The Complete Blood Cell Count (CBC) CBC - Part 1: The Hemogram

A Clinical Pathology 201 Study Module

by

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Special thanks is given to Angela Foley, MS, **MT(ASCP)**, Department of Clinical Laboratory Sciences, LSUHSC School of Allied Health in New Orleans, LA for the use of some of her images of blood cells and for her assistance in the art of creating image files...

...and to W. Douglas Scheer, PhD, Department of Pathology, LSUHSC School of Medicine in New Orleans, LA for converting the document for internet access.





This is the first module of a 4-part study exercise regarding the CBC. The four parts are entitled:

- CBC Part 1 The hemogram
- CBC Part 2 WBC differential & blood morphology
- CBC Part 3 RBC morphology & platelet estimate
- CBC Part 4 Post-test





Feedback as to the quality and usefulness of this exercise is solicited and suggestions for improvement are welcomed. Please forward your remarks by E-mail cwalte@lsuhsc.edu

or via US MAIL:

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The directions for navigating through the exercise are given on the next 3 pages. They are the same as those used in the other modules of this 4-part exercise. Click on:



to visit the directions before continuing with the exercise.





to bypass the directions.



Directions, d

The following directional icons are provided throughout the exercise for your convenience. You can click on:



in the <u>upper left</u> hand corner of every page to return to the <u>previous page</u>

menuin the upper right corner of the page toreturn to the Hemogram Menuselection.





You can click on:



in the <u>lower right</u> corner of the page to continue.



in the <u>lower right</u> corner of the <u>Main Menu</u> page to <u>Quit</u> (i.e., end the exercise).





Directions, d

"Hot points" (symbols, words, phrases) have been inserted on the pages as navigational tools and can be identified by their "gold" color. If it's "gold", click on it to move to the next text/data entry. Also, sounds have been added in a few places for emphasis.

<u>Caution</u>, failure to follow the structured order of the "hot points" may result in confusion. If you use the mouse without placing the cursor directly on the "gold hot point" or click without waiting for the "gold" to appear, you may skip over vital information.

Remember, if it's gold, click on it. Try it!



Special Comments

This exercise has numerous images. You may note that, when a page contains images, there may be a rather long delay before you regain control of the cursor. Please be patient. I think you will find the images are worth the wait.

NOTE:

Some animation and/or interactive affects may be lost if you attempt to replay a page by returning to the previous page and then advancing to that page again.

Now, click on the gold to begin.



CBC – Part 1

The Complete Blood Cell Count (CBC)

Part 1 - The Hemogram





Does the CBC have clinical value ?

A CBC (complete blood count) is one of the most frequently ordered laboratory procedures. It's useful in the diagnosis and clinical management of numerous diseases and disorders, such as: anemias

- **leukemias & other neoplasias**
- infections (bacterial & viral)
- inflammatory disorders (e.g., rheumatologic) inherited anomalies



A CBC (complete blood count) can be performed by automated electronic instruments or by manual methods on a whole blood specimen collected:

venipuncture

 in a tube containing EDTA anticoagulant
Or

capillary stick (e.g., finger, heel, ear)

ck - in a vial containing a measured volume of diluent appropriate for the method used





What is a CBC?

A CBC is a battery of hematologic tests. The values obtained provide valuable information regarding the three types of blood cells found in peripheral blood, which are red blood cells (RBC), white blood cells (WBC), and platelets (PLT).





Three types of cells in peripheral blood:

1. erythrocytes (RBC)



3. platelets (PLT)



2. leukocytes (WBC)

neutrophils

eosinophils basophils lymphocytes monocytes













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What information is provided by a CBC?

Basically, the CBC provides information regarding the:

- number of red cells, white cells, and platelets in circulating peripheral blood
- frequency distribution of white blood cells
- morphologic features of the blood cells
- hemoglobin content of red blood cells
- relationship of red blood cells to total blood volume and hemoglobin concentration



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What are the components of a CBC? In most laboratories, due to advanced technologies now available, an automated CBC is performed on a multi-channel instrument employing a variety of techniques. It usually includes a:

- **1** Hemogram
- **2** Differential WBC count
- **3** General description of blood cell morphology (WBC, RBC, and PLT)
- 4 Platelet estimate











HEMOGRAM MENU

- Introduction
- **Total WBC Count**
- Corrected WBC Count
- Total RBC Count
- **Hemoglobin**
- Hematocrit
- Erythrocyte (RBC) Indices
- Red Cell Distribution Width (RDW)
 - Platelet Count
- Mean platelet volume













What is a hemogram?

The hemogram components of a CBC are hematologic assays/procedures that provide useful information regarding the red blood cells (RBC), white blood cells (WBC), and platelets (PLT).

Automated electronic instruments are able to:

- enumerate the number of each of the three blood cell types
- differentiate normal from abnormal cells
- provide a variety of information related to each





What are the components of a hemogram?

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WBC - total number of white blood cells per μL of blood (SI units = per L)

- RBC total number of red blood cells per μL of blood (SI units = per L)
- HGB average number of grams of hemoglobin in red blood cells per dL of blood (SI units = per L)
- HCT hematocrit or % of packed red blood cells per unit of blood







Are the RBC indices included in a hemogram?

- The RBC indices is usually provided as part of an automated hemogram and includes.
- MCV (mean corpuscular volume) average size (μ³) of the red blood cells
- MCH (mean corpuscular hemoglobin) average hemoglobin content (μμg) in individual red blood cells
- MCHC (mean corpuscular hemoglobin concentration)
 - average hemoglobin concentration (%) per unit of packed red blood cells



What additional components are frequently included in a hemogram?

Depending upon the laboratory and available instrumentation, the hemogram may also include:

PLT - total number of platelets per μL of blood (SI units = per L)

MPV (mean platelet volume) - average size (μ³) of individual platelets

RDW (red cell distribution width) - an index of the variation in size of the red blood cells

Each parameter of the hemogram will be discussed later.

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Are the parameters of automated hemograms direct or indirect measurements?

Hemogram data obtained via an automated multi-channel instrument are obtained by one of two methods.

Direct counts/measurements

or

Indirect calculated measurements







Which components are:

Direct counts or measurements? WBC RBC RBC HGB PLT MCV Indirect calculated measurement based on direct measurements of other parameters? HCT **MPV RDW**







Are the RBC indices measured directly or indirectly?

A combination of direct and indirect measurements are used:

MCV (direct measurement of RBC volumes or sizes)

MCH (indirect calculation based on direct measurements for HGB and RBC)

MCHC (indirect calculation based on direct measurements for HGB and HCT)



- In some instances, interference factors can affect the validity of measurements.
- When evaluating the validity of direct measurements, interference factors affecting only the parameter measured must be considered.
- When evaluating the validity of indirect measurements, interference factors affecting each parameter used to calculate the indirect measurement must be considered.
- Interference factors are discussed briefly at the end of this exercise.



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How do you know if patient values are normal or abnormal?

A reference range of values for adults is provided for each parameter of the CBC. These values, which are generally considered to be normal, may be gender dependent for some parameters.

CBC patient values printed out by the multichannel instruments are usually flagged when higher or lower than the reference values.

Be aware that the reference ranges for children are different from adults and may vary according to age group (e.g., newborn, infants 10-17 months, child 1.5 - 4 years).





Always refer to the reference ranges established by the laboratory performing the CBC before evaluating patient results. Variation among laboratories may be seen due to different methodologies and/or patient populations.

every laboratory?

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Para	<u>meter</u>	<u>Units</u>	<u>Range</u>	
WBO		x 10³/μL	4.5 - 11.0	
RBC		<mark>x 10⁶/µL</mark>	m = 4.60 - 6.20 f = 4.20 - 5.40	
HGE	3	g/dL	m = 13.5 - 18.0 f = 12.0 - 16.0	
НСТ		%	m = 40.0 - 54.0 f = 38.0 - 47.0	



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Parameter	<u>r Units</u>	<u>Range</u>
MCV	μ ³	80 - 96
МСН	μμ g	26 - 34*
МСНС	%	31 - 37*
PLT	x 10³/μL	150 - 450
MPV	fL	6.5 - 12.0
RDW		11.6 - 14.6







End of Introduction

This concludes the Introduction to the Hemogram Section. Select one of the following:

Go to Total WBC Count, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.



Total WBC Count





A total white blood cell count is the number of leukocytes present per unit of peripheral blood (e.g., $6,000/\mu$ L).

WBC normally present in peripheral blood and included in the total WBC count are mature forms of neutrophils, eosinophils, and basophils; a few band neutrophils (about 0-5%); lymphocytes (mature and occasional atypical forms), and monocytes.

view WBCs or continue


WBC in Normal Blood (Adults):



neutrophils ____

eosinophil

basophil

menu







lymphocyte



*ATL (few) *atypical lymphocyte



In disease states, what other leukocytes may be included in the total WBC count ?

menu

When present in disease states, in addition to leukocytes normally circulating in peripheral blood, the total WBC count may also include:

Immature leukocytes

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 Leukoctyes with <u>abnormal</u> <u>morphologic</u> <u>alterations</u> (which may be acquired or inherited)

 Leukocytes associated with <u>neoplastic</u> disorders (e.g., leukemia)

view WBCs or continue

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Examples of immature WBC:



granulocytes (various stages)

myeloblasts



menu



lymphoblasts

monoblasts





Examples of WBC with acquired nonneoplastic abnormal alterations:

in megaloblastic

anemias

neutrophils

&

in bacterial infections



with Dohle bodies and/or toxic granulation with nuclear hypersegmentation (> 5 lobes)



menu

in viral infections



&

with reactive (atypical) changes





&

Examples of WBC with inherited nonneoplastic abnormal alterations:

&

&

Pelger-Huet Anomaly



hyposegmented neutrophils

May-Hegglin Anomaly



neutrophils w/ inclusions

Chediak-Higashi Syndrome

Alder-Reilly Anomaly



neutrophils w/ abnormal granules neutrophils and lymphocytes w/ inclusions









Examples of neoplastic WBC alterations:

hairy cell leukemia





hairy cell lymphocytes

acute myelocytic leukemias



myeloblasts w/ Auer rod(s)





Examples of WBC in acute leukemia:

AML -myeloblasts



AMML - monoblasts



ALL - lymphoblasts





Examples of WBC in chronic leukemia: Isuhsc

CML - granulocytes (various stages)

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caution

That's a nucleated RBC. more about NRBC later

CLL - mature lymphocytes



CLL - hairy cell lymphocytes







Are leukocytes the only cells included in the total WBC count?

Remember, all nucleated cells are included in the total WBC count. If present, nucleated red blood cells (e.g., in newborns and some disease states) will also be counted.





Examples of nucleated RBC in various stages of maturation:



proerythroblast



orthochromatophilic erythroblast



polychromatophilic erythroblast









If NRBC are included in the count, how can the number of WBC be determined?

When nucleated red blood cells are included, the total WBC count must be corrected for their presence. (Refer to the Corrected Total WBC Count Section on the Menu.)

A corrected total WBC count is determined manually and added to the report as a "corrected WBC".





Of what clinical importance are WBC counts?

WBC counts may be abnormal in disease states. WBC counts are characteristically:

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- e some viral infections
- acute leukemias (some
 - cases/phases)
- e during chemotherapy

- infections (bacterial and some viral)
 - and some viral)
- Ieukemoid reactions
- Ieukemias
- myeloproliferative disorders







Does the total WBC count differentiate WBC as to cell line?

No, the total WBC count is the total number of all nucleated cells. In the case of abnormal total WBC counts, a differential WBC count must be performed before it can be determined which cell line is decreased or increased.

It is also important to determine whether the increase/decrease is a relative percent or absolute number, which is discussed in the Differential WBC section presented later.



Isuhsc End of Total WBC Count

This concludes the Total WBC Count Section. Select one of the following:

Go to <u>Corrected WBC Count</u>, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.











While performing a differential WBC count on a Wright's or Wright's-Giemsa stained peripheral blood smear:

Count at least 100 WBC, e.g.,



PMNs bands eosinophils basophils monocytes lymphocytes & ATL



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...procedure for correcting the total WBC count for NRBC continued...

While counting the WBC, record the number of NRBC observed during the count, e.g., the number of...





proerythroblast

basophilic erythroblast



erythroblast



orthrochromatophilic erythroblast

Then apply the following formula...

True Total WBC = Uncorrected total WBC x 100 *100 + number of NRBC

i.e., 100 is the number of WBC counted





An example situation that requires a corrected (or true) total WBC count:

Patient: a 26-year-old female

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- total uncorrected WBC count = 10,000/μL
- 100 WBC are counted on a Wright's stained peripheral blood smear...(for example)









The corrected (or true) WBC count is calculated:

Uncorrected total WBC count = $10,000/\mu$ L

NRBC = 25/100 WBC

(i.e., 25 NRBC were noted per 100 WBC counted on a stained peripheral blood smear)

True/Corrected WBC Count $\frac{10,000 \times 100}{100 + 25} = \frac{1,000,000}{125} = \frac{8,000/\mu L}{125}$





What are the effects of NRBC on the total WBC count?



The effects of NRBC on the total WBC count depend upon the number of NRBC present <u>and</u> the total <u>un</u>corrected WBC count. For example, if the uncorrected WBC is low:

Before No. WBC	Correction Interpretation	NRBC /100 WBC	After <u>No. WBC</u>	Correction Interpretation
5,000	Normal	1	4,950	Normal
5,000	Normal	10	4,545	Marginally decreased
5,000	Normal	20	4,160	Decreased





What are the effects of NRBC on the total WBC count...

If the total <u>un</u>corrected WBC count is high? For example:

Before <u>No. WBC</u>	e Correction Interpretation	NRBC /100 WBC	After No. WBC	Correction Interpretation
12,000	Increased	1	11,880	Increased
12,000	Increased	10	10,900	Marginally increased
12,000	Increased	20	10,000	Normal







For example:

Before No. WBC	Correction Interpretation	NRBC /100 WBC	After <u>No. WBC</u>	Correction Interpretation
5,000	Normal	40	3,560	Decreased
12,000	Increased	40	8,570	Normal
50,000	Markedly Higl	า 400*	10,000	Normal

* not an uncommon finding in severe anemias





This concludes the Corrected WBC Section. Select one of the following:

Go to **Total RBC Count**, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.









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What is the total RBC count?

- The total red blood cell count is the number of erythrocytes present per unit of peripheral blood (e.g., / μ L).
- However, with the exception of platelets, all cells (i.e., red blood cells and white blood cells) are actually counted. In the case of healthy adults, the inclusion of the white blood cells is not clinically significant.





With the exception of newborns, the RBC count on normal peripheral blood is essentially a reflection of the number of erythrocytes, i.e..

primarily mature erythrocytes



& 0.5 - 1.5% polychromatophilic erythrocytes

Erythroblasts (nucleated RBC):



are <u>not</u> seen in normal blood.





Which cells are classified as erythrocytes?

Erythrocytes are red blood cell forms without a nucleus which includes:



mature erythrocytes

&



polychromatophilic erythrocytes







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orthochromatophilic erythroblast



polychromatophilic erythroblast



basophilic erythroblast



proerythroblast





Are NRBC ever seen in normal peripheral blood?

Erythroblasts (NRBC) are not seen in normal adult blood but may be seen in the peripheral blood of newborns and in adults in disease. However, the RBC count will reflect the number of all RBC, i.e.:

(anucleated RBC):

mature erythro<u>cytes</u>



& polychromatophilic erythro<u>cytes</u>

&, if present, nucleated RBC

erythroblasts





Why are other cells included in the total RBC count?

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> Although intended to reflect only the number of erythrocytes (red blood cells) in circulation, the RBC count in reality includes all blood cells except normal platelets, which are excluded because of their small size.







Neither size criteria nor lysing techniques can be used to exclude...

in the total RBC count?

white blood cells

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erythroid precursors



giant platelets & clumps











What affect does the inclusion of other cells have on the total RBC count?

Total WBC counts that are normal or only slightly increased will have little or no affect on the total RBC. WBC are reported in thousands/ μ L and RBC are reported in millions/ μ L.

For example: RBC = 4,520,000 (4.52 x 10⁶)/ μ L WBC = 8,000 (8.0 x 10³)/ μ L

If the RBC is corrected for the presence of the WBC, then RBC = 4,512,000 (4.51 x 10^6)/µL. The difference is <u>clinically insignificant</u>.





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Effect of increased WBC?

In disease states characterized by large numbers of <u>WBC</u>, it is important to remember that they too will be included in the RBC count. For example...

RBC = 4,00 CML patier

If the RBC | RBC = 4,00

In this ca than that have mor



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Effect of NRBC in disease states?

In disease states characterized by large numbers of NRBC (e.g., thalassemia), it is important to remember that they too will be included in the RBC count.

For examp

NRBC = 40 RBC = 3,00 Total WBC If the RBC uncorrecte RBC = 3,00

In this cas that indica clinical sig major)

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Effects of giant platelets and platelet clumps?

When giant platelets or platelet clumps are present in peripheral blood, they too will be included in the RBC count. For example...



- the automated platelet count (PLT) is inaccurate (the true PLT would be higher).
- estimated platelet counts from blood smears may also be inaccurate (the true PLT estimate would be higher).





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Of what clinical importance are RBC counts?

RBC counts (as well as hemoglobin concentration and HCT) may be abnormal in disease states. RBC counts are characteristically:

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- d &
- Anemia
 - Blood loss
 - Some leukemias

- n C
 - Polycythemia vera


Isuhsc End of Total RBC Count

This concludes the Total RBC Count Section. Select one of the following:

Go to <u>Hemoglobin</u>, the section, to with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.











Hemoglobin, a conjugated protein, is the main component of the red blood cell.

It's major function is to transport: O₂ from lungs to tissues and

CO₂ from tissues to lungs



LSUNSC HOW does HGB transport O₂ and CO₂?





Of what clinical importance are hemoglobin values?

Abnormal hemoglobin concentrations may be seen in disease states. Hemoglobin concentrations, as well as RBC count and HCT, are characteristically:

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- **c** Anemia
 - Blood loss
 - Some leukemias
- a s e

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Polycythemia vera
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This concludes the Hemoglobin Section. Select one of the following:

Go to <u>Hematocrit</u>, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.











The hematocrit is a reflection of the concentration of red cells, not of the total red cell mass.

Therefore, by definition, the hematocrit is the relative volume of packed erythrocytes expressed as a percentage of the volume of whole blood sample, e.g.:

40% packed cells or 40% HCT



& 60% plasma





Of what clinical importance are hematocrit (HCT) values?

Abnormal hematocrits may be seen in disease states. Hematocrit values, as well as RBC count and HGB concentrations, are characteristically:

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- Blood loss
- Some leukemias
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Polycythemia vera





How does the HCT correlate with the RBC count when RBC are normocytic?

When erythrocytes are **normocytic**, there is good correlation of the HCT and the total RBC count (i.e., when one is low, the other is low; when one is high, the other is high).



RBC of normal size and shape are called normocytic. They are about $6-8\mu m$ in diameter (slightly smaller than the nucleus of a normal, small **lymphocyte** which is about 8-10 μm in diameter).

Normal red cells are round and biconcave with a slight central pallor.





Isunsc How does the HCT correlate with the RBC count when the RBCs are macrocytic?

In macrocytosis, because the RBC are larger than normal,



(normocytic cells)



the total RBC count may be lower than expected based upon HCT values.





lsuhsc How does the HCT correlate with the RBC count when the RBC are microcytic?

In microcytosis, because the cells are smaller than normal,



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(normocytic cells)



the total RBC count may be higher than expected based upon HCT values.



How does the HCT correlate with the RBC count in a mixed RBC population?

The instrument electronically measures and determines a <u>mean</u> volume of the cell population, e.g.,

1 normal mature RBC
2 macrocytes
3 microcytes

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4 polychromatophilic erythrocyte

5 ovalocytes
6 target cells
7 RBC with inclusions
8 NRBC



Because of the variation in cell size and volume, correlations of RBC, MCV, and HCT becomes less predictable.







What are some examples of true HCT/RBC inconsistencies?

In hydremia of pregnancy:

the HCT is low although there is no reduction in the total number of circulating red cells.

and

In shock accompanied by hemoconcentration: the HCT may be normal or even high though blood loss may have caused a considerable decrease in the total red cell mass.





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What is the correlation between HCT and hemoglobin (HGB)?

HCT (expressed in per cent) is usually roughly 3 times the HGB (expressed in g/dL).

For example:

- An adult with hematocrit of 45% would normally have a hemoglobin of about 15 g/dL.
- **Further discussion:**

The relationships between the HCT and total RBC count and the HCT and HGB concentration are discussed further in the section on RBC indices.









This concludes the Hematocrit Section. Select one of the following:

- Go to Erythrocyte (RBC) Indices, the next section to continue with the exercise as designed.
- Return to the <u>Hemogram Menu</u> and make an alternate selection.



Erythrocyte (RBC) Indices







What are the erythrocyte (red blood cell) indices?

The indices are measurements of the:

- average volume or size of red cells (MCV)
- hemoglobin content or weight in the average red blood cell (MCH)
- hemoglobin concentration in a given volume of red blood cells (MCHC).







Of what clinical value are the erythrocyte (red blood cell) indices?

The indices are valuable tools in the study of anemias because they provide an objective quantitative standard for assessing the

- size of the red cells (MCV)
- relationship between individual blood cells and the hemoglobin concentration (MCH)
- red blood cell population as a whole and the hemoglobin concentration (MCHC).





What method is used to determine the erythrocyte (RBC) indices today?

When first introduced, all of the indices (MCV, MCH, and MCHC) were calculated values based on accurate direct measurements of the RBC, HGB, and HCT (packed cell volume).

Today, with the use of modern technology, it is the MCV that is now measured directly and the HCT that is calculated.

Let us look at the measurements included in the indices.





(mean corpuscular hemoglobin μμg) MCHC (mean corpuscular hemoglobin conc. %)

MCV (mean corpuscular volume μ³)

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MCH

What are the measurements in the erythrocyte indices?



What is the MCV?

MCV is the <u>average volume</u> (or size) of red cells expressed in μ^3 or femtoliters (fL).

Individuals with normal peripheral blood will have a normal MCV (i.e., 80-100 μ^3).

The normal small mature lymphocyte, because it is relatively consistent in size with a nucleus that is about $10-12\mu$ in diameter, is a useful tool in a visual assessment of red cell size on stained blood smears.)



lymphocyte
 (nucleus 10-12μ)
 normal RBC
 (6-8μ)







What affect do abnormally large or small red cells have on the MCV?

Individuals with red cells that are predominantly larger than normal (>8m diameter) will have an MCV >100 μ^3 .



Individuals with red cells that are predominantly smaller than normal (<6m diameter) will have an MCV < 80 μ^3 .







Usuhsc What if there are various sizes of red cells?

E.g., macrocytic and normocytic RBC: it is important to remember that the MCV is the <u>average volume (or size)</u> of the cells. For example, if an MCV is high, it does not mean that all of the red cells are larger. Some of the cells may be normal or even smaller than normal. It is an indication, however, that large cells are present in sufficient numbers to cause an increased MCV, e.g..









And conversely,

If the MCV is low, it does not necessarily mean that all of the cells are smaller than normal. It is an indication, however, that the number of small cells is sufficient to cause a decreased MCV, e.g.:







Does a normal MCV always indicate a normal RBC population?



Again, it is important to remember that the MCV is the <u>average volume (or size)</u> of the cells. A patient with a red cell population of varying sizes that include normocytic and/or microcytic and/or macrocytic cells may have a normal MCV. e.g.:



normocytic microcytic macrocytic







Microscopic examination of the peripheral blood smear provides a better evaluation of MCV when a mixed population of red cells is involved or when interference is suspected.





Of what clinical importance are MCV values?

MCV values may be abnormal in disease states. E.g., MCV is characteristically elevated in:



macrocytic anemias

and







polychromasia or reticulocytosis (e.g., hemolytic anemias, acute blood loss)







Clinical importance of decreased MCV values (continued)...

...and characteristically decreased in:



Microcytic hypochromic anemia and may be as low as 50 μ^3 (or fL).





What is the second measurement included in the RBC indices?

menu

In addition to the: MCV (mean corpuscular volume μ^{3})

There is the MCH.



What is the MCH?

MCH is the content (or weight) of hemoglobin of the <u>average red cell</u> (i.e., individual cells) expressed in micromicrograms (μμg) or picograms (pg).



The MCH is a calculated measurement based on the values obtained for the HGB concentration and the RBC count.

An individual whose red cells are normal (i.e., size, shape with slight central pallor) will have a normal MCH, even if the HGB and RBC values are decreased.





What about individuals with a decreased MCH?

A low MCH indicates a less than normal hemoglobin content in the average individual red cell (hypochromic erythrocytes). In some cases, the cells may also be microcytic (< 6 μm in diameter).



Again, this is an average.

though many RBC are microcytic hypochromic

__d/or normocytic hypochromic,

some may be normochromic





Of what clinical importance are MCH values?

MCH values may be abnormal in disease states. E.g., MCH is characteristically elevated in:



macrocytic anemias

and

may be as high as 50 μμg (or pg) if megaloblastic





polychromasia or reticulocytosis (e.g., hemolytic anemias, acute blood loss)





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Clinical importance of MCH values, (continued)

Characteristically decreased in:



Microcytic hypochromic anemia and may be as low as 15 $\mu\mu$ g (or pg).





What is the third measurement included in the RBC indices?

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In addition to the:

MCV (mean corpuscular volume μ³⁾ and MCH(mean corpuscular hemoglobin μμg)

There is the MCHC.



What is the MCHC?

MCHC is the average hemoglobin concentration in a given volume of packed RBC expressed in g/dL (or %).



MCHC is based on the relationship between the HGB concentration and the HCT and is calculated using the values obtained for those two measurements.

An individual whose red cells are normal (i.e., size, shape with slight central pallor) will have a normal MCHC even if the HGB and HCT are decreased.




Of what clinical importance is the MCHC value?

MCHC values may be abnormal in disease states.

A true elevated MCHC is seen <u>only</u> in spherocytosis.











How do spherocytes differ from normal RBCs ?



Spherocytes, (circled in blue) are round and have no central pallor.

Normocytic red cells (circled in red) have a slight central pallor (about 1/3 of the cell diameter) due to the biconcave shape





Why is a true elevated MCHC seen only in spherocytosis?



It's the only situation in which the cells are spherical and, therefore, have a greater capacity for hemoglobin than biconcave cells.



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In the absence of spherocytosis, an elevated MCHC may be an indication of

A falsely elevated MCV and decrease in HCT (e.g., caused by cold agglutinins)

or

Falsely elevated HGB (e.g., lipemia or some other interfering factor).







What disorders are characterized by a MCHC?

Normal or decreased: macrocytic anemias

Decreased: hypochromic anemia (usually no lower than 22 %)

Normal MCHC but reduced HGB & HCT: normochromic anemias



How are the RBC indices calculated?

When determined by automated electronic instruments, the MCV is a direct measurement.

A calculated indirect MCV measurment can be made using microhematocrit values.

Calculated MCV =	<u>Micro HCT (%) x 10</u>
	RBC (millions/µL)

MCH and MCHC are always calculated.

 $MCH = \frac{HGB (g/dL) \times 10}{RBC (millions/\mu L)}$

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 $\frac{\text{MCHC}}{\text{HCT}(\%)} = \frac{\text{HGB}(g/dL) \times 100}{\text{HCT}(\%)}$







This concludes the Erythrocyte (RBC) Indices Section. Select one of the following:

Go to <u>Red Cell Distribution Width (RDW)</u>, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.



Red Cell Distribution Width







What is the Red Blood Cell Distribution Width (RDW) ?

Normally

All red cells are approximately the same size and measure about 6-8 μ in diameter.

Anisocytosis

A "generic" term used to indicate a subjective visual assessment of abnormal variation in size of red cells





The RDW

An objective electronic measurement of the variation in the size of the cells in the RBC population





Based on data obtained by electronic measurement of the sizes of the red cells, the RDW is calculated by enumerating the number of erythrocytes that are:

than the reference (normal) cell volume (size)

or larger

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smaller



normal MCV



Thus, based upon objective measurements, the RDW provides an estimate of anisocytosis (i.e., variation in size of the red cells).







When are normal RDW values seen?

Normal RDW values of 11.6-14.6 (which may vary slightly among laboratories) are seen when the RBC are all about the same size.



(i.e., essentially homogeneous RBC population)





When are increased RDW values seen?

In disease, an increased RDW may be seen proportionate to the degree of variation in size of the red blood cell population (i.e., the greater the anisocytosis, the greater the RDW).



e.g.

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When are decreased RDW values seen?

The RDW is never decreased (i.e., less than the lower reference value). Think about what is being measured.

A normal RDW reflects an RBC population in which the cells are essentially the same size.



An increased RDW reflects a population with variation in size.



Those are the only two possibilities so the RDW can never be decreased.





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What causes the change in RDW?

Variability in the size of the cells in the RBC population causes the increased RDW. The increase may reflect an RBC population containing any combination of abnormal and normal (or less affected) red cells, for example...



normocytic and microcytic and/or macrocytic





What is the clinical importance of the RDW?

RDW is the most sensitive measurement involving red cells.

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RDW is the first to become abnormal (sometimes before anemia appears).

RDW is the first to become abnormal in iron deficiency due to chronic blood loss.







normocytic, & macrocytic, & microcytic

In this case, the MCV could be within reference range after averaging the normal size, large, and small cells included in the measurement.





How does the MCV differ from the RDW?

Whereas the MCV is the average cell size (volume)



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of a population that may include normocytes, macrocytes, and microcytes,

the RDW reflects the <u>degree of variation</u> in the sizes of those blood cells. In this case, while the MCV (average size) could possibly be within normal limits, the RDW would be increased.







Is there any relationship at all between the RDW and MCV?

Both the MCV and RDW are expected to be normal when the RBC population is relatively homogeneous in size (i.e., essentially one size).



What about abnormal RBC populations?





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Can a low or high MCV be expected if the RDW is normal?

Certain conditions are characterized by red cells that, although essentially the same size, are smaller (or larger) than normal. In those conditions, a normal RDW will be found even though the MCV is low (or high).



or

homogeneously microcytic & low MCV homogeneously macrocytic & high MCV

lymph





Can a normal MCV be expected if the RDW is abnormal?

Even in cases where the population of blood cells includes cells of varying size (i.e., abnormal RDW), the MCV may be normal because it reflects the <u>average</u> cell volume of all of the cells, e.g.:

[1] normocytic,

[2] macrocytic, and/or

[3] microcytic.





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In cases where the MCV is normal and the RDW is abnormal,



the RDW will be increased proportionately to the number of cells smaller or larger than normal and the degree of variability in cell size.







Examples in which both the RDW and MCV may be normal...

RDW	MCV	Conditions

Normal Normal good health (homogenous normocytic RBC population)

Normal Normal (homogenous normocytic RBC population) chronic disease (90% of cases are normocytic) uremia HbAS HbAC







Normal

Examples in which the MCV is abnormal but the RDW may be normal...

RDW MCV	Conditions
---------	-------------------

Lowthalassemia minor(homogenous
microcytic
red cells)iron deficiency
chronic disease (10% of cases are
microcytic)

Normal High (homogenous macrocytic red cells) aplastic anemia myelodysplastic syndrome anemia due to alcoholism







Examples of conditions in which the RDW is increased and MCV is likely to be normal...

<u>RDW</u>



Conditions

High

Normal

(average size of cells in a heterogenous RBC population) early factor deficiency HbSS disease HbSC disease





Examples of conditions in which the RDW is increased but the MCV may be decreased...

High L

RDW

Low

MCV

(average size of cells in severe anemias characterized by microcytic RBC and/or abnormalities associated with hemolytic alterations such as RBC fragments)

Conditions

HbS-thalassemia thalassemia major HbH disease hereditary spherocytosis TTP DIC





Examples of conditions in which both the RDW and the MCV may be increased...

<u>RDW</u> <u>MCV</u>

High High

(average size of cells in severe anemias characterized by macrocytic alterations in the RBC population)

Conditions

vitamin B12 deficiency folate deficiency reticulocytosis (e.g., acute blood loss, autoimmune hemolytic anemia newborn (due to presence of immature RBC) Cold agglutinins (due to false values)





Examples of conditions with a decreased RDW and their expected MCV.

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<u>RDW</u> <u>MCV</u> <u>Condition</u>

If you were expecting to see a list of conditions, you have not been paying attention. The RDW is never decreased.

You may review the section on RDW or .

review RDW or continue



End of Red Cell Distribution Width

This concludes the Red Cell Distribution Width (RDW) Section. Select one of the following:

Go to **Platelet Count**, the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> to review a section on one of the parameters of the Hemogram.











What is a platelet count (PLT)?

The total platelet count is the number of platelets per given volume of peripheral blood (e.g., $200,000/\mu$ L).

Platelets are difficult to count.

- They are small & must be differentiated from debris.
- They have a tendency to adhere to glass, to any foreign body, and to each other which may cause a false low count to be obtained.





Are there special specimen requirements for platelet counts?

Platelet counts must be performed within:

- <u>3 hours</u> after collection in an EDTA anticoagulated tube of blood.
- <u>1 hour</u> after collection in a unipette containing diluent (finger stick)

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The MPV (mean platelet volume) increases after 3 hours (at least in part as a result of the change in cell shape).





Questionable counts are verified by microscopic examination of a stained blood smear. Examine for:

- platelet clumps
- platelet distribution (evenly throughout smear)
- platelet estimate (1 platelet per oil immersion field on a smear is equivalent to approximately 15,000 to 20,000 platelets)

Extremely low electronic counts are verified by phase microscopy (i.e., manual hemacytometer count).







Of what clinical importance are platelet counts?

To be hemostatically effective, platelets must be present in sufficient numbers and must be functionally normal. Platelet disorders may be classified as:

Qualitative
of platelets)(i.e., defect in the functional ability

or

Quantitative (i.e., increase or decrease in the number of platelets), as determined by the platelet count.





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What disorders are associated with a quantitative <u>increase</u> in platelets?

Thrombocytosis, an abnormal increase in PLT, may be due to:

- **Reactive thrombocytosis:**
- A physiologic response that may be seen as a secondary phenomenon (eg, in trauma, hemorrhage, iron deficiency)

Temporary rise in platelets:

- May be seen following splenectomy (splenic pool is eliminated)
- Autonomous (primary thrombocythemia):
- A primary bone marrow disorder commonly seen in myeloproliferative disorders. Platelets may also have functional abnormalities.





What disorders are associated with a quantitative <u>decrease</u> in platelets?

Thrombocytopenia, an abnormal decrease in platelets may be attributed to:

Decreased production:

Due to a quantitative or hypoproliferative defect in megakaryocytes (e.g., marrow damage, replacement of normal marrow by metastatic tumor, intrinsic marrow disease [leukemia]),

Ineffective thrombopoiesis:

A normal number of marrow megakaryocytes, but platelet production is decreased (e.g., megaloblastic anemias)



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Quantitative platelet decreases, (continued)

With increased destruction of platelets, characterized by megakaryocytes in the bone marrow and low platelet counts in peripheral blood, PLT destruction may be due to:

Non-immune (consumptive) causes, e.g.:

- disseminated intravascular coagulation (DIC)
- thrombotic thrombocytopenic purpura (TTP)







- May also be due to immune causes, e.g.:
- drug-induced immune thrombocytopenia
- acute and chronic idiopathic thrombocytopenic purpura (ITP)

Decreases may also be due to increased spleen pooling.





This concludes the Platelet Count Section. Select one of the following:

Go to Mean Platelet Volume (MPV), the next section, to continue with the exercise as designed.

OR

Return to the <u>Hemogram Menu</u> and make an alternate selection.



Mean Platelet Volume





What is the Mean Platelet Volume (MPV)?

The mean platelet volume (MPV) is the <u>average</u> volume or size of the platelets in the population.

Once the platelet count and size distribution is determined, the mean platelet volume can be calculated from the arithmetic mean of the extrapolated histogram.

MPV reference values are about 6.5 to 12 fL.

Normally, the platelet size varies inversely with the platelet count.







Is platelet size affected if there is an abnormal PLT count but normal marrow function?

Platelet size also varies inversely with abnormal platelet counts when there is normal marrow function, e.g.:

Disorder

idiopathic thrombocytopenia

reactive thrombocytosis









Is platelet size affected if both PLT count and marrow function are abnormal?

If there is abnormal marrow function (e.g., folate deficiency or aplastic anemia), the MPV may be low even though there is thrombocytopenia.







This concludes the Mean Platelet Volume section and Part 1 of the study module, "The Complete Blood Cell Count (CBC)".

Click on <u>Hemogram Menu</u> to review a section.

OR







The following are additional exercises related to the CBC:

- CBC Part 2 WBC differential & blood morphology
- CBC Part 3 RBC morphology & platelet estimate
- CBC Part 4 Post-test

