

What Does My Bone Marrow Do?

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Table of Contents

What is Bone Marrow?	4
Stem Cells	4
Importance of the Circulatory System	10
Hemoglobin	10
Iron	12
Red Blood Cells	12
White Blood Cells	15
<i>Lymphocytes</i>	15
<i>Monocytes</i>	15
<i>Granulocytes</i>	15
<i>Neutrophils</i>	16
<i>Eosinophils</i>	16
<i>Basophils</i>	16
Platelets	17

How Does MDS Affect My Bone Marrow?	18
Effect on Red Blood Cells - Low Red Cell Count (Anemia)	19
Effect on White Blood Cells - Low White Cell Count (Neutropenia)	20
Effect on Platelets - Low Platelet Count (Thrombocytopenia)	20
Bone Marrow Examination	21
The Bone Marrow Aspirate	21
The Bone Marrow Biopsy	21
Processing the Sample	22
The Bone Marrow Biopsy Procedure	22
For More Information on MDS	24

What is Bone Marrow?

Bone marrow is a nutrient-rich spongy tissue located mainly in the hollow portions of long flat bones like the sternum and the bones of the hips. There are two types of bone marrow: red marrow and yellow marrow. Yellow marrow has a much higher amount of fat cells than red marrow. Both types of marrow contain blood vessels.

Stem Cells

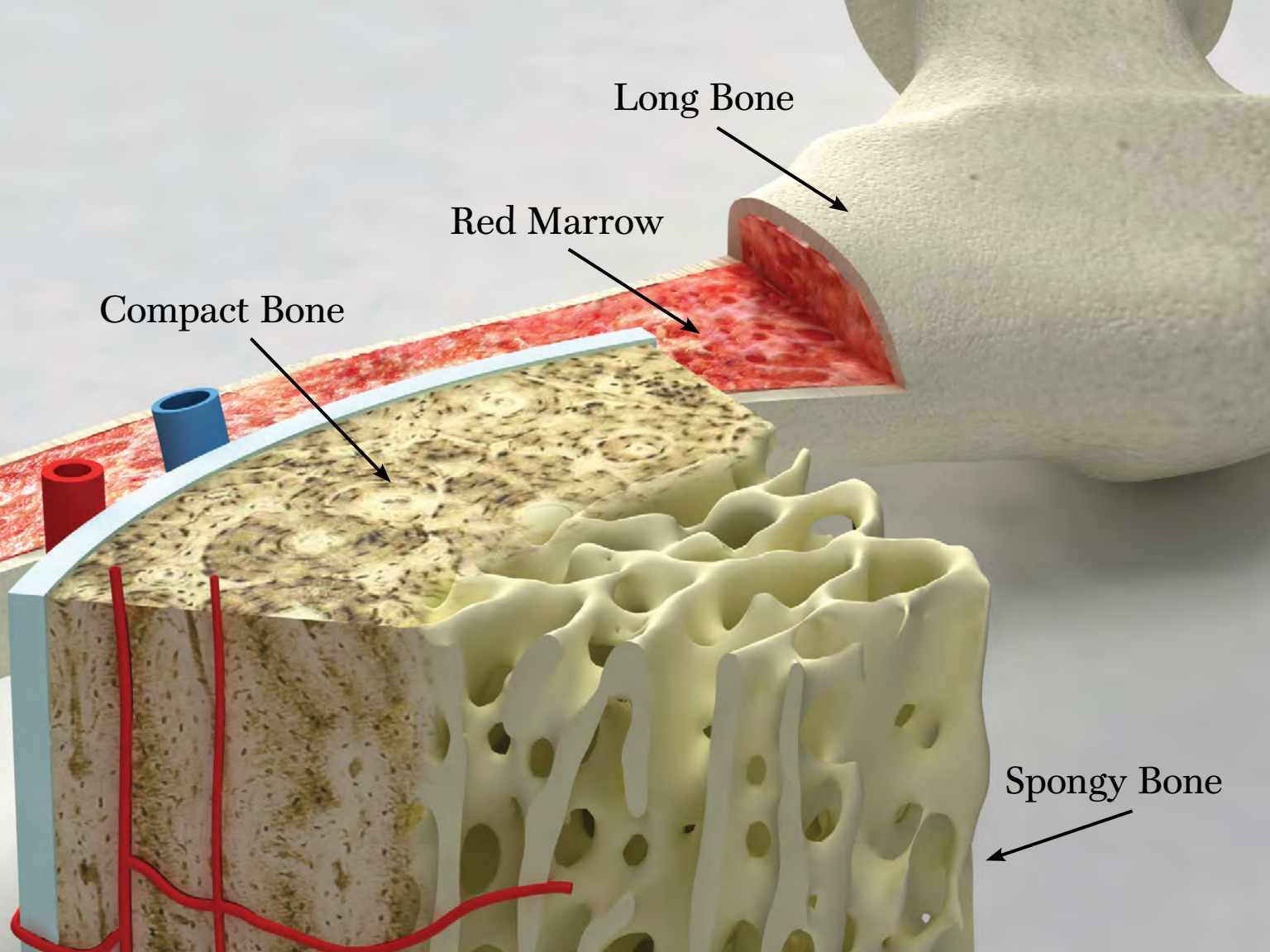
The bone marrow works like a 'factory' that produces all of the cells that are found in the bone marrow and in the peripheral blood stream. This factory is dependent on the function of the pluripotent stem cells. *Pluripotent* refers to the ability of a cell to become many different types of cells.

Did You Know?

At birth, all bone marrow is red. As we age, more and more of the marrow converts to yellow bone marrow. In adults, about half of the bone marrow is red and half is yellow.



Pluripotential is derived from the Latin *pluri* meaning more and *potential* meaning power.



Long Bone

Red Marrow

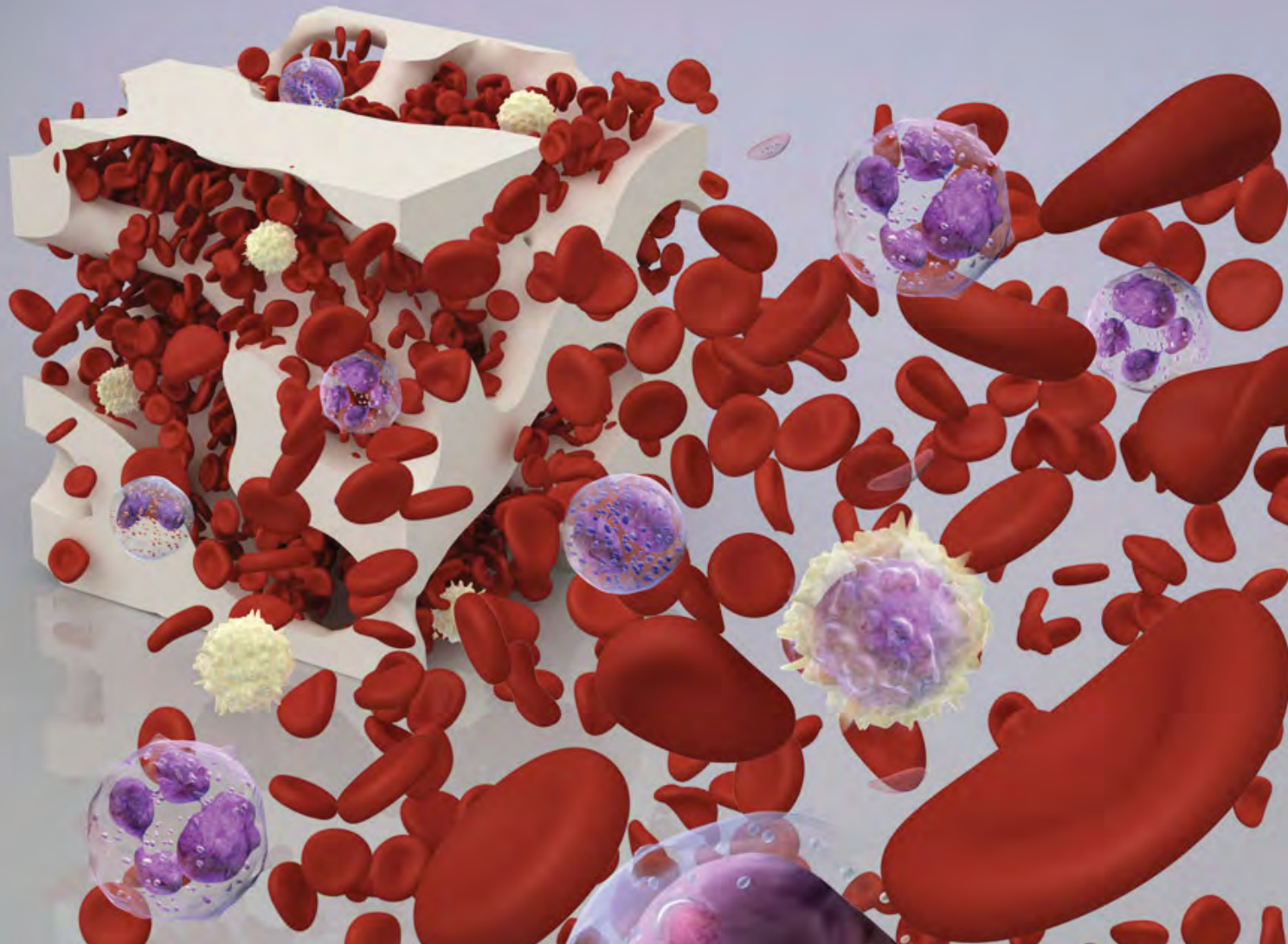
Compact Bone

Spongy Bone

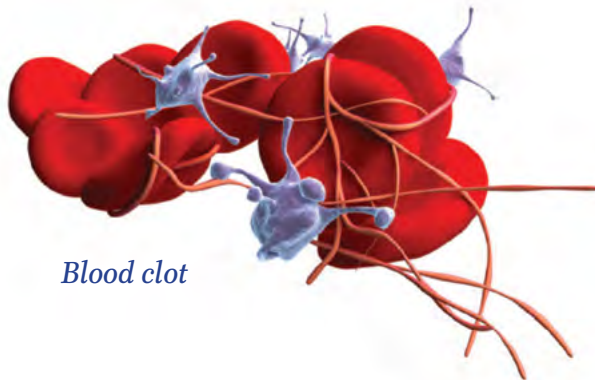
The bone marrow has two types of stem cells, *mesenchymal* and *hematopoietic*. This process of development of different blood cells from these pluripotent stem cells is known as hematopoiesis. Pluripotent hematopoietic cells can become any type of cell in the blood system. Under the influence of tissue and hormonal factors these cells develop into specific blood cell lines. When these cells differentiate or mature they become the cells that we can recognize in the blood stream.

Mesenchymal is embryonic tissue from which the connective tissue, blood vessels and lymphatic vessels are formed.

Hematopoietic is the formation and development of blood cells in the bone marrow.



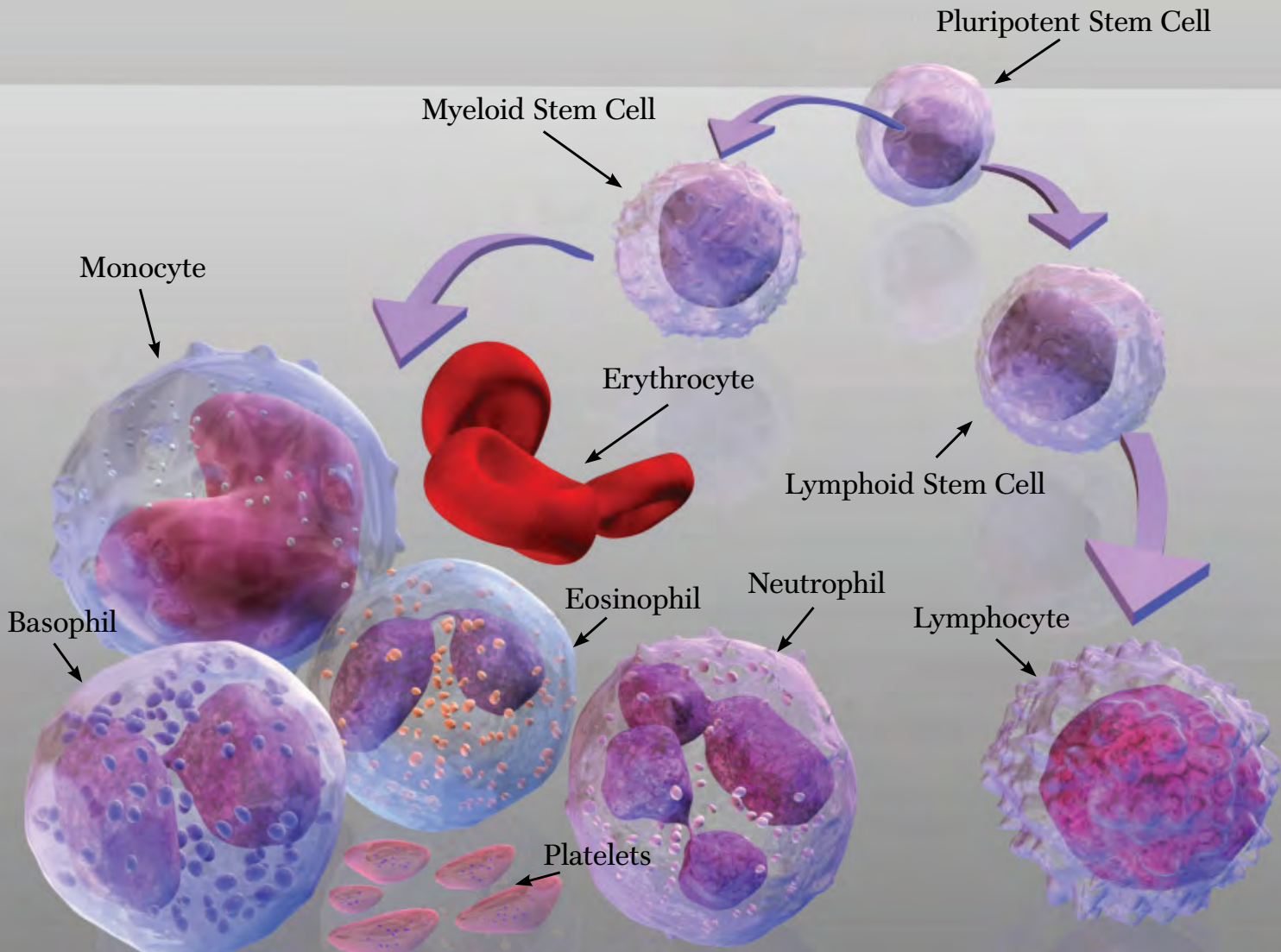
These include the erythroid or red blood cells (RBCs). RBCs are responsible for carrying oxygen from the lungs to all parts of the body. White blood cells (WBCs) include lymphocytes, the cornerstone of the immune system and myeloid cells which include granulocytes: neutrophils, monocytes, eosinophils, and basophils. WBCs fight infection by attacking and destroying bacteria or viruses and granulocytes are involved in a variety of immune processes. Platelets are fragments of the cytoplasm of megakaryocytes, another bone marrow cell.



Blood clot

Did You Know?

Platelet cells (platelets) control bleeding by forming blood clots when your body is injured.



The majority of RBCs, platelets, and most of the WBCs are formed in the red marrow while only a few of them are formed in the yellow marrow. Everyone needs a continuous production cycle of blood cells from our bone marrow throughout our lives since each blood cell has a set life expectancy. Healthy bone marrow produces as many cells as your body needs. Red cell production is increased when the body needs additional oxygen, platelets increase when bleeding occurs, and white cells increase when infection threatens.

Importance of the Circulatory System

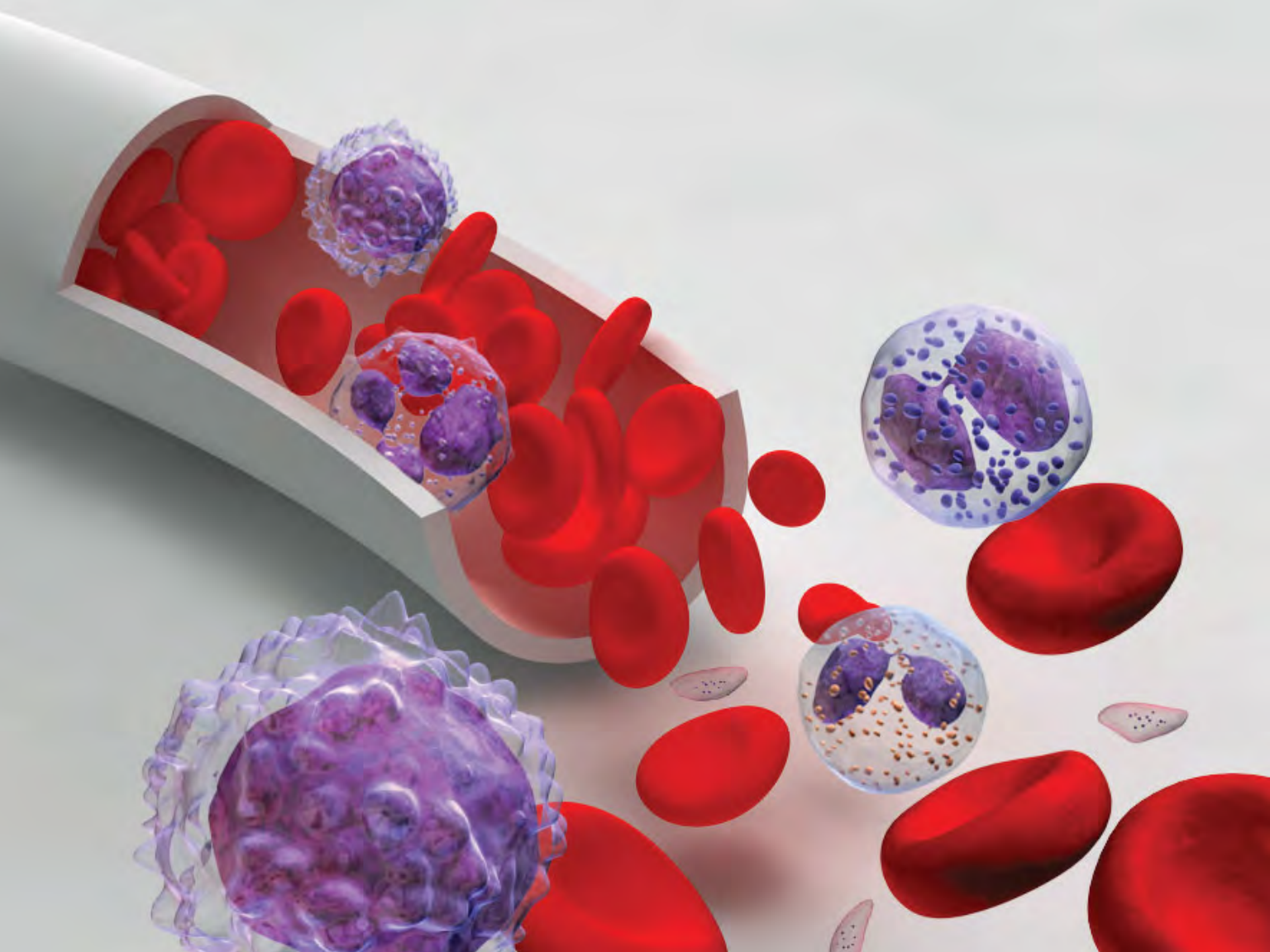
The circulatory system touches every organ and system in your body. Red cells flow in the blood circulation to transport oxygen. Every cell needs access to the circulatory system in order to function since oxygen is essential for proper cell function.

Hemoglobin

Hemoglobin (Hgb) is a protein that is found within red blood cells. This protein is what makes 'red cells' red. Hemoglobin's job is to pick up oxygen in the lungs, carry it in the RBCs, and then release oxygen to the tissues that need it like the heart, muscles, and brain. Hemoglobin also removes CO₂ or carbon dioxide and carries this waste product back to the lungs where it is exhaled.

Did You Know?

Red blood cells live an average of 120 days and platelets live 8 – 10 days. Some white blood cells are very short lived and live only hours, while others can live for many years.



Iron

Iron is an important nutrient in the body. It combines with protein to make the hemoglobin in red blood cells and is essential in the production of red blood cells (erythropoiesis). The body stores iron in the liver, spleen, and bone marrow. The storage form of iron is known as ferritin and ferritin can be measured through a blood test. Most of the iron needed each day for making hemoglobin comes from the recycling of old red blood cells.



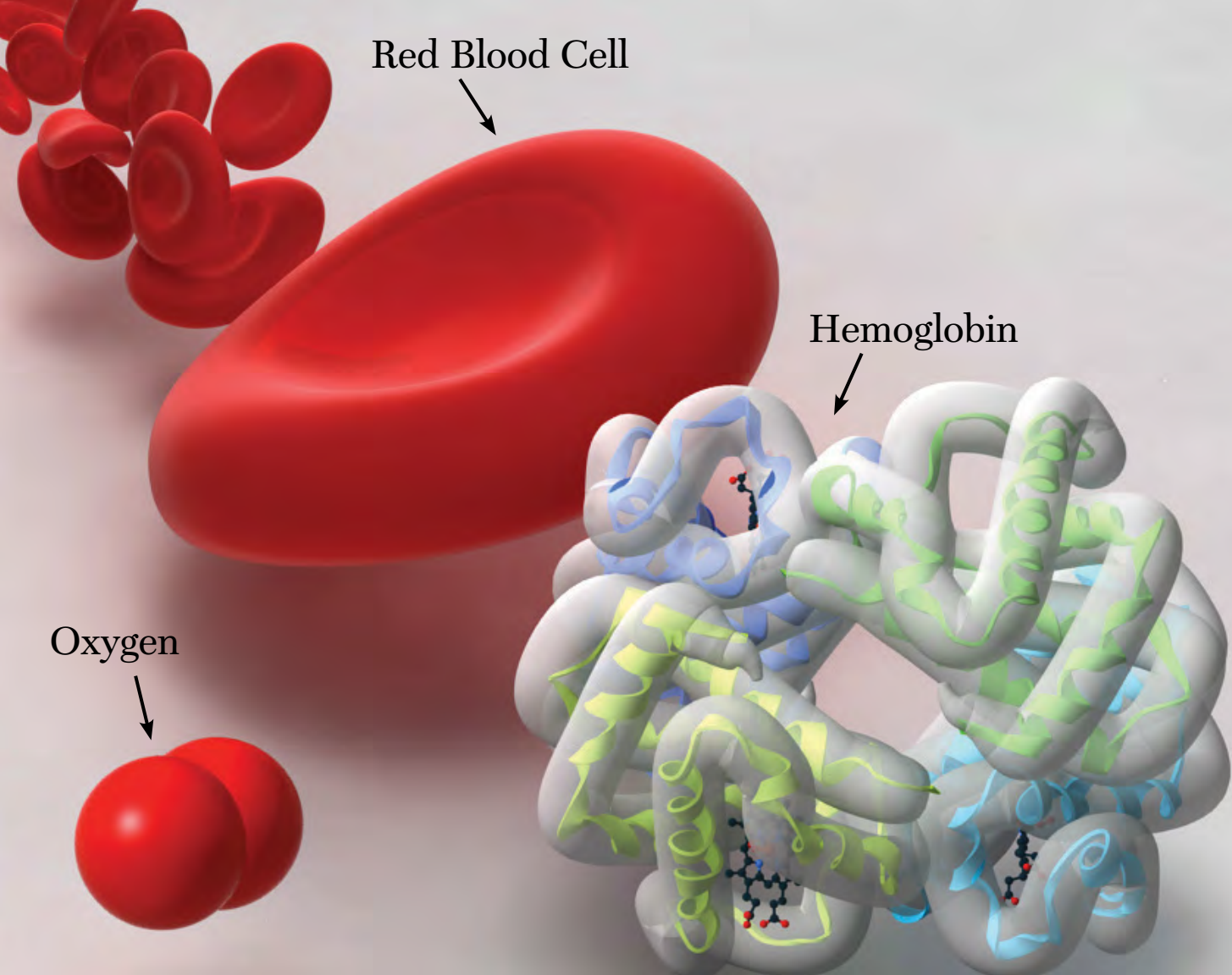
Red Blood Cells

The production of red blood cells is called erythropoiesis. It takes about 7 days for a committed stem cell to mature into a fully functional red blood cell. Red blood cells have a limited life span of approximately 120 days and must be continuously replaced by the body.

Did You Know?

The body has no active way to excrete unwanted iron, so little iron is lost from the body naturally.

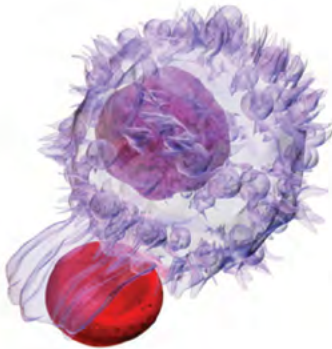
Erythropoiesis is stimulated by a lack of oxygen (hypoxia) in the body. This lack of oxygen tells the kidneys to produce a hormone, erythropoietin (EPO). EPO then stimulates the bone marrow to produce RBCs. Erythropoietin does this by entering the blood stream and traveling throughout the body. All the body's cells are exposed to erythropoietin, but only red bone marrow cells respond to this hormone. As these new red cells are produced they move into the blood stream and increase the oxygen-carrying ability of the blood. When the tissues of the body sense that oxygen levels are enough, they tell the kidneys to slow the secretion of



erythropoietin. This 'feedback' within your body ensures that the number of RBCs remains fairly constant and that enough oxygen is always available to meet the needs of your body.

As RBCs age, they become less active and become more fragile. The aging red cells are removed or eaten up by white blood cells (*macrophages*) in a process known as phagocytosis and the contents of these cells are released into the blood. The iron from the hemoglobin of the destroyed cells is carried by the blood stream to either the bone marrow for production of new RBCs or to the liver or other tissues for storage.

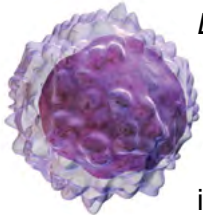
Normally, a little less than 1% of the body's total RBCs are replaced every day. The number of red blood cells produced each day, in the healthy person, is about 200 billion cells.



Macrophage is derived from the Ancient Greek: 'macro' meaning big and 'phage' meaning eat.

White Blood Cells

The bone marrow produces many types of white blood cells, which are necessary for a healthy immune system. These cells both prevent and fight infections. There are five main types of white blood cells, or leukocytes:



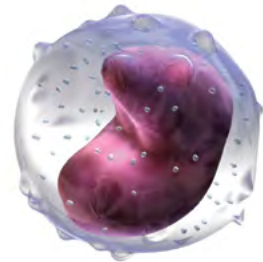
Lymphocytes

Lymphocytes are produced in bone marrow. They make natural antibodies to fight infection caused by viruses that enter your body through your nose, mouth, or cuts.

They do this by recognizing foreign substances that enter the body and then sending a signal to other cells to attack those substances. The number of lymphocytes increases in response to these invasions. There are two major types; B- and T-lymphocytes.

Monocytes

Monocytes are also produced in the bone marrow. Mature monocytes have a life expectancy in the blood of only 3-8 hours, but when they move into the tissues, they mature into larger cells called macrophages. Macrophages can survive in the tissues for long periods of time where they engulf and destroy bacteria, some fungi, dead cells, and other material foreign to the body.



Granulocytes

Granulocyte is the family or collective name given to three types of white blood cells: neutrophils, eosinophils and basophils. The development of a granulocyte may take two weeks, but this time

is shortened when there is an increased threat like a bacterial infection. The marrow also stores a large reserve of mature granulocytes. For every granulocyte circulating within the blood, there may be 50 to 100 cells waiting in the marrow to be released into the blood stream. As a result, half the granulocytes in the blood stream can be available to actively fight an infection in your body within 7 hours of recognizing that an infection exists! Once a granulocyte has left the blood it does not return. It may survive in the tissues for as long as 4 or 5 days depending on the conditions it finds, but it only survives for a few hours in the circulation.

Neutrophils

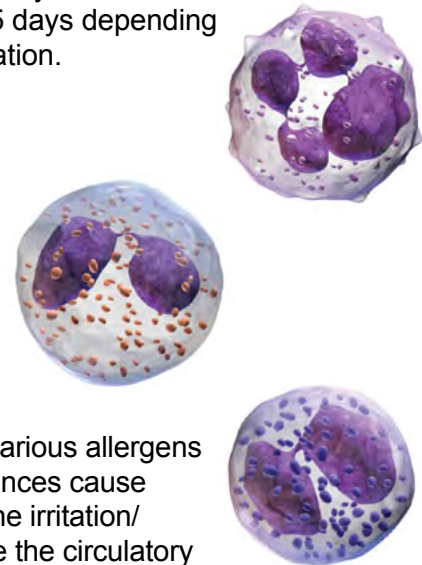
Neutrophils are the most common granulocyte. They can attack and destroy bacteria and viruses.

Eosinophils

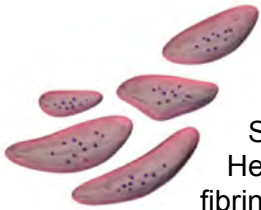
Eosinophils are involved in the fight against many types of parasitic infections and against the larvae of parasitic worms and other organisms. They are also involved in some allergic reactions.

Basophils

Basophils are the least common of the white blood cells and respond to various allergens that cause the release of histamines and other substances. These substances cause irritation and inflammation in the affected tissues. Your body recognizes the irritation/inflammation and widens (dilates) the blood vessels allowing fluid to leave the circulatory system and enter the tissue in an effort to dilute the irritant. This reaction is seen in hay fever, some forms of asthma, hives, and in its most serious form, anaphylactic shock.



Platelets



Platelets are produced in bone marrow by a process known as thrombopoiesis. Platelets are critical to blood coagulation and the formation of clots to stop bleeding.

Sudden blood loss triggers platelet activity at the site of an injury or wound. Here the platelets clump together and combine with other substances to form fibrin. Fibrin has a thread-like structure and forms an external scab or clot. Platelet deficiency causes you to bruise and bleed more easily. Blood may not clot well at an open wound, and there may be a greater risk for internal bleeding if the platelet count is very low.

Did You Know?

Healthy bone marrow normally manufactures between 150,000 and 450,000 platelets per microliter of blood, an amount of blood that fits on the head of a pin.

How Does MDS Affect My Bone Marrow?

In people with Myelodysplastic Syndromes (MDS) the bone marrow cannot produce enough healthy blood cells. It may affect only one of the cell lines or it may affect all three cell lines produced in the bone marrow. RBCs, WBCs, and platelets may not mature and all or some of them may not be released into the blood stream but accumulate in the bone marrow. These cells may have a shortened life span, resulting in fewer than normal mature blood cells in circulation. The cells may actually die in the bone marrow before they mature. This results in a higher than normal number of immature cells or blasts in the bone marrow and fewer than normal mature blood cells in the circulation. Low blood cell counts in any of these three cell lines (red cells, white cells, or platelets) are the hallmark feature of MDS. Low blood counts are responsible for some of the problems that MDS patients experience such as infection, anemia, easy bruising, or an increased chance of bleeding.

In addition to the lower number of blood cells in the circulation, the cells may be dysplastic. The formal definition of dysplasia is an abnormal shape and appearance (morphology) of a cell. The prefix myelo- is from the Greek and it means marrow. So myelodysplasia simply means that the mature blood cells found in the bone marrow or circulating in the blood 'look funny'. Dysplastic cells cannot function properly. In addition to the dysplasia, 50% of patients have an increase in very immature cells called "blasts".

Effect on Red Blood Cells

Low Red Cell Count (Anemia)

The bone marrow normally produces mature red blood cells and the hemoglobin in these cells carry oxygen to the tissues in your body. The percentage of red blood cells in the total blood volume is called the hematocrit. In healthy women the hematocrit is 36% to 46% while in healthy men the hematocrit is 40% to 52%. When the hematocrit falls below the normal range there are an insufficient number of healthy, mature red blood cells to effectively supply oxygen to all tissues of the body. This condition of below-normal numbers of red blood cells, low hemoglobin levels and low oxygen is called anemia, which can be relatively mild (hematocrit of 30% to 35%), moderate (25% to 30%), or severe (less than 25%). Anemia can also result from the inefficient transport of oxygen by dysplastic (mature but misshapen) red blood cells.



Healthy, mature red blood cells



Abnormal, ("dysplastic") red blood cells

Effect on White Blood Cells

Low White Cell Count (Neutropenia)

The bone marrow normally makes between 4,000 and 10,000 white blood cells per microliter of blood; in African-Americans the range is lower, between 3,200 and 9,000 white cells per microliter.

Some MDS patients develop neutropenia or a low white cell count. MDS patients with neutropenia usually have too few neutrophils. With fewer numbers of neutrophils, the risk of contracting bacterial infections such as pneumonia and urinary tract infections increases. Fever may accompany these infections. Sometimes infections occur despite adequate numbers of neutrophils because the WBCs are not able to function as well as they do in a person without MDS.

Effect on Platelets

Low Platelet Cell Count (Thrombocytopenia)

MDS can also cause a low platelet count, or thrombocytopenia. People with abnormal or low platelet counts may suffer from bruising or bleeding even after minor bumps, scrapes, or cuts.

Severe thrombocytopenia, which is uncommon, is defined as a platelet count below 20,000 and is associated with more serious bleeding problems.

Bone Marrow Examination

When blood tests indicate the presence of low blood counts (cytopenias), your physician may recommend a bone marrow examination. A bone marrow examination can reveal abnormalities in the cells of the marrow (e.g., dysplastic cells) and will allow evaluation of the chromosomes (cytogenetics).

These tests provide additional information that can help in establishing the diagnosis. There are two parts to a bone marrow examination: the aspirate and the core biopsy. Both procedures are usually performed at the same time.

The Bone Marrow Aspirate

The bone marrow aspirate is a sample of the liquid portion of the bone marrow. The sample provides information about the shape of the cells (morphology), how the cells are maturing (differentiation) and the number of blasts (immature cells) in the bone marrow. The aspirate may also be used for additional testing that may help to determine the cause of the cytopenias, such as cytogenetics.

The Bone Marrow Biopsy

The bone marrow biopsy is a small sample of the spongy center of the bone marrow. The bone marrow core is usually 1.5-2.0 cm in length. It provides information about the cellularity of the bone marrow (crowded = **hypercellular**, empty = **hypocellular**). It will also provide useful information about iron storage, scarring (fibrosis), and the presence of any other abnormal cells.

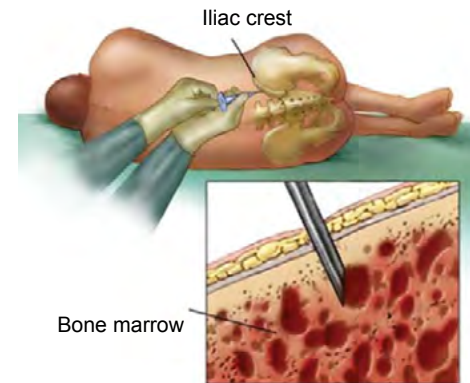
Processing the Sample

The bone marrow biopsy and aspirate samples are placed on glass slides and in various laboratory tubes. These are sent to a hematopathologist—a physician trained to evaluate blood and bone marrow samples to diagnosis diseases. This physician uses a microscope to examine the cells in the bone marrow aspirate and biopsy samples. The results of a bone marrow biopsy and aspirate generally take 2-4 days. Cytogenetic studies and other special studies may require up to 2 weeks.

The Bone Marrow Biopsy Procedure

A bone marrow examination can be performed in the physician's office usually in about twenty minutes. It can be performed with local anesthesia or, in some cases, mild sedation.

1. The patient lays on either their side or on their stomach.
2. The biopsy is taken from the right or left back side of the hip.
3. The skin above the site will be anesthetized (numbing the skin) using a form of lidocaine (numbing medicine).
4. Once the skin and bone has been anesthetized, a small incision may be made on the surface of the skin to allow



insertion of the bone marrow needle. The aspirate and the biopsy may be obtained with the same needle during the procedure.

5. After the procedure, the provider will apply pressure to the site to prevent any bleeding. A pressure dressing is generally applied.
6. You should not shower for 24 hours. No soaking in water (bath, swimming, hot tubs) for 48-72 hours. Ask your provider for instructions on how to care for the biopsy site.
7. Some patients may develop a bruise or swelling under the skin, particularly patients with a low platelet count or patients taking medication to thin the blood. ***Be sure to let your health care provider know if you are taking aspirin or other medications that thin the blood.***
8. Mild pain or discomfort may be experienced at the procedure site for two to three days after the bone marrow exam.
9. For safety reasons, the patient should have a friend, family member, or caretaker travel home with them. ***The patient should not drive.***

For More Information on MDS, Referrals to a Center of Excellence
or for a Second Opinion Contact:

Patient Liaison

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