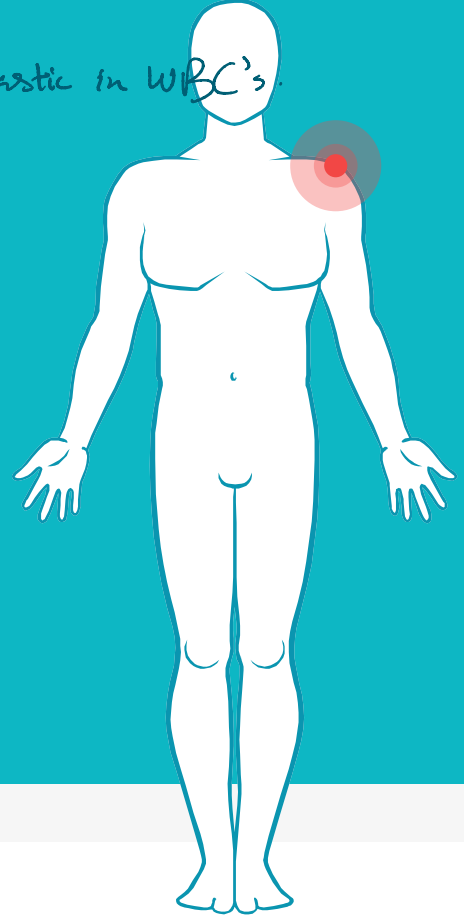


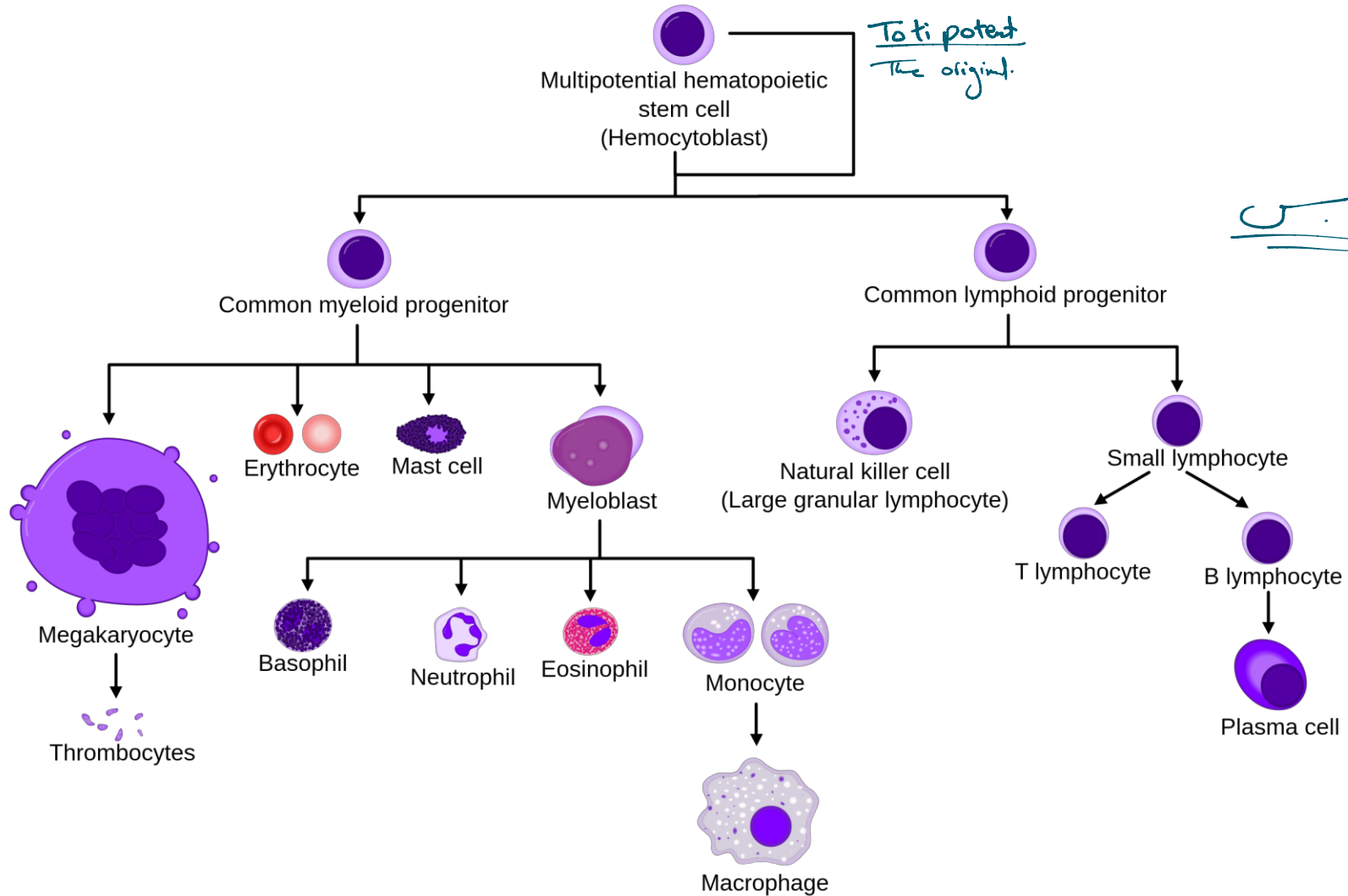
26/3/25

most neoplastic in WBC's.

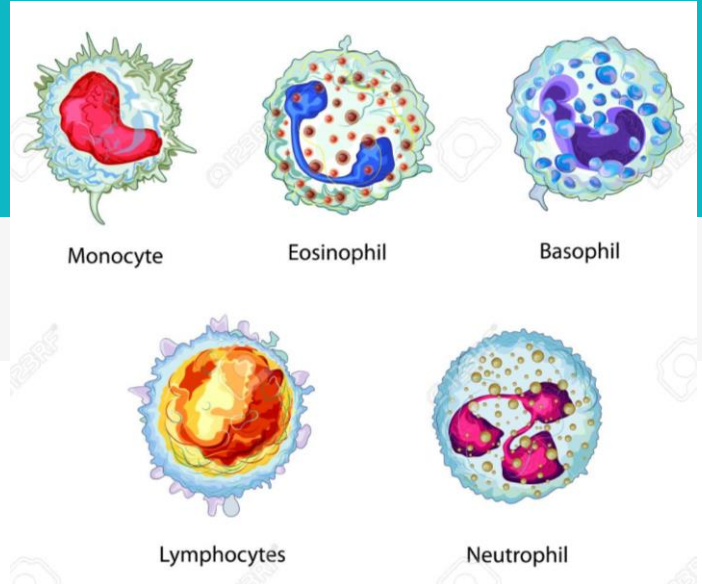
Hematopoietic & Lymphoid System

White Cell disorders





1. Nonneoplastic disorders of white cells



Peripheral blood
موسبودین بهم
ویس باره
4

- Disorders include **deficiencies** (abnormally low count → leukopenias) and **proliferations** (**leukocytosis**), which may be reactive or neoplastic.

زایه ست
Increased.

Cell Type	فشی هنزوری اسرغم للایحان بر ضروری الحیاة
White cells ($\times 10^3/\mu\text{L}$)	4.8-10.8
Granulocytes (%)	40-70
Neutrophils ($\times 10^3/\mu\text{L}$)	1.4-6.5 $\times 10^3$
Lymphocytes ($\times 10^3/\mu\text{L}$)	1.2-3.4 T/B/Natural killers
Monocytes ($\times 10^3/\mu\text{L}$)	0.1-0.6
Eosinophils ($\times 10^3/\mu\text{L}$)	0-0.5
Basophils ($\times 10^3/\mu\text{L}$)	0-0.2
Red cells ($\times 10^3/\mu\text{L}$)	4.3-5, men; 3.5-5, women
Platelets ($\times 10^3/\mu\text{L}$)	150-450

5 major types of WBCs

- ▶ Neutrophils
- ▶ Lymphocytes
- ▶ Monocytes
- ▶ Eosinophils
- ▶ Basophils

Neutropenia/Agranulocytosis

- ▶ Neutropenia: a reduction in the number of granulocytes in blood, when severe, agranulocytosis. (<500)
 - sever decrease.*
 - can include eosinophils, Basophils.*
- ▶ Most common Leukopenia.
 - neutrophils & platelets ← granulocytes*
- ▶ Neutropenic persons are susceptible to severe, potentially fatal bacterial and fungal infections.
 - innate immune protection*
 - neutrophils*
 - ~ phagocytosis process*
- ▶ Two major mechanisms:
 - ❖ Decrease production
 - ❖ Increase peripheral destruction of neutrophils

Neutropenia/Agranulocytosis

Decrease production

*decrease
in size.*

- ▶ Marrow hypoplasia in patients who receive chemotherapy or radiation therapy
- ▶ Leukemia or other tumors replacing the marrow
- ▶ Medications
- ▶ Certain types of neoplastic lymphocytic proliferations involving marrow.

*granulopiosis →
من قبيح عاكى الإنتاج*

~Lymphoma.

Neutropenia/Agranulocytosis

Increased peripheral destruction/consumption:

- ▶ Autoimmune destruction → *Anka that distract the macrophages.*
- ▶ Overwhelming bacterial, fungal or rickettsial infection (peripheral use) *بشكل تدريجي ما يقدر*
- ▶ Splenomegaly (sequestration & accelerated removal of neutrophils.) *استعمل Replicates. بـسـمـر الخـلايا أكثر.*

Neutropenia/Agranulocytosis

Clinical features:

- ▶ **Infections** ^{لأنه ما في كبر}
immunity
- ▶ Fever, chills, malaise.
- ▶ Mucocutaneous necrotizing ulcers
- ▶ High risk of sepsis ^{عشان}
ينقلو بالدم

Tx: broadspectrum antibiotics (bacterial and fungal),
granulocyte colony-stimulating factor (G-CSF).

Lymphopenia

- ▶ Much less common.
- ▶ Associated with rare congenital immunodeficiency diseases, advanced human immunodeficiency virus (HIV) infection, & high doses of corticosteroids Tx.
- ▶ Certain acute viral infections → stems from lymphocyte redistribution (to lymph nodes & increased adherence to endothelial cells) rather than a decrease in the number.

Adverse effect because in moderate high بترفع عدد الخلايا

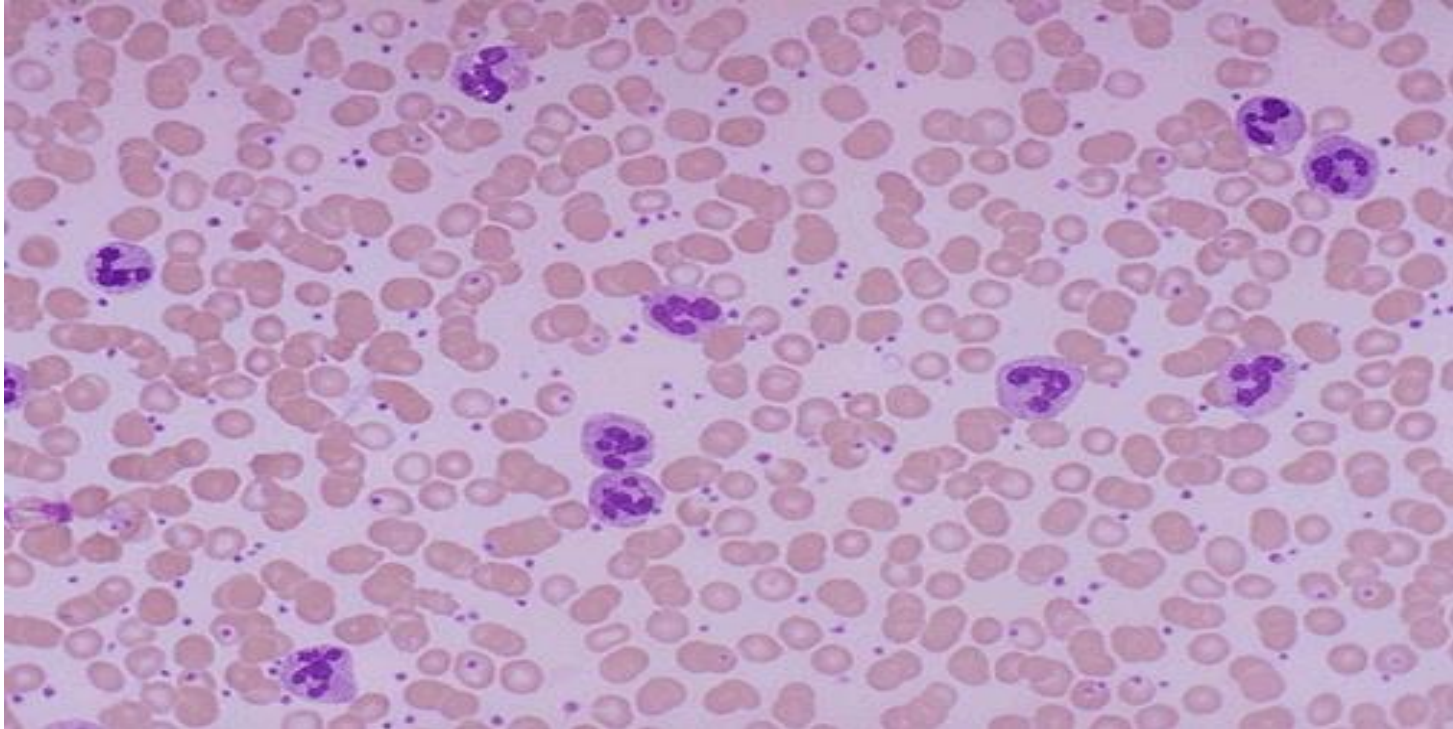
Leukocytosis

Case/All of the Following 

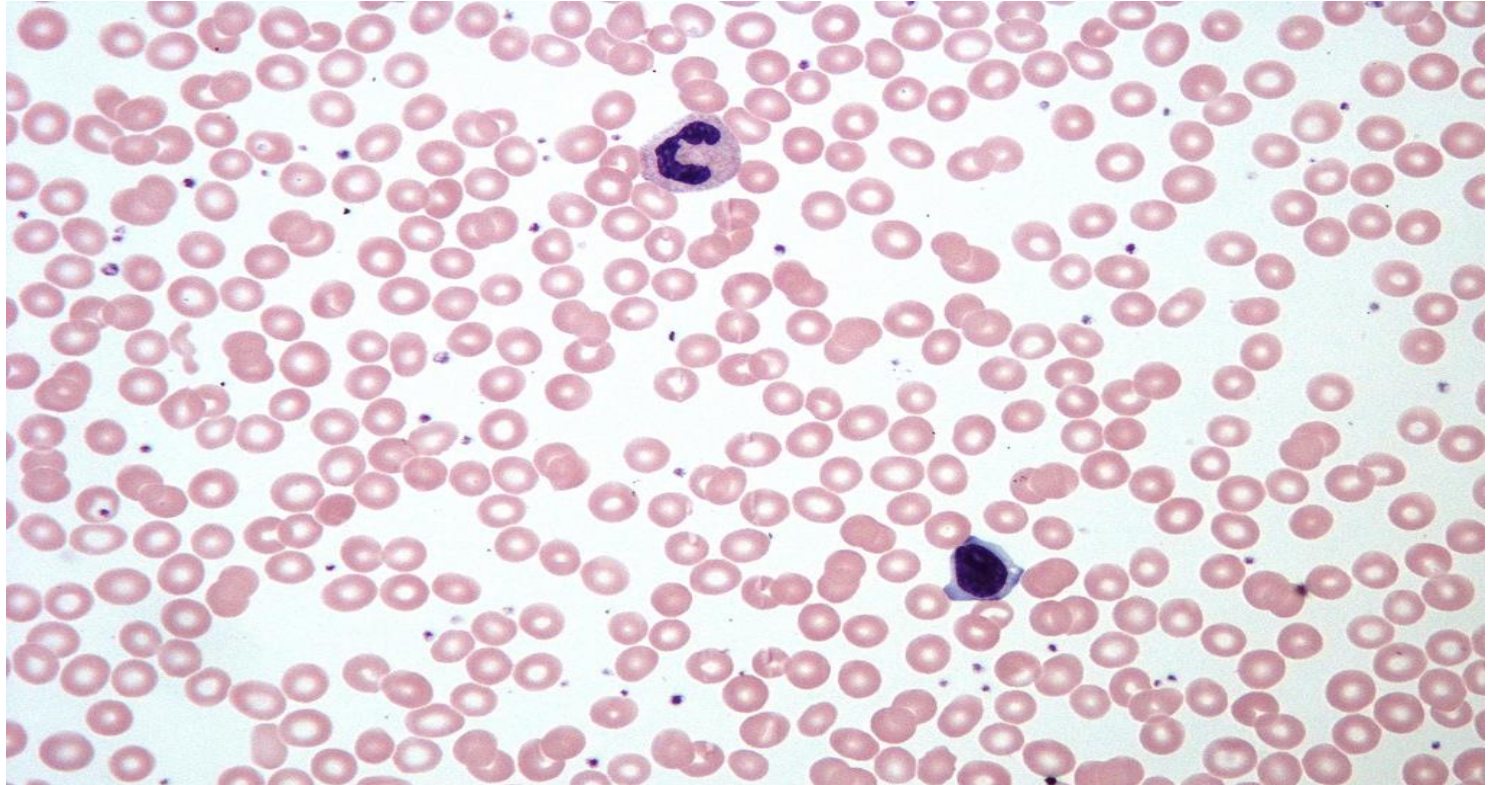
- ▶ Increase in the number of white cells in the blood.
- ▶ A common reaction to a variety of inflammatory states.
- ▶ Leukocytoses are relatively nonspecific and are classified according to the particular white cell series that is affected:
 - 1) **Neutrophilic**: Acute bacterial infections, sterile inflammation caused by tissue necrosis or burn.

Leukocytosis - neutrophilia

↑ neutrophils.
"Neutrophilia".



Normal Blood film



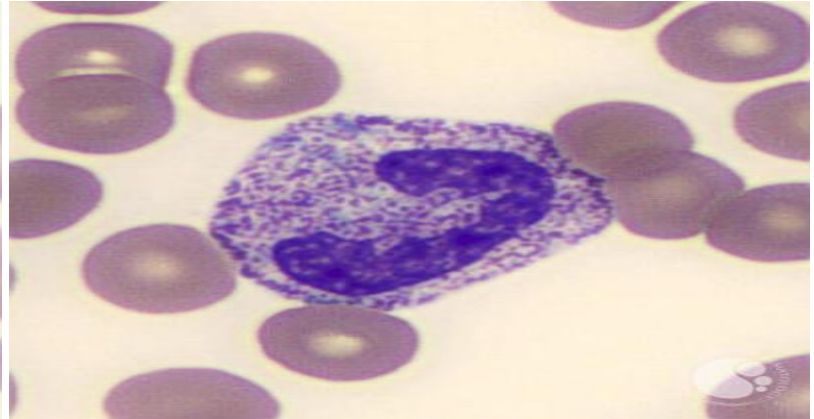
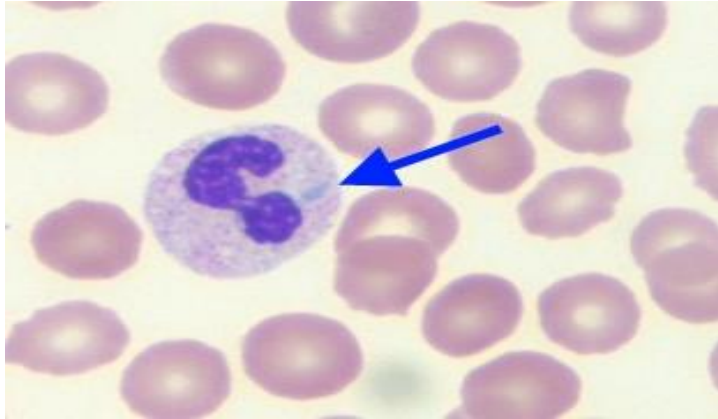
Leukocytosis - neutrophilia

In sepsis or severe inflammation neutrophilia is accompanied by

retention of neutrophils. ← morphologic changes: + cytoplasmic vacuoles → autophagic.

+ Toxic granules, coarser & darker than normal granules

+ **Döhle bodies**: patches of dilated ER (appear → sky-blue cytoplasmic “puddles.”)



Leukocytosis

- ▶ **Eosinophilic:** (eosinophilia) Allergic disorders (e.g., asthma), parasitic infestations or drugs. *للأشياء الغريبة
لبعض الجسم*
- ▶ **Basophilic:** (basophilia) Rare, often indicative of a myeloproliferative disease.
- ▶ **Monocytosis** *ex: giant cells, granuloma.* Chronic infections (e.g., tuberculosis), autoimmune disorders; inflammatory bowel diseases.
- ▶ **Lymphocytosis;** *→ Presently Ag to the immune system* chronic immunologic stimulation (e.g., tuberculosis, brucellosis); viral infections (e.g., HAV, CMV, EBV).

علاج بـ كل سينر

70-80%
of
Leukemia.

Leukemoid reaction

→ like patient in ICU that got infected.

بـ ارتفاع
count

infection

w/ $WBC > 25,000$

Happens in severe infections, many immature granulocytes appear in the blood, mimicking a myeloid leukemia

- ▶ must be differentiated from true white cell malignancies. (e.g., CML);
- + Younger age,
- + No BCR/ABL fusion gene
- + Subsides with treatment of underlying infection

Infectious mononucleosis

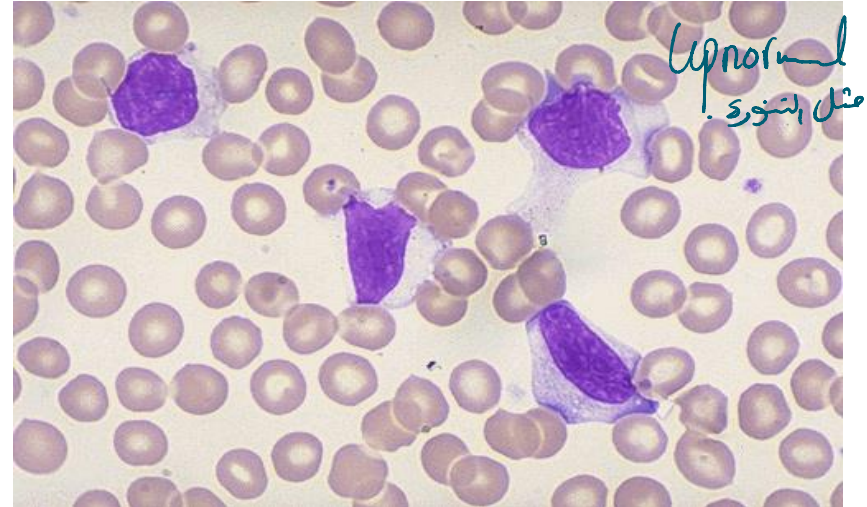
التي تنتج
lymphocyte
But will consider
a infection.

- ▶ Acute, self-limited disease of adolescents & young adults.
- ▶ Caused by Epstein-Barr virus (EBV), herpesvirus family.
- ▶ EBV invades B-cells & cause them to proliferate, cytotoxic (CD8+ T-cells) respond against B-cells.
- ▶ Usually present with: *Transient* ^{مؤقتة}
 1. Fever, sore throat, generalized lymphadenitis
 2. Lymphocytosis of activated CD8+ T cells. (up to 18,000)

خلایا ایمنه
DNA

Infectious mononucleosis

- ▶ More than half of these cells are large **atypical lymphocytes**; with an oval, indented, or folded nucleus & abundant cytoplasm with a few azurophilic granules

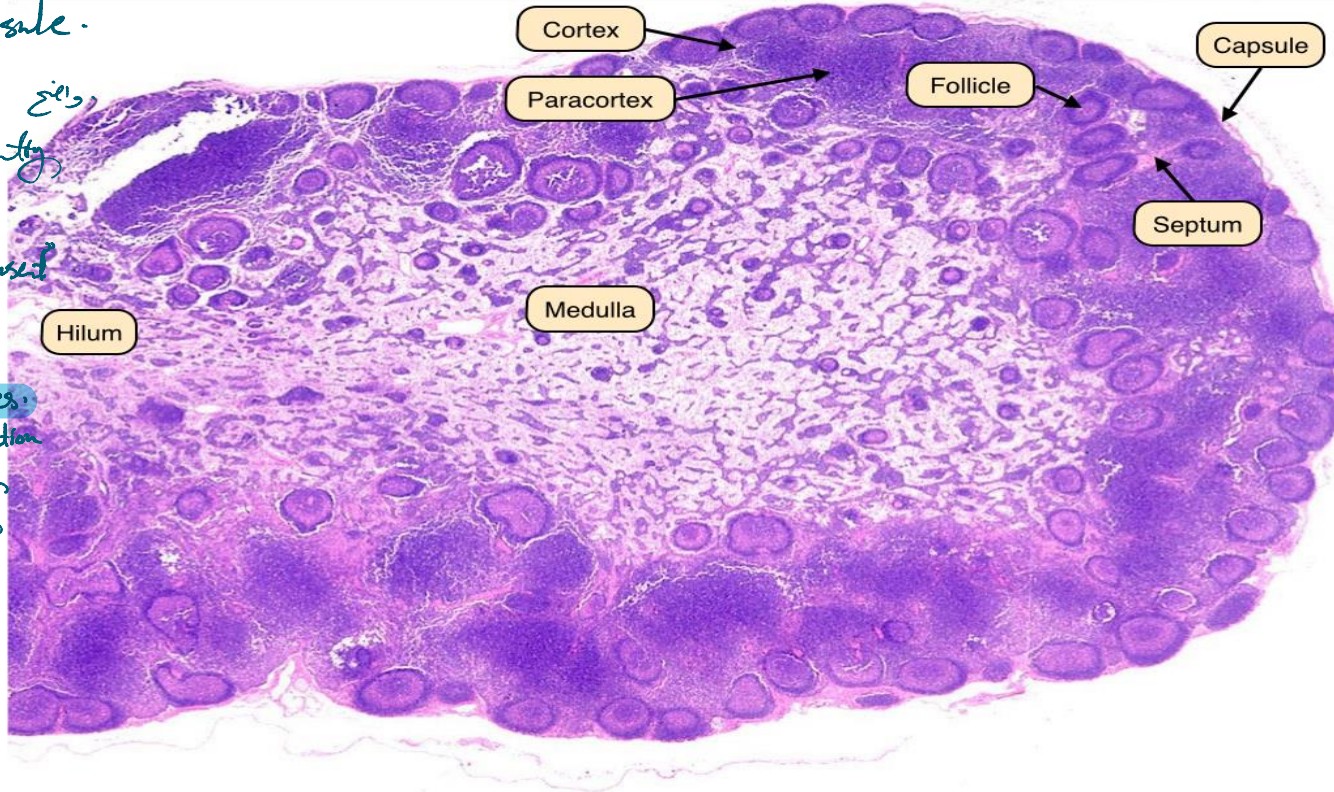


Reactive Lymphadenitis

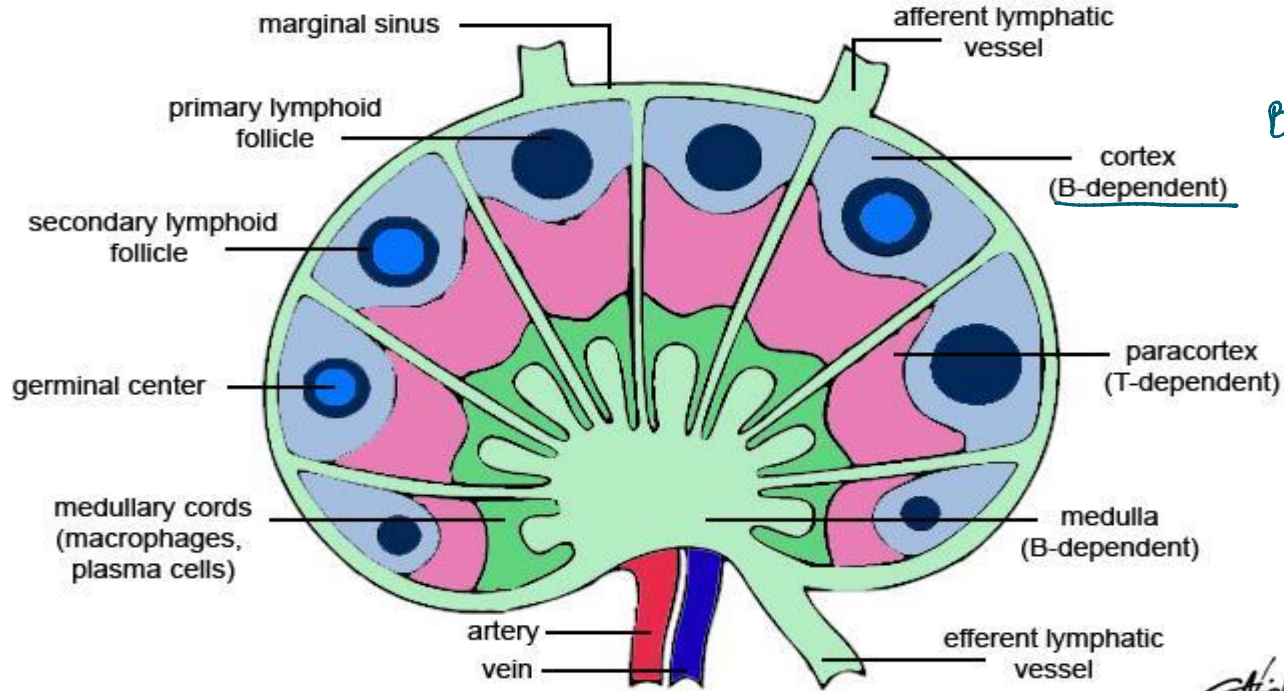
- Architecture preserved
- Thin capsule.
- Cortex *eng*
- Ag. presenting cell.
- primary "closed" follicles.

Secondary follicles:
Somatic hypermutation

"Class shifting"
It's not also
germline center.
↓
Busy area



Reactive Lymphadenitis



هم نفوذ کلا B/T
وین بترکز

E. Vigliani

Reactive Lymphadenitis

Acute Nonspecific Lymphadenitis *if painful not cancer*

- Can be:

in the location of infection. ← Localized → draining a local infection.

Generalized → systemic infectious & inflammatory conditions

- Inflamed nodes are swollen & tender. *→ painful in touch.*
- With infection control lymph nodes may revert to a normal appearance or, if damaged, undergo scarring.

*Severe necrosis.
will cause enlargement.*

Reactive Lymphadenitis

Histologic morphology:

- ▶ Large follicles with germinal center formation
ليس لأنه بأماكن محددة.
- ▶ Frequent GC mitoses & macrophages
- ▶ Sinus enlargement with histiocytes
- ▶ Parafollicular neutrophils, necrosis and possible pus formation (If pyogenic microbes)

In Bacterial infection
↓
neutrophils تكون
& Pus

Reactive Lymphadenitis

Chronic Nonspecific Lymphadenitis

- ▶ Depending on the causative agent, chronic nonspecific lymphadenitis can assume one of three patterns:
 - 1) Follicular hyperplasia. B-cells
 - 2) Paracortical hyperplasia. T-cells
 - 3) Sinus histiocytosis. Macrophages

Reactive Lymphadenitis

Chronic Nonspecific Lymphadenitis - Follicular hyperplasia.

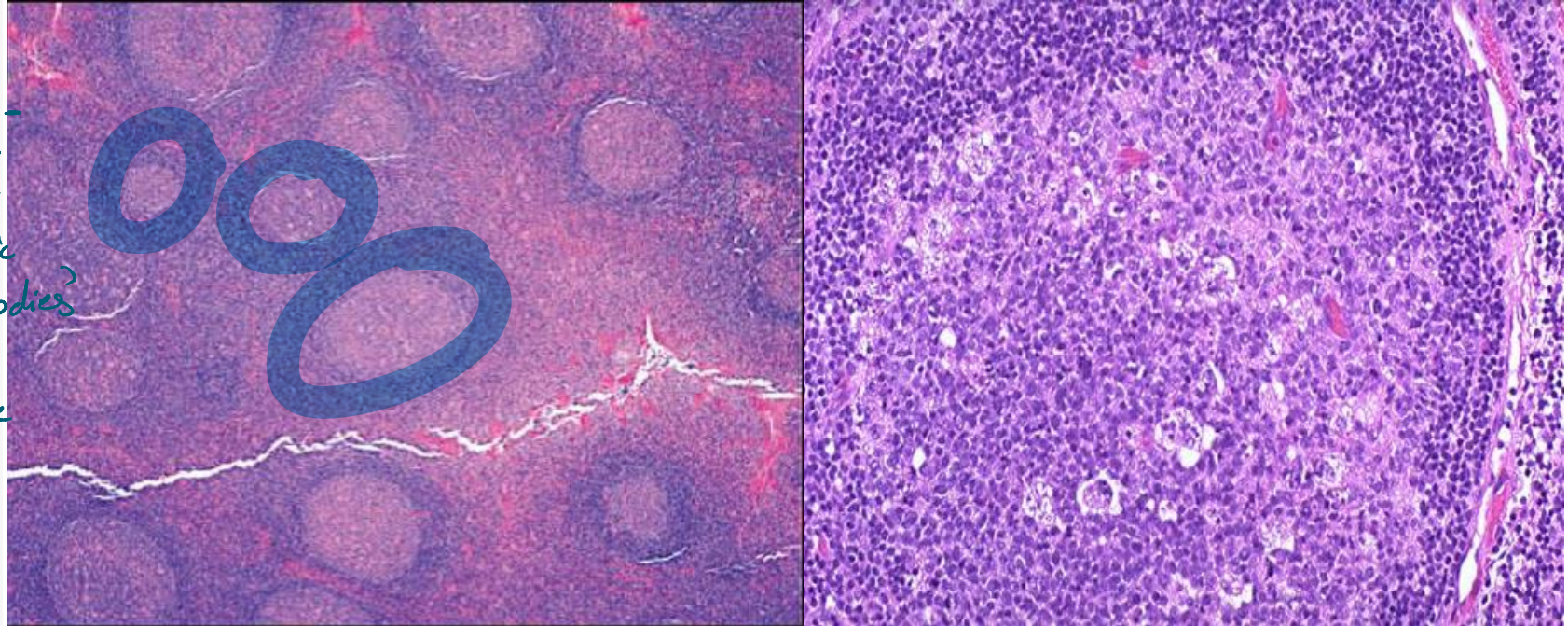
- ▶ Stimuli that activate humoral immune responses.
- ▶ defined by the presence of large germinal centers (secondary follicles), With tingible-body macrophages
- ▶ Causes include rheumatoid arthritis, toxoplasmosis, & early stages of infection with HIV

بمنطقة zone
عن آخر .

Anti-B
بدون ليا يكون
رابطه جرميه germinal

Reactive Lymphadenitis

Chronic Nonspecific Lymphadenitis - Follicular hyperplasia.



- محل واده
- حبيبه
- شكل

- Apoptotic
cells 'bodies'

- T cell
element
there -

Reactive Lymphadenitis

Chronic Nonspecific Lymphadenitis - Follicular hyperplasia.

This form of hyperplasia is morphologically similar to follicular lymphoma, Features favoring a reactive (nonneoplastic) hyperplasia include:

- (1) preservation of the lymph node architecture.
- (2) Variation in the shape and size of the follicles.
- (3) Frequent GC mitotic figures & phagocytic macrophages, & recognizable light and dark zones. (absent in neoplastic follicles)

الخصائص التي تميزها عن الليمفوما

elderly
→ capsule
→ Cortex
→ Medulla

Reactive Lymphadenitis

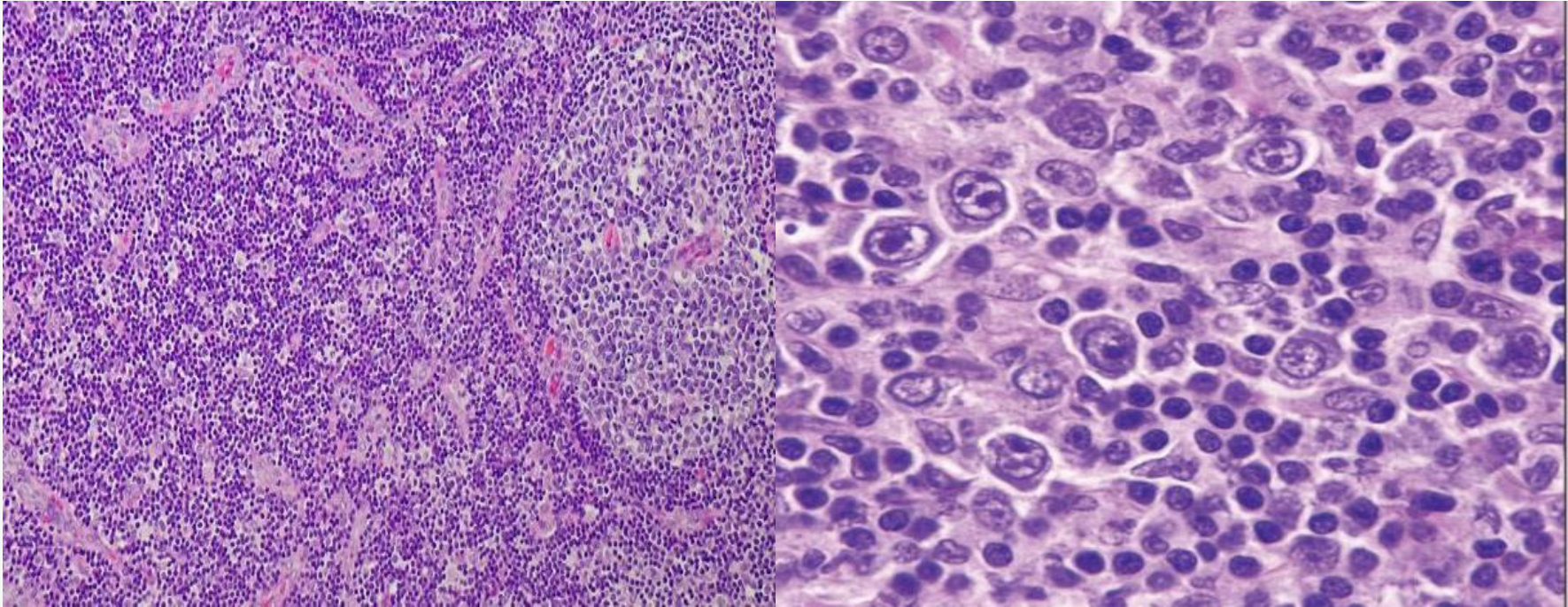
Chronic Nonspecific Lymphadenitis - Paracortical hyperplasia

- ▶ Caused by immune reactions involving the T cell regions.
- ▶ Activated parafollicular T cells transform into large proliferating immunoblasts → that efface B cell follicles.
- ▶ Encountered in:
 - 1) viral infections → activate paracortex.
 - 2) vaccinations (e.g., smallpox).
 - 3) Drugs induced immune reactions (phenytoin)

Reactive Lymphadenitis

فوليكولاته غير متغيرة → ليس
يتغير ولا تضخم.
ليس التفتت بينهم! - expansion

Chronic Nonspecific Lymphadenitis - Paracortical hyperplasia



Reactive Lymphadenitis

Specialised in BV.

Chronic Nonspecific Lymphadenitis - Sinus Histiocytosis

- ▶ Distention and prominence of the lymphatic sinusoids,
due to:
 - 1) Marked hypertrophy of lining endothelial cells.
 - 2) An infiltrate of macrophages (histiocytes).
 - ▶ In lymph nodes draining cancers. *Product of cancer
sinus expansion* بنتشر
 - ▶ Represent an immune response to the tumor or its products. *foreign material
like tattoo.*

Reactive Lymphadenitis

Chronic Nonspecific Lymphadenitis - Sinus Histiocytosis

