

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

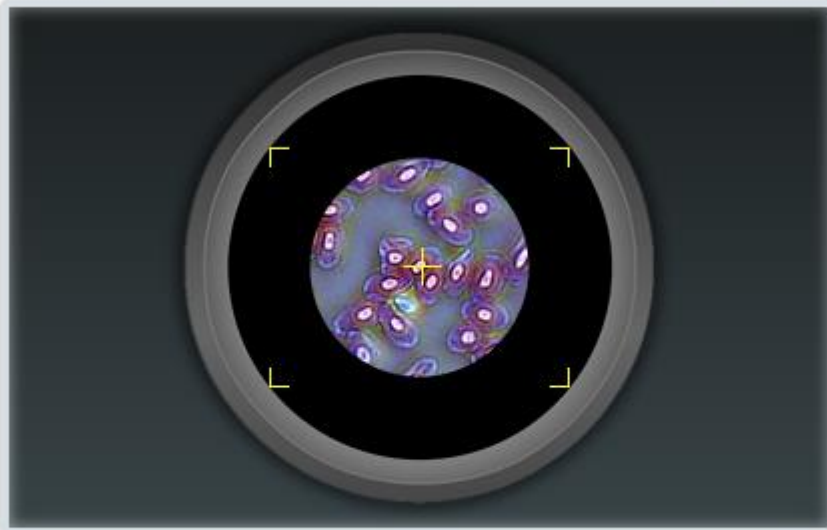
هما تومور فولوژی مقدماتی

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



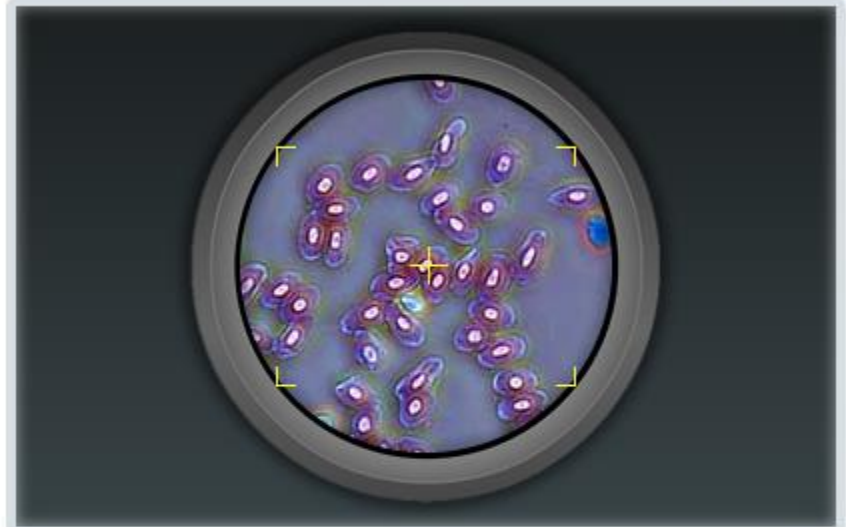
Choose a Specimen

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

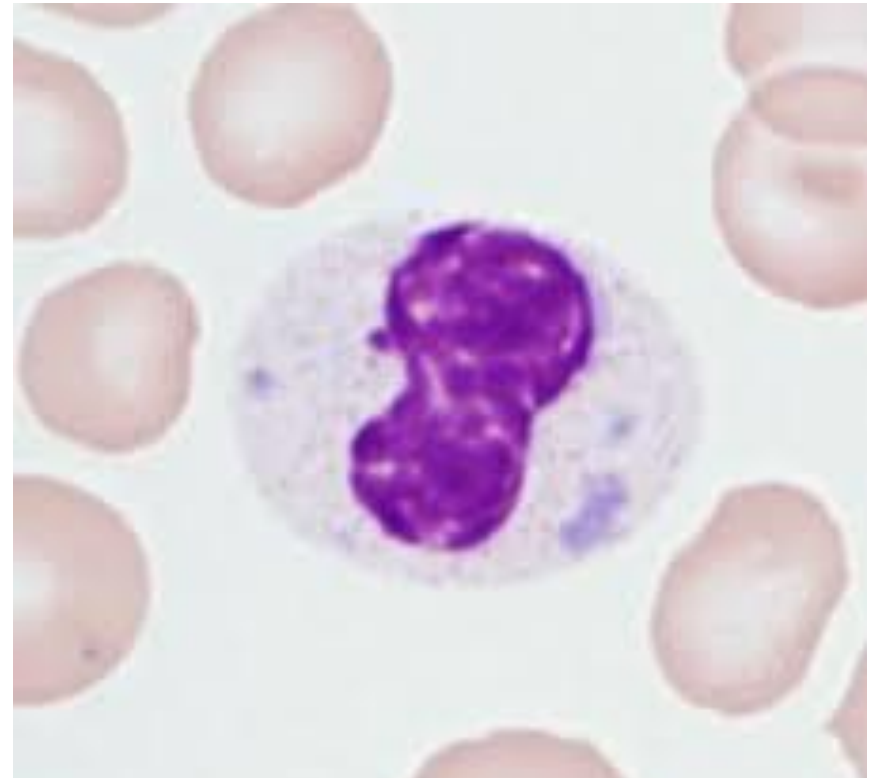
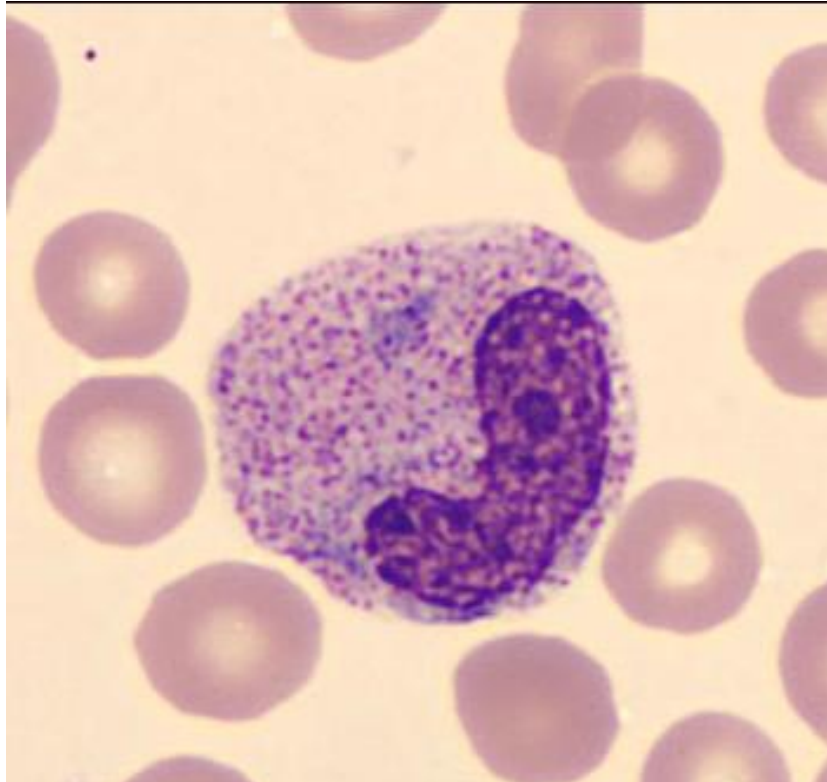
## Cytoplasmic abnormalities

### **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="text"/>	<input type="text"/>	<input type="text"/>
	Vacuolation (neutrophil)	<input type="text"/>	<input type="text"/>	<input type="text"/>
	Hypogranulation(neutrophil)	<input type="text"/>	<input type="text"/>	<input type="text"/>
	Hypergranulation(neutrophil)	<input type="text"/>	<input type="text"/>	<input type="text"/>
	Hypersegmented(neutrophil)	<input type="text"/>	<input type="text"/>	<input type="text"/>
Platelets	Giant Platelets	<input type="text"/>	<input type="text"/>	<input type="text"/>

# Qualitative abnormalities

## Cytoplasmic abnormalities

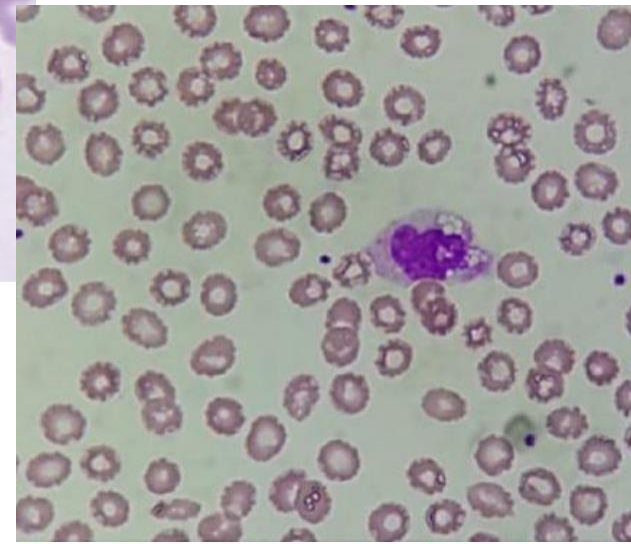
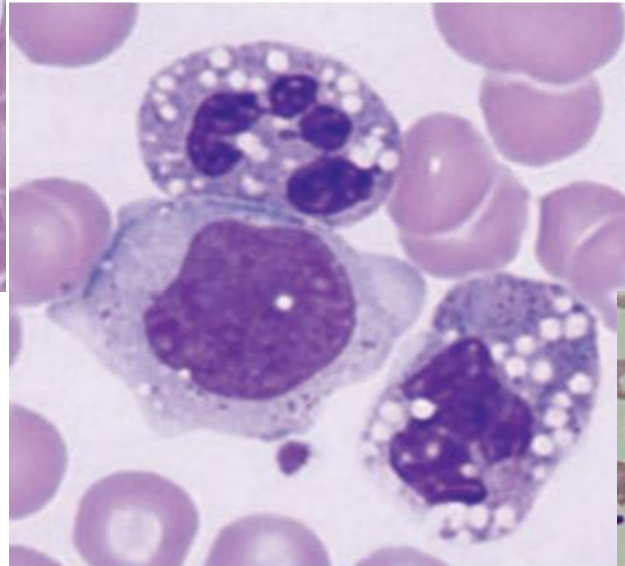
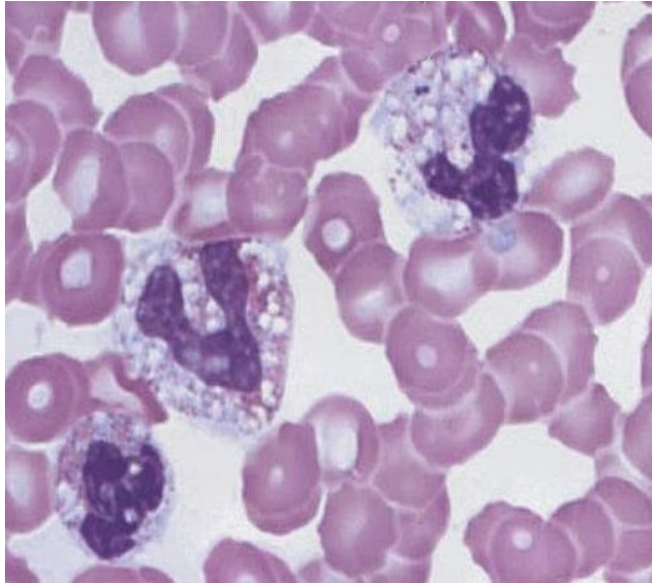
### **Vacuolation – neutrophil:**

- 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...

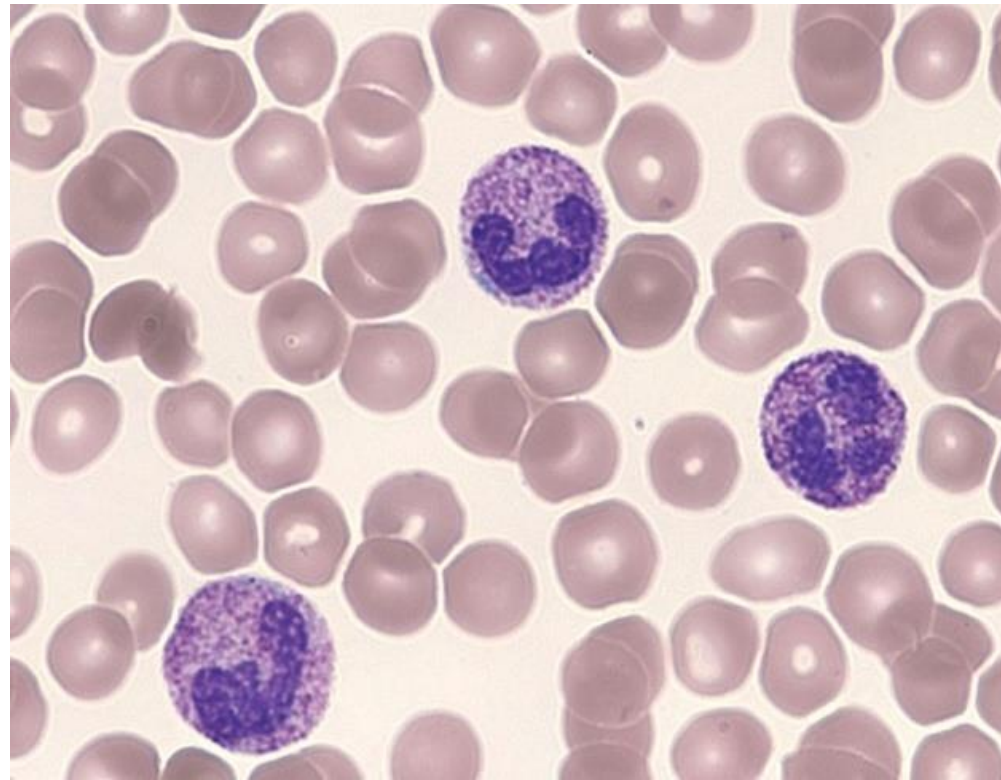


# Qualitative abnormalities

## Cytoplasmic abnormalities

### **Hypergranulation – neutrophil, (toxic granulation):**

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

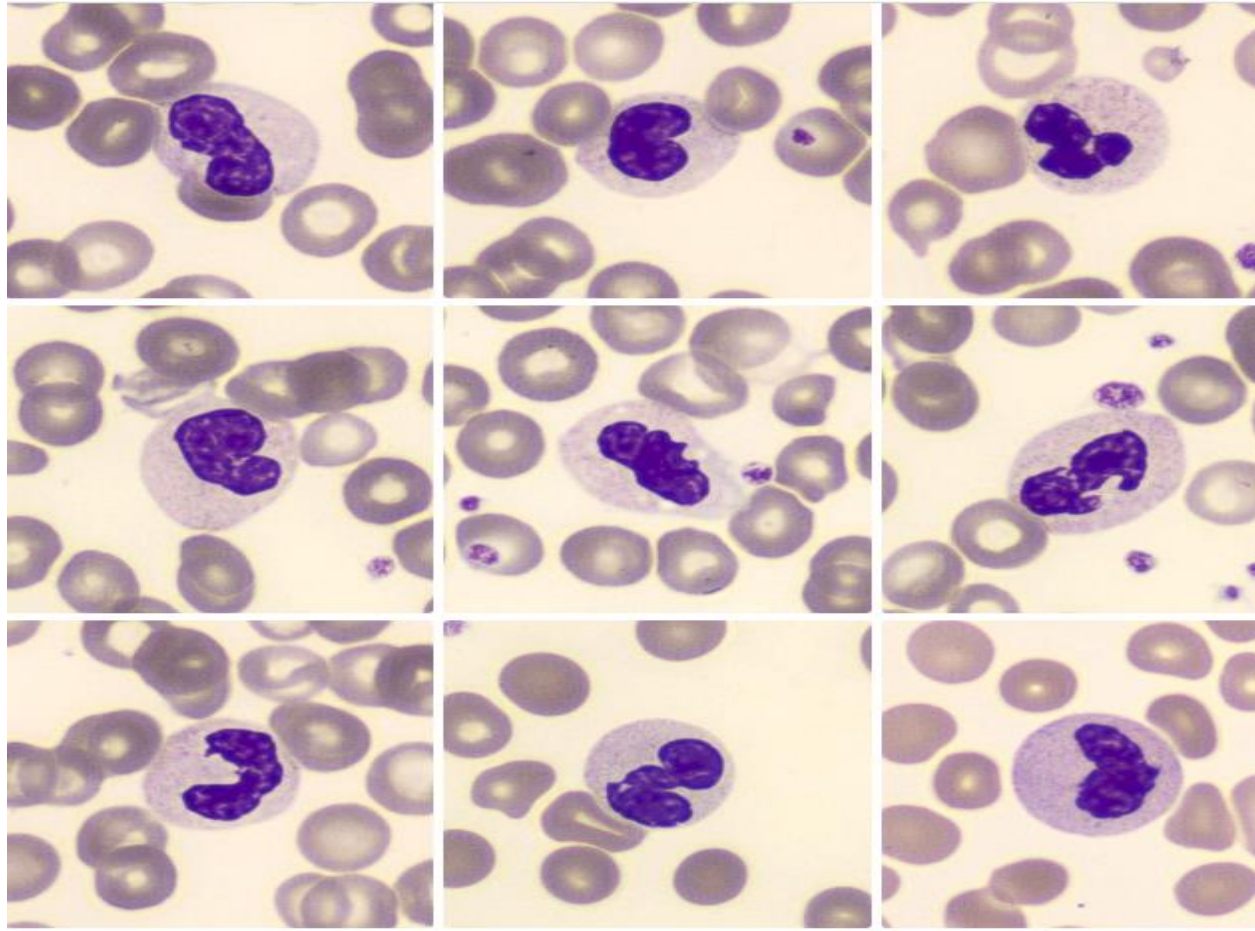


# Qualitative abnormalities

## Cytoplasmic abnormalities

### Hypogranulation – neutrophil:

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



# Qualitative abnormalities

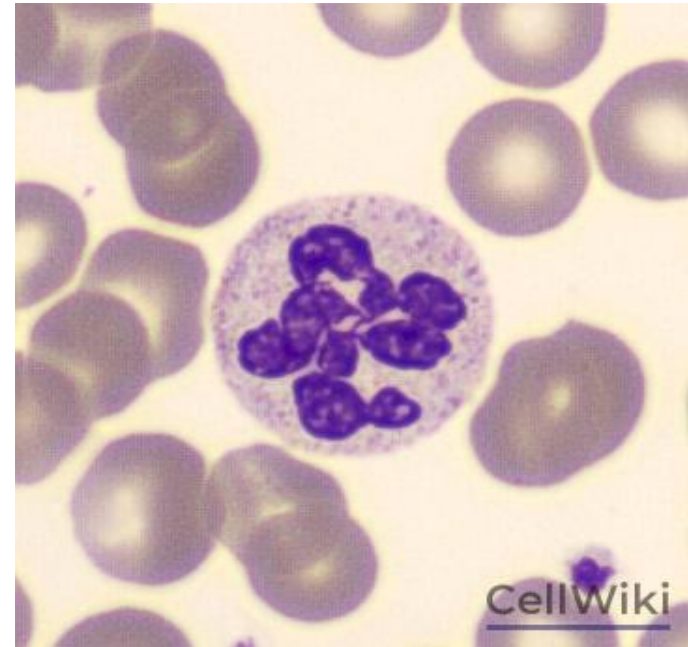
## Nuclear abnormalities

### 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:

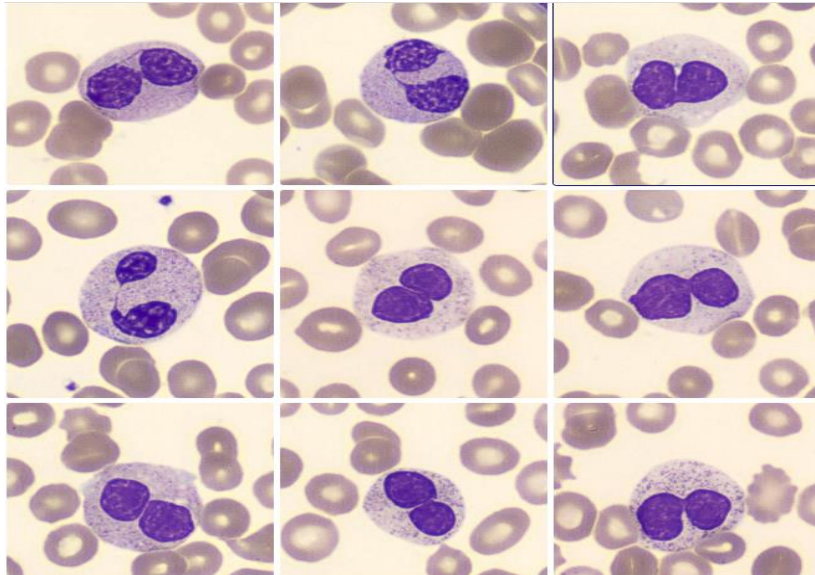
Counted and reported as mature segmented neutrophils but with a suitable interpretive comment.

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



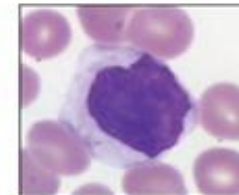
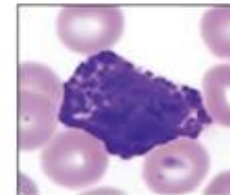
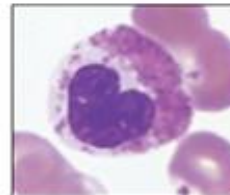
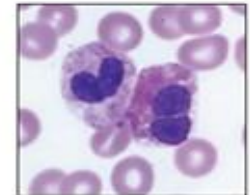
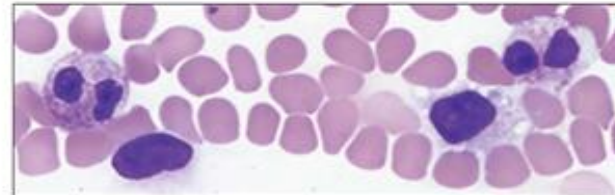
# Qualitative abnormalities

## Nuclear abnormalities CONT...



### Pelger-Huët anomaly

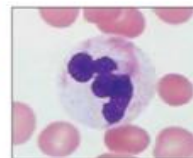
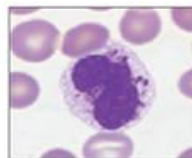
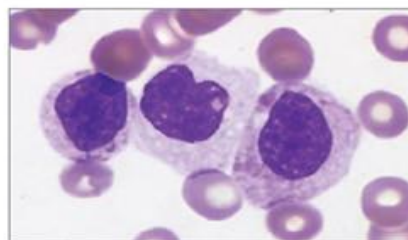
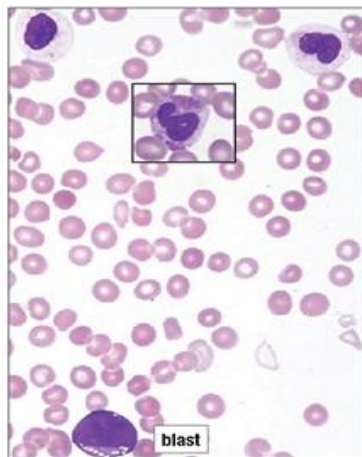
LBR protein amount determines degree of segmentation



Hyposegmentation  
and coarse clusters  
of nuclear chromatin

### MDS

Need neutrophils with normal granulation to confirm presence of hypogranulation

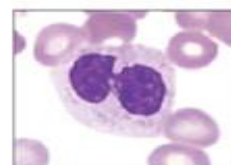


### MDS – Dysplastic features in blood



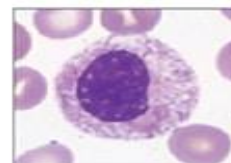
#### Red blood cells

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  $\mu\text{m}$  in diameter.
- **Large platelets** measure 3–7  $\mu\text{m}$  (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:**

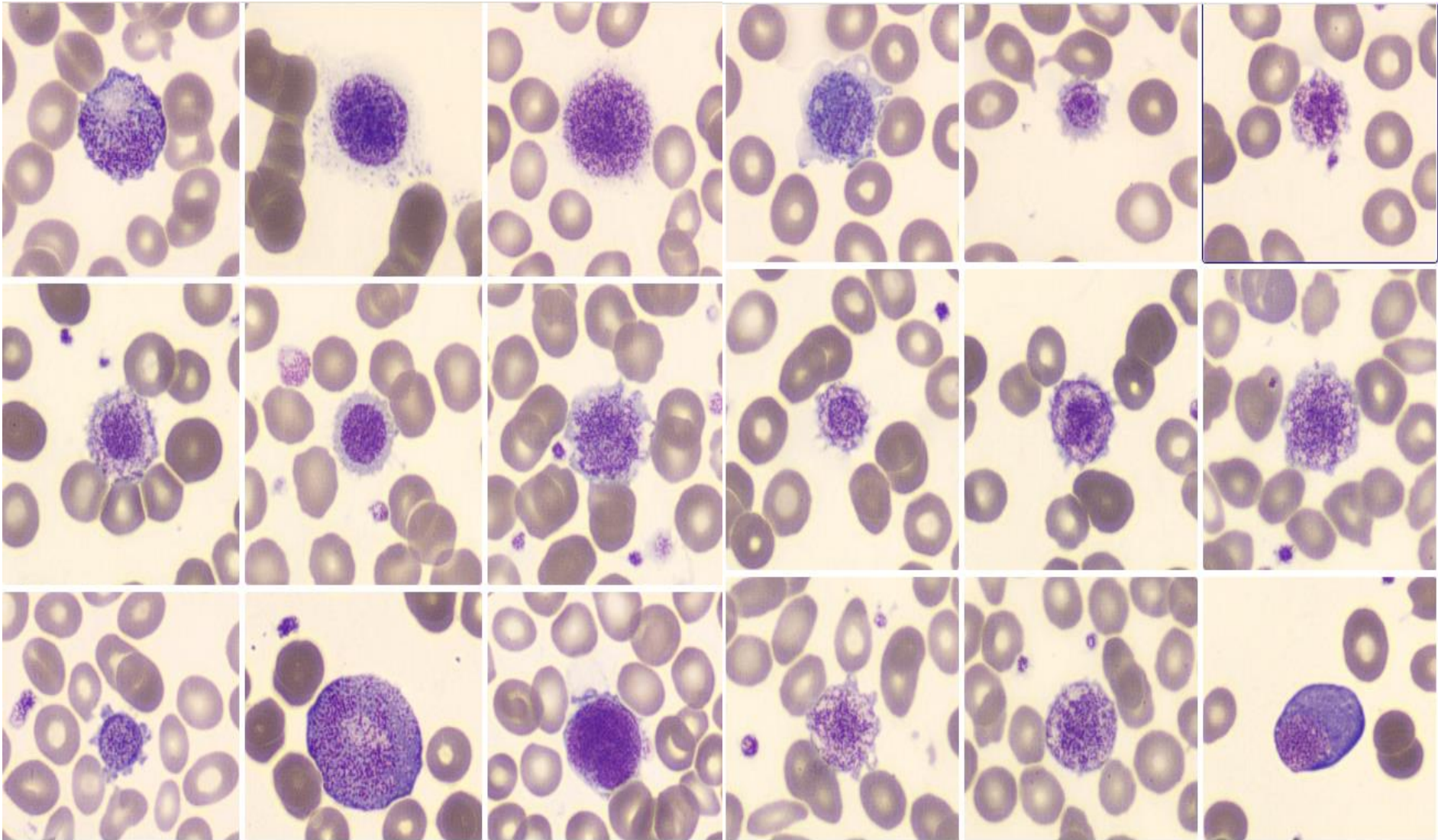
Giant platelets be graded.

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

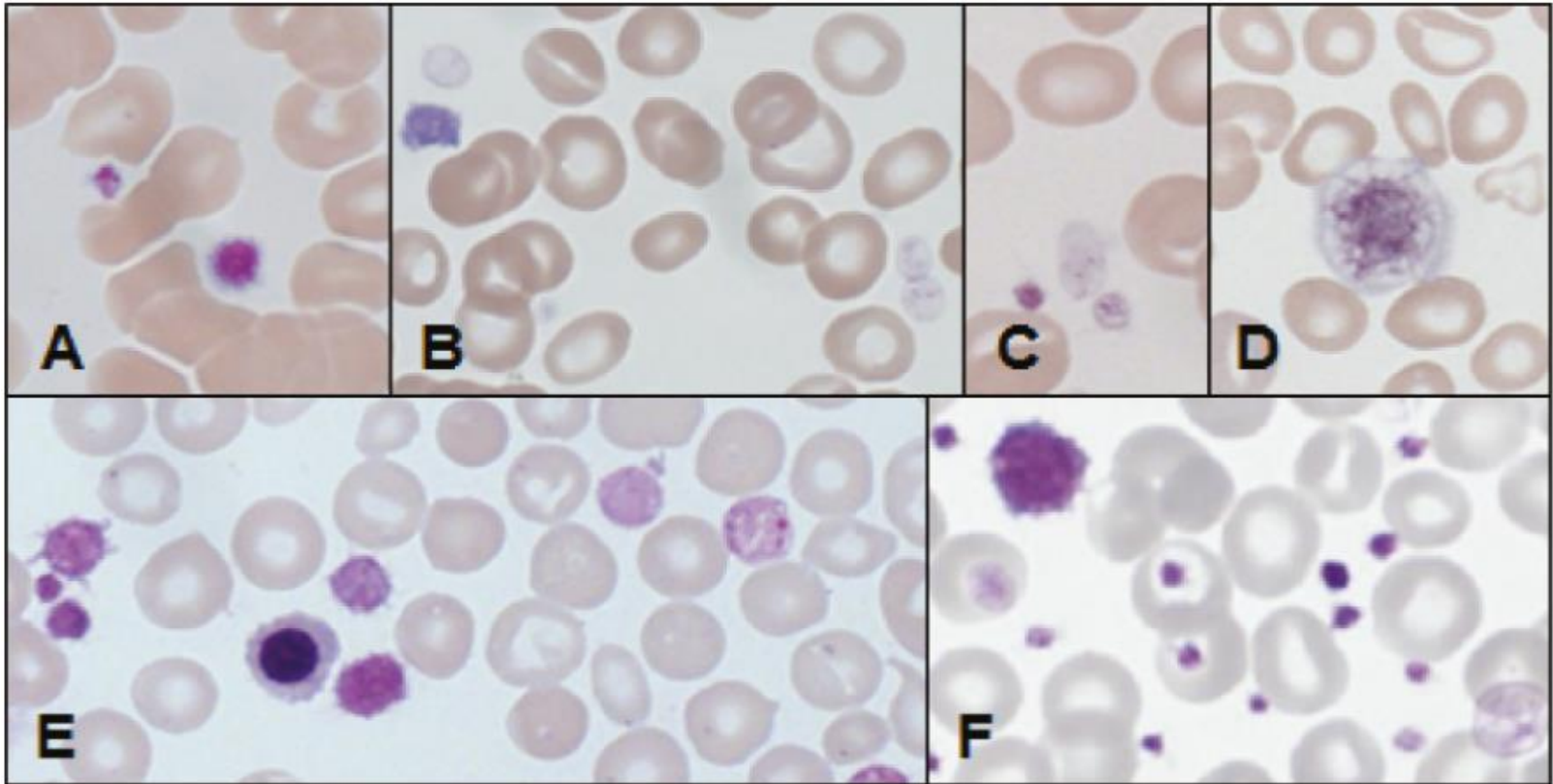
## PLATELETS CONT...

### 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
<i>GATA1</i> -related disease	<i>GATA1</i>	XL
Arthrogryposis renal dysfunction and cholestasis syndrome	<i>VPS33B, VIPAR</i>	AR
<b>Acquired</b>		
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;

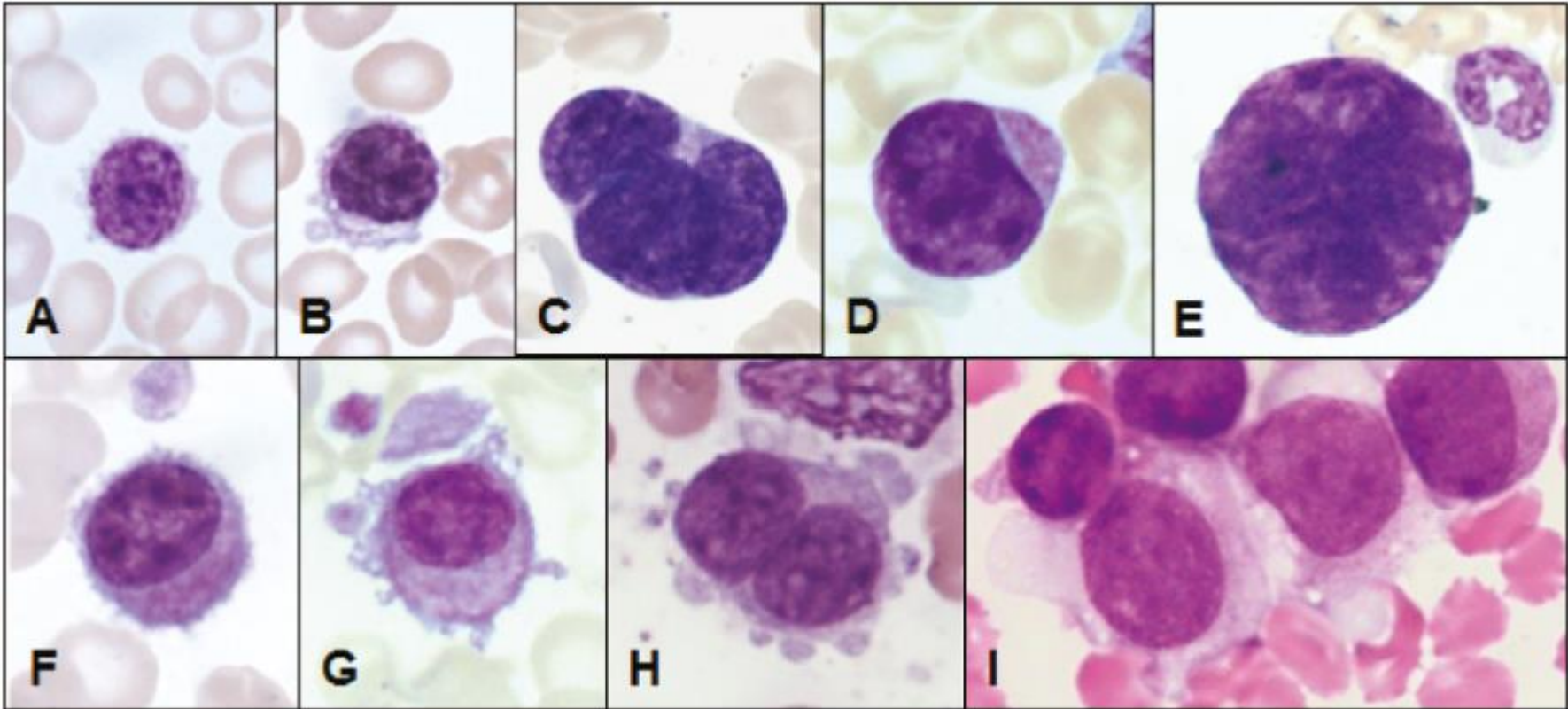
(B) and (C) agranular and hypogranular platelets in a patient with a myelodysplastic syndrome;

(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