

بِسْمِ اللّٰهِ الرَّحْمٰنِ الرَّحِیْمِ

هما تومور فولوژی مقدماتی

ناهنجاریهای خوش خیم گلبول های سفید و پلاکت

The grading of morphology
elements should provide the

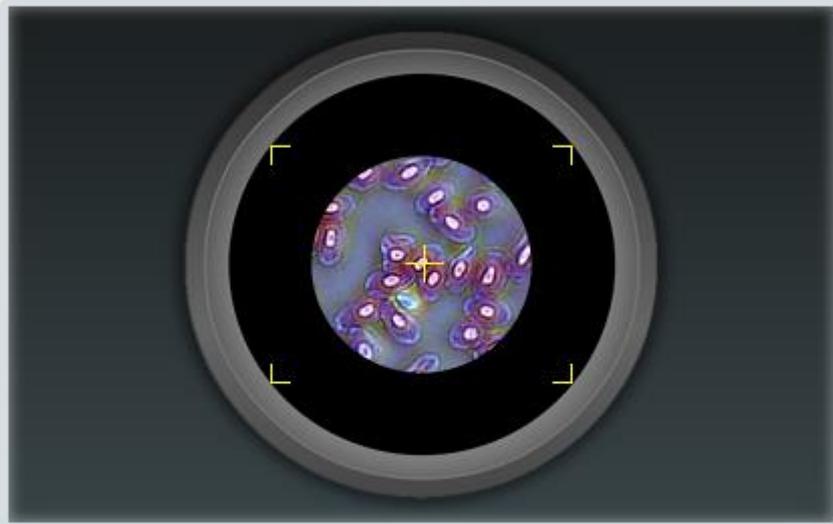
clinician

with **useful information** regarding
the status of

any abnormality

in the peripheral blood.

Field of View Diameter



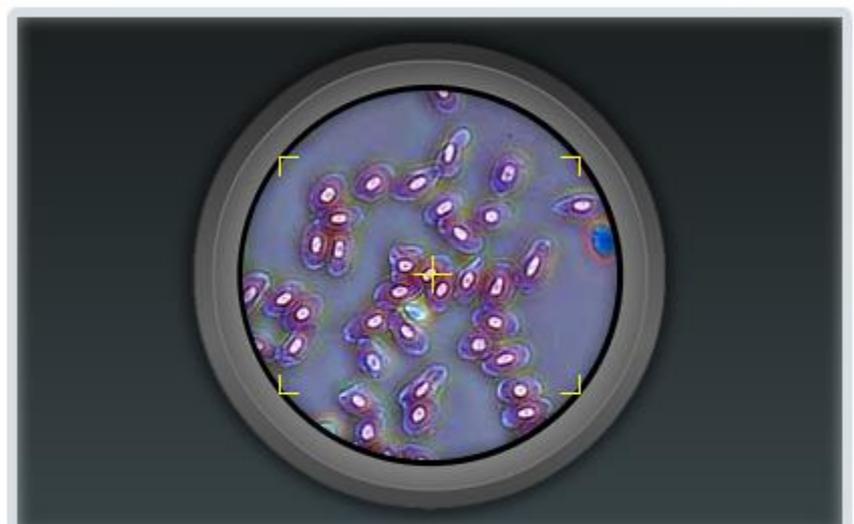
Choose a Specimen

Frog Blood

Field of View

16 mm

Field of View Diameter



Choose a Specimen

Frog Blood

Field of View

28 mm



Quantitative abnormalities

- In patients with normal cell populations in the absence of analyser flags or abnormal cell populations : **It is recommended that the automated analyser WBC differential count be reported**
- The automated differential may also be reported after viewing a blood film due to flags or other indicators where the automated values are found to be accurate.

Criteria for blood smear scan and/or blood smear examination at Thomas Jefferson University Hospital

		Adults	Infants
A. Based on CBC			
WBC ($\times 10^9/L$)	Initial*	<2.0 or >30.0 Or Delta failure of - 300% for WBC of 0.1 to 1.0 + 300% for WBC of > 10	
PLT ($\times 10^9/L$)	Initial	<100 or >999 or >30 with delta failure of -50% or greater	
B. Based on automated DIFF results			
Lymphocytes ($\times 10^9/L$)	Initial	>7.0 (for >14 yr old) >10.0 (for 1-14 yr old)	>14.0 (for <1 yr old)
Monocytes ($\times 10^9/L$)	Initial	>3.0	
Eosinophils ($\times 10^9/L$)	Initial	>2.0	
Basophils ($\times 10^9/L$)	Initial	>0.5	
Qualitative Flags		WBC abnormal scattergram, Immature granulocytes, Left shift, Atypical lymphocytes, Blasts, NRBC	

*Initial: first smear on a new patient per admission or an infrequent outpatient visit.

Qualitative abnormalities

Each laboratory and laboratory system should have policies in place to ensure the consistent application of the grading criteria.

Table 1. Morphology Grading Table

Cell Name	Grading System		
	Few/1+	Mod/2+, %	Many/3+, %
WBC			
Döhle bodies	N/A	2–4	>4
Vacuolation (neutrophil)	N/A	4–8	>8
Hypogranulation (neutrophil)	N/A	4–8	>8
Hypergranulation (neutrophil)	N/A	4–8	>8
Platelets			
Giant Platelets	N/A	11–20	>20

Qualitative abnormalities

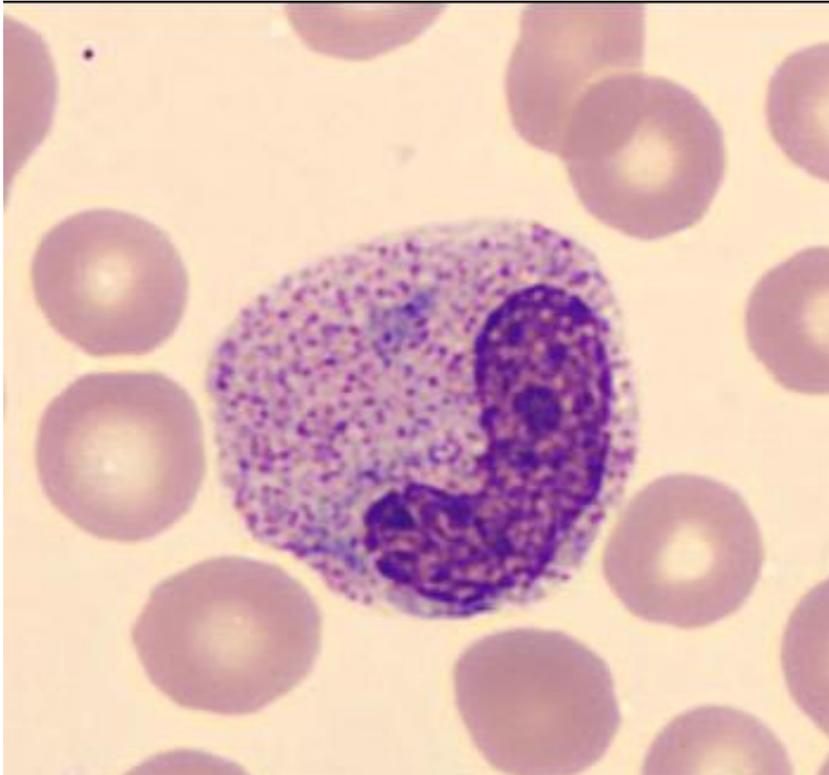
Cytoplasmic abnormalities

Dohle body:

- Pale light blue or grey
- Single or multiple
- Cytoplasmic inclusions found near the periphery of the neutrophil.
- Non-specific reactive change but may also indicate:
 - **Infection**: Toxic granulation, vacuolation, left shift
 - **May-Hegglin** anomaly: thrombocytopenia and giant platelets.
 - **G-CSF** administration
 - **CML**(infrequent)

Qualitative abnormalities

Cytoplasmic abnormalities CONT...



		Few/1+ %	Mod/2+ %	Many/3+ %
WBC	Dohle bodies	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
	Vacuolation (neutrophil)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
	Hypogranulation(neutrophil)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
	Hypergranulation(neutrophil)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
	Hypersegmented(neutrophil)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Platelets	Giant Platelets	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

Qualitative abnormalities

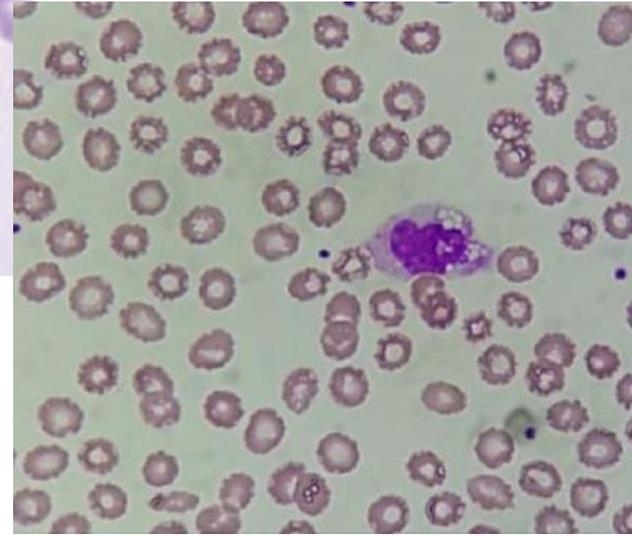
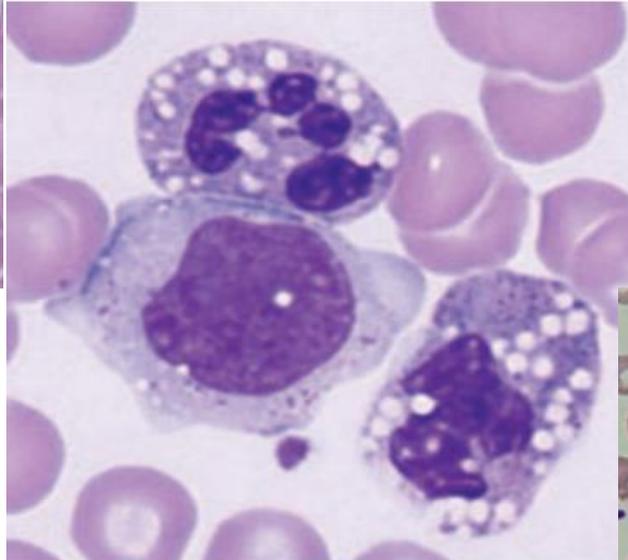
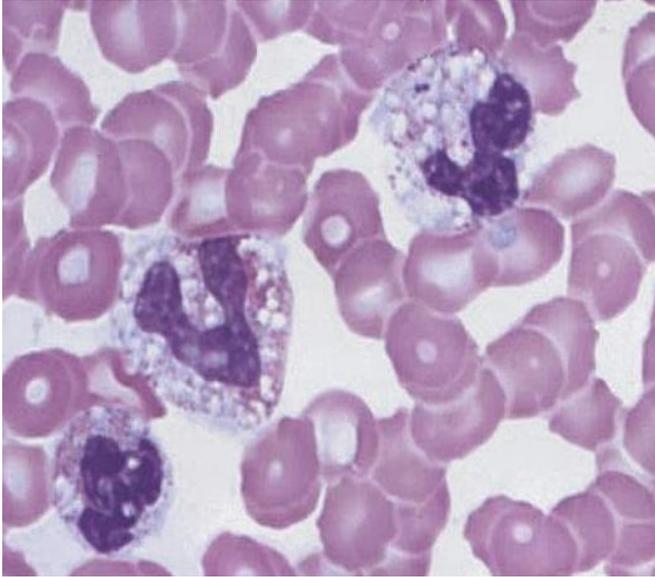
Cytoplasmic abnormalities

Vacuolation – neutrophil:

- I. **Infection** : ‘pin-hole’ vacuolation – small, discrete vacuoles, but the vacuoles may be larger
- II. **Alcohol toxicity**
- III. **Storage Artefact** : prolonged exposure to EDTA anticoagulant
- IV. **liver failure**
- V. **G-CSF**

Qualitative abnormalities

Cytoplasmic abnormalities CONT...

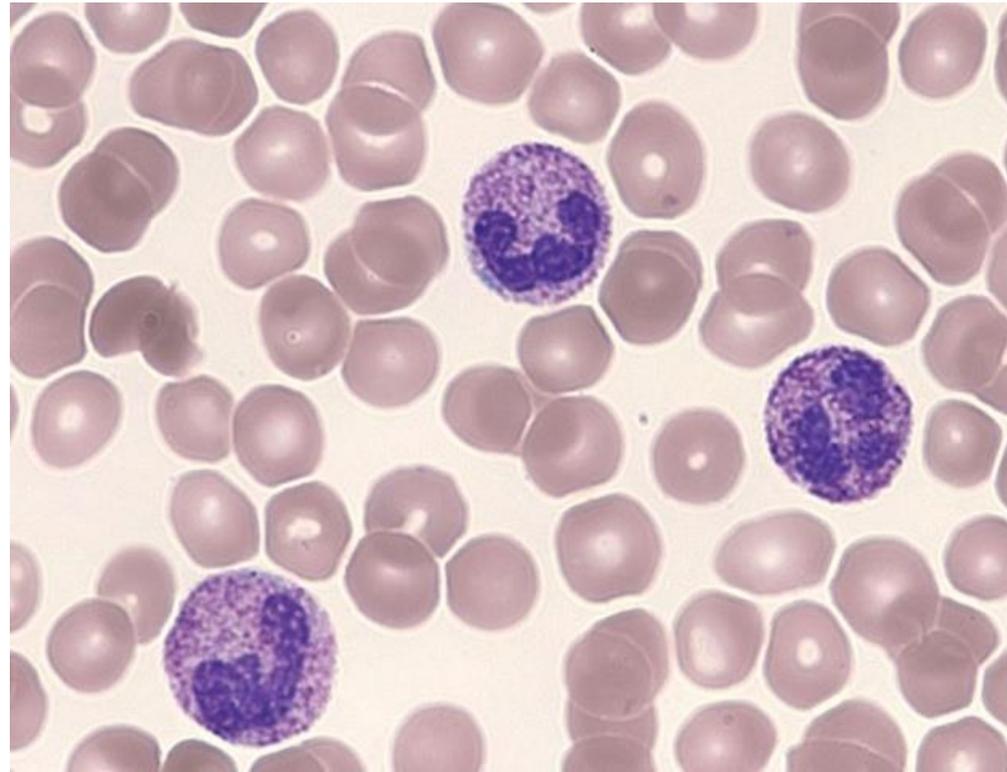


Qualitative abnormalities

Cytoplasmic abnormalities

Hypergranulation – neutrophil, (toxic granulation):

- Coarse, purple staining primary (azurophilic) neutrophil cytoplasmic granules
- Response to infection and inflammation.
- G-CSF
- Chemotherapy

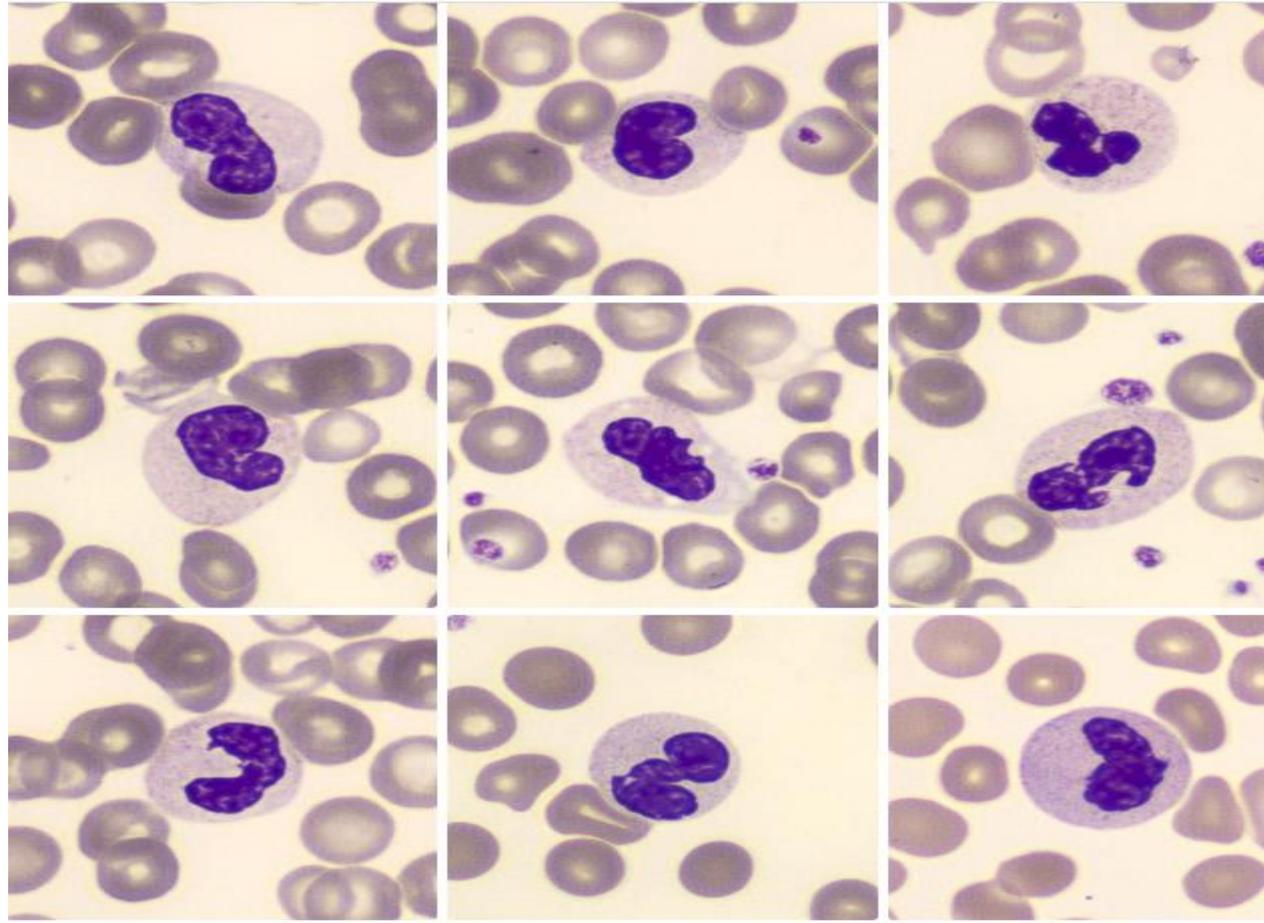


Qualitative abnormalities

Cytoplasmic abnormalities

Hypogranulation – neutrophil:

- Reduced presence of the normal granulation.
- All types of granulocytes → Dysgranulopoiesis and MDS



Qualitative abnormalities

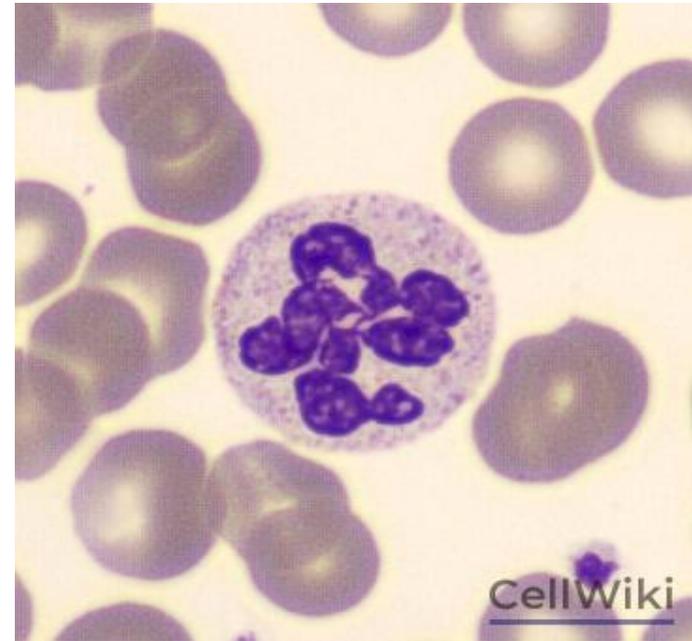
Nuclear abnormalities

Hypersegmented neutrophils

- Any neutrophil having 6 more lobes
- 100 neutrophils are examined → More than 3% of neutrophils having 5 lobes.

Associated with:

- Vitamin B12 / folate deficiency anemia
- MDS and MPD
- Cytotoxic chemotherapy
- long-term chronic infections.
- Hereditary hypersegmentation
- G-CSF administration
- Secondary to nitrous oxide anaesthesia
- Steroid therapy for immune thrombocytopenic purpura (ITP)



Qualitative abnormalities

Nuclear abnormalities

Pelger-Huet neutrophils

- Failure of normal nuclear lobe development during terminal differentiation and have coarse clumped nuclear chromatin.
- Dumbbell shaped, bi-lobed nuclei, 'pince-nez'

Importance:

Confused with: 1. Pseudo Pelger-Huet

2. Myelocytes, Metamyelocyte or Band neutrophils.

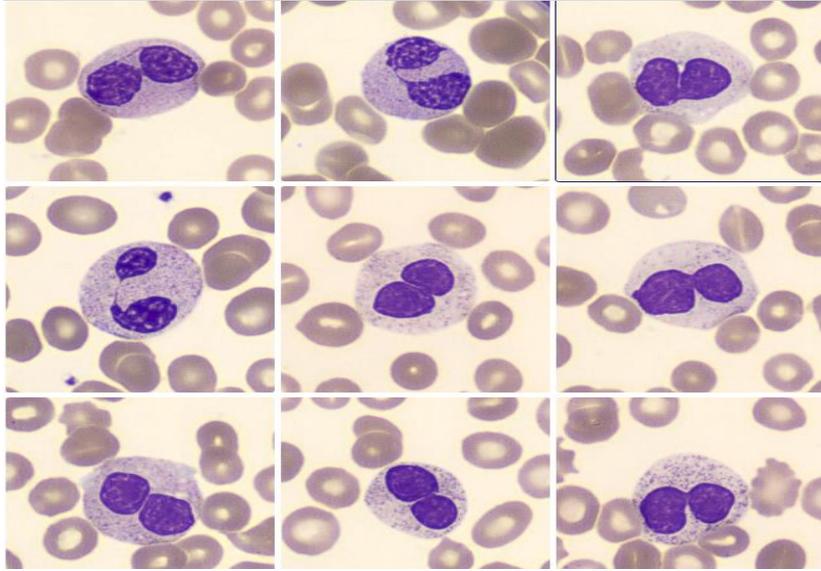
Recommendation:

Counted and reported as mature segmented neutrophils but with a suitable interpretive comment.

Pelgerhuet Anomaly	Seen	<input type="checkbox"/>	Not Seen	<input type="checkbox"/>
Pelgeroid morphology(Psdo Pelgerhuet)	Seen	<input type="checkbox"/>	Not Seen	<input type="checkbox"/>
Chediak Higashii Anomaly	Seen	<input type="checkbox"/>	Not Seen	<input type="checkbox"/>

Qualitative abnormalities

Nuclear abnormalities CONT...



Pelger-Huët anomaly

LBR protein amount determines degree of segmentation

Hyposegmentation and coarse clusters of nuclear chromatin

MDS

Need neutrophils with normal granulation to confirm presence of hypogranulation

blast
Basophil
Neutrophil or eosinophil?

MDS – Dysplastic features in blood

	Red blood cells	Anisocytosis, poikilocytosis (elliptocytes, small hypochromic irregular forms), dimorphic population, nucleated red cells, coarse basophilic stippling
	Neutrophils	Cytoplasmic hypogranulation, abnormal distribution or quality of cytoplasmic granules, pseudo-Pelger-Huët anomaly, nuclear hypersegmentation
	Monocytes	Dispersed chromatin, nucleoli, nuclear hyperlobulation
	Platelets	Large or giant forms, hypogranulation, bizarre shapes

Qualitative abnormalities

PLATELETS

Platelets are small, blue-grey granular fragments derived from megakaryocytic cytoplasm, containing many small, reddish-purple granules.

Platelet size:

Diagnostic significance particularly when considered in relation to the **platelet count**.

- A normal platelet measures 1.5–3 μm in diameter.
- **Large platelets** measure 3–7 μm (roughly the diameter of a normal sized red cell),
- **Giant platelets** are larger than normal sized

*Note: In a normal person, usually **less than 5%** of the platelets appear large.*

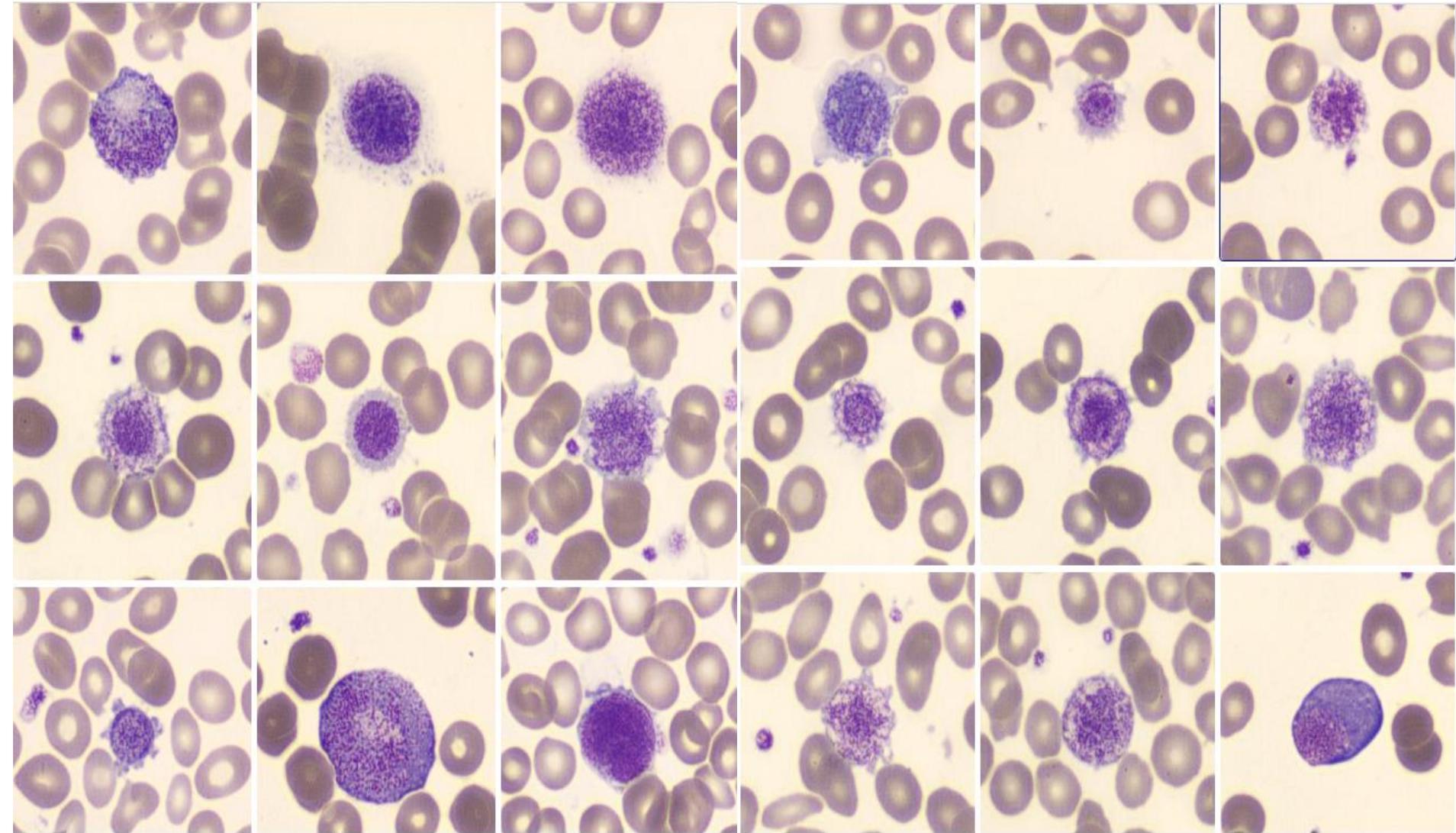
Recommendation:

Giant platelets be graded.

A comment about the platelet count and the presence of small, large and/or giant platelets can be made with an additional interpretive film comment if appropriate.

Qualitative abnormalities

PLATELETS CONT...



Qualitative abnormalities

PLATELETS CONT...

Acquired causes of large platelets :

Storage artefacts

Physiological platelets in neonates

Immune thrombocytopenia

Thrombotic microangiopathies

Disseminated intravascular coagulation

Myeloproliferative neoplasms (essential thrombocythaemia, polycythaemia vera, primary myelofibrosis, chronic myeloid leukaemia)

Myelodysplastic syndromes

Myelodysplastic/myeloproliferative neoplasms

Megakaryoblastic leukaemia

Postsplenectomy states and hyposplenism

Drug-induced (cholestyramine, erucic acid)

Inherited : Bernard–Soulier syndrome, gray platelet syndrome and May–Hegglin anomaly

Qualitative abnormalities

PLATELETS

PLATELETS granulation abnormality:

- Hypogranular
- Agranular platelets
- Presence of singular large granula

PLATELETS shape abnormality:

- Irregular to bizzare
- Cytoplasmatic blebs

Megakaryocytes and **megakaryoblasts** are rarely seen in normal peripheral blood.

- Small ***megakaryoblasts*** may be indistinguishable from lymphoblasts.
- ***Micromegakaryocyte***
 - Size of a promyelocyte or smaller with a nonlobated or a bilobed nucleus
 - Variable amount of weakly basophilic cytoplasm.
 - Nucleus may appear 'bare' but a small rim of cytoplasm

Qualitative abnormalities

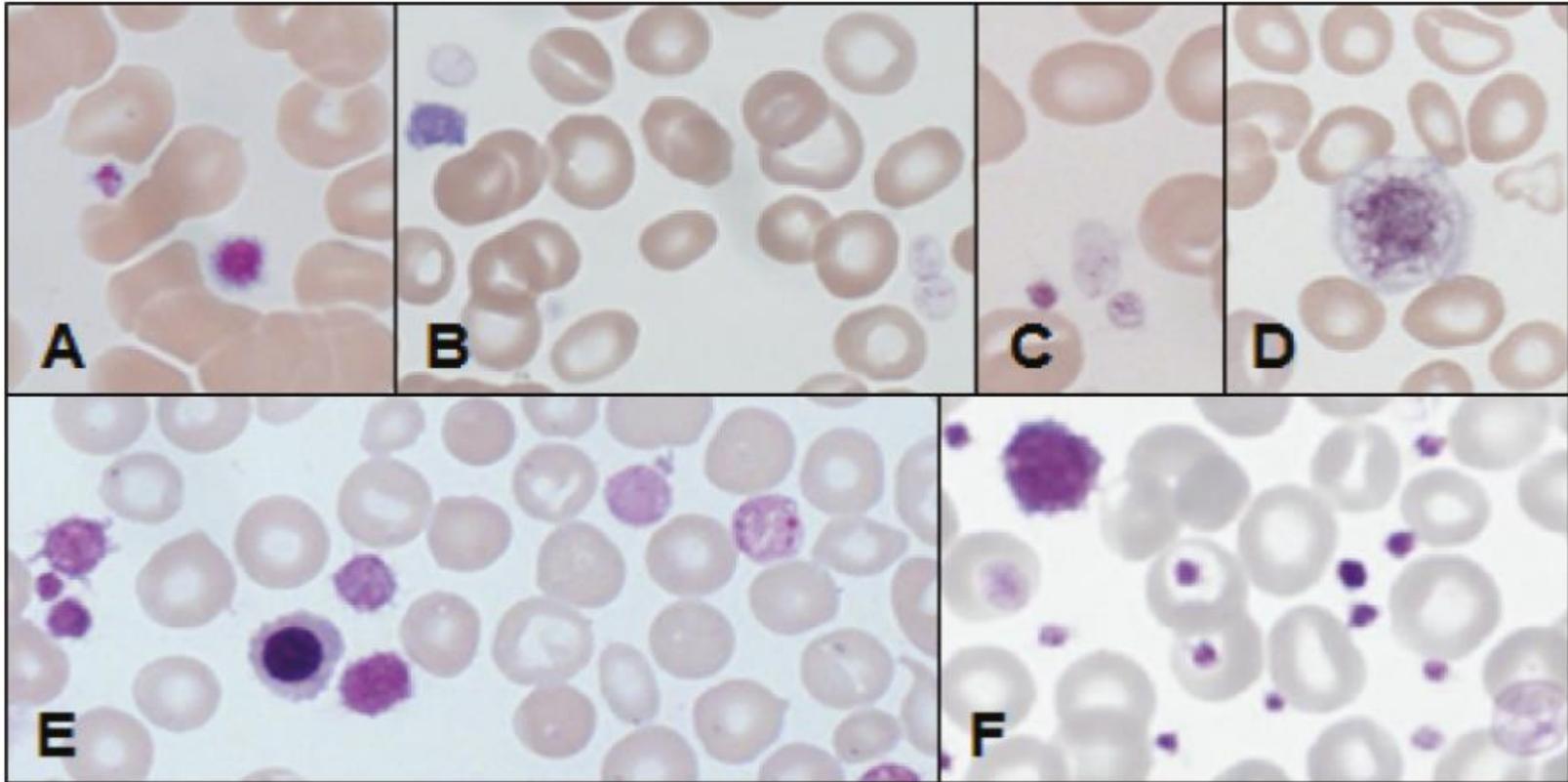
PLATELETS CONT...

Causes of hypogranular/agranular platelets

Inherited	Gene	Inheritance
Grey platelet syndrome	<i>NBEAL2</i>	AR
Chediak-Higashi syndrome	<i>LYST</i>	AR
Hermansky-Pudlak syndrome	<i>HPS</i>	AR
GrisCELLI syndrome	<i>MYO5A, RAB27A, MLPH</i>	AR
Wiskott-Aldrich syndrome	<i>WAS</i>	XL
Thrombopenia-absent radius syndrome	<i>RBM8A</i>	AR
GATA1-related disease	<i>GATA1</i>	XL
Arthrogryposis renal dysfunction and cholestasis syndrome	<i>VPS33B, VIPAR</i>	AR
Acquired		
In vitro degranulation (difficult venipuncture, platelet aggregation, EDTA-induced, cardiopulmonary bypass)		
In vivo degranulation (disseminated intravascular coagulation)		
Myelodysplastic syndromes		
Myeloproliferative neoplasms (essential thrombocythaemia, polycythemia vera, primary myelofibrosis)		

Qualitative abnormalities

PLATELETS CONT...



(A) Normal platelets;

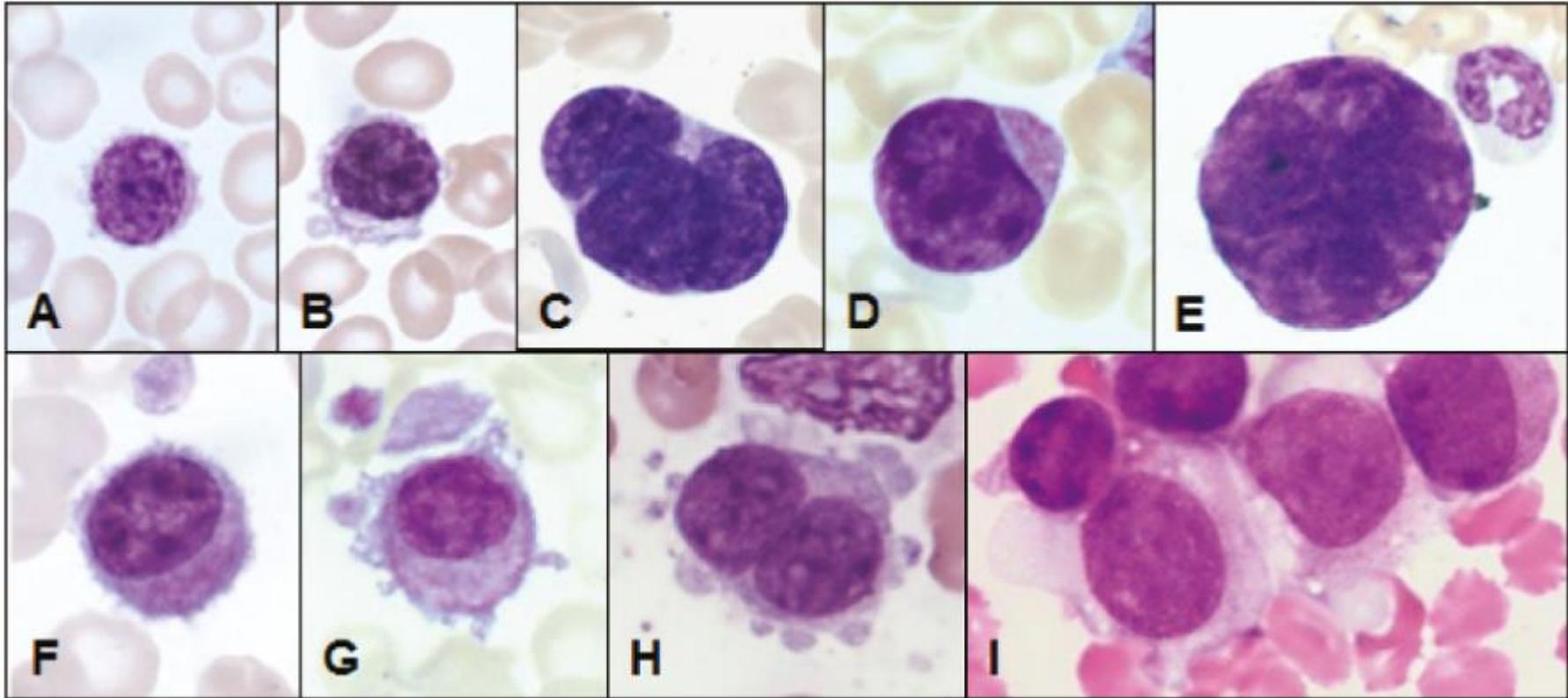
(B) and (C) agranular and hypogranular platelets in a patient with a myelodysplastic syndrome;

(D) giant platelet;

(E) platelet anisocytosis, large platelets and platelets with abnormal granulation in a patient with primary myelofibrosis; (F) platelet anisocytosis, a giant platelet and granulation anomalies in a patient with essential thrombocythaemia.

Qualitative abnormalities

PLATELETS CONT...



Circulating megakaryocytes, micromegakaryocytes and megakaryoblasts.
(A–E) Megakaryocytes; (F–H) micromegakaryocytes; (I) megakaryoblasts.

Recommendation:

Comment about the presence of hypogranular platelets, megakaryocytes, micromegakaryocytes and megakaryoblasts should be made if seen in the PB smear

 Thank You
For Your Attention