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• The SPO Virtual Classrooms offer many educational resources, including practice test questions, review questions, lecture PowerPoints, video tutorials, sample assignments and course syllabi. New materials are continually being developed, so check back frequently, or follow us on Facebook (Science Prof Online) or Twitter (ScienceProfSPO) for updates.

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• Images used on this resource, and on the SPO website are, wherever possible, credited and linked to their source. Any words underlined and appearing in blue are links that can be clicked on for more information. PPT files must be viewed in *slide show mode* to use the hyperlinks directly.

• Several helpful links to fun and interactive learning tools are included throughout the PPT and on the Smart Links slide, near the end of each presentation. You must be in *slide show mode* to utilize hyperlinks and animations.

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From the Virtual Microbiology Classroom on ScienceProfOnline.com

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Image: <u>Compound microscope objectives</u>, T. Port



From the Virtual Microbiology Classroom on ScienceProfOnline.com

Images:: <u>Castle</u>, S. Jervis,; Angelina Jolie as assassin in movie "Wanted"

It Isn't Easy Being a Pathogen

What a pathogen must do in order to cause disease:

1. Gain access to the body.

- 2. Attach to and/or enter cells of its host.
 - Receptors on pathogen must fit, lock-and-key, with receptor sites on host cell.
- 3. Reproduce while avoiding host's immune system long enough to produce harmful changes.



Images: <u>Helicobacter pylori</u>, Yutaka Tsutsumi, M.D; <u>Deep gastric ulce</u>r, Samir; <u>Histopathology of H.pylori</u> from a gastric

Normal Flora

 Protect the body by competing with potential pathogens.



- This is called microbial antagonism.
- Normal microbiota protect us by:
 - Consuming nutrients that would otherwise be available to pathogens.
 - Sometimes change the **pH** of the area they inhabit in ways that help them and hinder competing microbes.
 - Presence stimulates certain parts of the second line of immune defense, helping the body defend itself from invaders.
 - Normal flora of the intestines improve our overall health by producing several types of vitamins.

Innate Immunity

- First two lines of immune defense considered together.
- Q: Why do you think that they are called innate immunity?
- Innate immunity is nonspecific, meaning that these lines of defense work against a wide range of pathogens.



First Line of Defense Nonspecific

- Structures, chemicals, processes that work to prevent pathogens entering the body.
- Includes the skin and mucous membranes of the respiratory, digestive, urinary, and reproductive systems.



Skin – Physical Components of Defense

Two major layers:

1. epidermis

- Outer layer composed of multiple layers of tightly packed cells
 - Few pathogens can penetrate these layers
 - Shedding of dead skin cells removes attached microorganisms
- Epidermal dendritic cells phagocytize pathogens.
 - These cells extend out among other cells of the epidermis, forming a network to intercept invaders.





2. dermis

- Contains protein fibers called collagen
 - Give skin strength and pliability to resist abrasions that could introduce microorganisms

First Line of Defense

Image: "Skin" tattoo, Source unknown; <u>Skin diagram</u>, Daniel de Souza Telles

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Skin – Chemical Components of Defense

- perspiration secreted by sweat glands
 - Salt- inhibits growth of pathogen by drawing water from their cells
 - Antimicrobial peptides
 - Lysozyme- destroys cell wall of bacteria
- sebum secreted by sebaceous (oil) glands
 - Helps keep skin pliable and less likely to break or tear
 - Lowers <u>pH</u> of skin to a level inhibitory to many bacteria





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sebaceous

gland

Images: Cartoon of castle being defended, Source unknown; <u>Hair follicle</u>, Wiki

Mucous Membrane

- Line all body cavities open to the outside environment.
- Unlike surface epidermal cells, epithelial cells are living.
- Epithelial cells packed tightly to prevent entry of pathogens, but often only one cell layer thick, so pathogens sometimes breech the barrier.
- Continual shedding of cells carries attached microorganisms away
- Besides producing mucus, mucous membranes also produce lysozyme and other antimicrobial <u>peptides</u>.
- OMG U R Nasty > Every day you swallow and digest about 1 liter of mucus.



Images: Photo mucous membrane, Source unknown, <u>Drawing of mucous membrane</u>, Gray's Anatomy

Second Line of Defense Nonspecific

- Operates when pathogens penetrate skin or mucous membranes.
- Cells, antimicrobial chemicals, and processes, but no physical barriers.
- Many of these components are contained or originate in the blood.



Images: <u>Neutrophil bacterial phagocytosis</u>, Uwe Thormann; <u>Ingrown toenail inflammation</u>, Wiki

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Formed Elements

Three types of formed elements:

- erythrocytes red blood cell, carry oxygen & carbon dioxide in the blood.
- platelets involved in blood clotting (also called thrombocytes).
- leukocytes white blood cells; involved in defending the body against invaders.
 - 2 groups
 - Granulocytes
 - Agranulocytes



Scanning electron micrograph of formed elements

RBC (left)

platelet (center)

leukocyte (right)

Leukocytes > Granulocytes

Category of <u>white blood cells</u> characterized by presence of **granules** in their cytoplasm.

3 types:





Neutrophils - Most abundant white blood cell. Predominant cells in pus, accounts for its whitish appearance. Respond quickly following tissue injury. Hallmark of acute inflammation.



Basophils - Least common granulocyte. When activated, release histamine and other inflammatory chemicals.



Eosinophils - Main effecter cells in allergic responses & asthma. Also fight helminth (worm) colonization.

Neutrophils and eosinophils can *phagocytize* pathogens.



Image: <u>Neutrophil</u> engulfing *Bacillus anthracis*, Volker Brinkmann, Photos of <u>granulocytes</u>, Wiki; <u>Drawing of granulocytes</u>, US Gov

From the <u>Virtual Microbiology Classroom</u> on <u>ScienceProfOnline.com</u>

Leukocytes > Agranulocytes



2 types:



Lymphocytes - most involved in specific immunity (3rd line of immune defense),



Monocytes - leave the blood and mature into **macrophages** (phagocytic cells of the second line of defense).



Image: <u>Macrophage</u>, Wiki; <u>Lymphocyte,</u> Nicolas Grandjean; <u>Monocyte</u>, Bobjgalindo

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Components of the Second Line of Defense

- Leukocytes
 - Phagocytosis How phagocytes ingest and destroy foreign matter such as microorganisms or debris.
 - Extracellular killing by leukocytes
- Nonspecific chemical defenses
- Inflammation

THE Phagocytosis of a movie dude.

• Fever





Leukocytes: Extracellular Killing

3 Cell Types That Kill Extracellularly:

- natural killer lymphocytes (NK cells)
 - Secrete toxins onto surface of virally infected cells & tumors.
 - Differentiate normal body cells because they have membrane proteins similar to the NK cells.
- eosinophils
 - Mainly attack parasitic worms by attaching to their surface.
 - Secrete toxins that weaken or kill worm.
 - Elevated eosinophil levels, is often indicative of a helminth (parasitic worm) infection.
- neutrophils
 - Can create the active ingredient in bleach to kill nearby microbes.
 - Fibers called neutrophil extracellular traps (NETs) can ensnare and kill bacteria and fungi. Secrete antimicrobial proteins.



Components of the Second Line of Defense

- Leukocytes
 - Phagocytosis
 - Extracellular killing by leukocytes

• Nonspecific chemical defenses

- Lysozyme, Defensins & Cytokines (including interferons and interleukins).
- Augment phagocytosis
- Some attack pathogens directly
- Some enhance features of nonspecific resistance
- Inflammation
- Fever







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Components of the Second Line of Defense

- Leukocytes
 - Phagocytosis
 - Extracellular killing by leukocytes
- Nonspecific Chemical Defenses
 - Ex. Lysozyme, Defensins & Cytokines
- Inflammation
 - Nonspecific response to tissue damage.
 - Damages cells release histamines which increase vasodilation.
 - Heat, swelling, pain



• Fever

Components of the Second Line of Defense

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• Fever

- ✓ Body temp above normal range of 36.5-37.5 °C (98-100 °F).
- Results when chemicals called trigger the hypothalamus to increase body's core temperature.
- ✓ Various types of pyrogens
 - Bacterial toxins
 - Cytoplasm of bacteria released by lysis
 - Antibody-antigen complexes
 - Interleukin-I (IL-1 a cytokine)

✓ Benefits

- Speed of immune system reaction increased
- Inhibits growth of some temp sensitive microorganisms

Third Line of Defense Acquired

- The body's ability to recognize and defend itself against distinct invaders.
 - Is a "smart" system.
 - Also called **specific** and **adaptive** immunity.
 - "Memory" allows it to respond rapidly to additional encounters with a pathogen.
 - If nonspecific immune system has *warriors*, then acquired immunity has more sophisticated *special agents* and *assassins*.
 - Two types of specific immunity:
 - Naturally acquired = immune response against antigens encountered in daily life.
 - Artificially acquired = response to antigens introduced via vaccine.
- Q: How does the body recognize invaders?



Images: Child getting immunized, PHIL #9423

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Antigens

- Body does not direct immune response against whole bacteria, fungi, protozoa or viruses.
- Foreign molecules trigger a specific immune response.
- Include components of <u>bacterial</u> <u>cell walls</u>, capsules, pili, and flagella, as well as proteins of <u>viruses</u>, fungi and protozoa.
- Food and dust can also contain antigenic particles.
- Enter the body by various methods:
 - Through breaks in skin & mucous membranes
 - Direct injection, as with a bite or needle
 - Through organ transplants and skin grafts

HELLO my name is

Antigens Are Like Name Tags Antigenic particles are often associated with a specific characteristic of an organism, so are detected as foreign when they get inside another organism that doesn't have that characteristic.



Third Line of Defense

Images: **Blood cells**, National Cancer Institute

From the <u>Virtual Microbiology Classroom</u> on <u>ScienceProfOnline.com</u>

Antibodies

- Also called immunoglobulins (Ig).
- <u>Protein</u>aceous molecules that bind antigens at the antigenbinding site.
- Considered part of the humoral immune response since bodily fluids such as lymph and blood were once called humors.



Antibody

How Antibodies Work

- Some act as opsonins, markers to identify antigens for phagocytes and stimulate phagoctosis.
- Some work as antitoxins (i.e. they neutralize toxins for e.g. those causing diphtheria and tetanus).
- Some attach to bacterial flagella making them less active and easier for phagocytes to engulf.
- Some cause agglutination (clumping together) of bacteria making them less likely to spread

Q: But where do antibodies come from?



Image: <u>Antigen antibody complex</u>, Dr. Sanderson Immunology Page; <u>Opsoni</u>n, Graham Colm

From the <u>Virtual Microbiology Classroom</u> on <u>ScienceProfOnline.com</u>

Lymphatic System

- Screens tissues of the body for foreign antigens.
- Composed of lymphatic vessels and lymphatic cells.
- One-way system that conducts lymph from local tissues and returns it to the circulatory system.
 - Lymph is a liquid with similar composition to blood plasma.
 - Comes from fluid leaked from blood vessels into surrounding tissues.
- Lymph nodes house white blood cells called **lymphocytes** that recognize and attack foreign antigens present in lymph.





Lymphocytes

- WBCs of specific immunity. Smallest <u>leukocytes</u>. Have huge nucleus surrounded by thin rim of cytoplasm.
- Produced from blood stem cells in the red bone marrow.



Two main types:

- B cells mature in bone marrow, then concentrate in lymph nodes & spleen.
- T cells mature in thymus.
- B and T cells mature then circulate in the blood and lymph.
- Circulation ensures they come into contact with pathogens and each other.

T Lymphocytes (T cells)

- Produced in red bone marrow and mature in thymus.
- Circulate in the lymph and blood and migrate to the lymph nodes (and other areas of the lymph system).
- Part of the <u>cellular immune response</u> (aka cell-mediated immune response) because these cells act directly against various antigens
 - Endogenous invaders (intracellular pathogens inside the body's cells)
 - Abnormal body cells such as cancer cells
- Types
 - cytotoxic or killer T cells (Tc)
 - Destroy compromised body cells
 - helper T cells (T_{H})
 - Activate B-cells



What Is an Antigen Presenting Cell?

Consider your WBCs a security force for your body and any nonself antigens as pictures of a bad guy.

The larger the force, the more likely one of the officers will run into a "bad guy" and help the body apprehend it.

Any WBC that can grab and present an antigen to another, is called an antigen presenting cell (APC).



There are "professional" (WBC) APC cells, such as **B cells**, **macrophages** and **dendritic cells**.

There are also other cells in the body (non-WBCs) that are "non-professional" APC cells, such as fibroblasts (in skin), some epithelial and endothelial cells & glial cells (in brain).

Image: B cell differentiation, Source unknown

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B Lymphocytes (B cells)

Activated B-lymphocytes produce either:

- Plasma cells make antibodies to a pathogen.
- Memory cells remember the same pathogen for faster antibody production in future infections.







Animated lesson on Humoral Immune Response



Image: <u>Antigen antibody complex</u>; <u>Antibody</u>, Fvasconcellos; B cell differentiation, Source unknown

From the Virtual Microbiology Classroom on ScienceProfOnline.com

Confused?

Here are links to fun resources that further explain acquired immunity:

- <u>Immunology: Innate & Acquired Immunity</u> Main Page on the Virtual Cell Biology Classroom of <u>Science Prof Online</u>.
- **<u>Phagocytosis</u>** animation and quiz by McGraw-Hill.
- <u>Immune System</u> "Who Wants to Be a Millionaire" game.
- <u>Immune System</u> animation and guiz by McGraw-Hill.
- <u>"Fever</u>", song by Peggy Lee & "<u>Assassin</u>" song by John Mayer
- <u>Cellular Immune Response</u> & <u>Humoral Immune Response</u> narrated animation and quiz from W. H. Freeman.
- <u>Immune System Defender</u>, online game from the Nobel Prize website. Use your force of white blood cells to destroy invading bacteria, before they overpopulate and cause disease.
- <u>Immune System Game</u>, a collection of online fun and educational games about immunology.

(You must be in PPT slideshow view to click on links.)

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Are microbes intimidating you?

Do yourself a favor. Use the ...

Virtual Microbiology Classroom (VMC)

The VMC is full of resources to help you succeed, including:



- practice test questions
- review questions
- study guides and learning objectives

You can access the VMC by going to the Science Prof Online website <u>www.ScienceProfOnline.com</u>