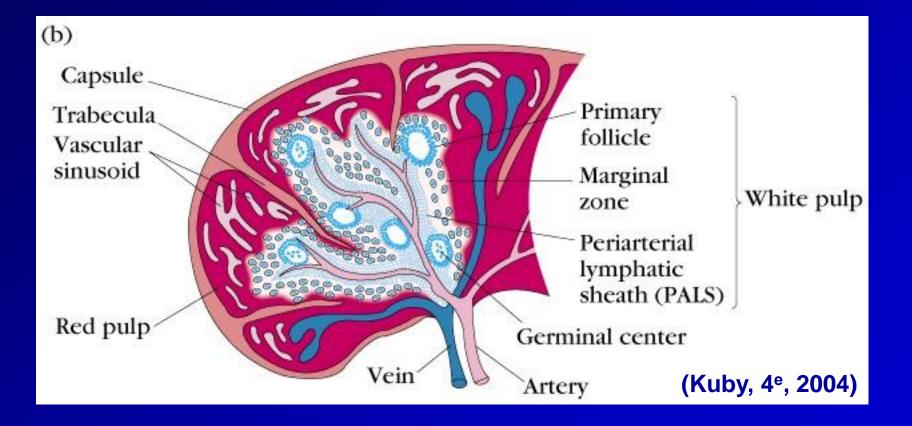
 the lymphatic follicles or( nodules ) :the Follicles similar to those in lymph nodes.

B-dependent area consists of lymphoid follicles

• T-dependent area consists of the peri arterial lymphatic sheath (PALS)

Immunological role: as for lymph nodes with germinal centres developing during immune responses

### **The Spleen**



## Mucosa-associated lymphoid tissue

 Diffusely distributed lymphoid tissues in lining (mucosae) of gastrointestinal, respiratory & urinogenital tracts

Gut-associated lymphoid tissue (GALT) and bronchus

 GALT made up of Peyer's patches and isolated follicles in colonic sub mucosa

 Lymphocytes also found in lamina propria, intestinal epithelium & in lumen of intestine

### **Payer's patches**

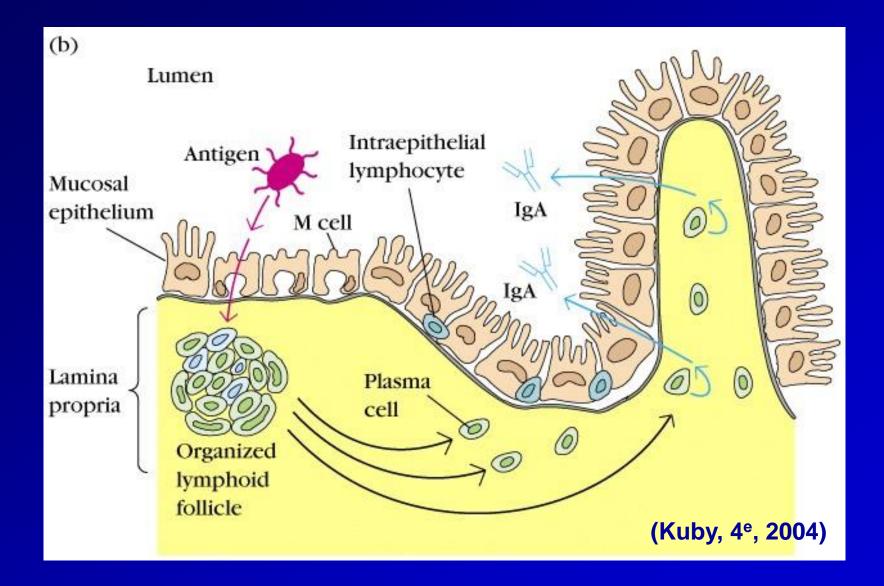
 Aggregates of lymphocytes; B cells form central follicle surrounded by T cells & macrophages

 Have efferent lymphatics that drain into mesenteric lymph nodes, but no afferent lymphatics

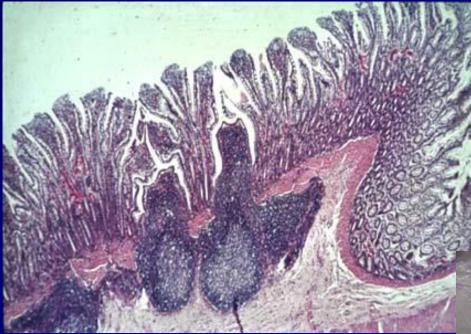
 Covered by specialised lymphoepithelium consisting of cells known as M cells

• Ag in gut enters via M cells that selectively take up particles & deliver them to lymphoid follicles also by specialised dendritic cells which 'reach' into the gut lumen!





# **Peyer's patches**



Peyer's Patch





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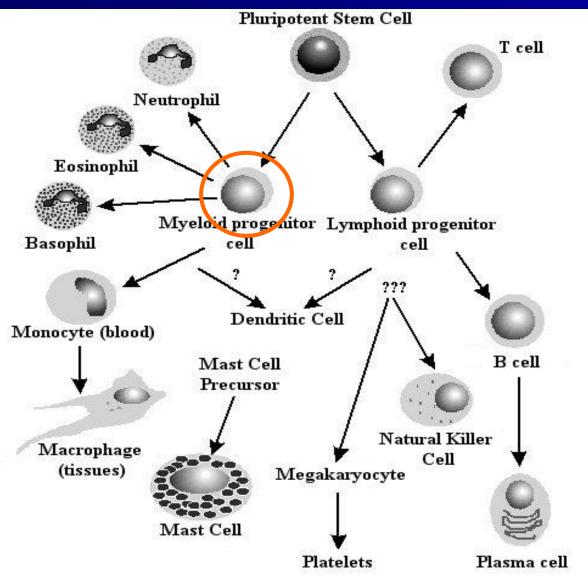
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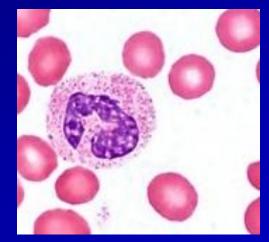
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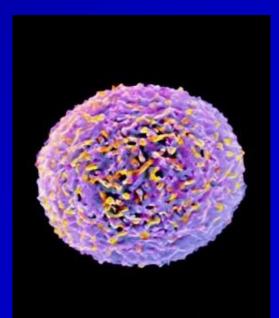
### **Cells involved in immunity**



# Neutrophils

- Large cells (10-20 microns); only live about 2-3 days
- 90% of granulocytes are neutrophils
- Neutral staining cytoplasmic granules containing enzymes
   e.g. lysozyme
- Phagocytic, kill bacteria by microbicidal mechanisms
- Most important cell in non-viral infections



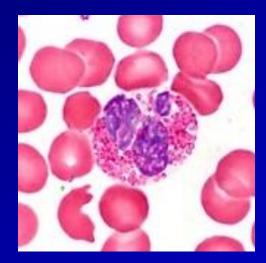


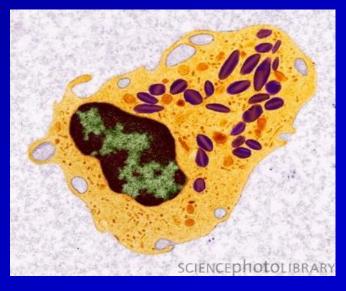
### **Myeloid progenitor cell**

- Gives rise to polymorphonuclear leukocytes and monocytes
- Polymorphs include neutrophils, eosinophils and basophils also called granulocytes (60-70% white blood cells)
- Monocytes are blood borne; in tissues they become macrophages (M $\Phi$ )

## **Eosinophils**

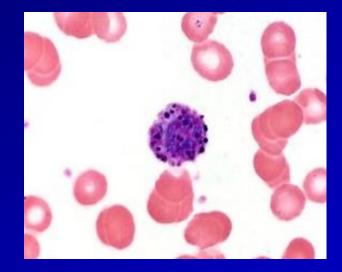
- Contain prominent granules, which stain red with eosin
- Granules contain a crystalline core cytotoxic for parasites; EOSINOPHIL BASIC PROTEIN
- Important in immunity to helminth infections
- Phagocytic, though this is not a major function





### Basophils

- <0.2% white blood cells, only go into tissues during inflammation
- When stimulated, release substances that promote inflammation
- Important in allergy
- Not thought to phagocytose

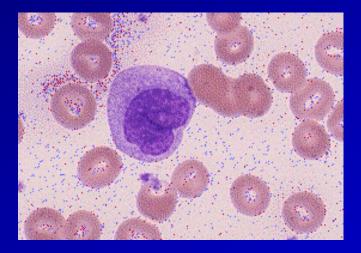




### **Monocytes and macrophages**

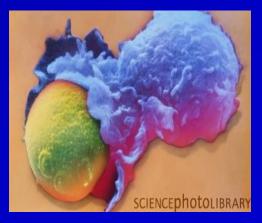
#### **Monocytes**

- in blood 1-2 days
- Mononuclear leukocytes
- Phagocytic



#### **Macrophages**

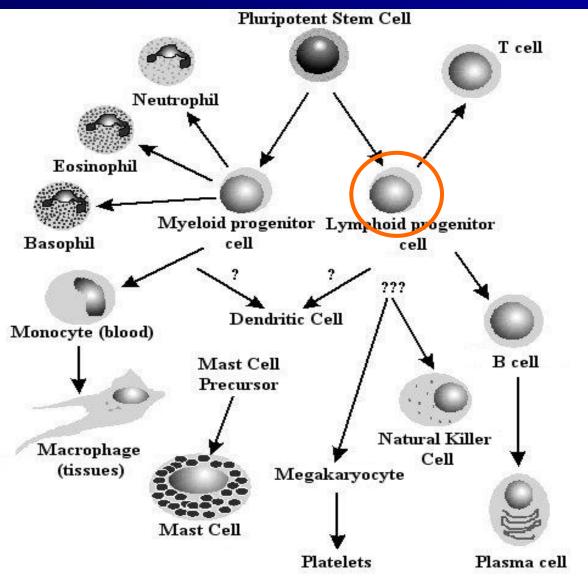
- MCs in tissues = macrophages (MΦ)
- Up to 10x larger than MCs
- can live months or years
- Characteristics of macrophages depend on tissue e.g. Kupffer cells in liver, microglia of brain
- Phagocytic (like neutrophils and eosinophils), adherent



### **Monocytes and macrophages**

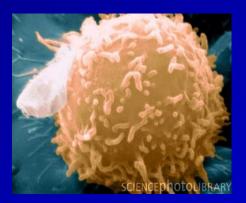
- MCs and MΦs have secrete a huge array of growth factors, chemicals and activation factors
- Can be identified by certain molecules on the surface (CD14 & CD15 on MCs)
- Good at engulfing 'foreign' material and 'showing ' it to other cells of the immune system

### **Cells involved in immunity**



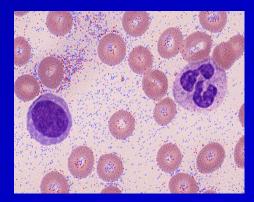
## Lymphoid progenitor cell

- Gives rise to Lymphocytes
- 20-30% peripheral blood white cells
- 6-10um in diameter with large nucleus, small halo of cytoplasm



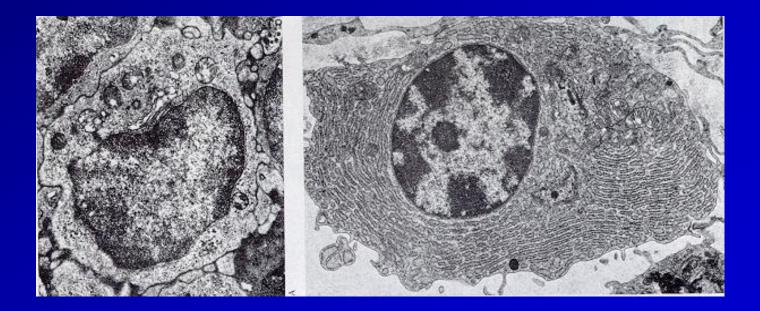
 Upon stimulation by Ag become EFFECTOR CELLS or MEMORY CELLS

2 main types: T cells and B cells (T-lymphocytes and B-lymphocytes)
Early developmental stage, cells pass to Thymus – become T cells or stay in Bone marrow – become B cells



# **B** Lymphocytes (**B** cells)

- Have immunoglobulin/antibody anchored in cell membrane (mlg) to form the B cell antigen receptor (BCR)
- When activated differentiate into Plasma cells and secrete antibody or memory cells

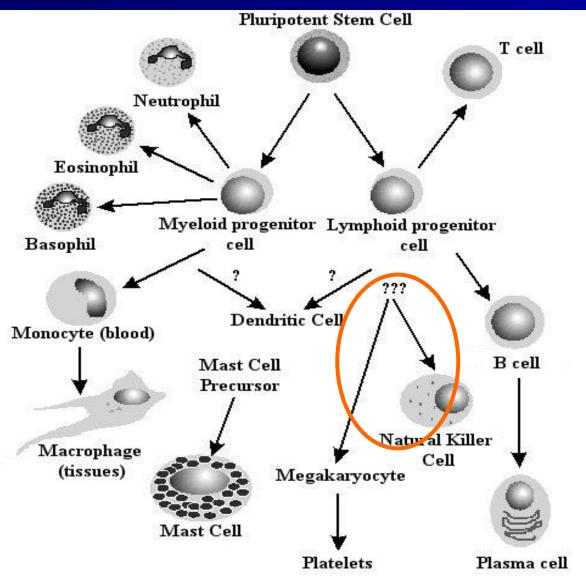


• Have T cell Receptor (TCR) on surface to 'recognise/bind' to foreign material (antigen)
• All T cells express the marker CD3 but there are 2 main sub-groups:
• Cytotoxic T cells (express CD8 and CD3)
• activated to kill infected targets or to become memory cells

Helper T cells (express CD4 and CD3)

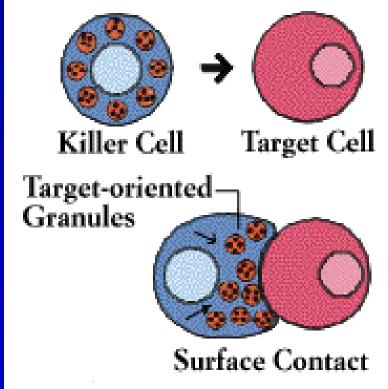
activated to secrete cytokines or to become memory cells

### **Cells involved in immunity**



### Natural Killer cells (Large granular lymphocytes)

- 4% white blood cells
- Collection of cells playing roles between innate & specific immunity
- Lymphocyte-like but larger with granular cytoplasm
- Kill certain tumour & virally infected cells
- Target cell destruction is caused by cytotoxic molecules called granzymes & perforins



### **Megakaryocytes & platelets**

Mature megakaryocytes are polyploid

 Megakaryocytes give rise to platelets by cytoplasmic fragmentation of cell

 Platelets are contractile, adhere to other cells & surfaces & play important roles in blood coagulation

