

# Lymphoma

# Overview

- Concepts, classification, biology
- Epidemiology
- Clinical presentation
- Diagnosis
- Staging
- Three important types of lymphoma

# How Cancer Develops

- Normal cells are programmed to multiply, and die when they're old
- Signals to multiply and die are controlled by specific genes
- Mutations can occur in these genes
- If enough mutations occur in genes controlling growth or cell death a cell begins to **multiply uncontrollably**
- The cell has then become cancerous or "malignant"

# Features common to cancer cells

- Growth in the absence of “go” signals
- Growth despite “stop” signals
- Locally invasive growth and metastasize to distant sites

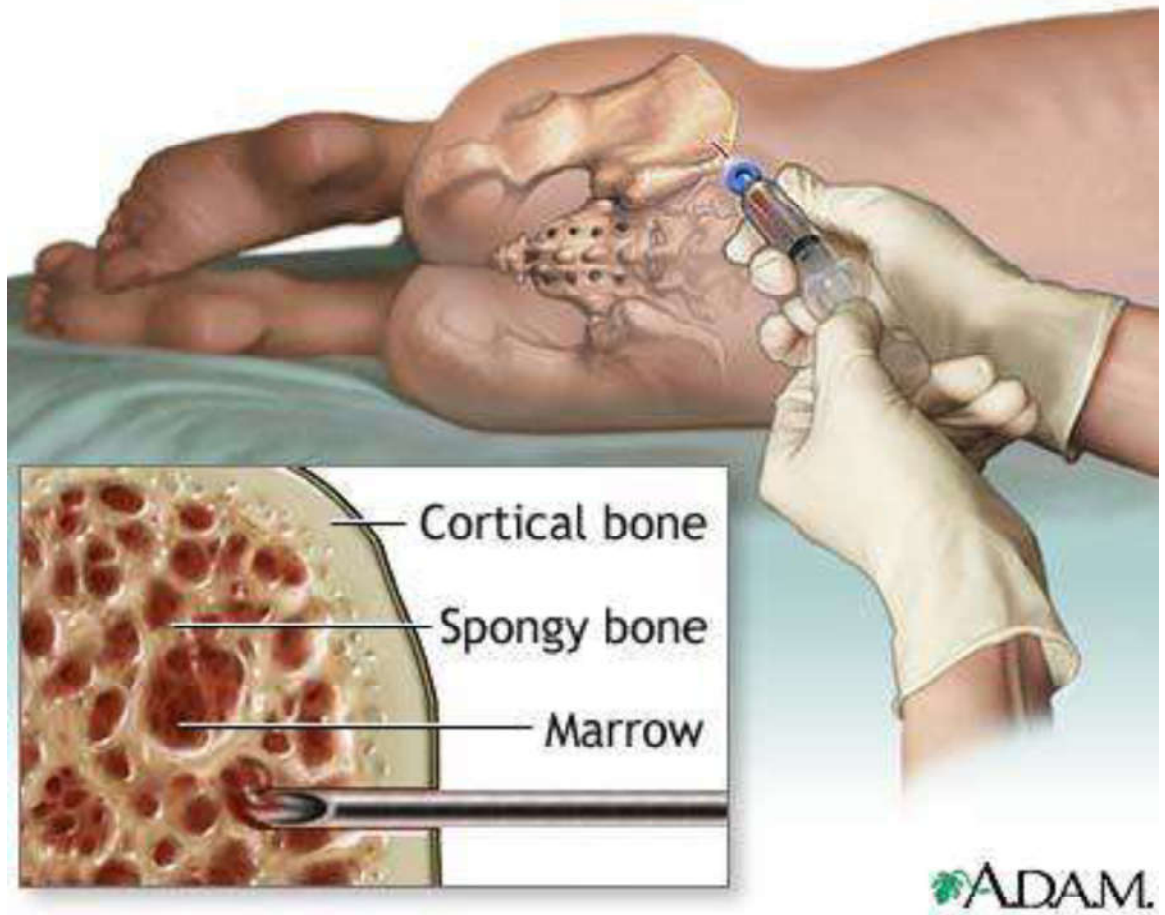
# Bone Marrow

- Present in the soft inner part of some bones such as the skull, shoulder, blade, ribs, pelvis, and backbones. (Occupies central cavity of bone)
- The bone marrow is made up of blood-forming stem cells, lymphoid tissue, fat cells, and supporting tissues that aid the growth of blood-forming cells.

# Bone Marrow

- Spongy tissue where development of a types of blood cells takes place
- All bones have active marrow at birth
- Adulthood - vertebrae, hip, shoulders, ribs, and skull contain marrow

# Bone Marrow Aspiration/Biopsy



# Hematopoietic Malignancies

- **Lymphoma is a general term for hematopoietic solid malignancies of the *lymphoid* series.**
- **Leukemia is a general term for liquid malignancies of either the *lymphoid* or the *myeloid* series.**



# Conceptualizing lymphoma

- neoplasms of lymphoid origin, typically causing lymphadenopathy
- leukemia vs lymphoma
- lymphomas as clonal expansions of cells at certain developmental stages

# What is Lymphoma

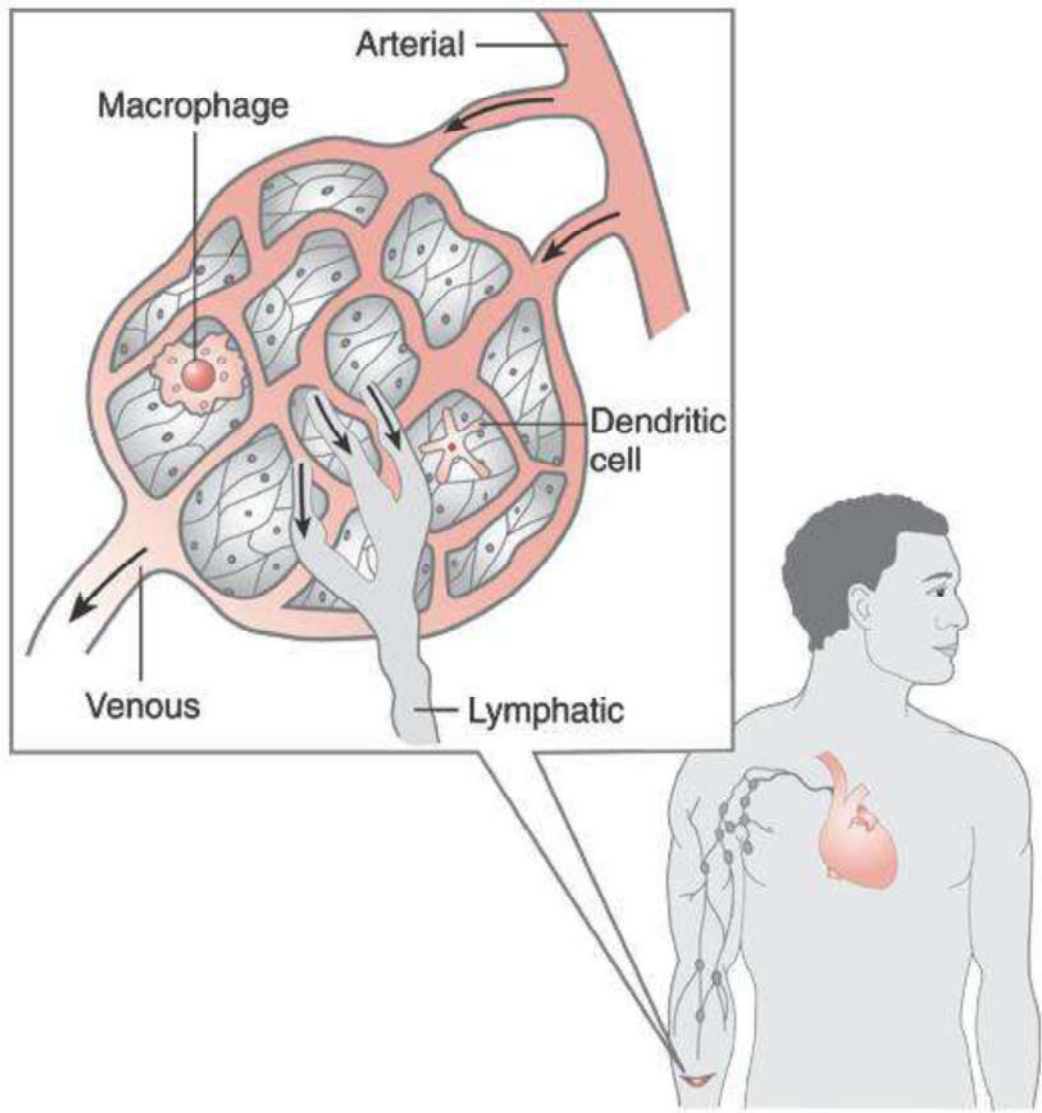
- Lymphomas are cancers that begin with the “malignant transformation” of a lymphocyte in the lymphatic system
- Many lymphomas are known to be due to specific genetic mutations
- Follicular lymphoma due to overexpression of BCL-2 (gene that blocks programmed cell death)

# What is the Lymphatic System

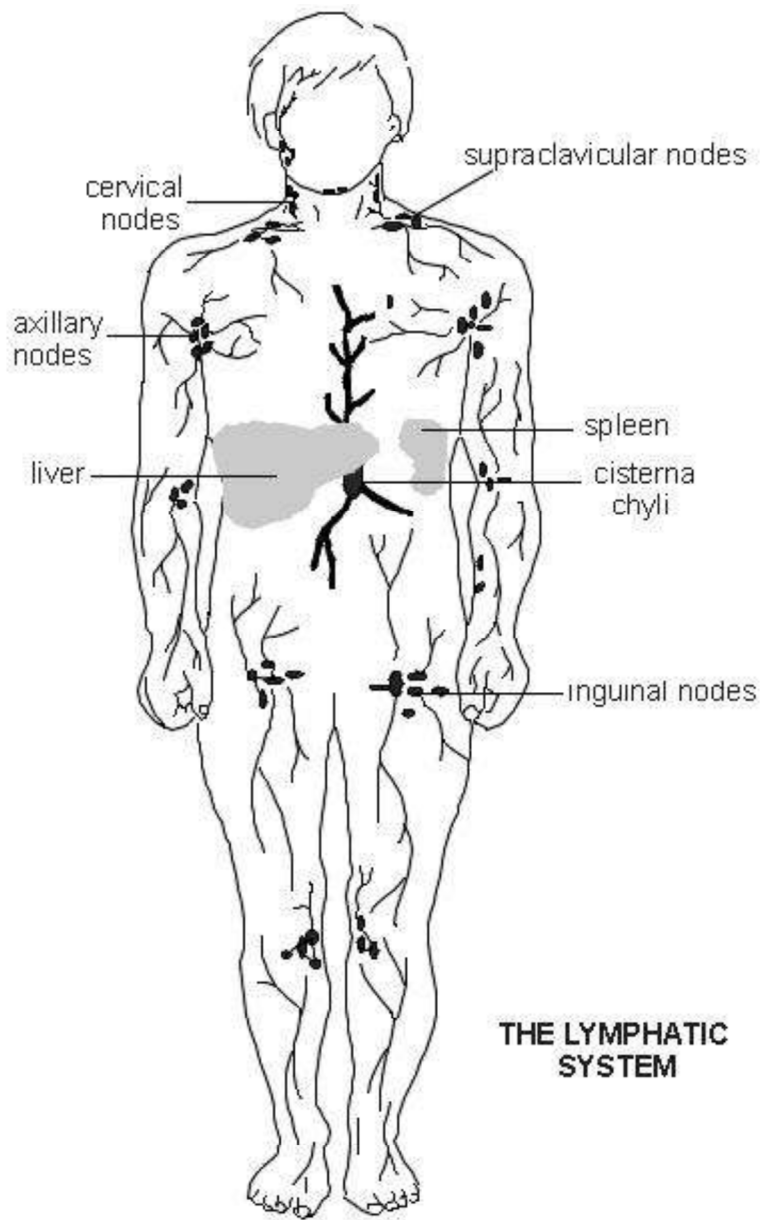
- Made up of organs, such as the tonsils, spleen, liver, bone marrow and a network of lymphatic vessels that connect glands, called lymph nodes
- Lymph nodes located throughout the body
- Lymph nodes filter foreign particles out of the lymphatic fluid
- Contain B and T lymphocytes

# Lymphatic System

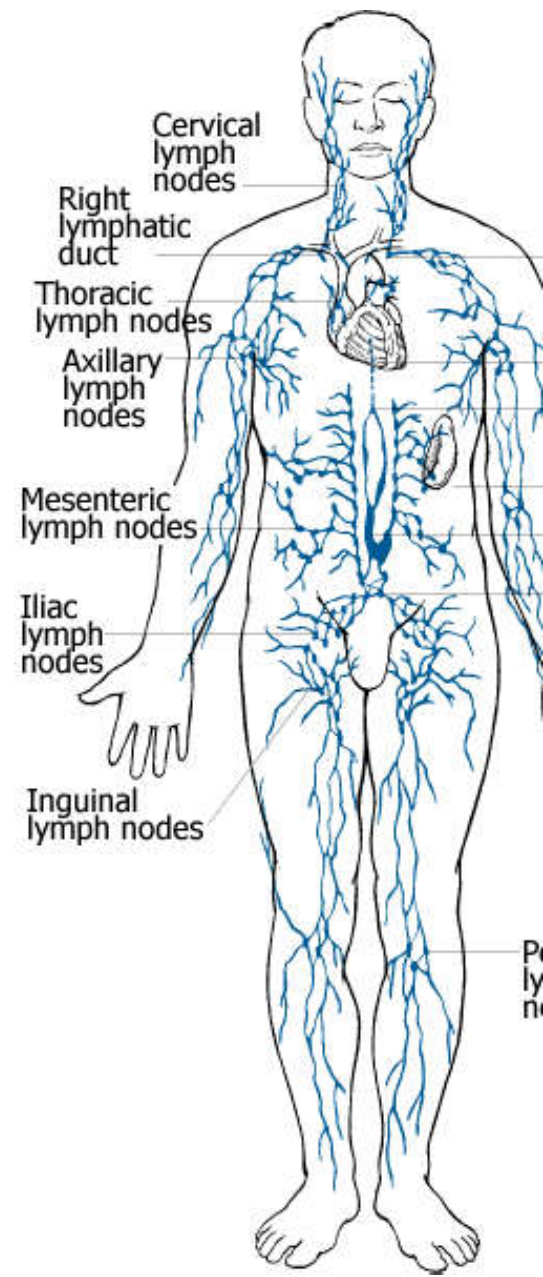
- Lymph nodes act as a filter to remove bacteria, viruses, and foreign particles
- Most people will have had “swollen glands” at some time as a response to infection



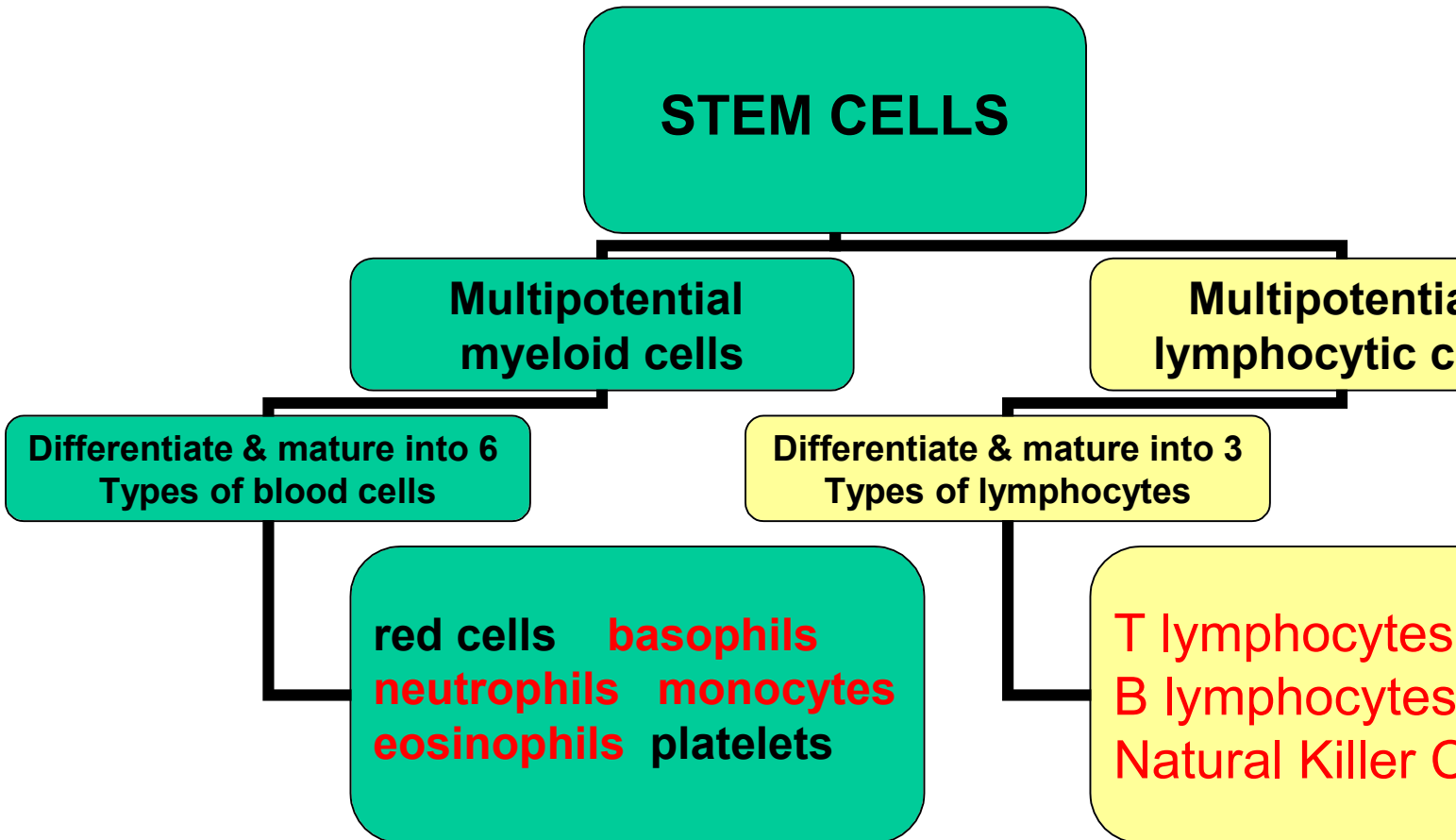
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**THE LYMPHATIC SYSTEM**



# Blood Cell and Lymphocyte Development



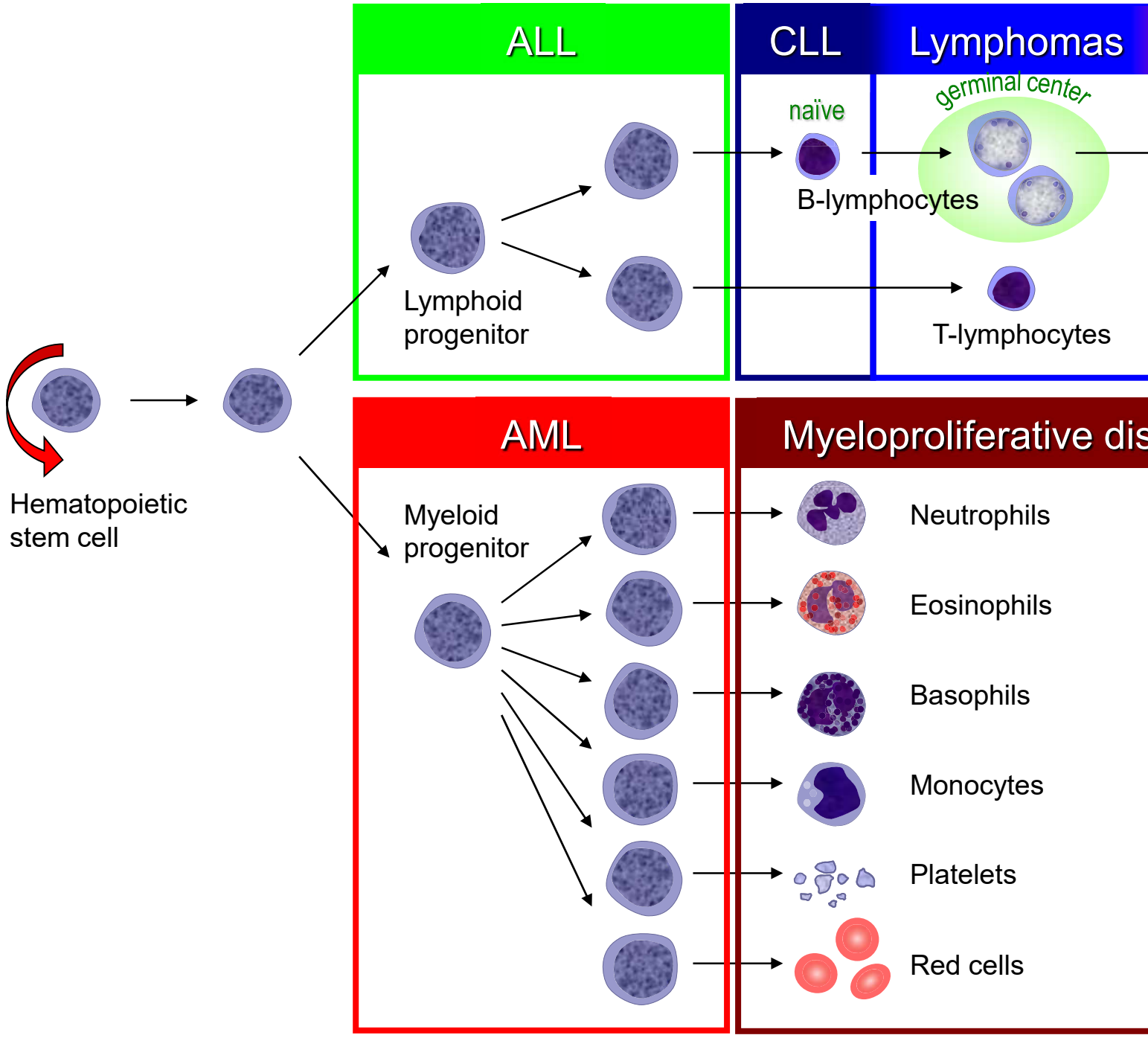
# Lymphocytes

- Most lymphocytes are in lymph nodes, spleen, bone marrow and lymphatic vessels
- 20% of white blood cells in blood are lymphocytes
- T cells, B cells, natural killer cells
- B cells produce antibodies that help fight infectious agents
- T cells help B cells produce antibodies and they fight viruses

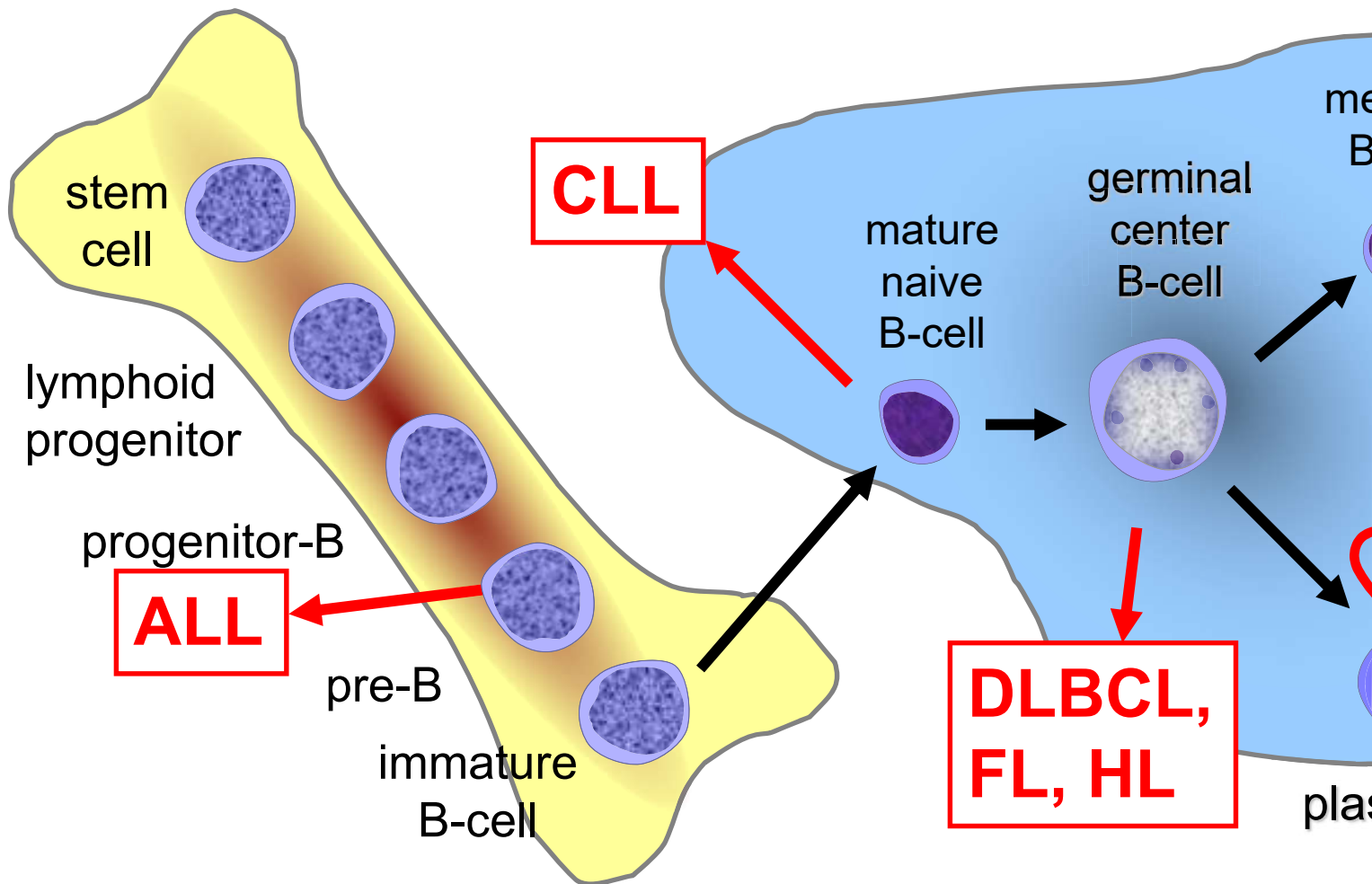


# T-Cells and B-Cells

- Immature lymphocytes that travel to the thymus differentiate into T-Cells
  - “T” is for thymus
- Immature lymphocytes that travel to the spleen or lymph nodes differentiate into B cells
  - “B” stands for the bursa of Fabricius, which is an organ unique to birds, where B cells mature.



# B-cell development



**Bone marrow**

**Lymphoid tissue**

# Classification

## Biologically rational classification

Diseases that have distinct

- morphology
- immunophenotype
- genetic features
- clinical features

## Clinically useful classification

Diseases that have distinct


- clinical features
- natural history
- prognosis
- treatment

# Classification

- Usually classified by how the cells look under a microscope and how quickly they grow and spread
  - Aggressive lymphomas (high-grade lymphomas)
  - Indolent Lymphomas (low-grade lymphomas)

# Lymphoma classification (2001 WHO)

- B-cell neoplasms
  - precursor
  - mature
- T-cell & NK-cell neoplasms
  - precursor
  - mature
- Hodgkin lymphoma

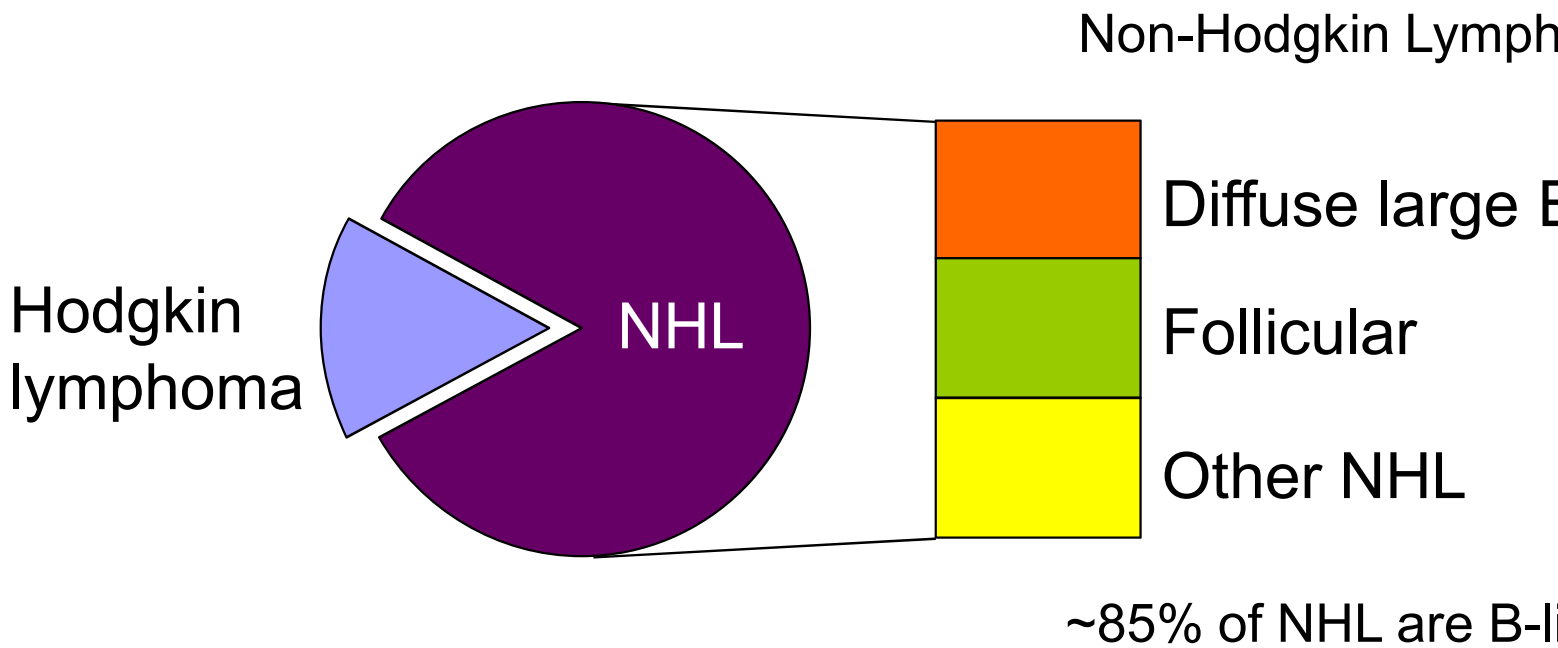


Non-  
Hodgk  
Lymph

# Three common lymphomas

- Follicular lymphoma
- Diffuse large B-cell lymphoma
- Hodgkin lymphoma

# Relative frequencies of different lymphomas





# Follicular lymphoma

- most common type of “indolent” lymphoma
- usually widespread at presentation
- often asymptomatic
- not curable (some exceptions)
- associated with BCL-2 gene rearrangement [t(14;18)]
- cell of origin: germinal center B-cell

- defer treatment if asymptomatic (“watch-and-wait”)
- several chemotherapy options if symptomatic
- median survival: years
- despite “indolent” label, morbidity and mortality can be considerable
- transformation to aggressive lymphoma can occur

# Diffuse large B-cell lympho

- most common type of “aggressive” lymphoma
- usually symptomatic
- extranodal involvement is common
- cell of origin: germinal center B-cell
- treatment should be offered
- curable in ~ 40%