

l-Farabi Kazakh National University Higher School of Medicine

Blood

Circulating Blood Cells



Figure 10–2 Cells and platelets of circulating blood.

The total volume of **blood** of an average adult is about 5 L, and it circulates throughout the body within the confines of the circulatory system. Blood is a specialized connective tissue composed of formed elements and a fluid component (the extracellular matrix) known as **plasma**.

The formed elements are composed of cells and cell fragments, known as platelets. Light microscopic examination of the formed elements is performed using either the Wright or Giemsa stains, and identification of blood cells is based on the colors produced by these stains.

The cells of blood are subdivided into two major components, **red blood cells** (RBC) and **white blood cells** (WBCs, leukocytes).

Red blood cells lose their nuclei and organelles during development, therefore mature, circulating RBC are anucleated cells whose cytoplasm is filled with hemoglobin.

White blood cells are subdivided into two categories, those without specific granules, agranulocytes and those housing specific granules, granulocytes. Lymphocytes and monocytes belong to the former and neutrophils, eosinophils, and basophils belong to the latter category.

Platelets are round to oval cell fragments derived from megakaryocytes.

For more information see Formed Elements in Chapter 10 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.

Hematocrit





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BLOOD HISTOLOGY RED AND WHITE BLOOD CELLS





Circulating Cells of Blood (cont.)



Figure 10–2 Cells and platelets of circulating blood.

Red blood cells are packed with **hemoglobin**, a large tetrameric protein composed of four polypeptide chains, each of which is covalently bound to an iron-containing **heme.** It is hemoglobin that provides the *unstained* cell with its pale yellow color. The globin moiety of hemoglobin releases CO_2 , and the iron binds to O_2 in regions of high oxygen concentration, as in the lung. However, in oxygen-poor regions, as in tissues, hemoglobin releases O_2 and binds CO_2 . This property of hemoglobin makes it ideal for the conveyance of respiratory gases. Hemoglobin carrying oxygen is known as **oxyhemoglobin**, and hemoglobin carrying carbon dioxide is called **carbaminohemoglobin** (or **carbamylhemoglobin**).

The extracellular surface of the red blood cell plasmalemma has specific inherited carbohydrate chains that act as antigens and determine the blood group of an individual for the purposes of blood transfusion. The most notable of these are the **A** and **B** antigens, which determine the four primary blood groups, **A**, **B**, **AB**, and **O** (Table 10–2).

Another important blood group, the **Rh** group, is so named because it was first identified in rhesus monkeys. Three of the Rh antigens (C, D, and E) are so common in the human population that the erythrocytes of 85% of Americans have one of these antigens on their surface, and these individuals are thus said to be **Rh**⁺.

For more information see Erythrocytes in Chapter 10 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.















Red blood cells look weird when they're squeezing through capillaries.



How Blood Cells Change Shape

Millions of times during their four-month lifespan, human red blood cells must squeeze through tiny capillaries to deliver their payload of oxygen and pick up waste carbon dioxide-functions essential to life.

How Blood Cells Change Shape

Red blood cells have a diameter of about eight microns, or millionths of a meter. As they flow through the body, they often encounter blood vessels, such as those in the brain, with a diameter of only about two microns. Each time the cells reach such a vessel, they must stretch into a bullet-like shape to squeeze through and then return to their original disc shape upon exiting the vessel.

How Blood Cells Change Shape

Do red blood cells change shape?

Human red blood cells rushing through the body to carry oxygen and carbon dioxide to and from the organs are forced to squeeze through smaller and smaller blood vessels. ... To change shape, the cells rearrange protein components of their internal scaffolding, called the cytoskeleton



Red Blood Cells In A Capillary Vessel is a photograph by David M Phillips which was uploaded on June 30th, 2014.

This image shows a healthy red blood cell (left) and a sickle cell (right).



sickle cell anemia



Platelets



Figure 10–10 Platelet ultrastructure. Note that the periphery of the platelet is occupied by actin filaments that encircle the platelet and maintain the discoid morphology of this structure.

Platelets are about 2 to 4 μ m in diameter in blood smears. In light micrographs, they display a peripheral clear region, the **hyalomere**, and a central darker region, the **granulomere**. The platelet plasmalemma has numerous receptor molecules as well as a relatively thick glycocalyx. There are between 250,000 and 400,000 platelets per mm₃ of blood, each with a life span of less than 14 days.

If the endothelial lining of a blood vessel is disrupted and platelets come in contact with the subendothelial collagen, they become **activated**, release the contents of their granules, adhere to the damaged region of the vessel wall **(platelet adhesion)**, and adhere to each other **(platelet aggregation)**. Interactions of tissue factors, plasma-borne factors, and platelet-derived factors form a blood clot.

For more information see Platelets in Chapter 10 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.





Тромбоциттің ультрамикроскопиялық құрылысы. (қан пластинкасы). А- горизонтальды кесінді; Б- көлденең кесінді.

1-гликокаликсы бар платмолеммасы; 2- платмолемма инвагинациямен байланысқан түтікшелердің ашық жүйесі; 3- актин филаменттері; 4- микротүтікшелердің колденең кесіндісі; 5- тығыз тубулин жүйесі; 6- α-түйіршіктері; 7- β-түйіршіктері; 8- митохондриялар; 9- гликоген түйіршіктері; 10- ферритин түйіршіктері; 11- лизосомалар; 12- пероксисомалар (Н.А.Юрина бойынша).

Circulating Cells of Blood (cont.)



Figure 10–2 Cells and platelets of circulating blood.

Neutrophils are the most numerous of the white blood cells, constituting 60% to 70% of the total leukocyte population. The lobes of their multilobed nucleus are connected to each other by slender chromatin threads. In females, the nucleus presents a characteristic small appendage, the "drumstick," which contains the condensed, inactive second X chromosome. but is not always evident in every cell. Neutrophils are among the first cells to appear in acute bacterial infections. They possess very small specific granules.

Eosinophils constitute less than 4% of the total white blood cell population. They are round cells containing large, salmon pink colored specific granules. They have a sausageshaped, bilobed nucleus in which the two lobes are connected by a thin chromatin strand and nuclear envelope. Eosinophils function in parasitic infections and phagocytosing antigen-antibody complexes.

Basophils constitute less than 1% of the total leukocyte population. They are round cells and have an **S**-shaped nucleus, which is commonly masked by the large darkblue to black specific granules present in the cytoplasm. Basophils have several surface receptors on their plasmalemma, including **immunoglobulin E (IgE) receptors.** Their function is very similar to those of mast cells.

For more information see Neutrophils, Eosinophils, Basophils in Chapter 10 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.



Lymphocyte

Monocyte

Many Erythrocytes

1 Platelet

Immature Band Neutrophil

> Mature Segmented Neutrophil

Monocyte -

Lymphocyte

Basophil

Eosinophil

Blood: Histology





















Circulating Cells of Blood (cont.)



Figure 10–2 Cells and platelets of circulating blood.

Monocytes are the largest of the circulating blood cells and constitute 3% to 8% of the leukocyte population. They have a large, acentric, kidney-shaped nucleus whose lobe-like extensions seem to overlap one another. The chromatin network is coarse but not overly dense, their cytoplasm is bluish gray and has numerous azurophilic granules, (lysosomes) and occasional vacuole-like spaces, but no specific granules. Macrophages are avid phagocytes, and as members of the **mononuclear phagocyte system** they phagocytose and destroy dead and defunct cells as well as antigens and foreign particulate matter (such as bacteria). They also have a major role in the immune response

Lymphocytes constitute 20% to 25% of the total circulating leukocyte population. They are round cells, somewhat larger than RBCs, and have a slightly indented, dense, round nucleus that occupies most of the cell. The peripherally situated cytoplasm stains a light blue and contains a few azurophilic but no specific granules. Lymphocytes can be subdivided into three functional categories, namely **B cells, T cells,** and **null cells.** Although morphologically they are indistinguishable from each other, they may be recognized by the differences in their surface markers. They function in the immune response.

For more information see Monocytes and Lymphocytes in Chapter 10 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.






























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HEMATO -POIESIS

Hemapoiesis (Hematopoiesis)

Hemo = Hemato = Blood Poiesis = Production

Hemopoiesis system

- Bone marrow
- Spleen
- Liver
- ThymusLymph node





Haematopoietic Family Photo

Daily Doodles

M. Michiko Maruyama, NMP 2014 mmichiko@interchange.ubc.ca



A GENERAL MODEL OF HEMATOPOIESIS





All blood elements develop from one origin cell – stem cell ,

Monophyletic theory, A. A. Maksimov The term **"stem cell"** Maksimov proposed in 1908.

Classification

Stem cells can be classified into four broad categories, based on their ability to differentiate:

Stem Cells

Totipotent stem cells

are found only in early embryos. Each cell can form a complete organism.

Pluripotent stem cells

Obtained from the inner cell mass of the blastocyst, able to differentiate into almost all cells of the three germ layers – but not into an embryo.

Multipotent stem cells

Found in most tissues, can produce a limited range of differentiated cell lineages appropriate to their location (e.g., Hematopoietic stem cells from the bone marrow). Limited in what the cells can become.

Unipotent cells

capable of generating only one cell type (epidermal stem cells, adult liver stem cells).

Stages of Hemopoiesis

Include:

- Proliferation (reproduction by neoplasms);
- Differentiation (sequential process of mitosis during which cells acquire specific morpho-functional traits)
- Maturation of cells (process when the cell continues to evolve but no longer divided)

Classes of stem cells

By degree of maturity the cells are divided into 6 classes:

- 1 class Stem cells (Pluripotent)
- 2 class half-stem cells (Polipotent)
- 3 class Unipotent cells (give rise to only one type formed elements)
- 4 class Blasts (large cells)
- 5 class Differentiating precursors
- 6 class Mature formed elements circulating in the bloodstream



Immature vs Mature cells

During maturation cells changing its properties.

The immature blast cell:

- Large cell, high Nucleocytoplasmic ratio
- Large nucleus, fine chromatin & nucleoli
- Small amount of dark blue
- cytoplasm



With maturation:

- Cells become smaller
- Nuclear chromatin clumps, nucleoli disappear
- Hgb or granules appear
- N:C ratio decreases









ERYTHROPOIESIS



Immature vs Mature cells

During maturation cells changing its properties.

The immature blast cell:

- Large cell, high Nucleocytoplasmic ratio
- Large nucleus, fine chromatin & nucleoli
- Small amount of dark blue
- cytoplasm



With maturation:

- Cells become smaller
- Nuclear chromatin clumps, nucleoli disappear
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- N:C ratio decreases







ERYTHROPOIESIS



Normoblast (erythroblast) is a type of red blood cell which still retains a cell nucleus. It is the immediate precursor of a normal erythrocyte. Develop into erythrocyte.

Human bone marrow: Normoblasts (b, d, f) differing in sizes and degree of chromatin clumping, late erythroblasts (c). W-G x 400.

Hemopoietic growth factors

A hormone-like substances (protein) that stimulate bone marrow to produce blood cell and promote the proliferation of blood cells.

Erythropoietin (EPO)

- produces primarily by cells in the kidney that lie between the kidney tubules (peritubular interstitial cells)
- Control erythropoiesis red blood cell production
- Required for a myeloid progenitor cell to become erythrocyte

Thrombopoietin (TPO)

- Hormone produced by by the liver
- Stimulates the formation of platelets from megakaryocytes (megakaryocytopoiesis)



•

Bone marrow smear shows developing cells differing in sizes, nuclear shapes, contour and chromatin pattern, cytoplasm for basophilia and presence or absence of granules. Wrights – Giemsa x 200



A myelocyte is a young cell of the granulocytic series, occurring normally in bone marrow, but not circulating in blood (except when caused by some disease). Develop into neutrophil.

Neutrophils are normally found in the bloodstream. During the beginning (acute) phase of inflammation, particularly as a result of bacterial infection, environmental exposure, and some cancers, neutrophils are one of the first-responders of inflammatory cells to migrate towards the site of inflammation.

Human bone marrow:

Neutrophilic myelocyte (h), metamyelocyte (a), and mature (e) and band (g) neutrophil. W-G x 400.



Promyelocyte (or progranulocyte) is a granulocyte precursor, developing from the myeloblast and developing into the myelocyte. In the end of the 4th phase develop into Basophil.

Human bone marrow: Very early neutrophilic promyelocyte (a) is very basophilic & chromatophilic, proerythroblast (c). Wrights-Giemsa x 400



A metamyelocyte is a cell undergoing granulopoiesis, derived from a myelocyte, and leading to a band cell. Develop into eosinophil, neutrophil or basophil. Eosinophils are white blood cells and one of the immune system components responsible for combating multicellular parasites and certain infections in vertebrates. Lymphocytes are cellular components of the adaptive immune response.

Human bone marrow: Neutrophilic metamyelocytes (f, g), mature eosinophil (h), neutrophilic myelocyte (i), eosionophilic myelocyte (j), lymphocyte (k). W-Giemsa x 400



A metamyelocyte is a cell undergoing granulopoiesis, derived from a myelocyte, and leading to a band cell. Develop into eosinophil, basophil or neutrophil.

Human bone marrow: Band neutrophil (b), neutrophilic metamyelocytes (e), late neutrophilic myelocyte (d), normoblast (f). Wrights-Giemsa x 400.





Human bone marrow: A megakaryocyte shedding its cytoplasm to form platelets (arrow). W-G x 400.






Bone marrow Soft tissue within bones where blood cells are formed. Bone marrow samples are obtained from the back of the hipbone using a needle.

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RESPIRATORY SYSTEM

Olfactory Epithelium

Figure 15–2 The olfactory epithelium, displaying basal, olfactory, and sustentacular cells.

The respiratory system has two major portions, the **conducting portion**, situated both outside and within the lungs, conveys air from the external milieu to the lungs and the **respiratory portion**, located strictly within the lungs, functions in the actual exchange of oxygen for carbon dioxide (external respiration).

The roof of the nasal cavity, the superior aspect of the nasal septum, and the superior concha are covered by an **olfactory epithelium**. The underlying lamina propria houses serous fluid–secreting Bowman's glands, a rich vascular plexus, and collections of axons that arise from the olfactory cells of the **olfactory epithelium**. The olfactory epithelium is composed of three types of cells: olfactory, sustentacular, and basal cells.

Olfactory cells are bipolar neurons whose apical aspect, the distal terminus of its slender dendrite, is modified to form a bulb, the **olfactory vesicle**, which projects above the surface of the sustentacular cells. Six to eight long, nonmotile olfactory cilia extend from the olfactory vesicle and lie on the free surface of the epithelium. The basal region of the olfactory cell is its **axon**, which penetrates the basal lamina and joins similar axons to form bundles of nerve fibers that synapse with secondary neurons in the olfactory bulb.

The tall, columnar **sustentacular cells** have secretory granules housing a yellow pigment characteristic of the color of the olfactory mucosa. These cells are believed to provide physical support, nourishment, and electrical insulation for the olfactory cells.

Basal cells have considerable proliferative capacity and can replace both sustentacular and olfactory cells. In a healthy person, the olfactory and sustentacular cells have a life span of less than a year.

For more information see Olfactory Epithelium in Chapter 14 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.

TRACHEA

Кон қатарлы кірпікшелі энитегий. А-баңғы: генатоқсалақ-зазақ ж 190; Б-улсігі;

1-жасуны кірнікцалері; 2-бокал тәріхлі беллі жасуналар; 3-кірнікшелі жасуналар; 4-ұлык кыстырма жасуналар; 5-кысқа кыстырма жасуналар; 6-базальды мембрана; 7-дәнекер тіні (И.В. Алмазов, Л.С. Сутузов бойынна).

Рис. 296. Эпителиальные клетки слизистой оболочки воздухоносных путей (схема по Ю.И.Афанасьеву).

реснитчатые клетки; 2 — нейроэндокринные клетки; 3 — бокаловидные клетки; 4 — камбиальные клетки; 5 — безреснитчатые клетки; 6 — нервное волокно; 7 — секреторные клетки (клетки Клара); 8 — базальная мембрана; 9 — хемочувствительные клетки.

Basal cells

Brush cells

Endocrine (small dranule) cells

Экзокринді, экзоэпительазды бездердің құрылысы мен секрециясы. Өзектер мен эпителидің устіңгі беткейі кулгіп-кок түске, секреторлық бөлімдері қызыл түске боялган. А- эпителий, Б- дәнекер тін.

 карапайым тармақталған түтікше безі, 2- карапайым тармақталмаған альвеолалы без, 3- секреторлық бөлімі тармақталған қарапайым түтікше безі, 4- секреторлық бөлімі тарамдалған қарапайым альвеолалы без, 5күрделі альвеолалы түтікше безі; 6- мерокринді секреция; 7- апокринді секреция; 8- голокринді секреция; абазальді клеткалар; 6- тіршілігін жойған клеткалар; в- ыдырау сатысындағы клетка

AIRWAYS- BRONCHI

•large bronchi 5 - 15 мм middle size bronchi 2 - 5 MM •small bronchi -**1 - 2 MM** terminal bronchioles **0,5** MM

Bronchi and Bronchioles

Figure 15–7 The respiratory system, displaying bronchioles, terminal bronchioles, respiratory bronchioles, alveolar ducts, alveolar pores, and alveoli.

The **bronchial tree** (conducting portion) begins at the bifurcation of the trachea, as the right and left primary bronchi, which arborize. The bronchial tree is composed of airways located outside the lungs, the primary bronchi, and airways located inside the lungs, the intrapulmonary bronchi, bronchioles, and terminal bronchioles.

Primary bronchi are identical to the trachea, except that bronchi are smaller in diameter and their walls are thinner.

Intrapulmonary bronchi are similar to primary bronchi, except that the cartilage C-rings are replaced by irregular plates of hyaline cartilage that completely surround the lumina of the intrapulmonary bronchi. The smooth muscle is located at the interface of the fibroelastic lamina propria and submucosa as two distinct smooth muscle layers spiraling in opposite directions. Elastic fibers, radiate from the adventitia to connect with elastic fibers arising from other parts of the bronchial tree.

Each **bronchiole** supplies air to a pulmonary lobule. Their epithelial lining ranges from ciliated simple columnar with occasional goblet cells in larger bronchioles to simple cuboidal (many with cilia) with occasional **Clara cells** and no goblet cells in smaller bronchioles.

Terminal bronchioles are lined by Clara cells and cuboidal cells. The thin lamina propria consists of fibroelastic connective tissue and is surrounded by one or two layers of smooth muscle cells. Elastic fibers radiate from the adventitia and bind to elastic fibers radiating from other members of the bronchial tree.

For more information see Bronchial Tree in Chapter 14 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.

TERMINAL BRONCHIOLES

Respiratory Portion

Figure 15–11 A, A respiratory bronchiole, alveolar sac, alveolar pore, and alveoli. **B**, Interalveolar septum. **C**, Carbon dioxide uptake from body tissues by erythrocytes and plasma. **D**, Carbon dioxide release by erythrocytes and plasma in the lung. (Compare **A** with the alveolar duct shown in Fig. 15-10.)

The **respiratory portion** of the respiratory system is composed of respiratory bronchioles, alveolar ducts, alveolar sacs, and alveoli.

Respiratory bronchioles are similar to terminal bronchioles, but their wall is interrupted by **alveoli**, where gaseous exchange (O_2 for CO_2) can occur. Subsequent to several branchings, each respiratory bronchiole terminates in an alveolar duct

Alveolar ducts do not have walls of their own. Each alveolar duct usually ends as a blind outpouching composed of two or more small clusters of alveoli, in which each cluster is known as an **alveolar sac.** These alveolar sacs thus open into a common space, which some investigators call the **atrium.**

Alveoli are small air sacs composed of highly attenuated type I pneumocytes and larger type II pneumocytes.

The region between adjacent alveoli is known as the **interalveolar septum.** It is occupied by an extensive capillary bed composed of **continuous capillaries.**

The thinnest regions of the interalveolar septum where gases can be exchanged are called the **blood-gas barriers** The narrowest blood-gas barrier, where the type I pneumocyte is in intimate contact with the endothelial lining of the capillary and the basal laminae of the two epithelia become fused, is most efficient for the exchange of O_2 (in the alveolar lumen) for CO_2 (in the blood). These regions are composed of surfactant (manufactured by type II pneumocytes), type I pneumocytes, basal lamina, endothelial cells.

For more information see Respiratory Portion of the Respiratory System in Chapter 14 of Gartner and Hiatt: Color Textbook of Histology, 3rd ed. Philadelphia, W.B. Saunders, 2007.

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URINARY SYSTEM









Fig. 15-3. Drawing of a "typical" nephron.



Fig. 15-2. A cutaway drawing of a nephron, straightened out and shortened, that shows the relative positions of the parts of a nephron along its length.





















d Cross-section of renal medulla

Proximal and Distal Convoluted tubules



Collecting Ducts









Ureters



Urinary Bladder



Urinary Bladder

