

ABC OF CBC

DR SUDHIR MEHTA MD,MNAMS,FICP

Senior Professor & Head Medical Unit

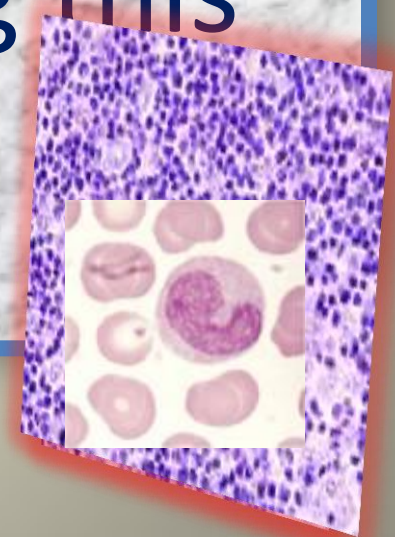
SMS Medical College & Hospital

Jaipur

s.smehta@hotmail.com



CBC.....What is the
UTILITY of performing this
basic Hematology Test?



Half of our population
suffer from anemia

#...Up to 15% of population in
certain communities are
carriers of
Thalassemia gene.....!!

Infectious diseases

Acquiring epidemic proportions

Malaria on the rise.....!!

Cases of Dengue fever
&

Platelet & DIC related problems
are reported more frequently

ON TOP OF EVERYTHING...

The preliminary diagnostic
information this single test can
generate and make available

AS GUIDE

for proceeding further towards the
Final Diagnosis is
UNIQUE

AUTOMATION & INTERPRETATION

Automation has changed principles and methodologies, approaches and conclusions of various disciplines of medicine

But...

Few branches are modified to the extent that their entire philosophy is so much reoriented that it needs to be re-written

and

HEMATOLOGY IS ONE OF THEM.

PARAMETERS LIKE RDW & HISTOGRAM

AUTOMATION & INTERPRETATION

BUT ..the fact remainsautomation is

NO REPLACEMENT

for the study of **Peripheral smear**

it just **COMPLEMENTS**

Manual Microscopy



AUTOMATED BLOOD CELL COUNTERS

WHAT A DOCTOR SHOULD KNOW FOR INTERPRETATION OF RESULTS

PARAMETERS PROVIDED BY AUTOMATED HEMATOLOGY ANALYZERS

Measured parameters

- Hemoglobin (**Hb**)
- Hematocrit (**HCT/PCV**)
- RBC count (**RBC #**)
- Platelet count (**PLT #**)
- WBC count (**WBC #**)
- WBC diff. (**WBC %**)

Alerts/Flags

Derived parameters

- **MCH** (Hb/RBC #)
- **MCV** (HCT/RBC #)
- **MCHC** (Hb/HCT)
- **RDW** (RBC volume)
- **MPV** (Plt TV/PLT)

Histograms & Scatter Plots (RBC, WBC & PLT)

RED CELLS

- Remarkable & dramatic improvement in the reproducibility of blood cell counters .

These parameters could be put to practical use in anemia classification.

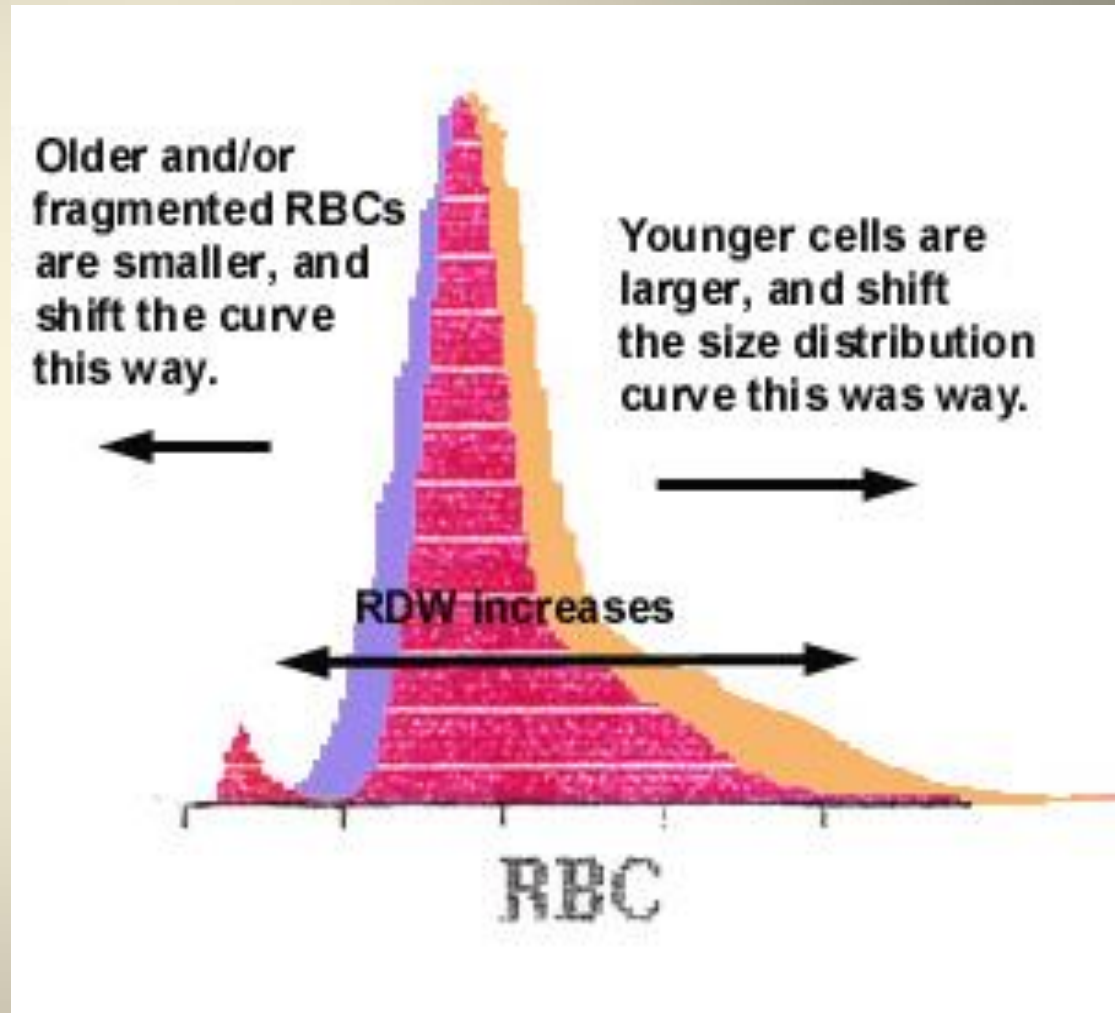
- **MCV – Mean Cell Volume (microcytes, normocytes, macrocytes)**
- **MCH – Mean Cell Hemoglobin (hypochromia etc.)**
- **MCHC – Mean Cell Hemoglobin Concentration (hemoglobinisation)**

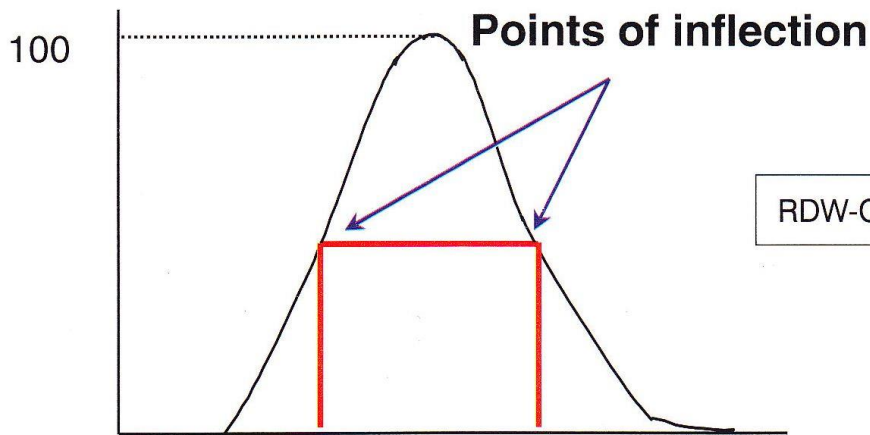
RED CELL DISTRIBUTION WIDTH (RDW)

- RDW is a measure of variation in red cell size - **anisocytosis**
- Expressed as coefficient of variation (CV 11-16%) or in standard of deviation (SD 37-46fl)
- **RDW- CV** : ratio of standard deviation(SD) of the distribution width divided by the “mean red cell volume” (MCV).
- **RDW-SD**: actual distribution width of RBC population at 20% above the baseline.
- Unlike most variables, in which there are abnormally high and abnormally low values – **No disorder is known to have abnormally low RDW.**

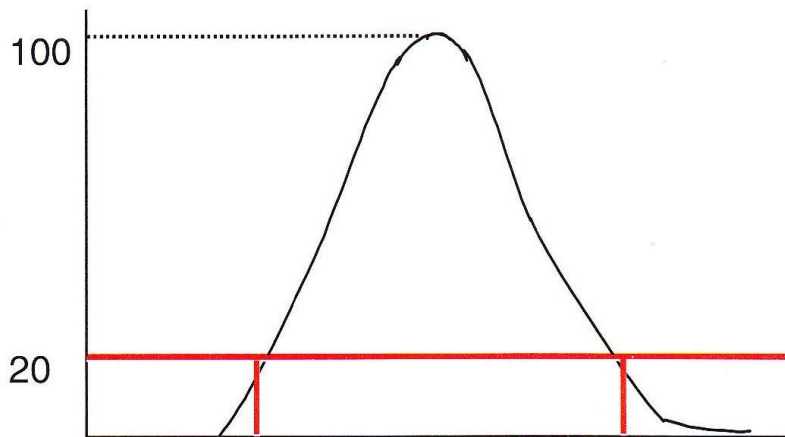
RDW

- RDW is an expression of the homogeneity of the RBC population size.
- **A large RDW says there's a wide variation in the RBC diameters within the test pool.**
- It doesn't say the cells are large or small, rather that the population is not homogenous.
- Younger cells are larger (reticulocytes).
- Older, and generally beat up, RBCs are smaller.





σ μ σ
 ↑ 68,26 %
 of all









- RDW-CV:
 Calculated by $SD/MCV \times 100$
 Normal range: 10.4-15.2%

- RDW-SD:
 Not a statistical SD
 Obtained from the RBC histogram

FIG 4: RDW

CLASSIFICATION OF ANEMIAS BASED ON MCV & RDW

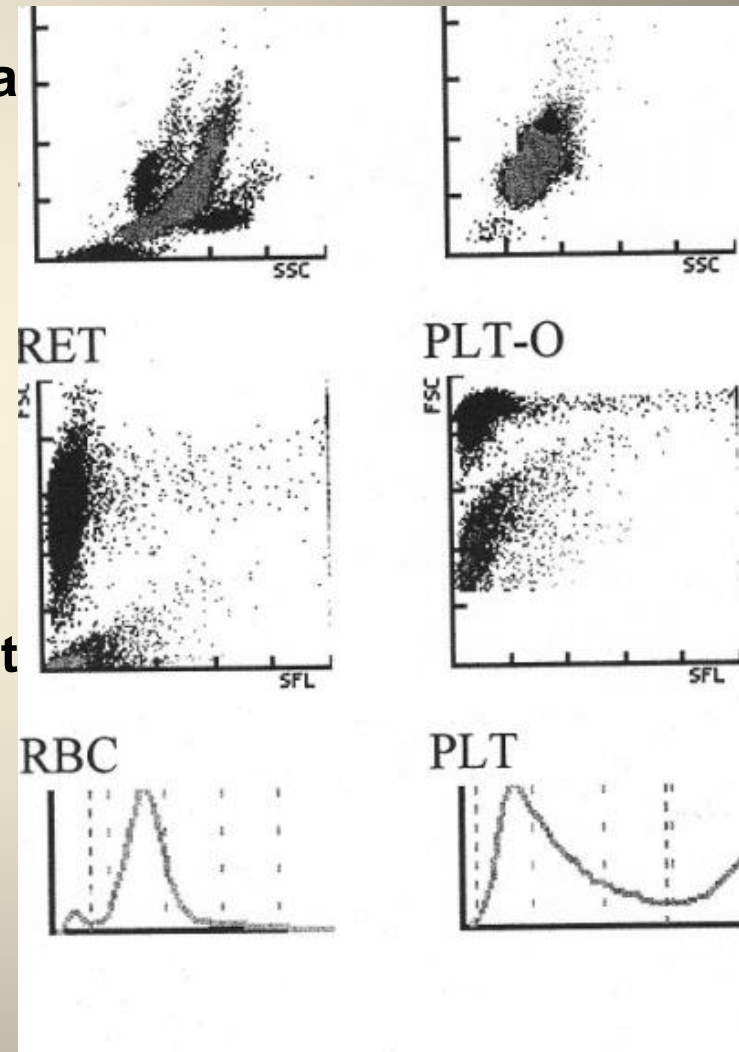
MCV Low		MCV Normal		MCV High	
RDW Normal	RDW High	RDW Normal	RDW High	RDW Normal	RDW High
Chronic disease Nonanemic heterozygous thalassemia Children	Iron deficiency Hb S- α or β thalassemia Hb H	Normal Chronic disease Nonanemic hemoglobin or enzyme abnormality Splenectomy CLL (except extreme high lymphocyte number) Acute blood	Early or mixed nutritional deficiency Anemic abnormal hemoglobin Myelofibrosis Sideroblastic Myelodysplasia	Aplastic anemia	Folate or vitamin B ₁₂ deficiency Sickle cell anemia (1/3 of cases) Immune hemolytic anemia Cold agglutinins Preleukemia Newborn
					

DISTRIBUTION CURVES/HISTOGRAMS



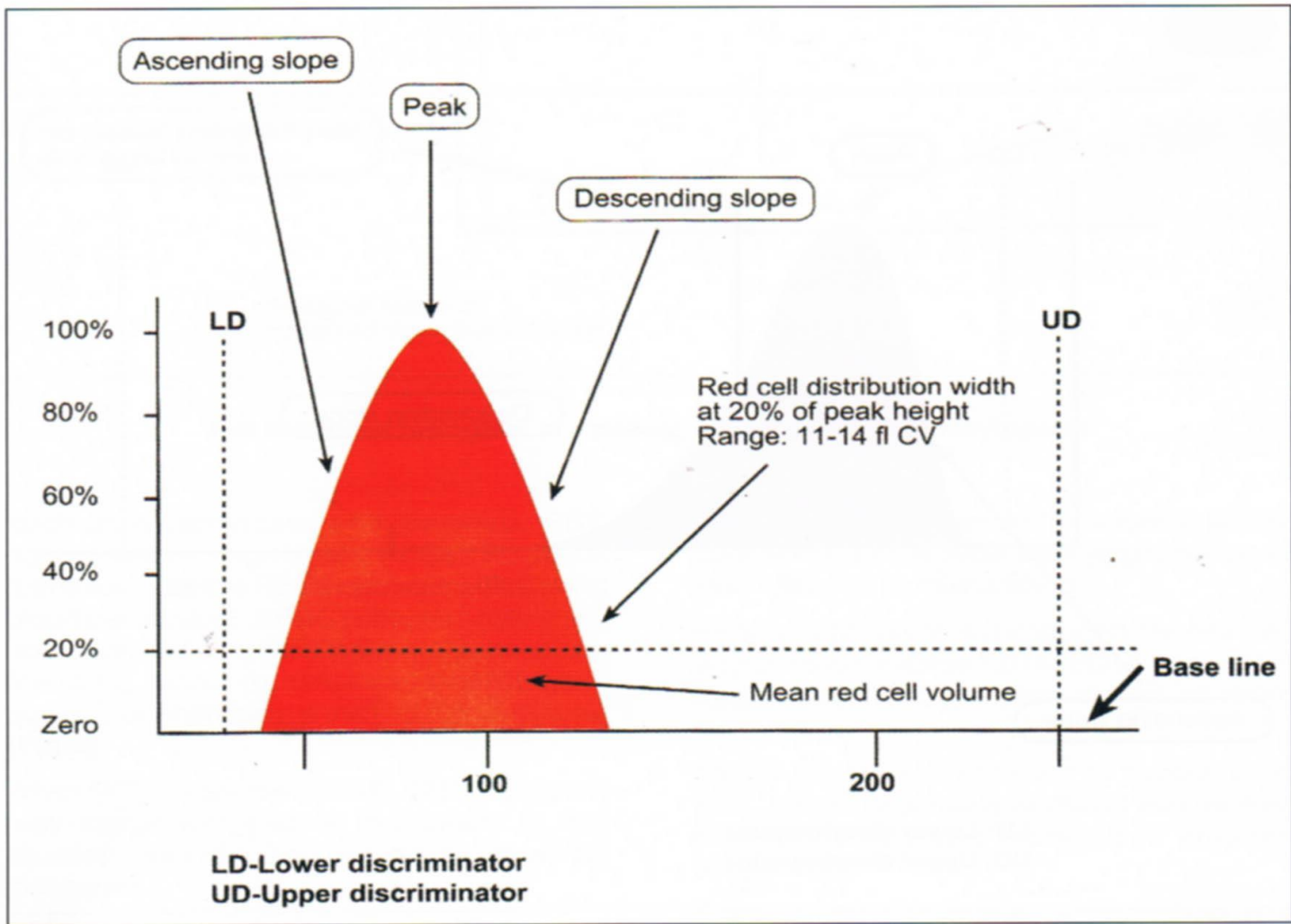
Histograms - Scatter plots

- Histograms are size distribution curves
- Graphical representation of numerical data about different cell populations
- Created for RBC, WBC and platelet populations
- Based on **cell volume and relative cell number**
- Allows visualization of:
 - Subpopulation of cells
 - Their average size in relation to the rest of the population
 - Their relative number

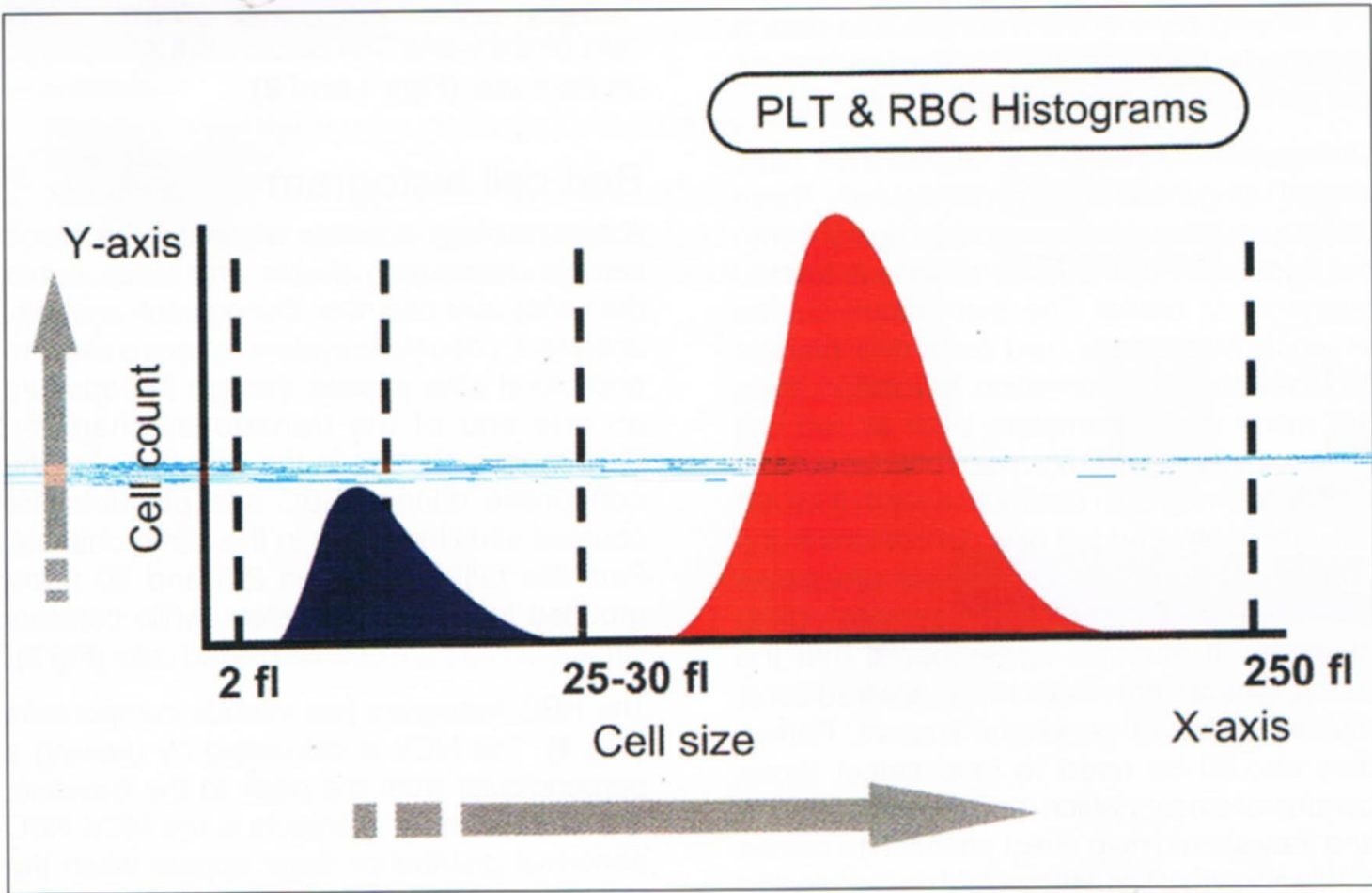


RBC HISTOGRAM





Components of RBC histogram



PLT & RBC histograms

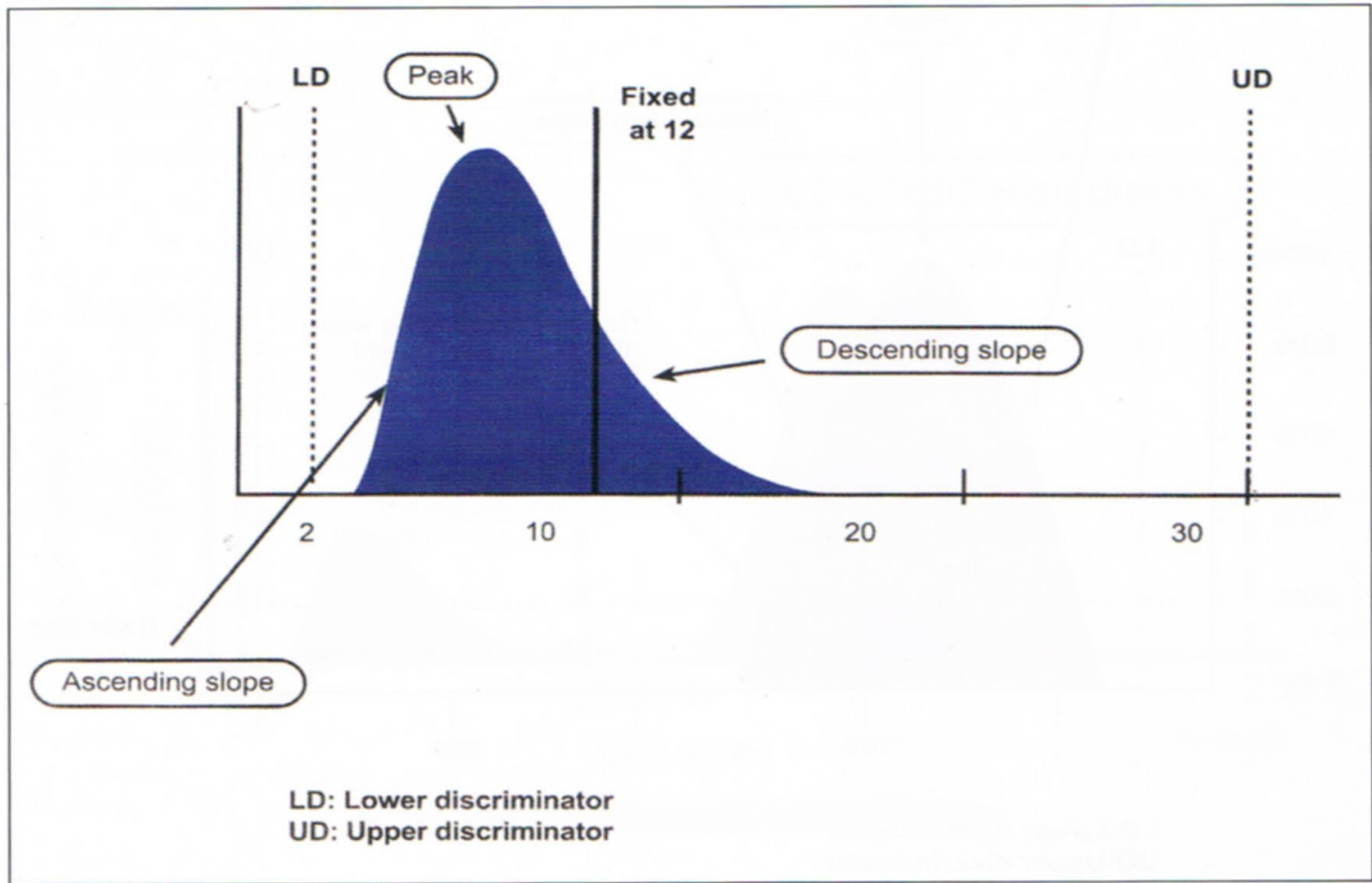
RBC ALARMS

When MCV very low <55fl, RBC histogram may merge with PLT histogram- both RBC & PLT counts inaccurate

- Falsely high RBC count: large PLTs, marked thrombocytosis, fragmented red cells.
- Falsely low RBC count: cold agglutinins, EDTA-dependent agglutination, RBC lysis d/t mishandling, extreme microcytosis
- Falsely high MCV: red cell agglutination, excess EDTA, EDTA-dependent agglutination or samples stored at room temp.
- Falsely low MCV: hypochromic RBC, severe anemia with marked thrombocytosis, increase in ambient temp.

PLATELET HISTOGRAM



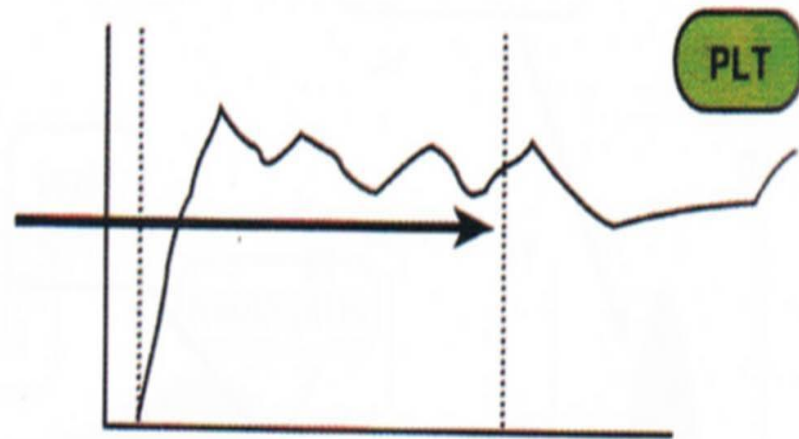


Histogram showing ascending slope, peak and descending slope

PLATELET ALARMS

- **Falsely high APC: microcytic RBCs, fragmented RBCs, HbH disease, cryoglobulin**
- **Falsely low APC: giant PLTs, PLT satellitism, EDTA-induced PLT agglutination, activation of PLT during venipuncture, partially clotted blood sample.**
- *If PLT flag generated, Scan PBF*
- *Verify PLT count manually*
- **MPV: 8-12 fl**
- **PDW at 20% of peak height, 9-14 fl**

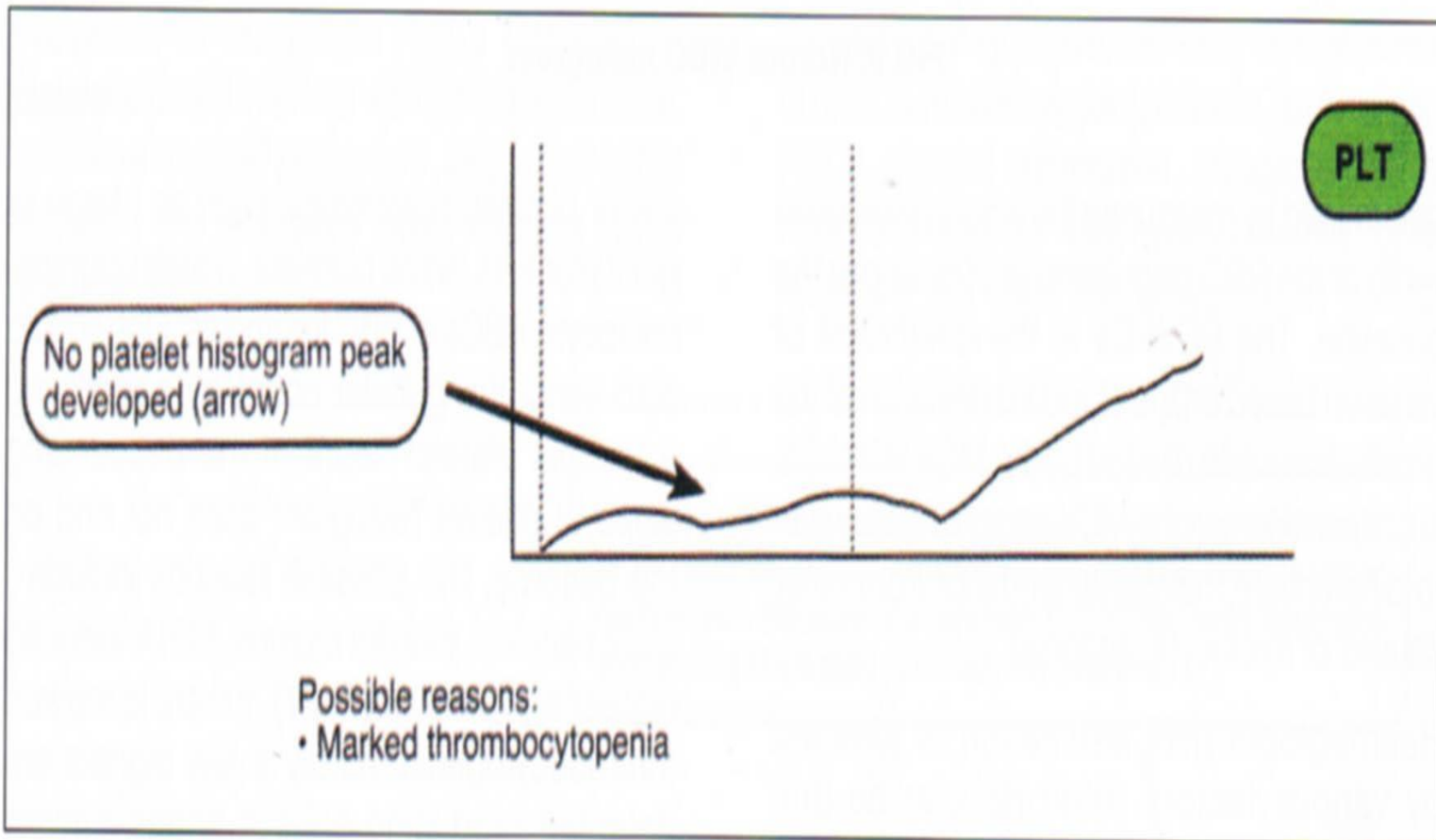
Descending slope of platelet histogram does not end on the baseline (arrow)



Possible reasons:

- EDTA induced platelet agglutination
- Giant platelets
- Platelet clumps

Platelet histogram: The descending slope is not ending on the baseline

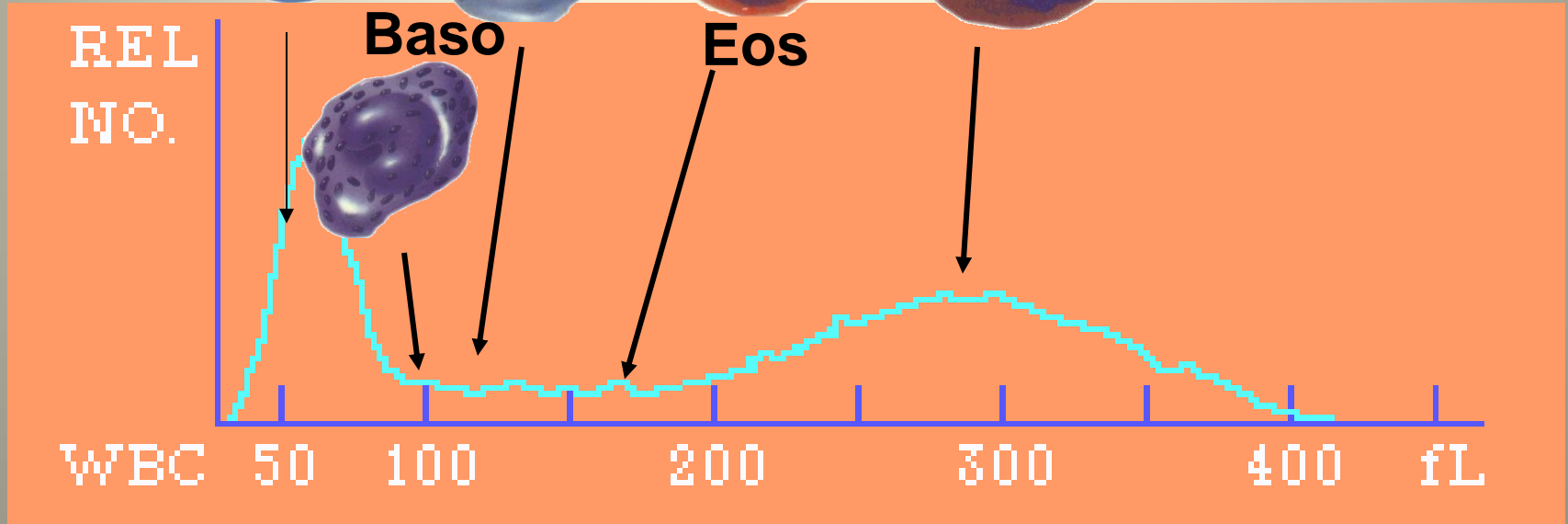
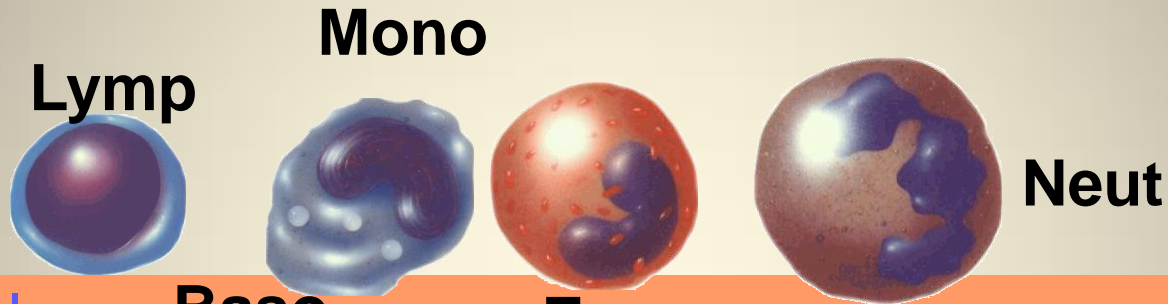


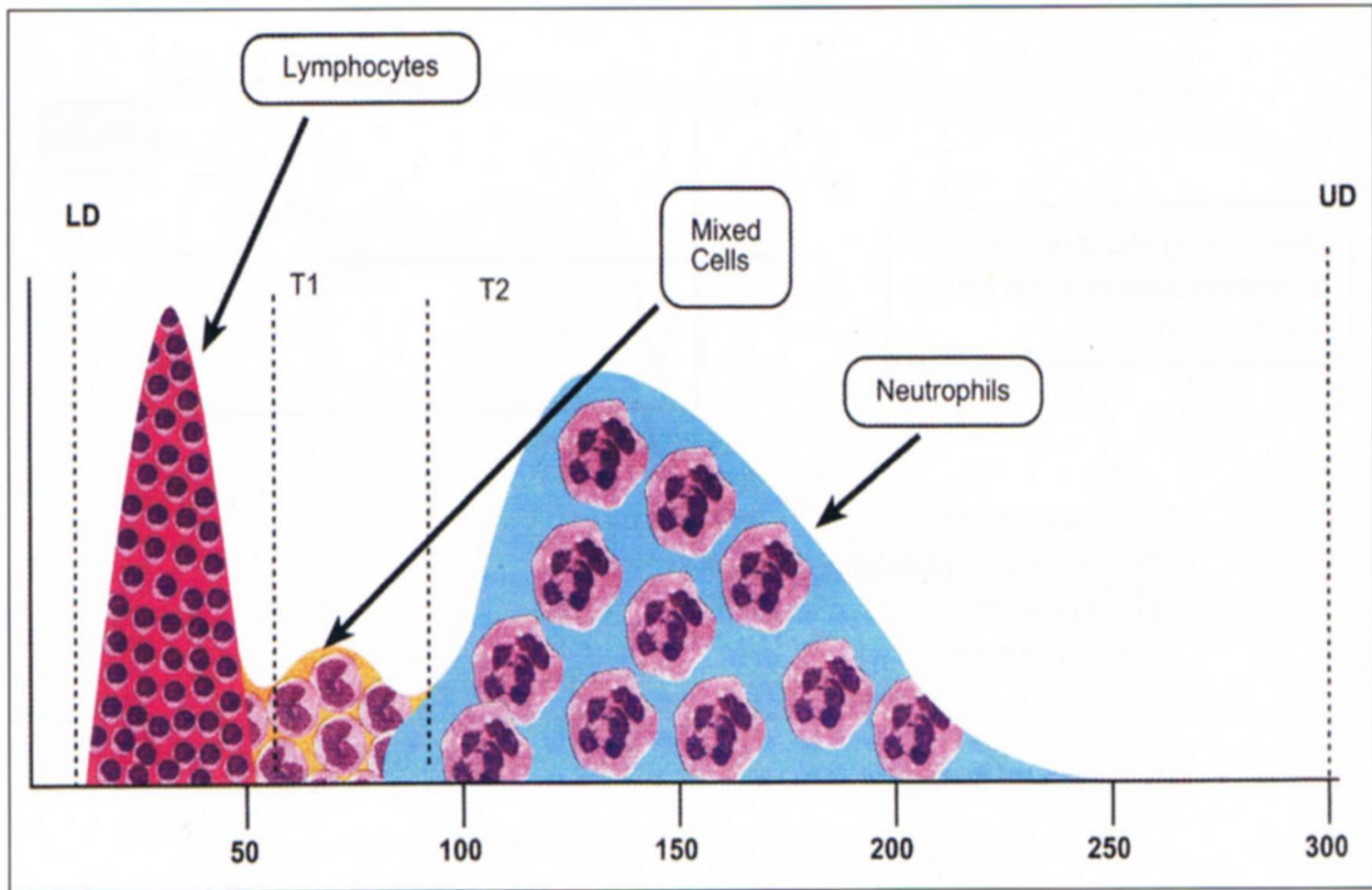
Platelet histogram in a patient with marked thrombocytopenia

WBC HISTOGRAM



WBC HISTOGRAM



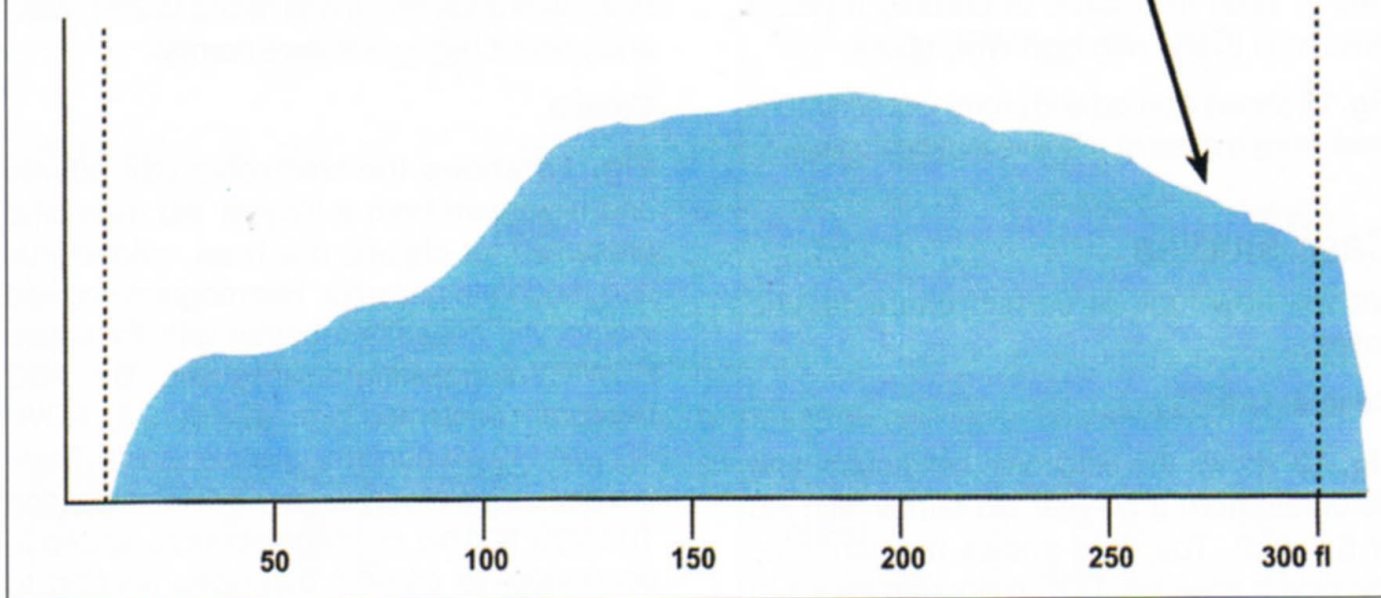


Normal WBC histogram

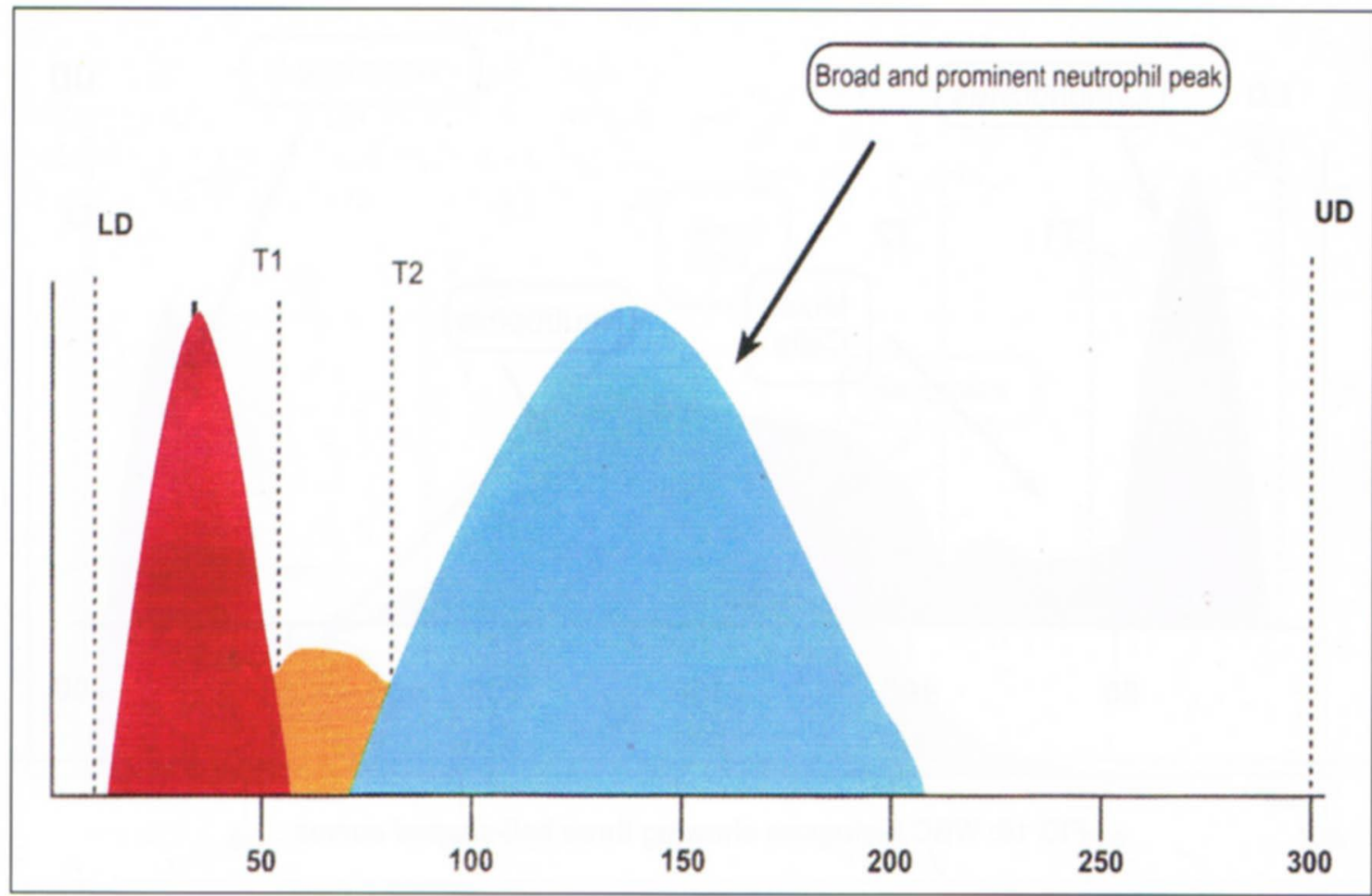
WBC HISTOGRAM: ALARMS

- Presence of NRBCs or PLT clumps interfere with lympho peak
- Immature myeloid precursors, large PMNs shift neutro peak towards UD
- Falsely high WBC count: NRBCs, giant PLT, lyse-resistant RBCs, PLT aggregates, malarial parasites, hyperlipidemia
- Falsely low WBC: WBC lysis after prolonged storage, WBC/PLT aggregates, cold agglutinins

Immature myeloid precursors, eg, myelocytes, promyelocytes & larger neutrophils, etc, shifts the neutrophil peak towards upper discriminator, Because of a very high TLC $>200 \times 10^9/l$, neutrophil descending slope may not touch the baseline.



WBC histogram from a case of CML showing one large dome shaped curve without any troughs



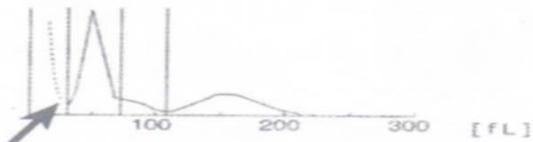
WBC histogram from a case of reactive leukocytosis showing a broad and prominent neutrophil peak

INTERPRETATION OF CBC REPORTS IN CLINICAL SETTINGS

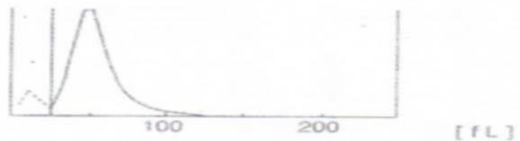


No. 12-46255
 Date 30/4/02 12:18
 Mode Whole Blood

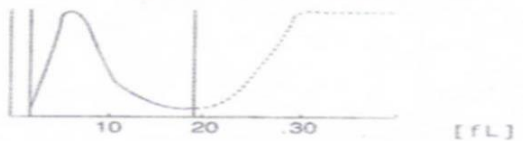
EBC	10.4	x10 ⁹ /μL
RBC	4.54	x10 ⁶ /μL
HGB	6.8	g/dL
HCT	24.2	%
MCV	53.3	fL
MCH	15.0	pg
MCHC	28.1	g/dL
PLT	667	x10 ³ /μL



LYMPH%	55.0	%
MXD%	9.1	%
NEUT%	35.9	%
LYMPH#	5.7	x10 ⁹ /μL
MXD#	1.0	x10 ⁹ /μL
NEUT#	3.7	x10 ⁹ /μL



RDW-CV + 20.5 %









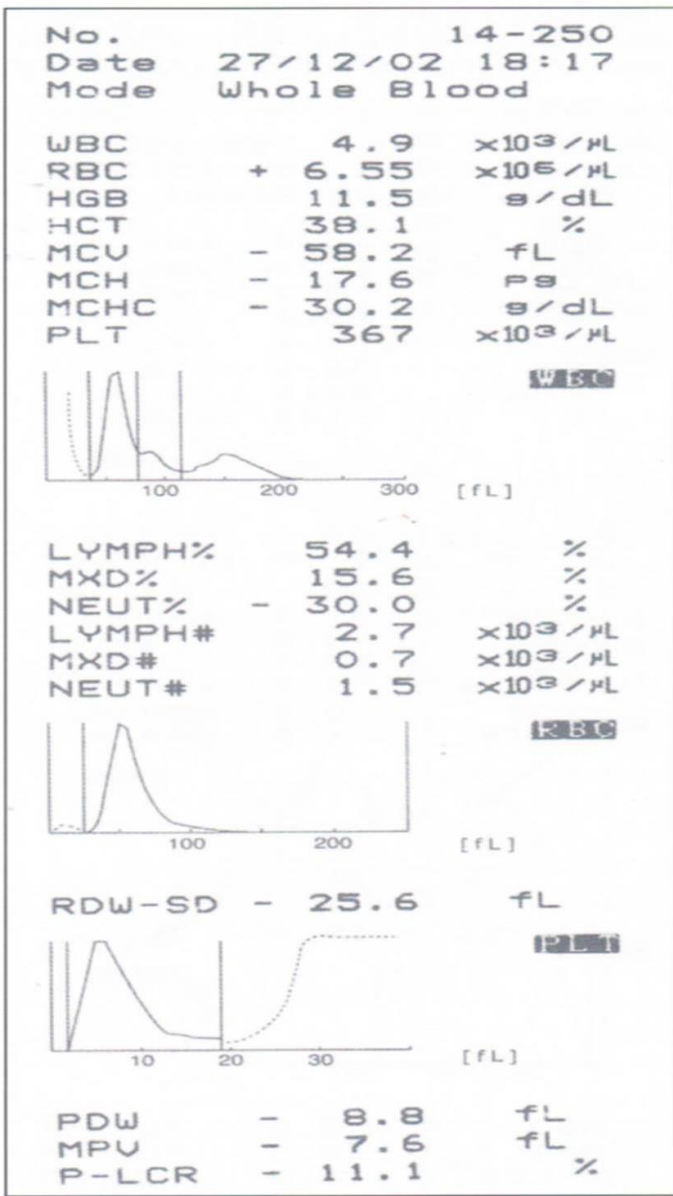
PDW	9.9	fL
MPV	8.0	fL
P-LCR	13.8	%

Iron deficiency anaemia

- 50 yr old man with marked pallor
- Microcytic hypochromic indices
- RBC histogram peaking to left, low MCV
- RDW-CV high s/o anisocytosis
- WBC & PLT histograms normal
- Diagnosis:IDA
- Confirmed by Iron studies
- Stool for occult blood

CLASSIFICATION OF ANEMIAS BASED ON MCV & RDW







MCV Low		MCV Normal		MCV High	
RDW Normal	RDW High	RDW Normal	RDW High	RDW Normal	RDW High
Chronic disease Nonanemic heterozygous thalassemia Children	Iron deficiency Hb S- α or β thalassemia Hb H	Normal Chronic disease Nonanemic hemoglobin or enzyme abnormality Splenectomy CLL (except extreme high lymphocyte number) Acute blood	Early or mixed nutritional deficiency Anemic abnormal hemoglobin Myelofibrosis Sideroblastic Myelodysplasia	Aplastic anemia	Folate or vitamin B ₁₂ deficiency Sickle cell anemia (1/3 of cases) Immune hemolytic anemia Cold agglutinins Preleukemia Newborn
					



β -Thalassaemia Minor

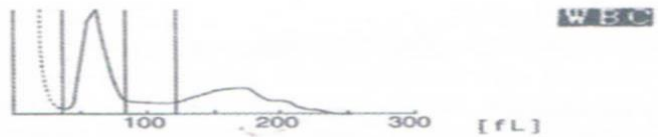
- 20 yr old primi gravida
- Hb normal but microcytic hypochromic indices disproportionately low
- Disproportionately high TRBC
- RDW-SD of 25.6 fl suggests a homogenous red cell population
- PS: uniform microcytosis
- HbA2 5.4%

CLASSIFICATION OF ANEMIAS BASED ON MCV & RDW

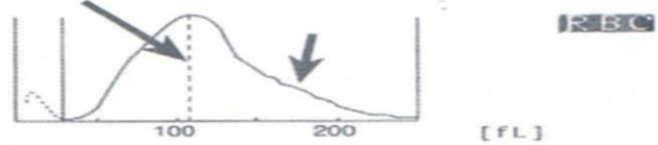
MCV Low		MCV Normal		MCV High	
RDW Normal	RDW High	RDW Normal	RDW High	RDW Normal	RDW High
Chronic disease Nonanemic heterozygous thalassemia Children	Iron deficiency Hb S- α or β thalassemia Hb H	Normal Chronic disease Nonanemic hemoglobin or enzyme abnormality Splenectomy CLL (except extreme high lymphocyte number) Acute blood	Early or mixed nutritional deficiency Anemic abnormal hemoglobin Myelofibrosis Sideroblastic Myelodysplasia	Aplastic anemia	Folate or vitamin B ₁₂ deficiency Sickle cell anemia (1/3 of cases) Immune hemolytic anemia Cold agglutinins Preleukemia Newborn
					

No. 25-4958
 Date 9/12/02 15:22
 Mode Whole Blood

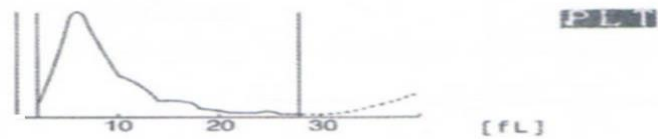
HBC 4.4 X10⁹/μL
 RBC - 2.24 X10⁶/μL
 HGB 8.2 g/dL
 HCT - 25.4 %
 MCV +113.4 fL
 MCH 36.6 pg
 MCHC 32.3 g/dL
 PLT 149 X10³/μL



LYMPH% 51.1 %
 MXD% 8.8 %
 NEUT% - 40.1 %
 LYMPH# 2.2 X10⁹/μL
 MXD# 0.4 X10⁹/μL
 NEUT# 1.8 X10⁹/μL



RDW-SD +119.4 fL









PDW 9.8 fL
 MPU - 8.5 fL
 P-LCR 18.2 %

Macrocytic anaemia due to megaloblastosis

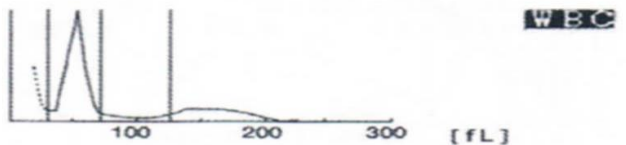
- Young male with mild icterus, moderate anemia
- Macrocytic anemia with high RDW
- Shift of RBC histogram to right- population of very large red cells 150-200 fl
- Serum B12 & RBC folate

CLASSIFICATION OF ANEMIAS BASED ON MCV & RDW

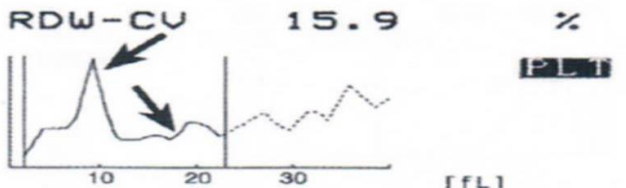
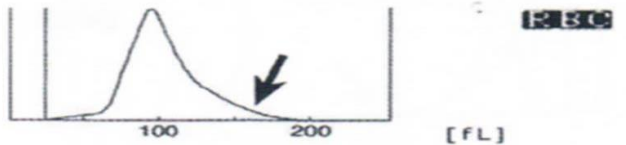
MCV Low		MCV Normal		MCV High	
RDW Normal	RDW High	RDW Normal	RDW High	RDW Normal	RDW High
Chronic disease Nonanemic heterozygous thalassemia Children	Iron deficiency Hb S- α or β thalassemia Hb H	Normal Chronic disease Nonanemic hemoglobin or enzyme abnormality Splenectomy CLL (except extreme high lymphocyte number) Acute blood	Early or mixed nutritional deficiency Anemic abnormal hemoglobin Myelofibrosis Sideroblastic Myelodysplasia	Aplastic anemia	Folate or vitamin B ₁₂ deficiency Sickle cell anemia (1/3 of cases) Immune hemolytic anemia Cold agglutinins Preleukemia Newborn
					

No. 15-44902
 Date 26/ 4/02 18:00
 Mode Whole Blood

WBC	-	1.9	$\times 10^9 / \mu\text{L}$
RBC	-	1.98	$\times 10^6 / \mu\text{L}$
HGB	-	6.7	g/dL
HCT	-	19.9	%
MCV		100.5	fL
MCH		33.8	pg
MCHC		33.7	g/dL
PLT	-	14	$\times 10^9 / \mu\text{L}$



LYMPH%	+ 66.1	%
MXD%	7.4	%
NEUT%	- 26.5	%
LYMPH#	1.3	$\times 10^9 / \mu\text{L}$
MXD#	0.1	$\times 10^9 / \mu\text{L}$
NEUT#	0.5	$\times 10^9 / \mu\text{L}$









PDW	+ 18.9	fL
MPV	12.0	fL
P-LCR	41.4	%

Aplastic anaemia

- 12 yr old boy with purpura, marked pallor, fever
- Pancytopenia
- MCV 100.5, RDW 15.9%
- RBC histogram skewed to right
- WBC histogram: lymphocyte peak, faint dome of neutrophils
- PLT histogram- abn shape, descending slope not touching baseline
- BM Bx confirmed AA

CLASSIFICATION OF ANEMIAS BASED ON MCV & RDW

MCV Low		MCV Normal		MCV High	
RDW Normal	RDW High	RDW Normal	RDW High	RDW Normal	RDW High
Chronic disease Nonanemic heterozygous thalassemia Children	Iron deficiency Hb S- α or β thalassemia Hb H	Normal Chronic disease Nonanemic hemoglobin or enzyme abnormality Splenectomy CLL (except extreme high lymphocyte number) Acute blood	Early or mixed nutritional deficiency Anemic abnormal hemoglobin Myelofibrosis Sideroblastic Myelodysplasia	Aplastic anemia	Folate or vitamin B ₁₂ deficiency Sickle cell anemia (1/3 of cases) Immune hemolytic anemia Cold agglutinins Preleukemia Newborn
					

**Anemia is not yet
apparent**

**MCV still is in the normal range and
Peripheral Smear shows mild
Anisocytosis**

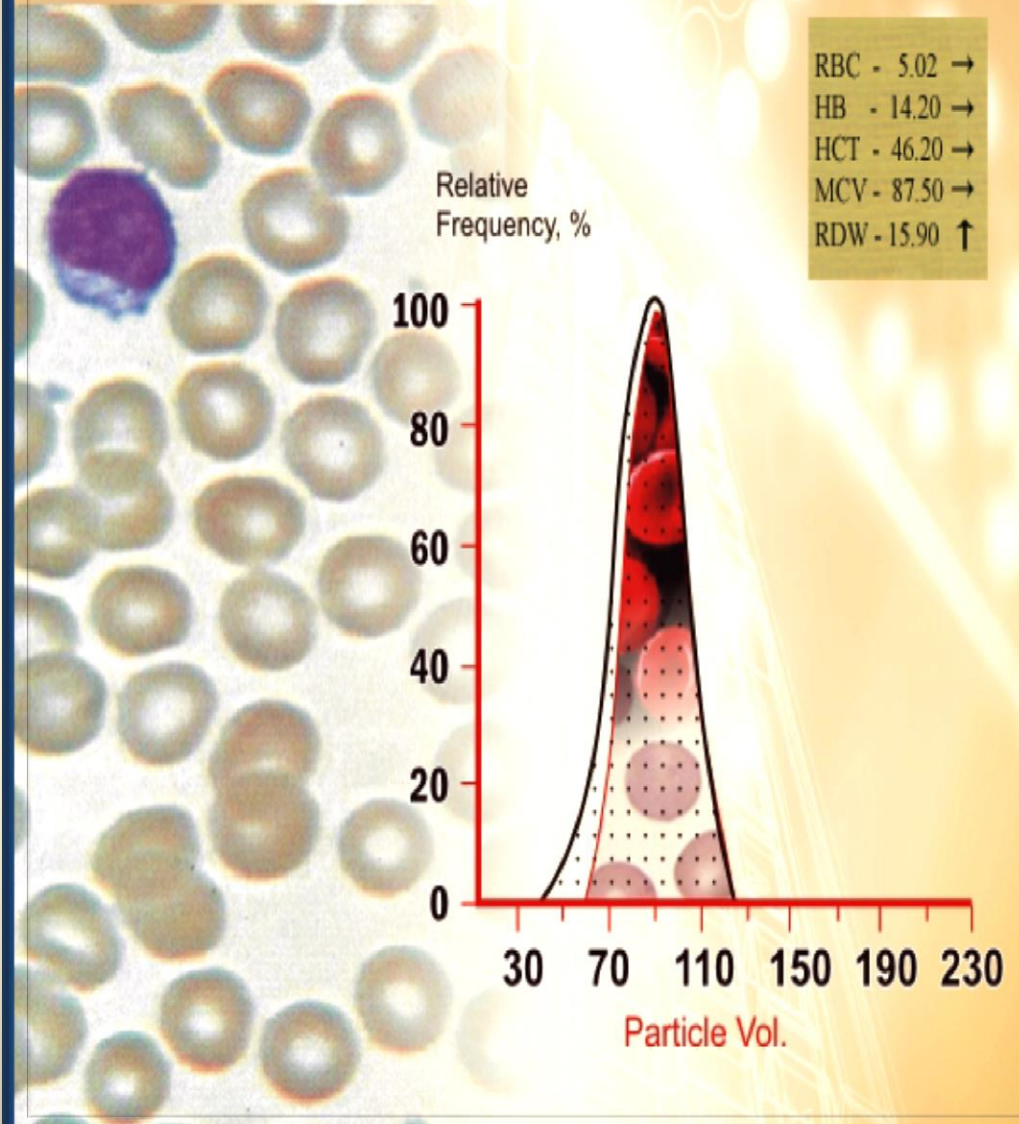
BUT

**RDW is increased (Earliest
Indicator)**

**Histogram is Unimodal
but is wider**

**Increased RDW combined
with normal RBC values
(MCV , Hb , Hct) distinguishes
Iron deficiency from Normal
Subjects**

Early Iron Deficiency

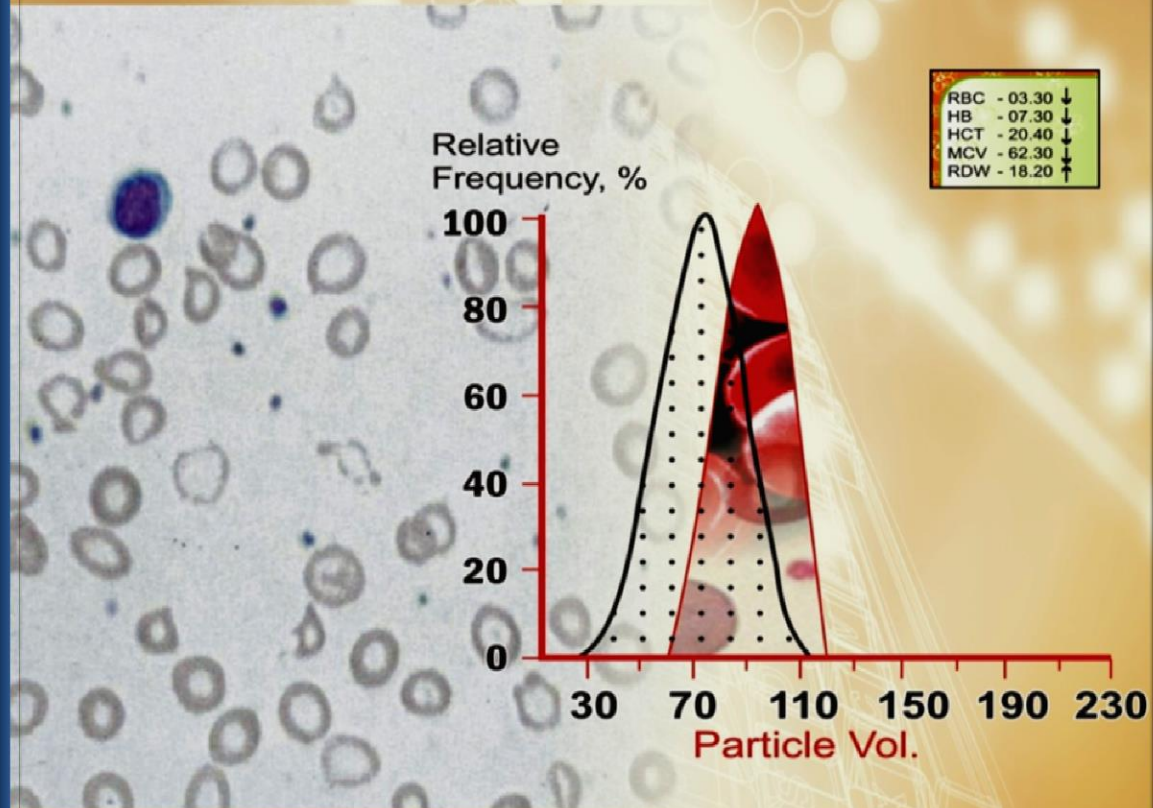


Advanced Iron Deficiency

Anemia is present, MCV is very low, and the smear is very abnormal

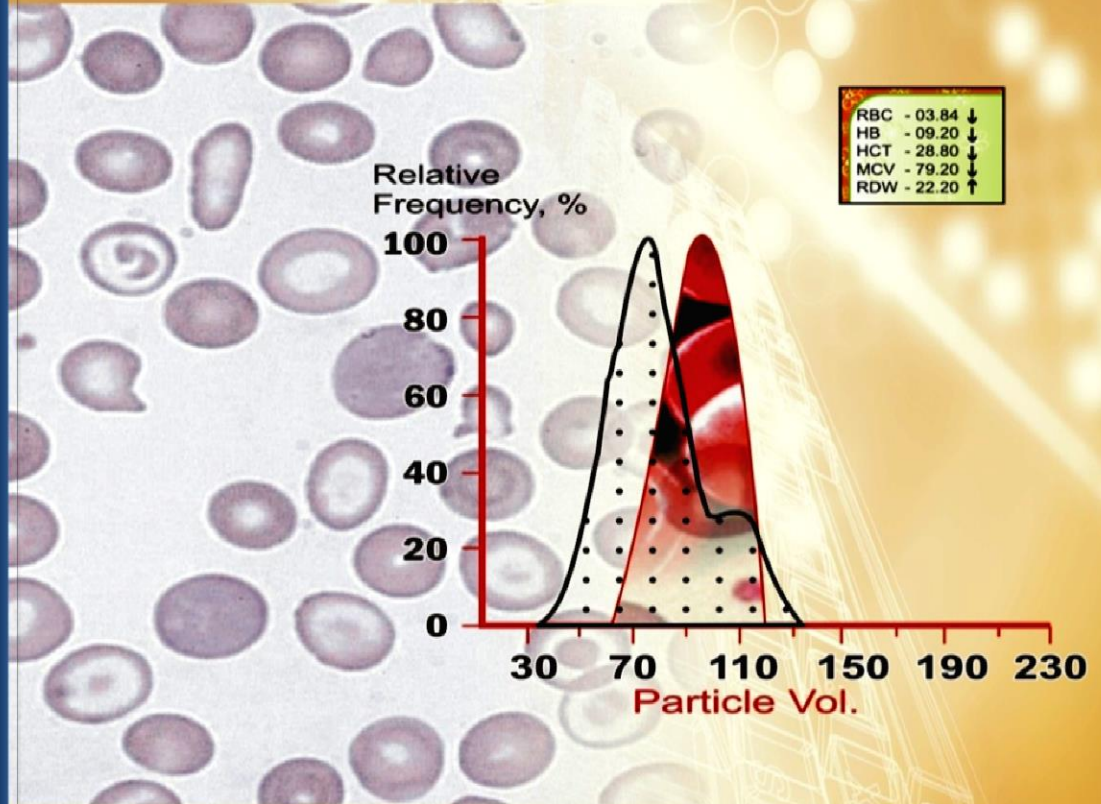
RDW is abnormally high;
Histogram remains abnormal.

The diagnosis is easily made at this point, but earlier identification would improve management



Recovery from Iron Deficiency

The red cell count is increasing,
MCV is not yet normal, and
Two populations of red cells
are seen—preexisting
microcytes, and newly
formed normocytes.
*The two populations are
distinguished easily on the
red cell histogram but not so
easily on the peripheral
blood smear.*



In contrast to earlier figure, in this case, the new cells are macrocytic.

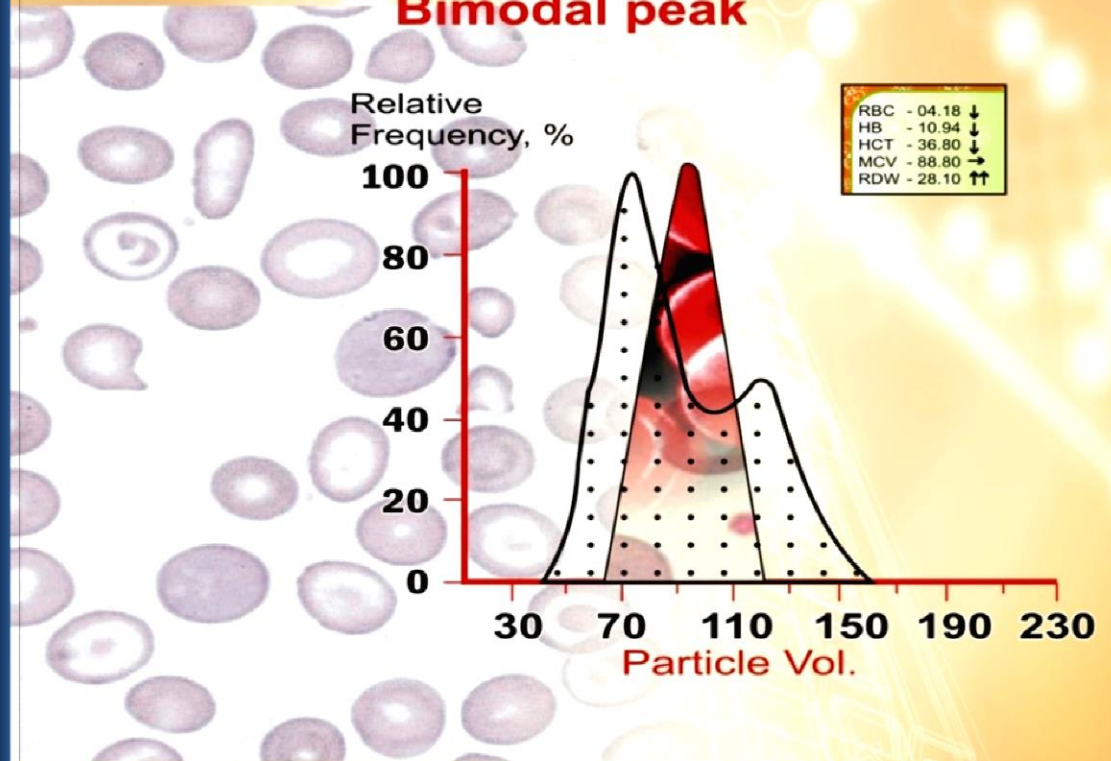
Note the right peak has a mean value of 117 fl.

This macrocytic response indicates an unmasked underlying macrocytic disorder.

Use of the histogram allows this analysis even though the MCV is only 86.8fl.

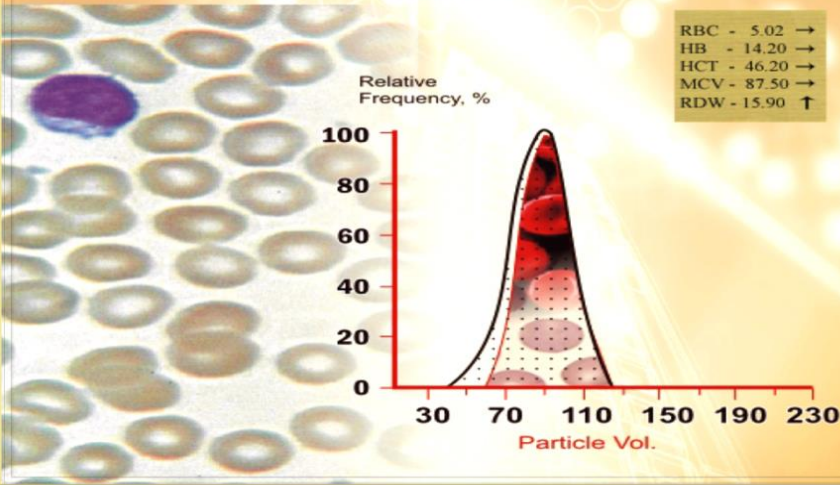
The two populations cannot be well distinguished from the blood smear.

Recovery from Iron Deficiency Bimodal peak

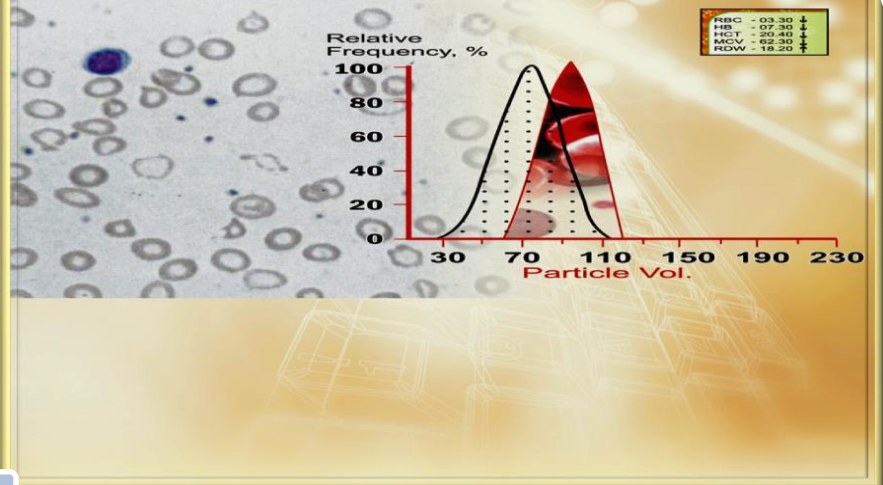


IRON DEFICIENCY ANAEMIA

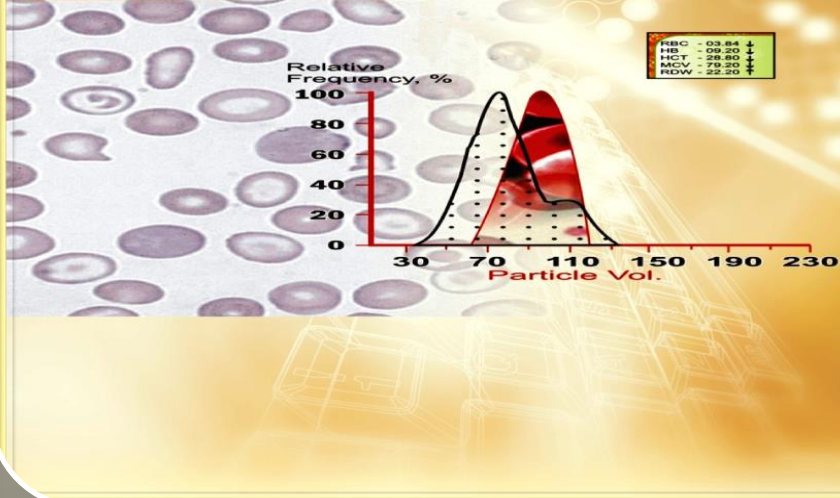
Early Iron Deficiency



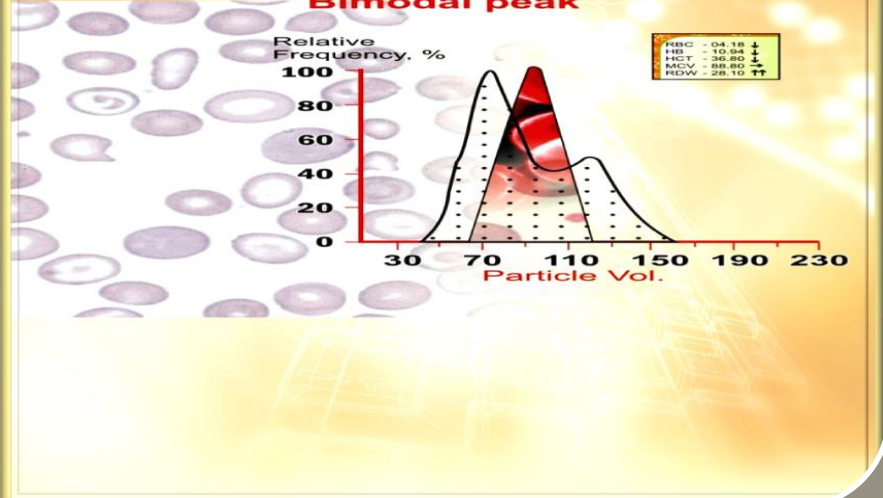
Advanced Iron Deficiency



Recovery from Iron Deficiency



Recovery from Iron Deficiency Bimodal peak



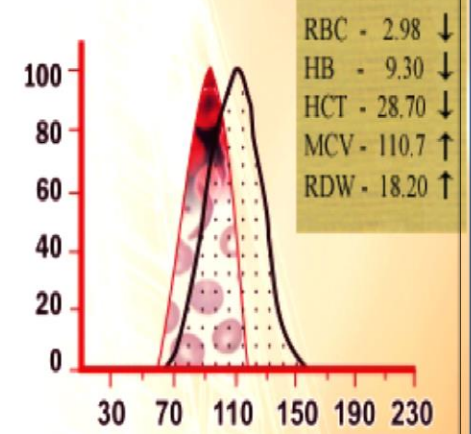
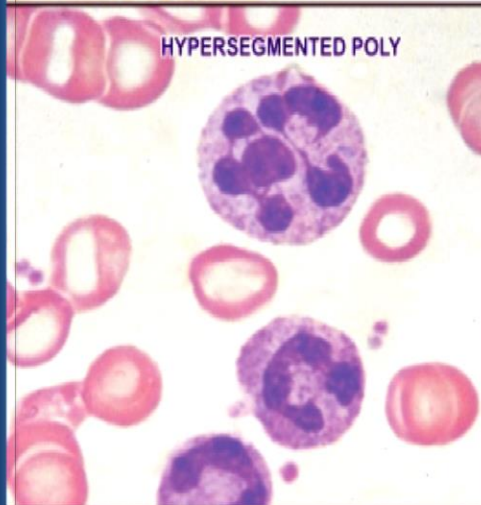
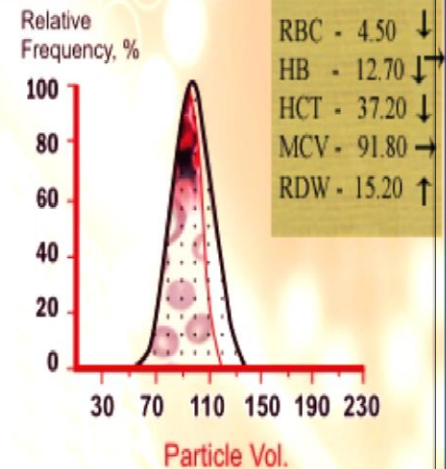
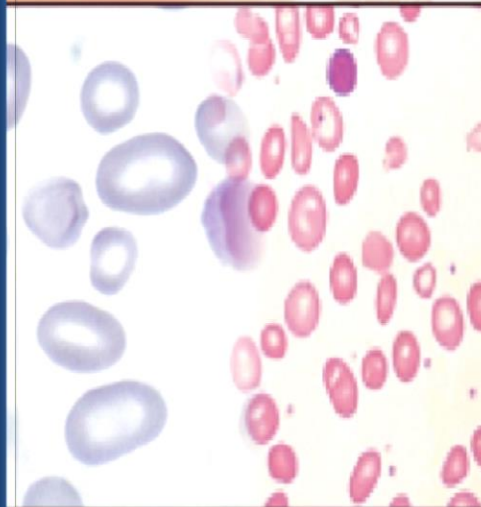
EARLY FOLATE DEFICIENCY-

- The MCV is still normal RBC count and Hb slightly reduced but
- RDW is clearly increased , even before apparent anemia.

SEVERE FOLATE DEFICIENCY –

- RBC Count is low.
- MCV is high.
- RDW is increased

EARLY MEGALOBLASTIC ANEMIA



ADVANCE MEGALOBLASTIC ANEMIA

Normocytic recovery

a small peak of cells in the normal range

- RDW is higher than untreated megaloblastic anemia due to two cell population contributing to the heterogeneity.

Microcytic recovery

Two Cell population is clearly seen in this histogram – old macrocytes and newly produced microcytes .

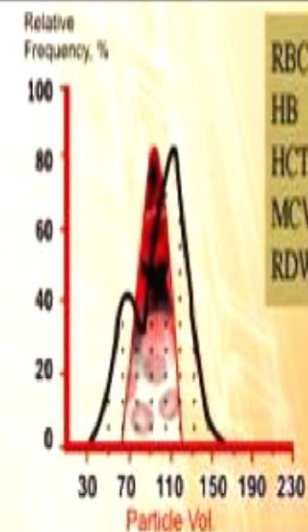
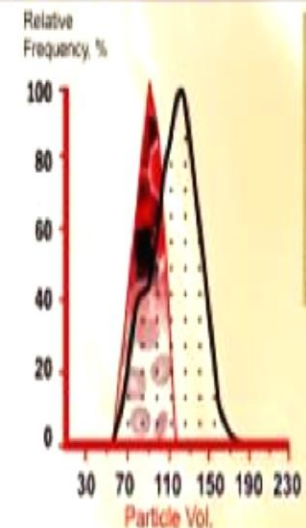
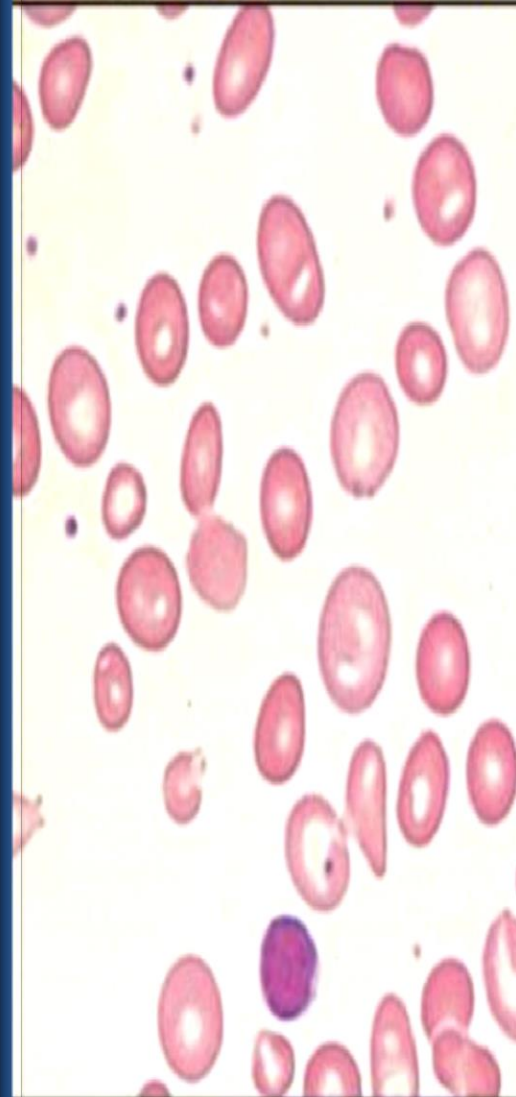
Concomitant iron deficiency has been unmasked.

RDW is markedly increased..

MCV is normal only because it reflects the average of two abnormal populations

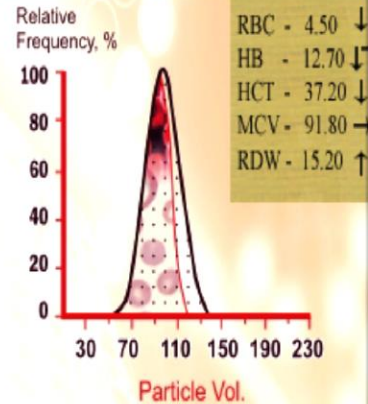
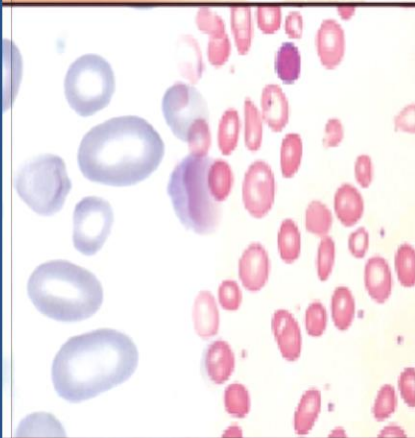
There is no population of RBC with normal MCV

RECOVERY - MEGALOBLASTIC ANEMIA

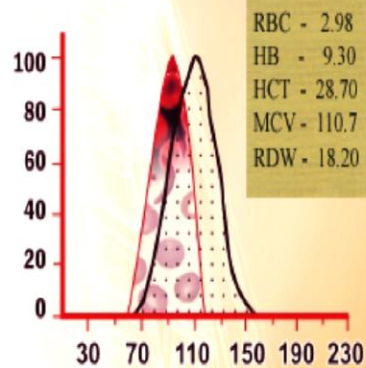
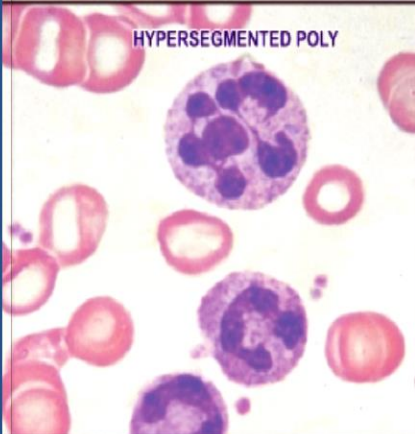


MEGALOBLASTIC ANEMIA

EARLY MEGALOBLASTIC ANEMIA

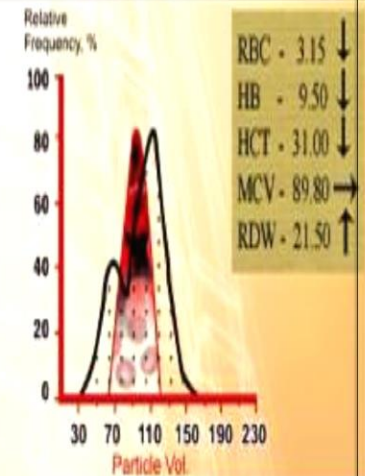
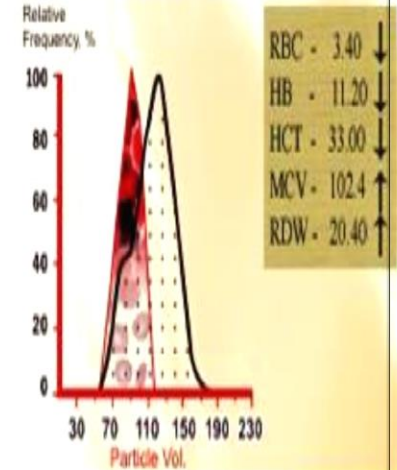
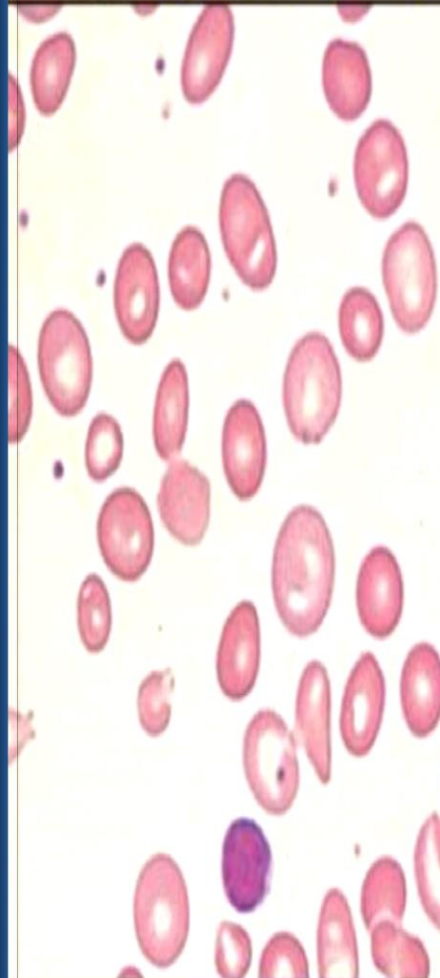


HYPERSEGMENTED POLY



ADVANCE MEGALOBLASTIC ANEMIA

RECOVERY - MEGALOBLASTIC ANEMIA



CARRY HOME MESSAGE

- **Graphics generated by hematology analyzers are a neglected piece of information; until recently, very little emphasis has been given to the graphs**
- **Often information generated by hematology analyzers in form of numerics & graphs is available at the time of initial consultation & hence becomes a guiding force in making the initial diagnosis and in planning further investigations**
- **Numerical data & graphs and manual exam of blood smear are complementary to each other in hematological work-up**



"Okay your father
managed to get a mouse.
Now how do we use it?"

ORGANIZING COMMITTEE

PATRONS

- **Prof Subhash Nepalia**
Principal & Contoller
SMS Medical College &
Attached Hospitals
- **Prof S R Mehta**
Emeritus Professor of Medicine
Chairman, Four Decade Trust

ORGANIZING CHAIRPERSONS

- **Prof C L Nawal**
Chief of Medicine
- **Prof Hemant Malhotra**
Chief of Medical Oncology
- **Prof Ajay Yadav**
Chief of Pathology
- **Dr V D Maheshwari**
Chairman, Jaipur Chapter of API

PROGRAM DIRECTOR

- **Prof Sudhir Mehta**

PROGRAM CO-DIRECTORS

- **Dr Upendra Sharma**
- **Dr Seema Mehta**
- **Dr Sandhya Gulati**
- **Dr Nidhi Sharma**

REGISTRATION DETAILS

- **Registration Fee : Rs 300/-** only payable by **Cash/Demand Draft** in favor of **"SMS Medical College Four Decade Trust, Jaipur"** (No cheques will be entertained)
- Duly filled Registration Form (enclosed herewith) along with registration fee (by demand draft/cash) should reach the Program Director on or before 8th February, 2014
- Registration fee is **non-refundable**.
- All registrations after 8th February 2014 will be treated as spot registration. **On-spot registration fee is Rs 600/-** subject to availability of seats.

**Last Date for submission of duly filled Registration Form
Saturday, 8th February, 2014**

ADDRESS FOR ALL CORRESPONDENCE & SUBSCRIPTION :

Dr Sudhir Mehta

Program Director

5th Regional CME in Haematology

16 Ganesh Colony, Opposite Soni Hospital, JLN Marg, Jaipur-302004

Tel : 0141-2620710 • Mobile : 094140-42033

Email : s.smehta@hotmail.com

5th Regional

CME IN HAEMATOLOGY 2014



Sunday, 16th February , 2014

**Venue : Shri Mahaveer Digamber Jain School Auditorium
Mahaveer Marg, 'C' Scheme, Jaipur**

UNDER THE AUSPICES OF

- SMS MEDICAL COLLEGE FOUR DECADE TRUST
- DEPARTMENT OF MEDICINE, SMS MEDICAL COLLEGE
- DEPARTMENT OF PATHOLOGY, SMS MEDICAL COLLEGE
- JAIPUR CHAPTER OF API

Invitation

Dear Friends,

It is our pleasure & privilege to invite you all to the 5th Regional CME in Haematology being organized under the auspices of SMS Medical College Four Decade Trust. Departments of Medicine & Pathology, SMS Medical College & Jaipur Chapter of Association of Physicians of India. With each passing year, this CME is growing & making its presence felt in neighboring states and in North India. All this has been possible due to your enthusiasm & support.

The biggest & most precious catch of this CME is the world renowned hematologist. Dr. Near Young. He is not unknown to those who have passion for hematology. Dr. Young is the Chief of the Haematology branch of the National Heart, Lung & Blood Institute and Director of the Trans-NIH Centre for Human Immunology, Auto-immunity & Inflammation, Bethesda, Maryland. The national faculty includes stalwarts in the field of Haematology- Dr M B Agarwal, Dr R Manchanda, Dr Pritesh Junagade & Dr Pankhi Dutta. We are making great efforts in offering you a high-quality scientific program and the opportunity to interact one-to-one with the invited faculty. The selected topics are of immense interest and everyday relevance to physicians, pediatricians, pathologists & gynecologists.

Wishing you all a very happy & prosperous New Year 2014




Dr C L Nawal




Dr Hemant Malhotra



Dr Ajay Yadav



Dr V D Maheshwari



Dr Sudhir Mehta

FACULTY



Dr Neal Young
(USA)



Dr M B Agarwal



Dr R Manchanda



Dr Pankhi Dutta



Dr Pritesh Junagade

PROPOSED TOPICS

- ABC of CBC (Case-based)
- Interpreting Lab data in anemia (panel discussion)
- Life without platelets
- XYZ of coagulation
- Approach to evidence management of DVT
- Role of Haematologist in PUO
- Advances in understanding & treating aplastic anemia
- Antiphospholipid syndrome-then & now
- PS-a picture speaks thousand words
- Intricacies in lab diagnosis of lupus anticoagulants
- Role of bone marrow trephine biopsy in benign haematology (case-based)

**LAST DATE FOR SUBMISSION OF DULY FILLED REGISTRATION FORM
SATURDAY, 8TH FEBRUARY 2014**



