

HLS - Final

① ITP: IgG against GPIIb/IIIa } Acquired → Acute: children
 → chronic: Adults F > M

② Heparin-Induced thrombocytopenia: IgG against PF-4 } Acquired

③ TTP: ADAMST 13 (vWF) } Genetic
 ④ HUS: Shiga-toxin "E. coli" } Acquired (children) } Schistocytes

⑤ Glanzmann: Def. of GPIIb/IIIa receptors } Autosomal recessive
 (CD41/CD61 complex) } AI "acquired" } flow cytometry

⑥ Bernard Soulier: Def. of Gb1B (CD42b) } Autosomal recessive } flow cytometry

⑦ vWD: ↓↓ vWF ⊕ ↓↓ Factor VIII } Autosomal Dominant } prolonged PTT only

⑧ Hemophilia A: ↓↓ Factor VIII or Non-functional } X-Linked } *No skin petechia
 *prolonged PTT
 *specific assay test } Mixing study

⑨ Hemophilia B: ↓↓ Factor IX } X-Linked } prolonged PTT
 Factor assay test } Mixing study

⑩ DIC: consumption of clotting factors } Acquired } prolonged PT & PTT ⊕ schistocytes

⑪ HLH:

- Type 1: Homozygous in PRF1 / Infants & children
- Type 2: X-Linked SLAM protein / Adults / EBV
- Type 3: Heterogenous in CD8 + T-cells
- Type 4: T-cell lymphoma

WBC's Disorders

① ALL: *mutations in transcription factors* involved in maturation ⊕ RAS & Tyrosine kinases / Flow cytometry

- ↳ B-ALL: PAX5 / Hyperdiploidy
 - ↳ Children: t(12;21) ETV6 ⊕ RUNX1
 - ↳ Adults: t(9;22) ABL ⊕ BCR
- ↳ T-ALL: NOTCH1 / PTEN ⊕ CDKN2A] CD34, tD ⊕ CD2-8 ⊕ CD10
 - CD34, tD
 - ⊕ CD19, 20, 22
 - ⊕ CD10

② AML: RAS & Tyrosine kinase & P53 & mutated IDH / Auer Rods] → CD34, MPO ⊕ CD13, 33
"Anemic ⊕ Neutropenic ⊕ Thrombocytopenic"

- ↳ A APL: t(15;17) / Auer Rods] → ⊖ for CD34
 PML → RARA / "8" nuclei
- ↳ B MDS: chromosomal aberration
 - ↳ Monosomy 5
 - ↳ Monosomy 7
 - ↳ Deletion of 5q, 7q, 20q
 - ↳ Trisomy 8

⊕ Mutation in DNA methylation & Histone modif. ⊕ P53 ⊕ RNA splicing factor] old age
 ↓
 RNA Sideroblasts "سحب الحديد"]

③ CLL: ↑ Bcl2 ⊕ *Rare translocation of chromosomes* ⊕ Active BCR ↓ Active Bruton Tyrosine Kinase (BTK)

⊕ Bcl2 ⊕ CD19, 20 ⊕ CD5] *Smudge cells "old age" *Richter transformation to DLBCL (prolymphocyte)

④ Hairy Cell Leukemia: *Serine/threonine kinase (BRAF gene)*] → old age ⊕ prominent cytoplasmic projections ⊕ BM fibrosis

⑤ Adult T-cell leukemia/lymphoma: Tax protein \oplus
 PI3 kinase & cyclin D \ominus
 \downarrow
 CDK inhibitor \oplus \rightarrow NF- κ B

] Hypocalcemia
 &
 CD 25 \oplus

⑥ Mycosis fungoides: cerebriform \oplus Sezary syndrome
 nucleus

⑦ peripheral T-cell lymphoma: CD 2, 3, 5, 7

⑧ MPN: Tyrosine kinase / Extramedullary hem.

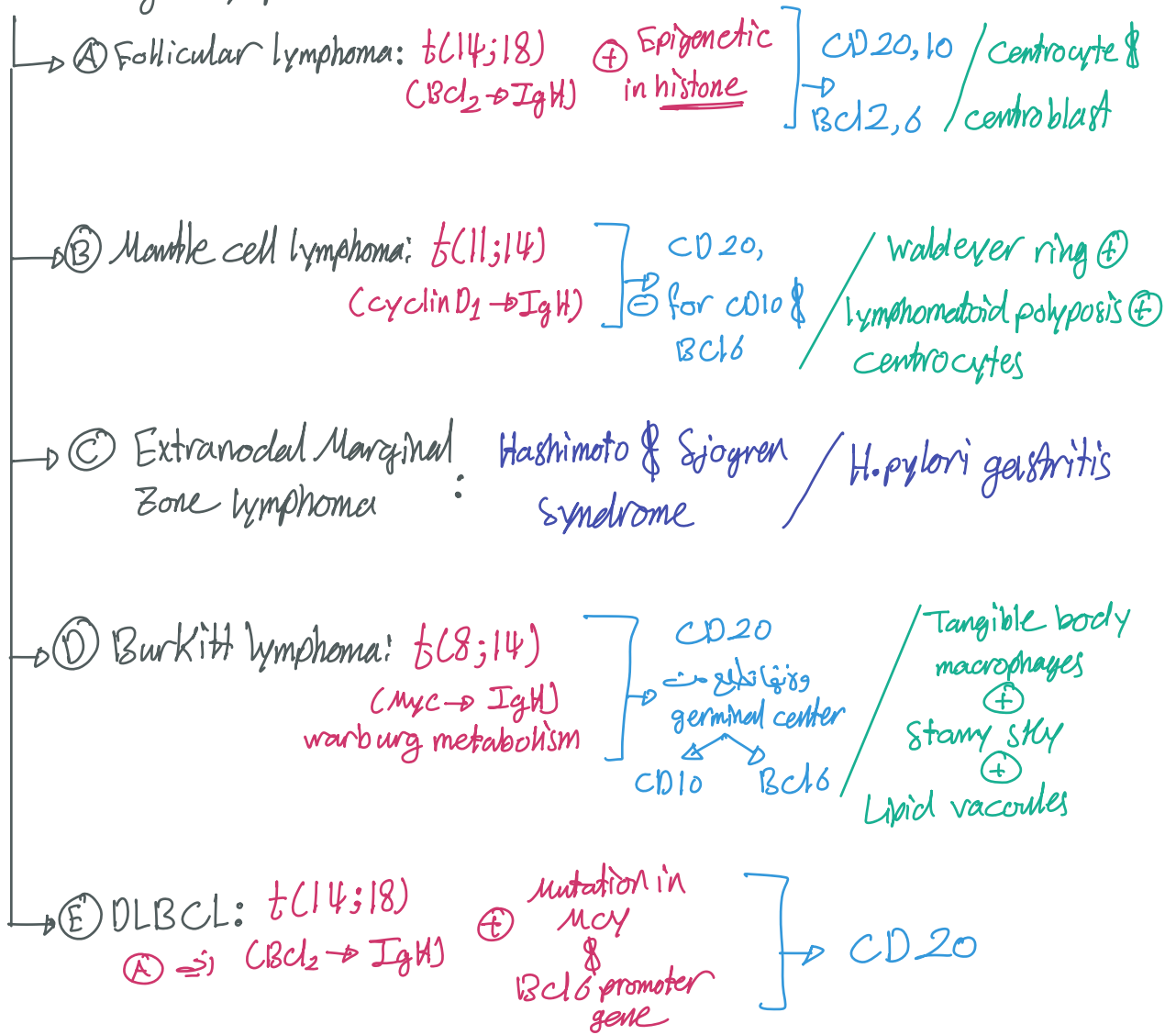
\rightarrow ① CML: Harbor (4; 22) BCR/ABL \rightarrow New Tyr. kin.] \rightarrow Basophilia

\rightarrow ② polycythemia vera: Tyrosine kinase JAK2] \rightarrow panmyelosis \oplus \downarrow Epo \oplus Basophilia

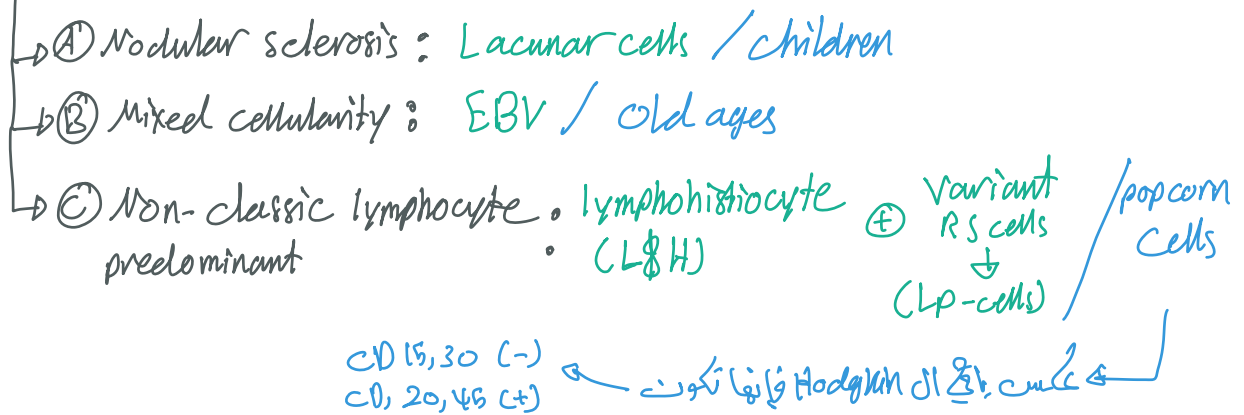
\rightarrow ③ primary myelofibrosis: JAK-STAT pathway \oplus MPL gene \oplus JAK2] \rightarrow Tear Drop \oplus Leucoerythroblastic Anemia

\rightarrow ④ Essential thrombocythemia: JAK2 mutation

④ Non-Hodgkin lymphomas:



① Hodgkin lymphoma: RS cells → CD15, 30 (+) / PD / IL-5
CD 3, 20, 45 (-)



① plasma cell myeloma, t(Ch;14) or Myc
 (Multiple Myeloma) • Cyclin D1 or D3 → Igh

* ↑↑ ESR
 * CRAB
 * Amyloidosis
 * RANCL
 * IL-6

Bence Jones protein (+)
 Rouleaux formation (+)
 cartwheel chromatin

② LCH: CD1a & Langerin → Birbeck granules "Tennis Racket shape" → mutation in serine/threonine kinase (BRAF)

④ Multisystemic LCH: osteolytic (+) pulmonary lesions / children

③ Unisystemic LCH: Eosinophilic granuloma / Diabetes (+) Exophthalmos (+) osteolytic lesion

Hamel-Schuller-Christian Triad

Notes

* X-Linked Hemophilia A, B / HLH Type-2

* Schistocytes: TTP, HUS, DIC

* pancytopenia: HLH / MDS / Hairy cell leukemia / MDS / myelofibrosis / plasma cell myeloma / multisystem LCH

* EBV: HLH-2 / Burkitt lymphoma / DLBCL / Mixed cellularity

* Thrombocytopenia: ALL, AML, MDS, CLL, Accelerated phase of CML

* Cytosis: MPN

* Anemia: ITP, DIC, ALL, MDS, plasma cell myeloma / multisys. LCH / ~~AML~~ / primary myelofib.

TTP
 AML
 CLL

* Leukocytosis: CLL, MPN

- Good Luck -