














# ***Cracking the Code***

## ***CBC Interpretation in Children and Adolescents***

***A Case-Based and Evidence-Based Teaching Approach***

***Prof. Mohammad Faranoush***  
***Iran University of Medical Sciences***

## CBC

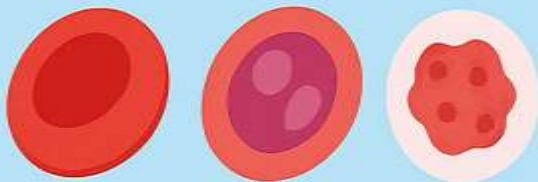
WBC	8.1	
RBC	4,76	
HQB	13.5	
MCV	84	
MCH	23.4	
RDW	33.8	
PLT	255	
MPV	9.1	
NEUT%	56	
LYMPH%	34	
MONO%	6	
EO%	3.5	
BASO%	0,5	

# COMPLETE BLOOD CUNT (CBC) INTERPRETATION IN CHILDREN AND ADOLESCENTS

Understanding Pediatric Hematology through Cases

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June 2025 | Pediatric Hematology Seminar Series

CBC



Interactive & Case-Based

# *Objectives*

- Interpret pediatric CBCs across age groups
- Recognize patterns of anemia, leukocyte abnormalities, and thrombocytopenia
- Apply systematic reasoning through clinical cases

# ***PRACTICE GAPS***

- Interpretation of the complete blood count (CBC) is not a simple task.
- Laboratory reports include a variety of useful values that may be overlooked.
- It is important for all practitioners to develop an adequate foundation in CBC interpretation early in their career.
- Possessing a thorough understanding of the results reported in the CBC allows the practitioner to focus subsequent evaluation and reduce unnecessary laboratory testing.

# *Introduction*

## CBC...

- is one of the most common laboratory tests in medicine.

Typically, it includes the following:

- *White blood cell count (WBC or leukocyte count)*
- *WBC differential count*
- *Red blood cell count (RBC or erythrocyte count)*
- *Hematocrit (Hct)*
- *Hemoglobin (Hb)*
- *Mean corpuscular volume (MCV)*
- *Mean corpuscular hemoglobin (MCH)*
- *Mean corpuscular hemoglobin concentration (MCHC)*
- *Red cell distribution width (RDW)*
- *Platelet count*
- *Mean Platelet Volume (MPV)*

# ***The Code***

## ***Why Pediatric CBCs Are Unique***

- ***Age-Specific Reference Ranges***
- ***Physiological Processes***
  - Anemia of prematurity
  - Physiologic anemia of infancy
  - Benign neutropenia of childhood (especially certain ethnicities)
  - Stress leukocytosis
- ***Common Benign Variations***
  - Benign ethnic neutropenia
  - Transient reactive changes (lymphocytosis, thrombocytosis)

# *Cracking the Code*

## *The Systematic Approach*

- *Step 1:*

- Verify & Contextualize: Is the sample valid (clots, hemolysis)?
- MOST CRITICAL: What is the patient's age? What is the clinical context (reason for test, symptoms, signs, history, medications, family history, ethnicity)?

- *Step 2:*

- Scan for Critical Values: Blasts? Severe pancytopenia? Severe anemia/thrombocytopenia? Extreme leukocytosis? (Flag these immediately).

## *Step 3*

- *Analyze Line-by-Line with Age in Mind:*
  - **RBC Line:**Anemia (Hb/Hct)? Is it microcytic (MCV low), normocytic, macrocytic (MCV high)? RDW? Reticulocyte count? (Think Iron Deficiency, Thalassemia, Hemolysis, Chronic Disease, B12/Folate).
  - **WBC Line:**Total WBC high/low? Differential is KEY: Neutrophils (ANC), Lymphocytes (ALC), Monocytes, Eosinophils, Basophils, Bands? Any atypical/abnormal cells? (Think Infection [bacterial, viral, parasitic], Inflammation, Allergy, Stress, Leukemia/Lymphoma, Benign Ethnic Variation).
  - **Platelet Line:**Thrombocytosis? Thrombocytopenia? (Think Reactive, ITP, Infection, Malignancy, Bone Marrow Failure, Artifact).



## *Step 4*

- *Synthesize:*
  - Does the picture fit the clinical context? Is it a single-line issue or multi-line? Is it likely reactive/benign or concerning for a primary hematologic disorder?

## *Step 5*

- *Next Steps:*

- Repeat test? Further investigations (peripheral smear, iron studies, B12/folate, LDH, haptoglobin, Coombs, flow cytometry, bone marrow)? Referral? Observation?

***PART 1***  
***AGE-SPECIFIC CBC REFERENCE RANGES***  
***EVIDENCE-BASED REVIEW***

## *WBC Count by Age*

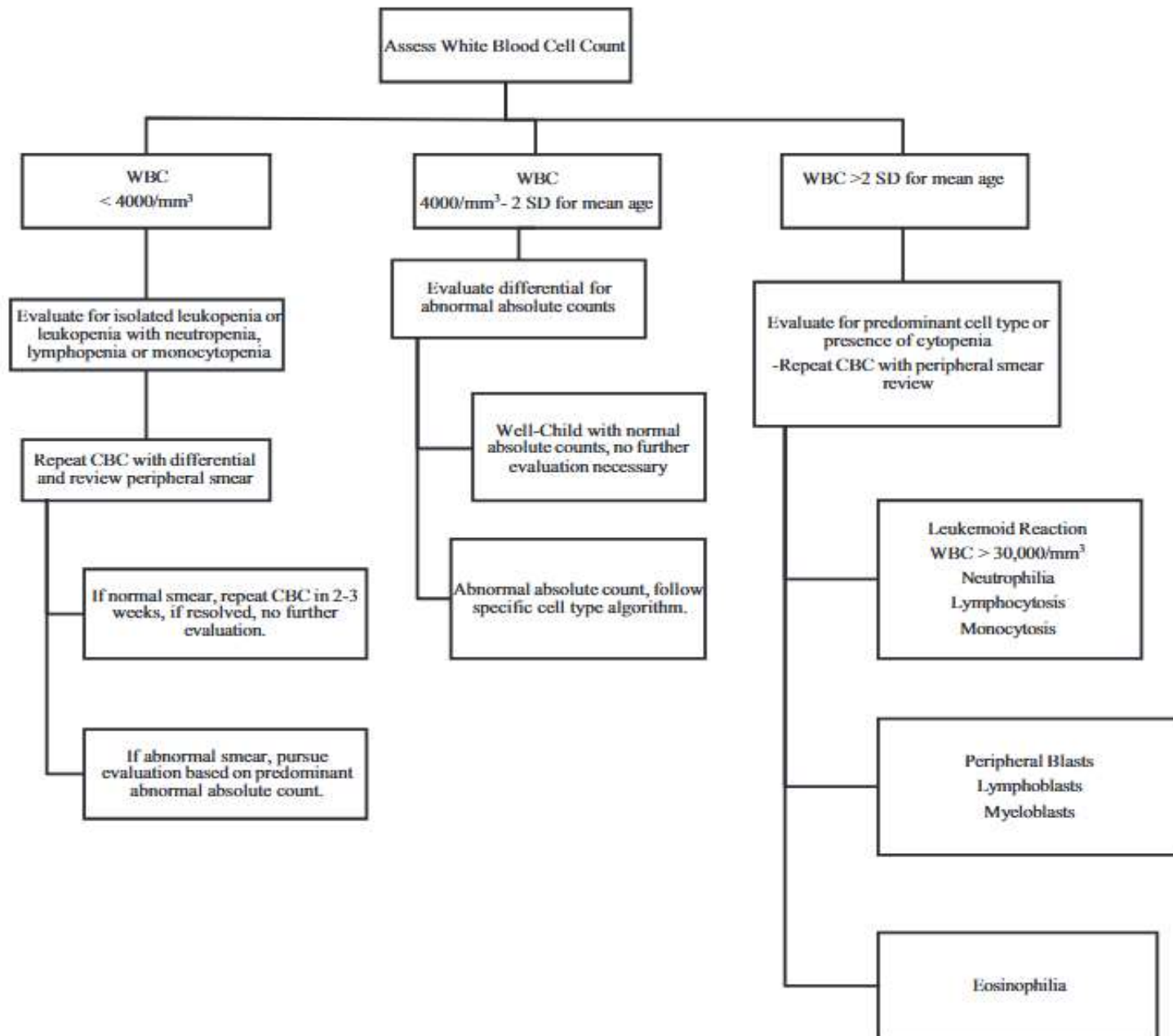
- Neonates:  $9\text{--}30 \times 10^9/\text{L}$  (transient leukocytosis at birth)
- Infants (1–12 months):  $6\text{--}17 \times 10^9/\text{L}$
- Children (1–10 years):  $5\text{--}14.5 \times 10^9/\text{L}$
- Adolescents:  $4.5\text{--}13.5 \times 10^9/\text{L}$
- Note: Differential varies—neutrophil predominance after infancy

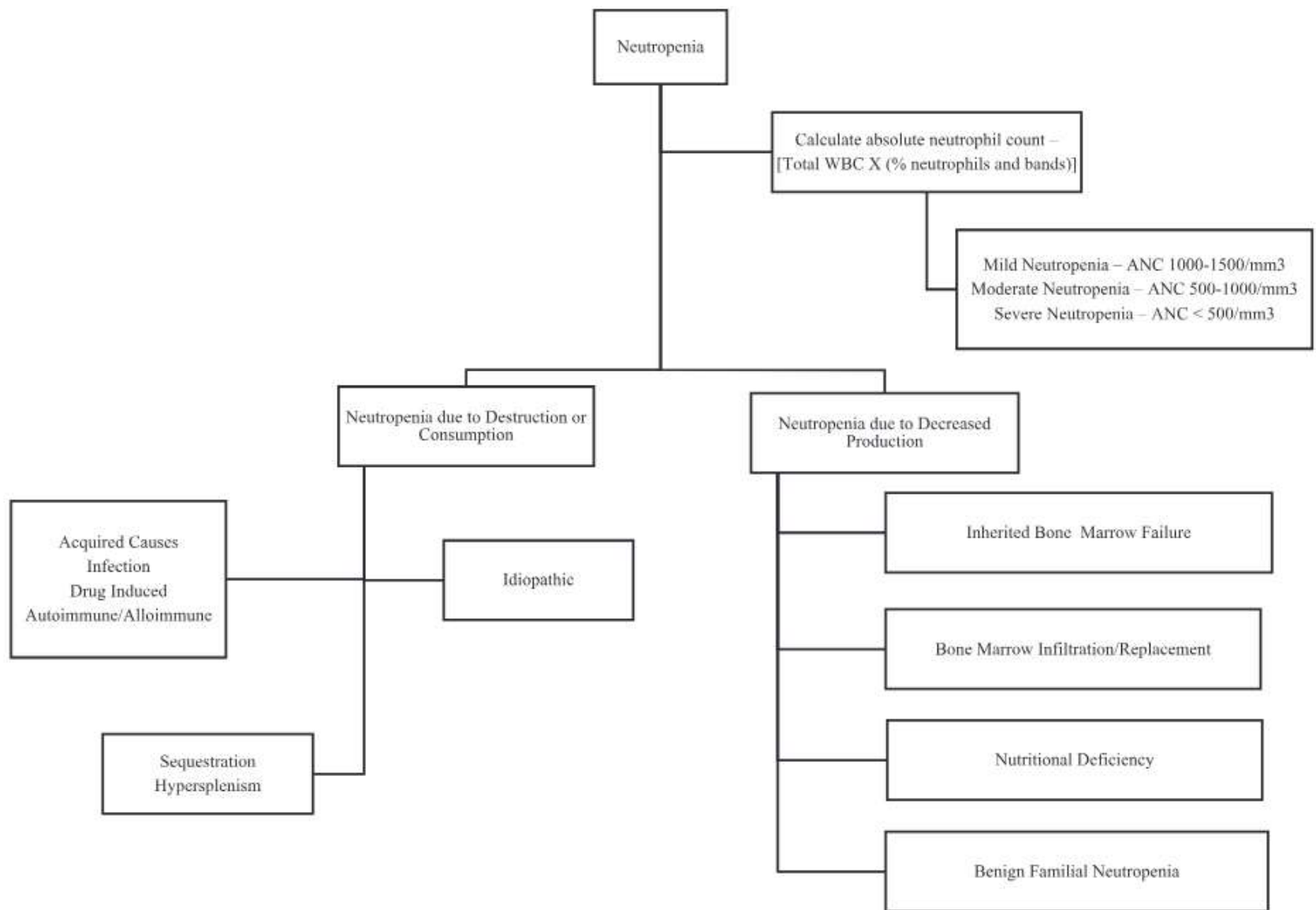
**Table 3.** WBC Parameters In the Pediatric Population

Age	WBC Count, 10 <sup>3</sup> /μL		Neutrophil Count, 10 <sup>3</sup> /μL			Lymphocyte Count, 10 <sup>3</sup> /μL			Eosinophil Count, 10 <sup>3</sup> /μL		Monocyte Count, 10 <sup>3</sup> /μL	
	Mean	Range	Mean	Range	%	Mean	Range	%	Mean	%	Mean	%
Birth	18.1	9.0–30.0	11.0	6.0–26.0	61	5.5	2.0–11.0	31	0.4	2	1.1	6
12 h	22.8	13.0–38.0	15.5	6.0–28.0	68	5.5	2.0–11.0	24	0.5	2	1.2	5
24 h	18.9	9.4–34.0	11.5	5.0–21.0	61	5.8	2.0–11.5	31	0.5	2	1.1	6
1 wk	12.2	5.0–21.0	5.5	1.5–10.0	45	5.0	2.0–17.0	41	0.5	4	1.1	9
2 wk	11.4	5.0–20.0	4.5	1.0–9.5	40	5.5	2.0–17.0	48	0.4	3	1.0	9
1 mo	10.8	5.0–19.5	3.8	1.0–9.0	35	6.0	2.5–16.5	59	0.3	3	0.7	7
6 mo	11.9	6.0–17.5	3.8	1.0–8.5	32	7.3	4.0–13.5	61	0.3	3	0.6	5
1 y	11.4	6.0–17.5	3.5	1.5–8.5	31	7.0	4.0–10.5	61	0.3	3	0.6	5
2 y	10.6	6.0–17.0	3.5	1.5–8.5	33	6.3	3.0–9.5	29	0.3	3	0.5	5
4 y	9.1	5.5–15.5	3.8	1.5–8.5	42	4.5	2.0–8.0	50	0.3	3	0.5	5
6 y	8.5	5.0–14.5	4.3	1.5–8.0	51	3.5	1.5–7.0	42	0.2	3	0.4	5
8 y	8.3	4.5–13.5	4.4	1.5–8.0	53	3.3	1.5–6.8	39	0.2	3	0.4	4
10 y	8.1	4.5–13.5	4.4	1.8–8.0	54	3.1	1.5–6.5	38	0.2	2	0.4	4
16 y	7.8	4.5–13.0	4.4	1.8–8.0	57	2.8	1.2–5.2	35	0.2	3	0.4	5
21 y	7.4	4.5–11.0	4.4	1.8–7.7	59	2.5	1.0–4.8	34	0.2	3	0.3	4

WBC=white blood cell.

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# Lymphocytes

Lymphopenia  
ALC < 1000/mm<sup>3</sup>

Assess patient age and if well or sick

Newborn  
Immunodeficiency  
Syndromes

Infant or Child  
Assess for history of infections, Immunology referral

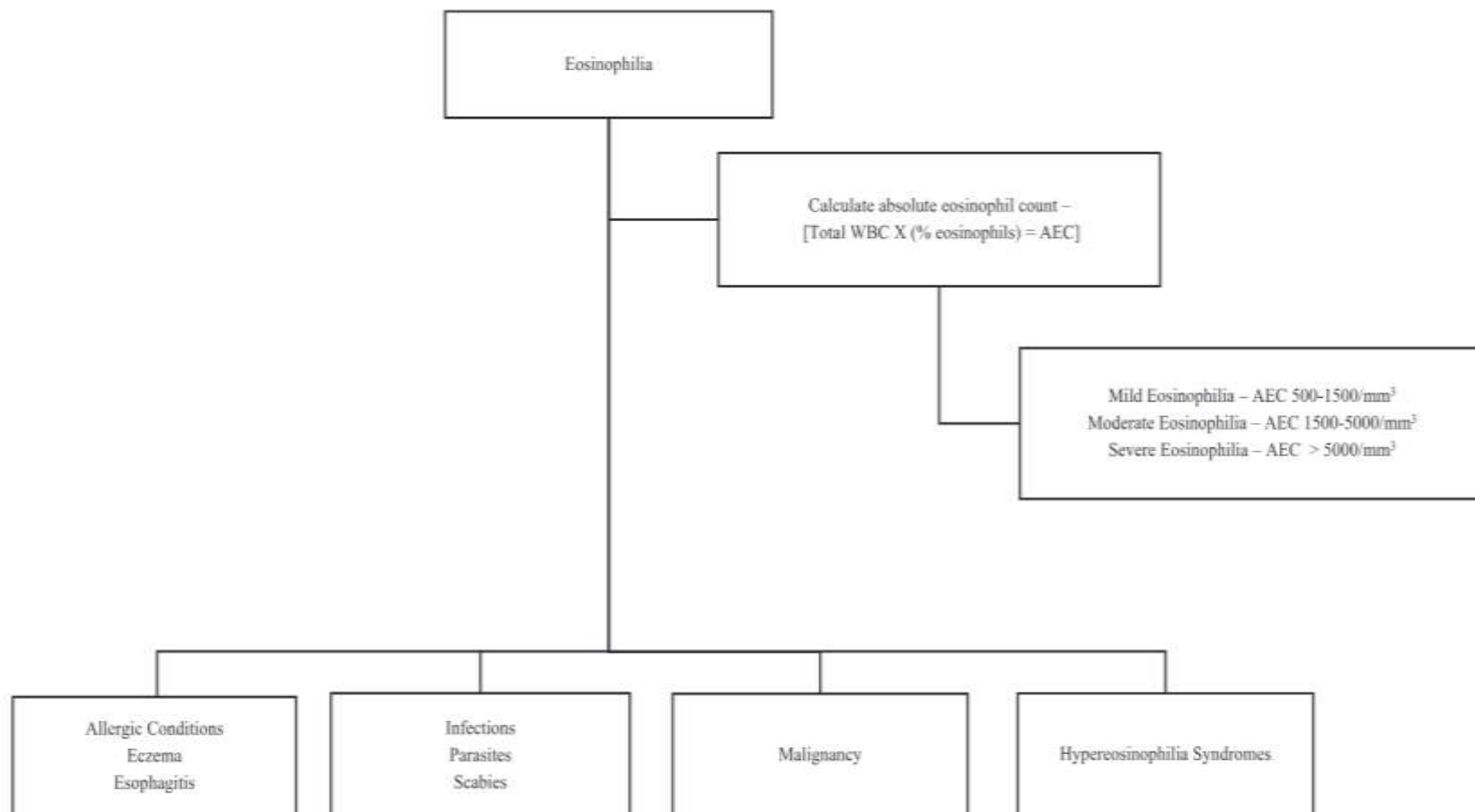
Lymphocytosis  
Review Peripheral Smear

Evaluate for infectious etiology

Other cytopenias (thrombocytopenia or anemia)  
Evaluate for malignancy

Reactive Lymphocytes =  
Infection





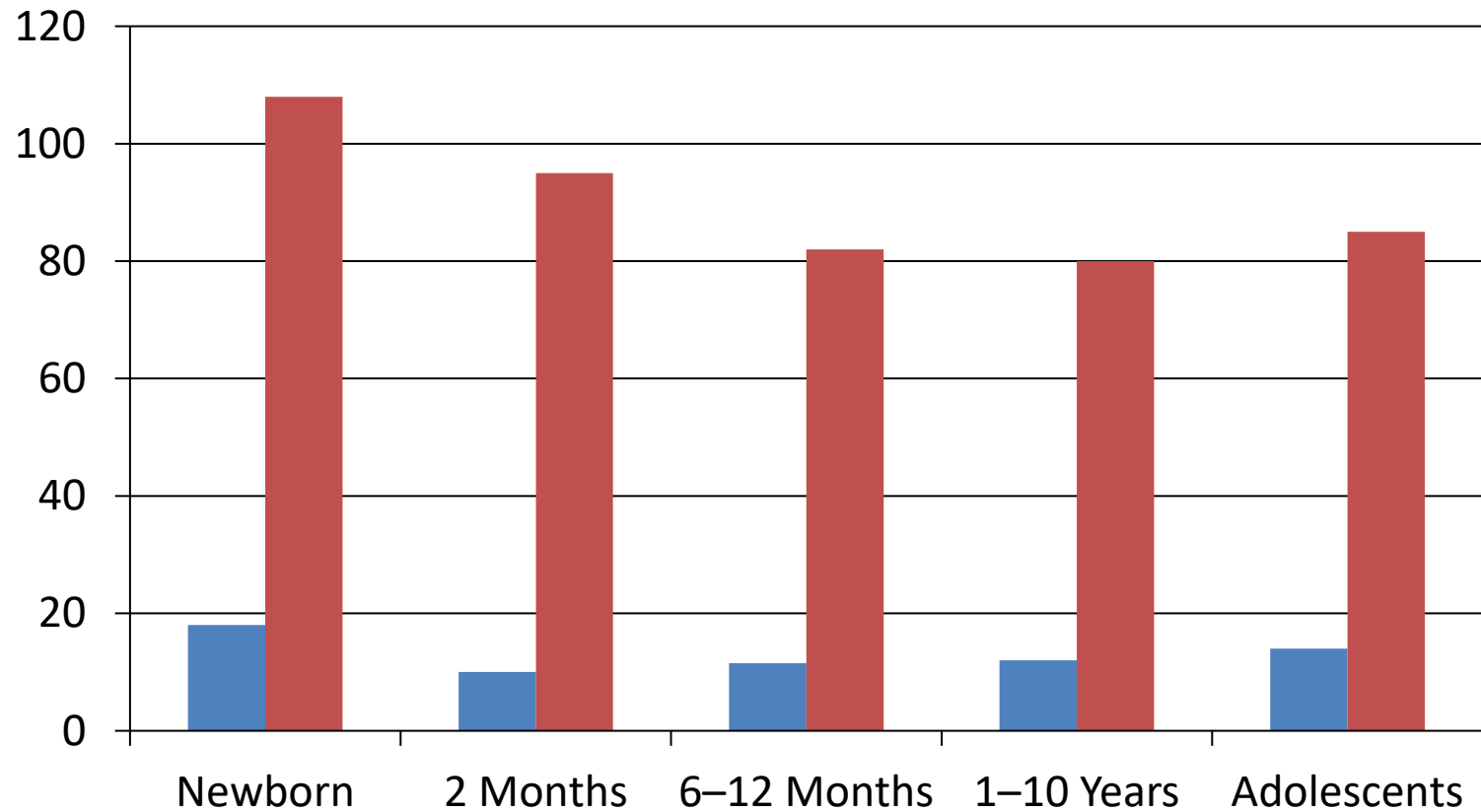
# *Hemoglobin & MCV by Age*

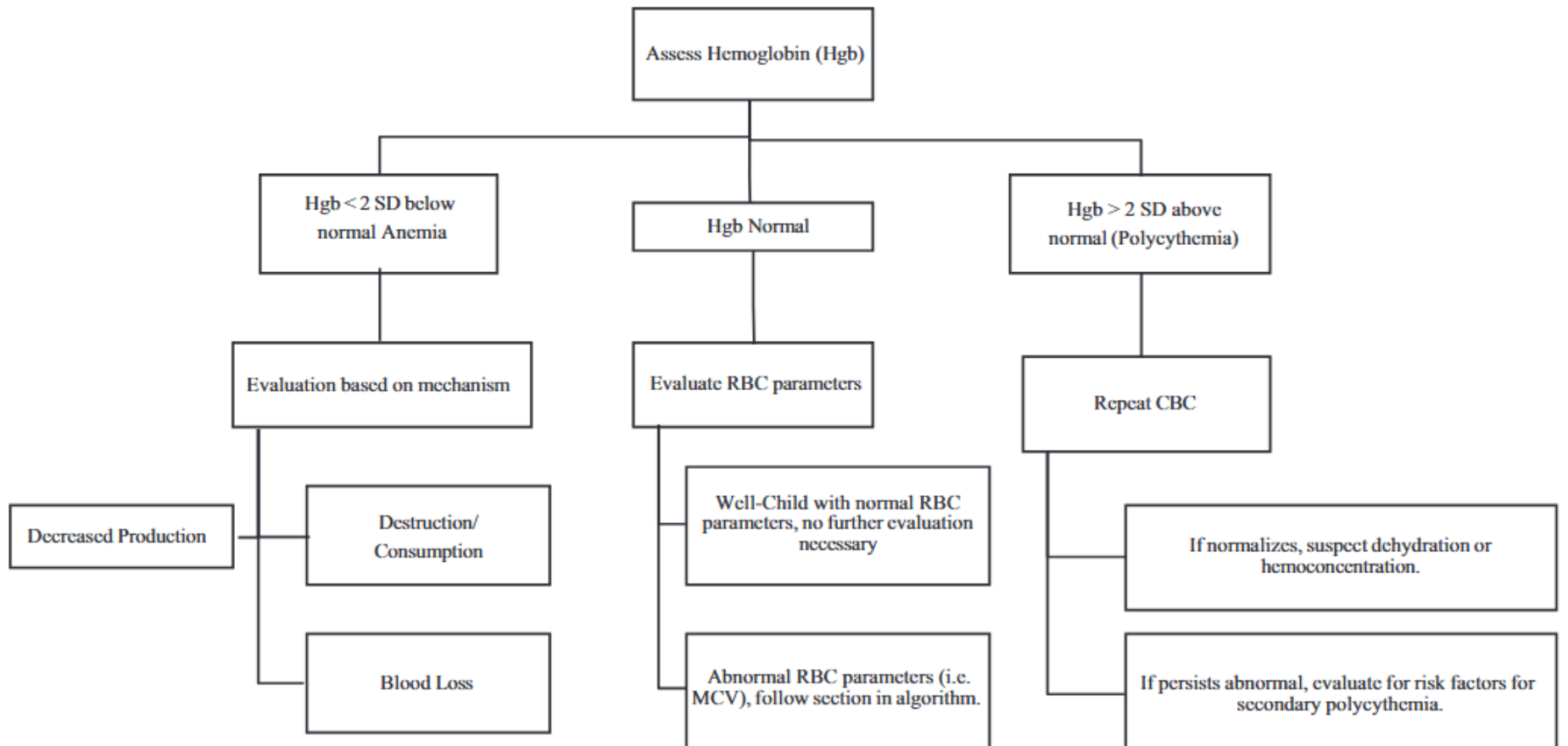
- Neonates: Hb 14–24 g/dL, MCV ~108 fL (physiologic polycythemia)
- 2 months: Hb 9–11 g/dL (physiologic nadir), MCV ~95 fL
- 6–12 months: Hb 10.5–13.5 g/dL, MCV 72–88 fL
- Adolescents: Hb 12–16 g/dL (f), 13–17 g/dL (m); MCV 80–96 fL

# *RBC indices*

<b>Age</b>	<b>Lowest Normal Hemoglobin (g/dL)</b>	<b>Normal RBC Size Mean Corpuscular Volume (fL)</b>	<b>Fetal Hemoglobin (%)</b>	<b>Hemoglobin A2 (%)</b>
Birth	14.0	100–130	55–90	<1
6 months–1 year	11.0	70–85	<5	1.5–3.5
1–4 years	11.0	70–85	<2	1.5–3.5
4 years–puberty	11.5	75–90	<2	1.5–3.5
Postpubertal female	12.0	80–95	<2	1.5–3.5
Postpubertal male	14.0	80–95	<2	1.5–3.5

# Normal Hemoglobin and MCV by Age





Age	Hemoglobin, g/dL		Hematocrit, %		MCV, fL		RDW, % 2 SD	RBC Count, $\times 10^6/\mu\text{L}$ Mean	MCH, pg Mean	MCHC, g/dL Mean	Reticulocytes, % Mean
	Mean	2 SD	Mean	2 SD	Mean	2 SD					
Overall											
1–3 d	18.5	14.5–22.5	56.0	45.0–67.0	108	95–121	13.0–18.0	5.3	34	33	3.0
1 wk	17.5	13.5–19.5	54.0	42.0–66.0	107	88–126	13.0–18.0	5.1	34	33	0.5
2 wk	16.5	12.5–20.5	51.0	39.0–63.0	105	86–124	13.0–18.0	4.9	34	33	0.5
1 mo	14.0	10.0–18.0	43.0	31.0–55.0	104	85–123	11.5–16.0	4.2	34	33	0.8
2 mo	11.5	9.0–14.0	35.0	28.0–42.0	96	77–115	11.5–16.0	3.8	30	33	1.6
3–6 mo	11.5	9.5–13.5	35.0	29.0–41.0	91	74–108	11.5–16.0	3.8	30	33	0.7
0.5–2 y	12.0	10.5–13.5	36.0	33.0–49.0	78	70–86	11.5–16.0	4.5	27	33	1.0
2–6 y	12.5	11.5–15.5	37.0	34.0–45.0	81	75–87	11.5–15.0	4.6	27	34	1.0
6–12 y	13.5	11.5–15.5	40.0	35.0–45.0	86	77–95	11.5–15.0	4.6	29	34	1.0
Female sex											
12–18 y	14.0	12.0–16.0	41.0	33.0–51.0	90	78–102	11.5–14.0	4.6	30	34	1.0
$\geq 18$ y	14.0	12.0–16.0	41.0	33.0–51.0	90	80–100	11.5–13.1	4.6	30	34	1.0
Male sex											
12–18 y	14.5	13.0–16.0	43.0	36.0–51.0	88	78–108	11.5–14.0	4.9	30	34	1.0
$\geq 18$ y	15.5	13.5–17.5	47.0	37.0–53.0	90	80–100	11.5–13.1	5.2	30	34	1.0

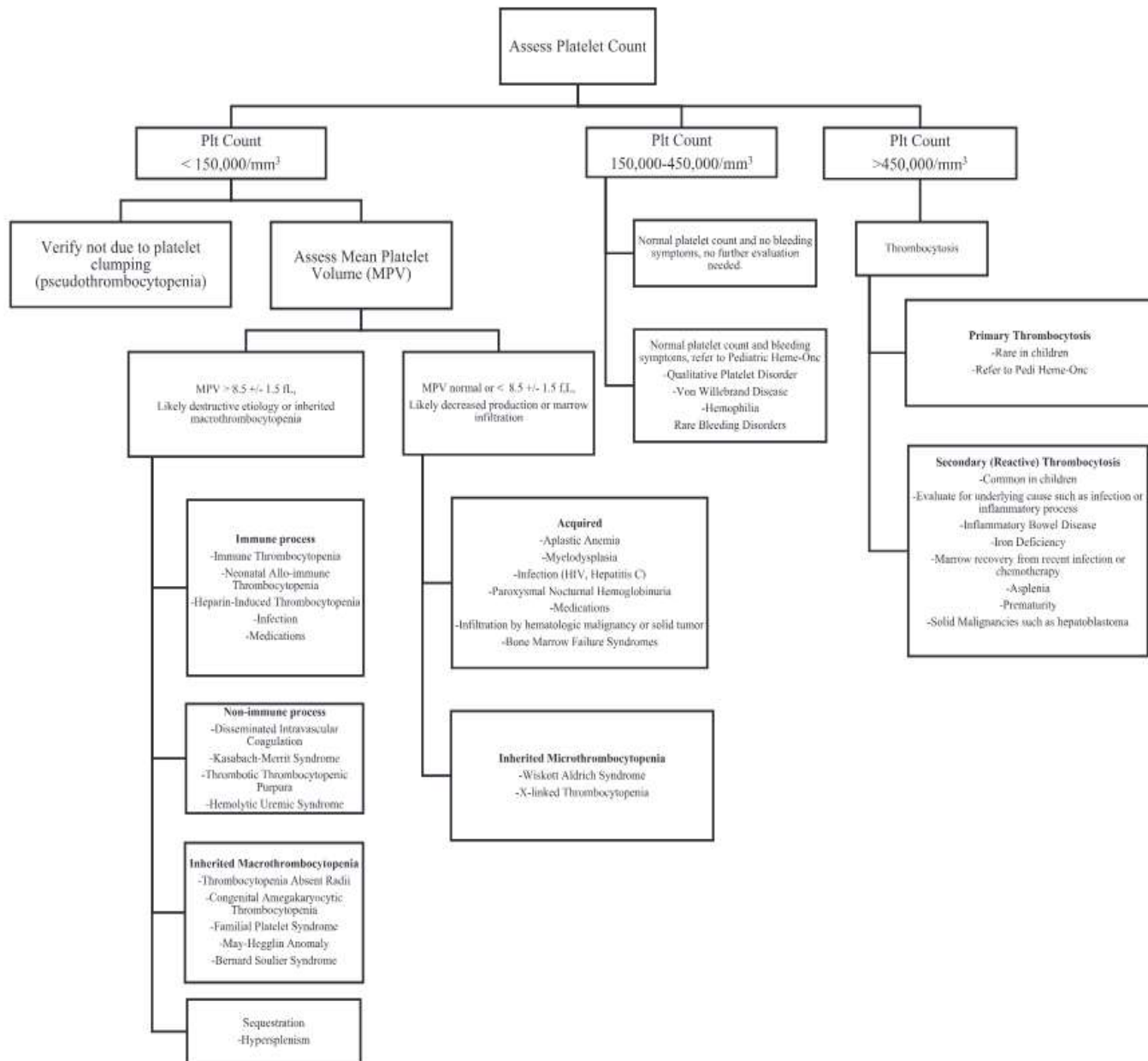
## ***Reticulocyte Count & RDW***

- Reticulocytes: Higher in neonates (up to 4–7%), normalizes to 0.5–2.5%
- RDW: Normal 11.5–14.5%, elevated in iron deficiency or recent recovery

# *Platelet Count by Age*

- All ages: Platelets 150–450  $\times 10^9/L$
- Mild transient thrombocytosis common after infections or iron deficiency
- Neonates may show mild thrombocytopenia due to perinatal factors





***PART 2***  
***CLINICAL CASES***  
***INTERACTIVE TEACHING WITH***  
***DISCUSSION***

## *Case 1 – Toddler with Pallor*

- 2-year-old with pallor, poor appetite, irritable mood
- RED FLAGS

## *Lab Data*

- CBC
- Hb 8.2, MCV 62 fL, high RDW
- WBC/PLT :Normal
- Retic: Low

## *Interactive poll*

- Iron Deficiency Anemia
- Thalassemia trait
- Lead poisoning
- Sideroblastic anemia

# *Diagnosis*

- Iron deficiency anemia

<i>Cause</i>	<i>Key Features</i>
<i>Iron deficiency anemia</i>	<i>Low MCV, increase RDW, low ferritin, poor diet, pica, pallor</i>
<i>Thalassemia trait</i>	<i>Low MCV, normal RDW or increase RBC, family history</i>
<i>Lead poisoning</i>	<i>Microcytic, basophilic stippling, environmental exposure</i>
<i>Sideroblastic Anemia</i>	<i>Rare in children, dimorphic RBCs, ringed sideroblasts on BM</i>



## *Useful Indices*

- RDW: High in iron deficiency; normal in thalassemia trait
- Mentzer Index =  $MCV \div RBC \text{ count}$ 
  - 13  $\rightarrow$  likely iron deficiency
  - $<13$   $\rightarrow$  likely thalassemia trait
- This child likely has high RDW and high Mentzer Index  $\rightarrow$  favors iron deficiency



## ***When to Check Lead Levels?***

- If child has risk factors (old home, pica, low SES, poor nutrition)
- Consider lead level if anemia persists after iron trial, or if neuro signs are present





## ***Next Steps in Management***

- Order serum ferritin and CRP
- Trial of oral iron for 4–6 weeks
- Dietary counseling (meat, fortified cereals, reduce cow's milk)

## *Teaching Points*

- Most common cause of anemia in this age group
- High RDW distinguishes from thalassemia trait
- Confirm with ferritin, treat with iron and dietary guidance



***Case 2***  
***Febrile 3-Month-Old with***  
***Neutropenia***



## *Presentation*


- Age: 3-month-old infant
- Symptoms: 38.5°C fever, irritable but feeding okay
- CBC:
  - WBC:  $2.3 \times 10^9/\text{L}$  (low)
  - ANC: 400/ $\mu\text{L}$  (severe neutropenia)
  - Hb/Platelets: Normal
- No clear source of infection

## ***Interactive Poll Question***

- What is the most appropriate next step?
  - A. Discharge with oral antibiotics and follow-up
  - B. Start IV ceftriaxone and observe at home
  - C. Admit for IV broad-spectrum antibiotics
  - D. Bone marrow aspiration and biopsy



## ***Correct Answer***

- C. Admit for IV broad-spectrum antibiotics
-  ***Teaching Points:***
  - This infant has severe neutropenia (ANC = 400) → high risk for serious bacterial infection (SBI).



## ***Fever + Neutropenia in Infants***

- <3–6 months with fever and ANC <500 → treat like immunocompromised host
- High risk of bacteremia, especially Gram-negative rods, Staph aureus, and Pseudomonas
- Workup includes: blood cultures, urine culture, LP if indicated



## *Initial Management*

- Admit
- Start IV broad-spectrum antibiotics
- Cefepime, ceftazidime, or piperacillin-tazobactam
- Monitor for signs of sepsis or marrow suppression





# ***Differential Diagnosis for Isolated Neutropenia in Infants***

- Viral suppression
- Benign ethnic neutropenia
- Congenital neutropenia
- Auto immune neutropenia



## ***When to Investigate Further***

- Recurrent neutropenia
- Persistent ANC  $<500$  for  $>3$  weeks
- Recurrent bacterial infections
- Growth failure or other cytopenias → consider bone marrow exam

## *Teaching Points*

- Emergency in infants: admit for IV antibiotics
- Consider viral suppression vs congenital neutropenia
- ANC <500 = high risk for sepsis



## ***Case 3***

# ***The Tired Teen with Arthritis***



## *Presentation*

- Patient: 13-year-old girl with juvenile idiopathic arthritis (JIA)
- Complaints: Fatigue, pallor, poor concentration
- Exam: Pale conjunctivae, active arthritis in wrists



## ***CBC***


- Hb: 9.5 g/dL
- MCV: 78 fL
- WBC/Plt: Normal
- Retic: Low
- Ferritin: 150 ng/mL (normal-high)
- CRP/ESR: Elevated

## *Interactive Poll*

- What is the most likely cause of her anemia?
  - A. Iron deficiency anemia
  - B. Thalassemia trait
  - C. Anemia of chronic disease
  - D. Sideroblastic anemia



## ***Correct Answer***

- C. Anemia of chronic disease
-  Teaching Points





## ***What is Anemia of Chronic Disease (ACD)?***

- Also called anemia of inflammation
- Common in chronic infections, autoimmune disease (e.g. JIA, SLE), IBD, CKD
- Mild to moderate normocytic or microcytic anemia
- Caused by hepcidin-induced iron sequestration and blunted erythropoietin response



# ***Distinguishing ACD vs Iron Deficiency***

<i>Test</i>	<i>Iron Deficiency</i>	<i>ACD</i>
Ferritin	Low	Normal or High
TIBC	High	Low
Serum Iron	Low	Low
RDW	High	Normal
CRP/ESR	Normal	High
Retic count	Low	Low

*Ferritin is an acute-phase reactant: elevated in inflammation*  
*In mixed ACD + iron deficiency, ferritin can be misleading*

## *key message*

- Always assess inflammation markers (ESR, CRP) in children with anemia and chronic disease
- Iron therapy is not effective unless combined with treatment of underlying inflammation
- Newer markers (e.g., soluble transferrin receptor, hepcidin) may help in complex cases



## *Clinical Pearls*

- ACD is often underrecognized in pediatric patients with chronic disease
- In kids with chronic arthritis and borderline anemia → think ACD
- Do not give iron routinely unless iron deficiency is confirmed

# *Case 4 – Adolescent with Bruising*

- Case: 10-year-old with fatigue, bruising, leg pain. CBC: WBC 32,000 with 70% blasts
- Diagnosis: Acute leukemia
- Teaching Points:
  - Blasts = red flag; confirm with flow cytometry
  - Thrombocytopenia + leukocytosis + anemia = marrow failure
  - Don't start steroids before diagnosis confirmed

# *Case 5 – Reactive Thrombocytosis*

- Case: 11-year-old recovering from viral illness, Platelets 720,000
- Diagnosis: Reactive thrombocytosis
- Teaching Points:
  - Common and benign after viral illness or inflammation
  - Differentiate from essential thrombocythemia with smear and history
  - No intervention required unless sustained  $>1$  million

## ***Case 6 – Macrocytic Anemia***

- Case: 15-year-old vegan adolescent, Hb 9.2, MCV 108 fL, low retic
- Diagnosis: Vitamin B12 deficiency
- Teaching Points:
  - Always consider dietary history in adolescents
  - Macrocytosis + low retic + neurologic signs = B12
  - Rule out other causes (hypothyroidism, meds, marrow disease)

# *Summary*

- The identification of abnormal complete blood count (CBC) requires clinicians to use an appropriate set of reference ranges, acknowledging that CBC will vary based on a child's age and sex. (Based on consensus)
- Consensus recommendation now advocates against using age- race-based CBC reference ranges.
- An evidence-based approach to interpreting the CBC is critical to avoid missing important diagnoses or pursuing unnecessary evaluation and referral.





# ***Thank You!***

- 🩺 Questions & Discussion
- 💬 How do you approach a child with anemia when CBC indices are borderline?
- 📊 What is your favorite 'first step' test after seeing a low hemoglobin in a child?
- 🔍 Have you encountered mixed anemia (e.g. ACD + IDA) in practice?
- Let's discuss your experiences and insights in interpreting pediatric CBCs!

# MCQ 1

- A 9-month-old boy, who is new to your practice, is seen for a health supervision visit. This child is exclusively breast fed and has received no other nutritional supplementation since birth. A screening hemoglobin level is low (10 g/dL [100 g/L]). You order additional laboratory studies, including a complete blood cell count. Which of the following sets of values/results would be most consistent with this clinical presentation.*
  - A. Mean corpuscular volume (MCV) 5 68 mm<sup>3</sup> (68 fL); platelet count 5 450 × 10<sup>3</sup> /mL (450 × 10<sup>9</sup> /L); reticulocytes 5 0.5%; red blood cell distribution width (RDW) 5 16.5%.**
  - B. MCV 5 90 mm<sup>3</sup> (90 fL); platelet count 5 300 × 10<sup>3</sup> /mL (300 × 10<sup>9</sup> /L);reticulocytes 5 1.1%; RDW 5 11%.**
  - C. MCV 5 68 mm<sup>3</sup> (68 fL); platelet count 5 450 × 10<sup>3</sup> /mL (450 × 10<sup>9</sup> /L);reticulocytes 5 1.1%; RDW 5 11%.**
  - D. MCV 5 68 mm<sup>3</sup> (68 fL; platelet count 5 300 × 10<sup>3</sup> /mL (300 × 10<sup>9</sup> /L);reticulocytes 5 0.5%; RDW 5 11%.**
  - E. MCV 5 90 mm<sup>3</sup> (90 fL; platelet count 5 450 × 10<sup>3</sup> /mL (450 × 10<sup>9</sup> /L); reticulocytes 5 0.5%; RDW 5 16.5%.**

# MCQ2

- You are evaluating a 3-year-old child referred to your office by the local Woman, Infants, and Children (WIC) program for anemia. On obtaining additional history, you learn that this child was born prematurely and had “some kind of intestinal surgery,” as indicated by an abdominal scar on physical examination. There are no unusual dietary preferences reported. Repeated laboratory studies show the following results: hemoglobin, 10.0 g/dL (100 g/L); hematocrit, 30.0%; MCV, 105 mm<sup>3</sup> (105 fL); and reticulocytes, 0.7%. A peripheral blood smear shows megaloblasts and hypersegmented neutrophils. Based on this history and the laboratory results, nutritional deficiency of which of the following is the most likely reason for this child’s anemia?*
  - A. Vitamin C.*
  - B. Iron.*
  - C. Vitamin B12.*
  - D. Vitamin D.*
  - E. Vitamin E*

# MCQ3

- *A 15-year-old boy has become increasingly tired/fatigued during the past 3 to 4 weeks. He denies any changes in mood or drug/alcohol use. On physical examination he exhibits cervical adenopathy, mild jaundice, and some hepatosplenomegaly. Laboratory studies confirm the suspected diagnosis of Epstein-Barr virus infection, but the patient is also noted to be anemic. Which of the following best describes the expected characteristics to be seen in the anemia in this patient?*
  - A. Microcytic with a decreased reticulocyte count.*
  - B. Microcytic with an increased reticulocyte count.*
  - C. Normocytic with a decreased reticulocyte count.*
  - D. Normocytic with an increased reticulocyte count.*
  - E. Macrocytic with an increased reticulocyte count.*

# MCQ4

- ***A 2-year-old child is being evaluated for recurrent fevers in association with mouth ulcers and gingivitis. The parents report that she is repeatedly ill with respiratory illnesses but is currently well. She is frequently sent home from the out-of-home child care she attends. She is not taking any long-term medications, she is growing well, and there is no contributory family history. Laboratory studies (complete blood cell count) report the following results: hemoglobin, 12 g/dL (120 g/L); hematocrit, 37%; normal red blood cell indices; platelet count,  $370 \times 10^3$  /mL ( $370 \times 10^9$  /L); white blood cell count, 4,100,000/mL ( $4,100 \times 10^9$  /L); absolute neutrophil count, 700/mL; lymphocyte count, 3,100/mL ( $3.10 \times 10^9$  /L); eosinophil count, 100/mL ( $0.10 \times 10^9$  /L); monocyte count, 200/mL ( $0.20 \times 10^9$  /L). To develop the most appropriate evaluation and treatment plan, these studies should be repeated in which of the following time frames?***
  - A. 2–4 wk.***
  - B. 4–6 wk.***
  - C. 6–8 wk.***
  - D. 10–12 wk.***
  - E. 12–16 wk.***

# MCQ5

- You follow in your clinic a 7-year-old girl with cerebral palsy (Gross Motor Function Classification System II) and a poorly controlled seizure disorder treated with valproic acid. Falls associated with her seizures result in easy bruising and mucosal bleeding. Laboratory studies report the following results: hemoglobin, 13.2 g/dL (132 g/L); MCV, 90 mm<sup>3</sup> (90 fL); mean corpuscular hemoglobin, 30 pg; reticulocytes, 1%; white blood cell count, 8,500/mL ( $8.5 \times 10^9$  /L); neutrophils, 52%; lymphocytes, 40%; eosinophils, 3%; monocytes, 5%; platelet count,  $330 \times 10^3$  /mL ( $330 \times 10^9$  /L). Which of the following is the most likely etiology for her easy bruising and mucosal bleeding?***
  - A. Autoimmune reaction.***
  - B. Chronic infection.***
  - C. Drug reaction.***
  - D. Functional platelet disorder.***
  - E. Malignancy.***