

A course on Basic and Translational Immunology, with emphasis on immunologic diseases and therapeutic strategies

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Themes of the course

- The nomenclature of immunology
- Basic principles: mechanisms underlying immune responses
- Emerging concepts, and their potential clinical and therapeutic implications

What does the immune system do?

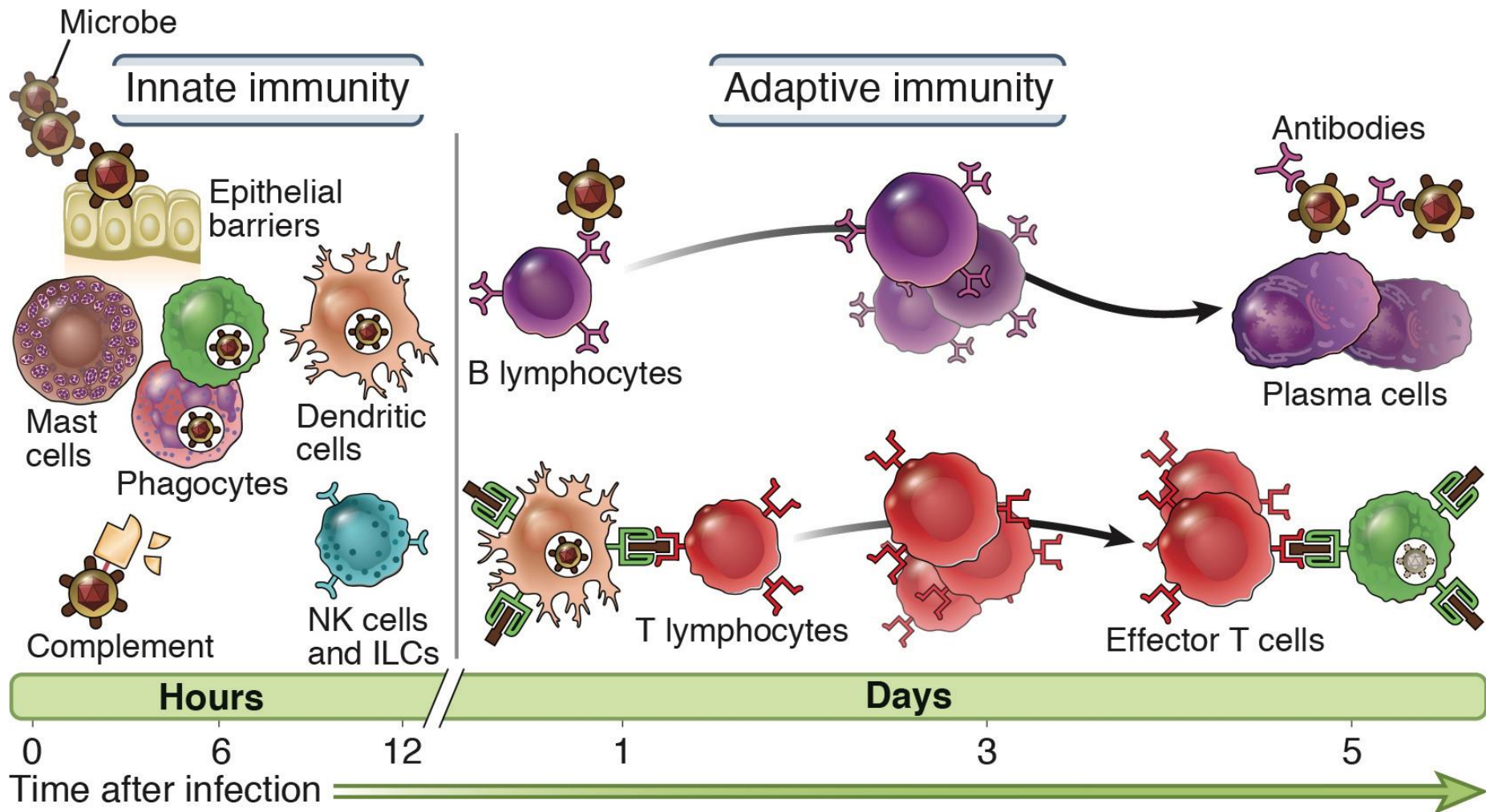
Normal functions

- Defense against infections
- Defense against some tumors

Disease and therapeutic implications

- Cause of disease (autoimmunity, allergy)
- Barrier to transplantation, gene therapy

Innate and adaptive immunity








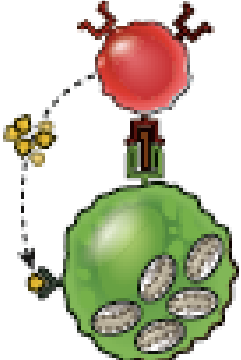



Abbas, Lichtman and Pillai. *Basic Immunology*, 5th edition, 2016, Elsevier

Innate immunity: always present (ready to attack); many pathogenic microbes have evolved to resist innate immunity

Adaptive immunity: stimulated by exposure to microbe; more potent

Types of adaptive immunity

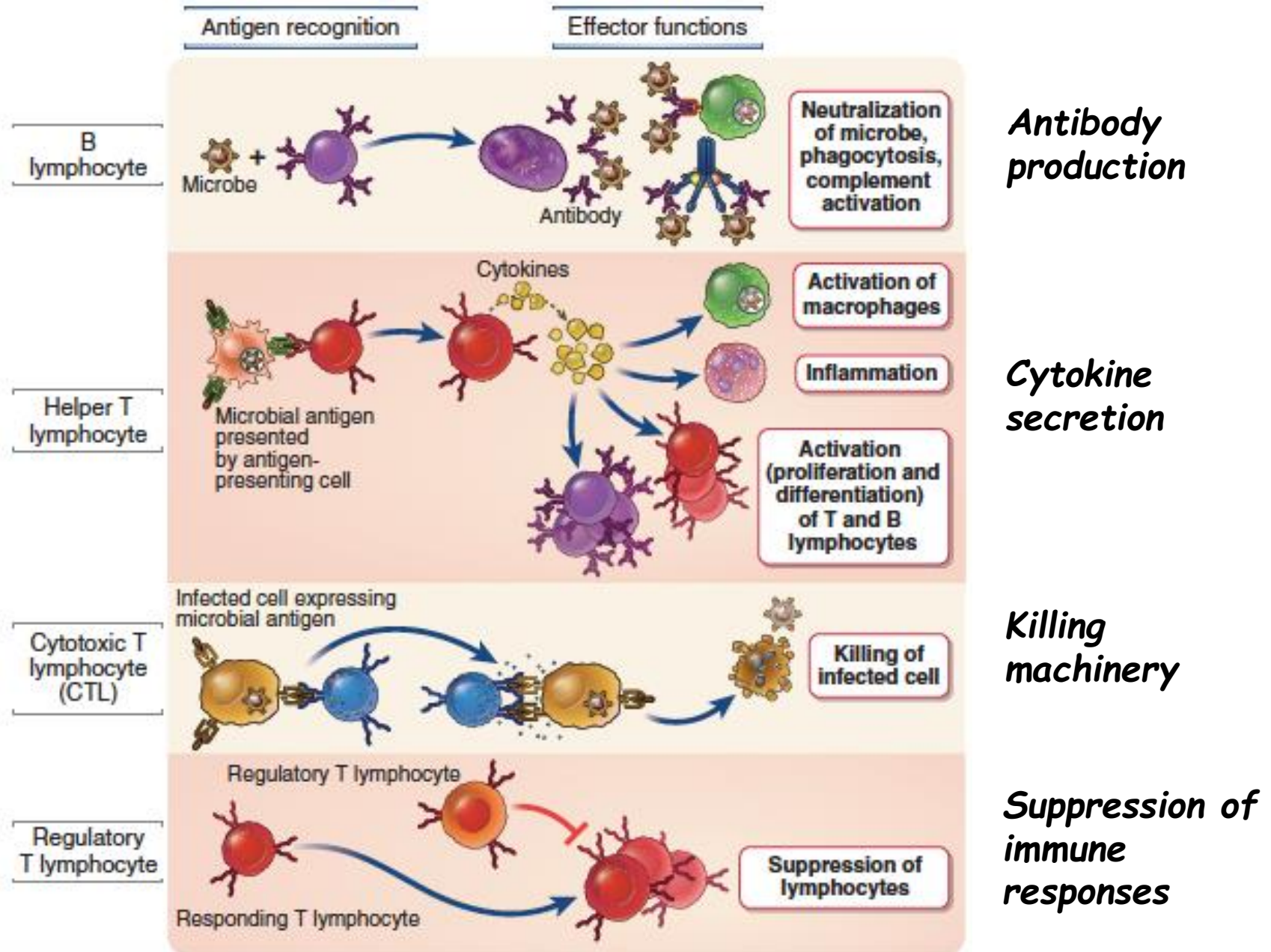
	Humoral immunity	Cell-mediated immunity	
Microbe	 <p>Extracellular microbes</p>	 <p>Phagocytosed microbes that can live within macrophages</p>	 <p>Intracellular microbes (e.g., viruses) replicating within infected cell</p>
Responding lymphocytes	 <p>B lymphocyte</p>	 <p>Helper T lymphocyte</p>	 <p>Cytotoxic T lymphocyte</p>
Effector mechanism	 <p>Secreted antibody</p>	 <p>Activated macrophage</p>	 <p>Killed infected cell</p>
Functions	Block infections and eliminate extracellular microbes	Elimination of phagocytosed microbes	Kill infected cells and eliminate reservoirs of infection

Different types of immune responses are mediated by different classes of lymphocytes and defend against different types of microbes

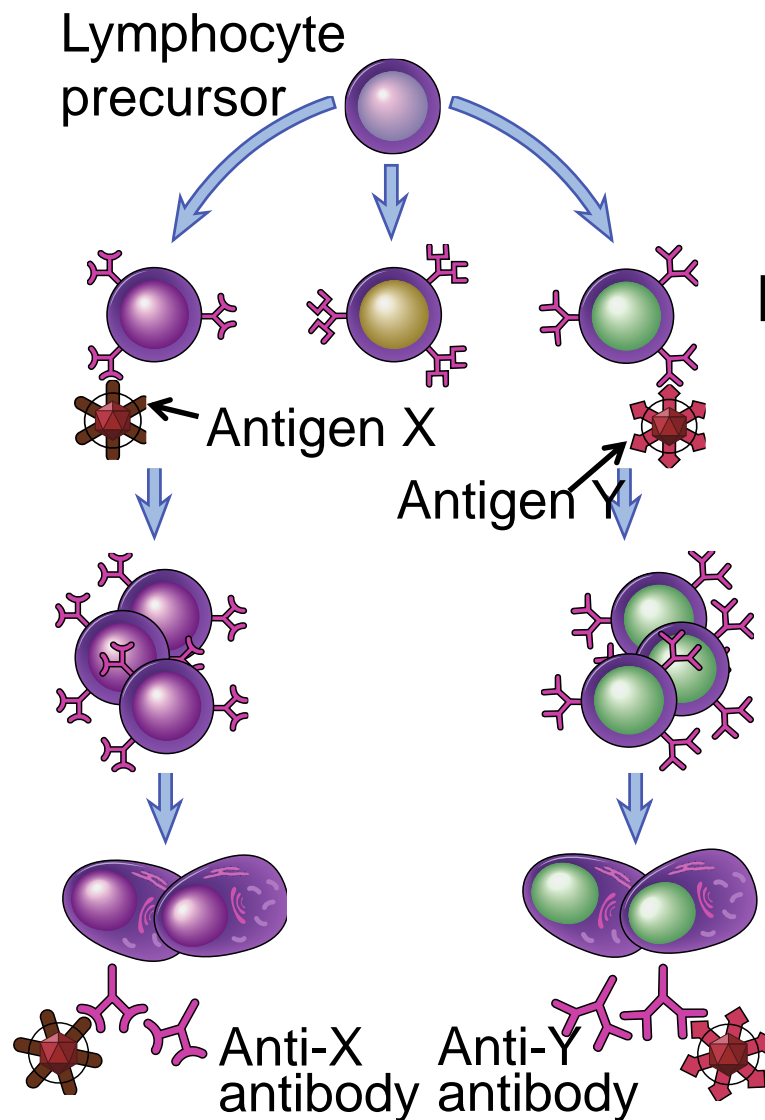
Cells of the immune system

- **Lymphocytes:** the cells of adaptive immunity; recognize antigens and develop (differentiate) into cells that perform the defense functions
- **Antigen-presenting cells:** cells that capture antigens and display them to lymphocytes
- **Effector cells:** leukocytes (white blood cells) that eliminate microbes (the “effect” of the immune response); may be lymphocytes, but are often other leukocytes

Classes of lymphocytes



Lymphocyte diversity and clonal selection



Generation of mature lymphocytes with many different antigen receptors

Naïve lymphocytes circulate through lymphoid organs

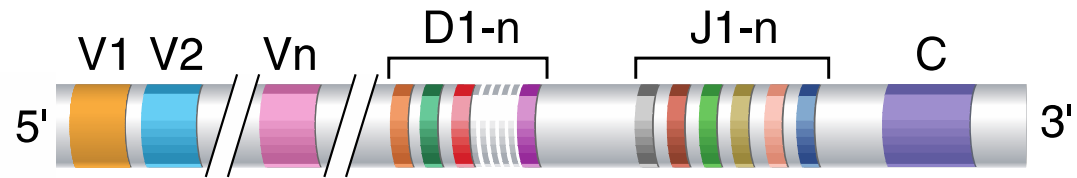
Specific lymphocytes recognize antigens

Lymphocytes are activated to proliferate and to differentiate into effector cells

Lymphocytes with highly specific and diverse antigen receptors develop prior to exposure to antigens

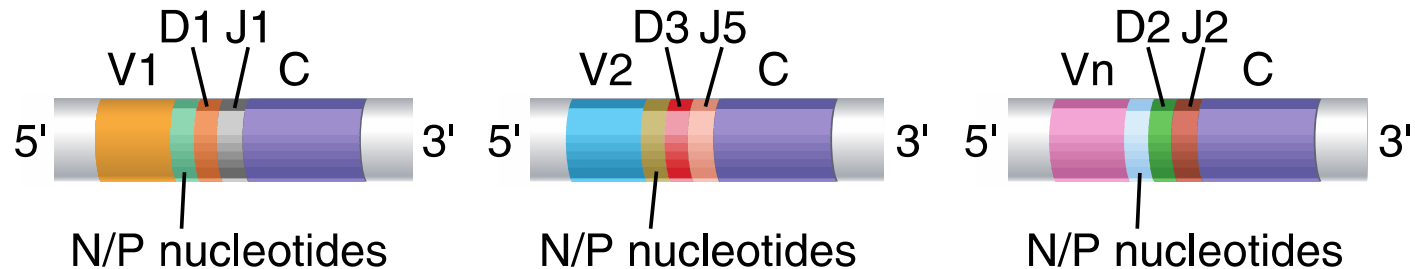
Generation of diversity

Germline
DNA

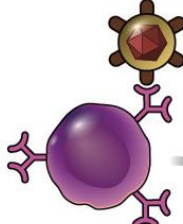

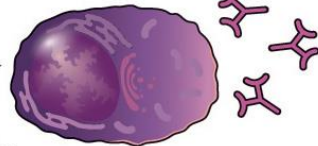
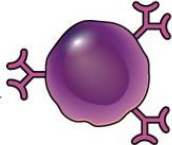
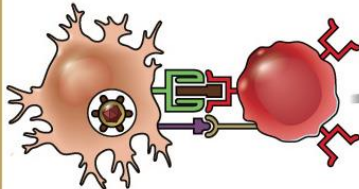
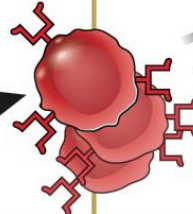

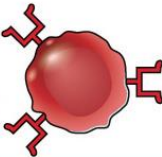


Somatic recombination (V-D-J joining), addition of N and P nucleotides, transcription and RNA processing in three B cell clones

Expressed
mRNA in
three
lymphocyte
clones



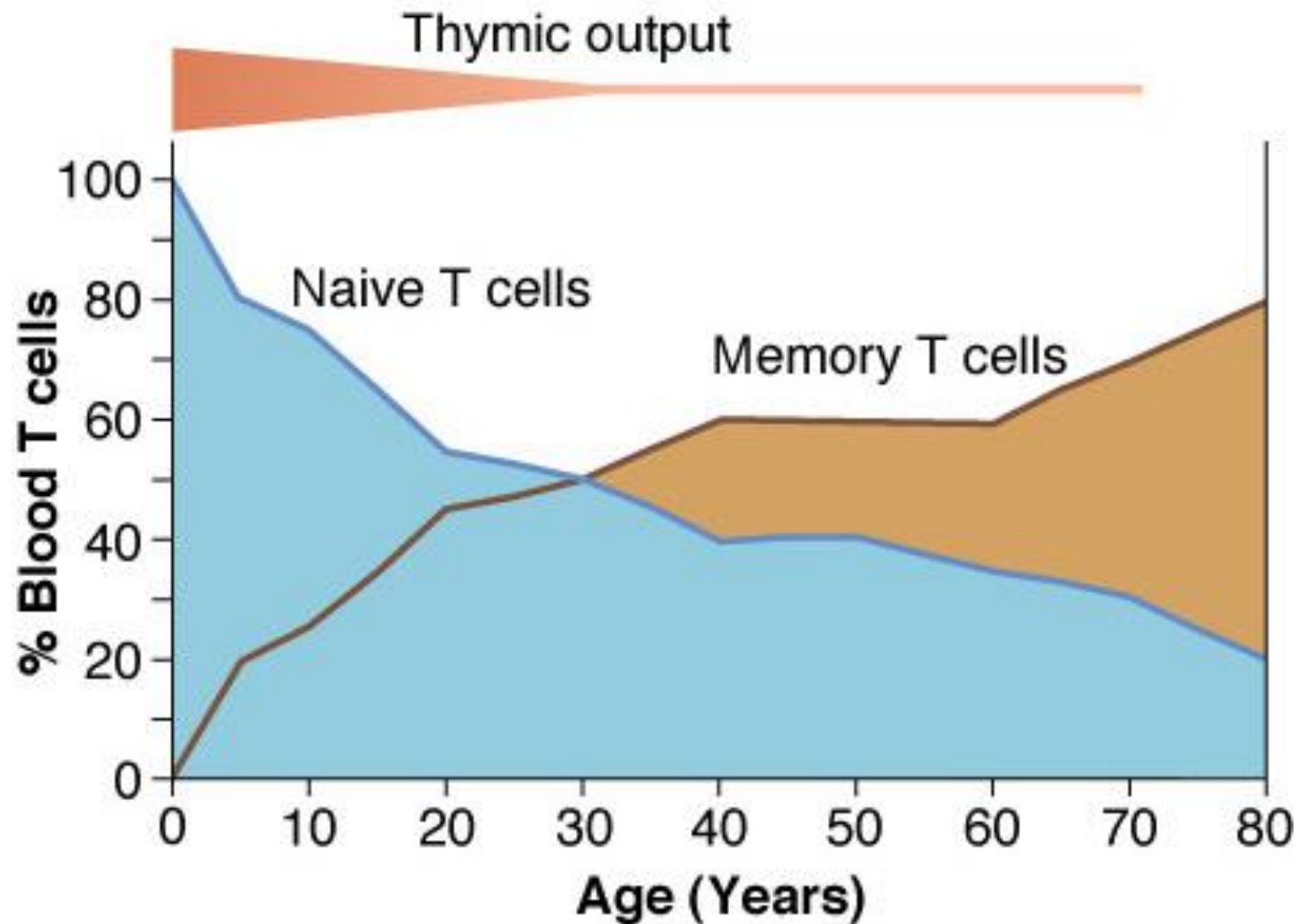
Stages in the life history of lymphocytes

Cell type	Stage			
	Naive cell	Activated or effector lymphocyte	Memory lymphocyte	
B lymphocytes	Antigen recognition 	Proliferation 	Differentiation 	
T lymphocytes	Antigen recognition 	Proliferation 	Differentiation 	

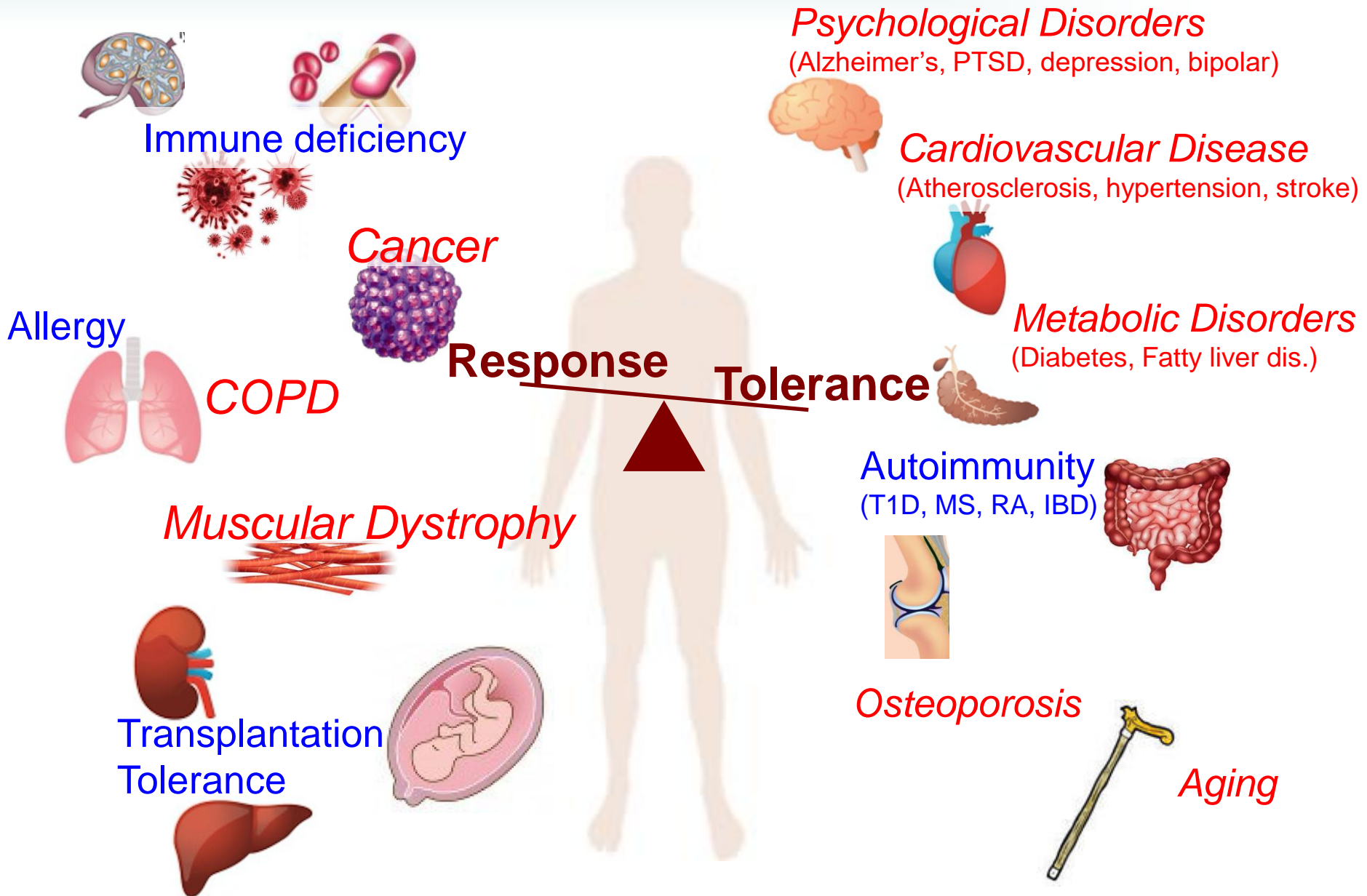
Proliferation: expands number of antigen-specific cells

Differentiation: converts lymphocytes into effective defenders

Accumulation of memory T cells with age



Immunological basis of human disease



The significance of recent advances

- Provides a solid foundation of basic principles
- Improved understanding of disease mechanisms
- Development of novel therapies
- Appreciation of the role of the immune system in non-immune diseases

Challenges in Immunology

- Explosion of information creates complexity
 - “Big data” is difficult to interpret, has not yet provided many useful answers
 - Many complex cell populations and pathways
- Translating results from mouse to human
 - Co-housing with dirty mice makes the immune system of lab mice more like humans
- Translating results from cell cultures to in vivo