

CLL SOCIETY

Smart Patients Get Smart Care™

Learning to Decode Your Blood Test Results for CLL

June 29, 2021

10:00 AM PT, 11:00 AM MT, 12:00 PM CT, 1:00 PM ET

This program was made possible by grant support from





Bristol Myers Squibb

Speakers





Welcome: Patricia Koffman Co-Founder and Communications Director CLL Society



Moderator: Brian Koffman, MDCM (retired), MS Ed Executive Vice President and Chief Medical Officer CLL Society



Speaker: Susan Leclair, PhD, CLS (NCA) Chancellor Professor Emerita University of Massachusetts Dartmouth

A QUICK RUN DOWN OF ALL 25,000!



- Not really
- But I did want to start off by saying this is a BIG subject and we will only look at a few of the more common tests.

But first – a word from reality

Tests live in a real world that is bounded by

Preanalytical Aspects

patient preparation issues time of collection conditions of collection transport &storage confounding meds wrong test requested fingerstick vs. venous

Analytical Aspects

instrumentation reagent quality specificity & sensitivity technique/method location patient population.

Post-Analytical Aspects

reporting mechanisms (to whom, when) reflex testing protocols presence/absence of interpretation wrong test requested

Each of these attributes influences the value of the test result – consistency is key.

- We do not count white cells.
 - We count nuclei so nucleated red blood cells are counted here – corrected white cell count
 - 5 major cell lines present in the peripheral blood
 - Neutrophils present all the time
 - Lymphocytes present all the time
 - Monocytes present occasionally
 - Eosinophils present occasionally
 - Basophils present rarely















- The number will bounce around all day long in response to your environment.
 - Want your granulocytes to increase?
 - Exercise (walk up the stairs) for a few minutes before getting your blood drawn
 - Half of your granulocytes usually marginate along the walls of the blood vessels. Exercise "shakes" them off putting them into the circulating pool for about 15-20 minutes.
 - Have a panic attack reaction to stress
 - Adrenaline will also take cells away from the marginating pool
 - Be on Steroids



- Two ways to report white cells by type
 - Circa 1900 the traditional differential
 - Look at the first 100 random white blood cells you see using a microscope and your own trained eyes.
 - Some people are better than others
 - Some days and better than others
 - If there are 7500 white cells in a microliter of blood and you count 100 of them report in percentages
 - What are the odds that you will fin what is important?
 - How can you tell which cell line is increased or decreased?
 - Circa 1980s with the advent of multi-channel instruments
 - Counts the exact number of cells in a specific volume of blood.
 - Count somewhere between 20,000 and 50,000



- The best differential is the ABSOLUTE differential.
 - Counts the exact number of cells in a specific volume of blood.
 - Percentages cannot tell which cell line is increased or decreased.

White cell count	% neutrophils	% lymphocytes	A.N.C. Absolute neutrophil Count	A.L.C. Absolute lymphocyte count
2.0x10 ⁹ /L	63	37	1.2x10 ⁹ /L	0.74x10 ⁹ /L
4.0x10 ⁹ /L	63	37	2.5x10 ⁹ /L	1.4x10 ⁹ /L
8.0x10 [/] /L	63	37	5.0x10 ⁹ /L	2.8x10 ⁹ /L
16.0x10 ⁹ /L	63	37	<mark>10.0x10⁹/L</mark>	3.7x10 ⁹ /L



Then why use both?

Absolute

 You get a real number of cells by cell line. And there is NO way to confuse which cell line is increased/decreased

Percentage

- There is nothing better to assess the quality of the cells than having someone who knows what they are doing look at them.
- So doing both gives you a more complete picture of the cells and what they have been doing



- For example
- Both of these are the same cell. One is exhausted and on the brink of death itself. No instrument can tell them apart.



Basophil



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WHITE CELL VALUES

Is anyone better than another?

- Neutrophils (once known as granulocytes)
 - Most common cell
 - Exists in the marrow, the two pools in the peripheral blood and in the tissues
 - Phagocytize dead/dying cells and any foreign item (particle or droplet)
 - Incites, controls, and participates in the inflammatory process
 - Determines acute or chronic inflammation
 - Most varied morphology









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WHITE CELL VALUES

Is anyone better than another?

- Neutrophils (once known as granulocytes)
 - Fully functional
 - Polys, segs, PMNs, bands
 - Seen in peripheral blood
 - Minimally functional
 - Metamyelocytes (Meta) and myelocytes (Myelo)
 - Should not be seen in peripheral blood
 - Not functional
 - Promyelocyte (Pro) and Myeloblast (Blast)
 - Must not be seen in peripheral blood



Is anyone better than another?

Monocytes

- Usually found in the tissues uses blood stream to move from one place to another
- Exists in the marrow, only one pool in the peripheral blood and in the tissues
- Phagocytize dead/dying cells and any foreign item (particles only)
- Processes antigens for the T cell to recognize
- Will be increased after any trauma









Is anyone better than another?

- Lymphocytes
 - Found in lymph nodes, lymphatics, peripheral blood, and bone marrow
 - Circulate freely between the nodes/lymphatics and the peripheral blood/marrow
 - Cannot be differentiated using light microscopy
 - Could comment on size (small, medium, large)
 - Assumed an unusual looking lymphocytes WAS damaged in some fashion (atypical)
 - Did not have a known function until the mid 1960s (Robert Good Minnesota Nobel Prize)
 - Were separated by function T, B, and NK cells
 - Realized that "atypical" cells were in fact reacting to the presence of a foreign antigen and were defenders not the illness – new Name Reactive Lymphocytes
 - Sadly many people refuse to update to the correct name apathy?

- Lymphocytes
 - Small lymphocytes
 - Usually B cells
 - Resting from any action so can be naïve or memory
 - Medium lymphocytes
 - Can be T, B or NK cells
 - If it has granules, more likely to be t or NK
 - Large lymphocytes
 - Can be T or NK
 - Reactive lymphocyte
 - If B cell, than larger cytoplasm for antibody production
 - If T or NK cell, less cytoplasm and more granules









- Smudge cell
 - Can be any cell line
 - Usually
 - A cell that is very fragile and cannot withstand the collection and processing.
 - Frequently seen in CLL
 - A cell that has died in the course of some reaction/inflammatory response etc.
 - If a monocyte or neutrophil, can just have died if in a circle.
 - If a neutrophil, then it is called a neutrophil trap or net. The cell has exploded itself in order to make the largest area filled with protein braking or killing enzymes.









PLATELETS



- The number will bounce around all day long in response to your environment.
 - When performed manually very difficult so the accepted range is +/-50,000.
 - When performed by instrument, the accepted range is = +/- 20,000
 - As the instruments got better, the acceptable range has moved from 150 500 to 150 – 450 to 130 – 400 to even smaller ranges for some facilities (150 – 350)



PLATELETS

- PDW platelet distribution width
 - Similar to the RDW
 - Mathematical description of size variation
 - Do we care?
 - Partially
 - Larger platelets suggest some type of inflammation, overuse, or drug response
 - Smaller platelets suggest deficiencies similar to microcytic red cells (iron, B₆, hypothyroidism)
 - Why not?
 - The most important things about platelets is their function. We have a few, not very
 precise tests for platelets because their function is in such a complex situation we cannot
 replicate it size of capillary damage, type of damage (smooth vs. ragged), integrity of
 vessel walls, signaling from localized cells, eternal conditions (heat/cold, pressure, etc.)

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PLATELETS

- Bone marrow cell megakaryocyte, get larger and then breaks off pieces of the cytoplasm
 - Those pieces then need to reorganize themselves
 - Large platelets are usually not well organized and function less efficiently



PLATELETS

When not stimulated

- Platelets are small disc shaped pieces of cytoplasm
- When stimulated, they wring themselves out like a sponge, spreading contents into the area.
 - Their protrusions interlace forming a lattice structure.
 - They ADHERE to damaged walls.
 - Once "laced" they AGGREGATE and tighten
 - Other Contents are stimulants for the clotting process.
- Platelet lack or lessened function is seen in
 - Small blood vessel bleeding (gums, mucous membranes, skin, etc.
 - Not big clots
- Drugs aspirin, clopidogrel, ticagrelor, ticlopidine









PLATELETS

Anti- Platelet Drugs

- Aspirin usually low dose (81mg) but can be 325mg.
- Clopidogrel Plavix
- Ticagrelor Brilinta
- Prasugrel (Effient)
- Dipyridamole/aspirin (Aggrenox)
- Ticlopidine (Ticlid)
- Eptifibatide (Integrilin)



PACKS-4's $8^{1/2}$ x 11" color reports make it easy to interpret results from multi-reagent profiles and are suitable for direct posting to patient charts.



Normal Lab Values

Find Information on CLLSOCIETY.ORG



🛠 Chronic Lymphocytic Leukemia Toolbox





Normal Lab Values

Complete Blood Count (CBC)

Test Acronym	Meaning	Normal Range Values (Male)	Normal Range Values (Female)
WBC	Number of white blood cells	3.5-10.5 x 109/L	3.5-10.5 x 109/L
RBC	Number of red blood cells	4.7 to 6.1 million cells/mcL	4.2 to 5.4 million cells/mcL
HGB	Hemoglobin level	13.8-17.2 g/dL	12.1-15.1 g/dL
НСТ	Hematocrit	40.7-50.3%	36.1-44.3%
MCV	Mean corpuscular volume	80-100 fL	80-100 fL
MCH	Mean corpuscular hemoglobin	27-31 pg	27-31 pg
MCHC	Mean corpuscular hemoglobin concentration	32-36 g/dL	32-36 g/dL
RDW	Red cell distribution width	11.8-15.6%	11.9-15.5%
PLT	Number of platelets	150-450 x 109/L	150-450 x 109/L



Normal Lab Values

White Blood Cell Differential (Diff)

Test	Meaning	Normal Range Values			
Neuts.%	Percentage of Neutrophils	40% to 60%			
Lymphs%	Percentage of Lymphocytes	20% to 40%			
Monos.%	Percentage of Monocytes	2% to 8%			
Eos.%.	Percentage of Eosinophils	1% to 4%			
Baso.%	Percentage of Basophils	0.5% to 1%			
Neuts.# (ANC)	Absolute Neutrophil Count	1.70-7.00 x 109/L			
Lymphs# (ALC)	Absolute Lymphocyte Count	1.00-4.80 x 10 ⁹ /L			
Monos#	Number of Monocytes	0.30-0.90 x 109/L			
Eos#	Number of Eosinophils	0.05-0.50 x 109/L			
Baso#	Number of Basophils	0.00-0.30 x 109/L			



Keeping Track of Your Lab Results

Download the Template to Keep Track of Your Lab History



🛠 Chronic Lymphocytic Leukemia Toolbox





How to use the Keeping Track Patient Records Spreadsheet

Keeping Track of Your Lab Results



Allows for a broader view of your long-term *trending* history for all key CLL lab components (CBC, Absolute Lymphocytes, and more)



Example Lab Tracking Form



															ULL DO CIETT
Jane Doe											© Copyright 2020 CLL Society, Inc. All Rights Reserved.				
					Click on a ?										
						fo	r oddit	ional h	ole inf	ormati	o p				
	HOME CHARTS					for additional help information					on				
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CBCIN	formati	on						Bernsteint	Alexandrate						
Date	WBC	RBC	HGB	нст	Platelets		Absolute Lymphs	Percent Neuts	Absolute Neuts	MCV	МСН	МСНС	RDW	MPV	NOTES
?	?	?	?	?	?	?	?	?	?	?	?	?	?	?	?
Range=>	3.5-10.5	4.2-6.1	12.1-17.2	36-50	150-450	28-55%	0.85-4.1	25-70%	1.5-7.8	80-100	28-32	32-36	11%-15%	7.5-11.5	
1/1/15	5.5	4.2	12.1	36.0	150.0	28.0%	0.85	25.0%	1.5	80.0	28.0	32.0	11.0%	7.5	
5/1/15	6.0	7.0	18.0	4	154.0	-	0.90	26.0%	Quelle		33.0	37.0	20.0%	11.0	
1/1/16	7.6	and the second	15.0	a second and			0.95		4.2	81.5	28.0	32.0		7.5	
5/1/16	5.5	4.2	12.1	36.0			1	25.0%	3.0		33.0	37.0	11.0%	8.4	
1/1/17	11.0	7.0	18.0	50.0	160.0		0.9	27.0%	1.5	80.0	28.0	34.3	12.0%	8.5	
5/1/17	9.0		15.0			40.0%	2.00	26.0%	2.0	1000	33.2	36.0	13.4%	9.2	
1/1/18	5.5		12.1	36.0			2.15		3.2	83.0	28.0	42.0		7.5	
5/1/18	11.0	7.0	18.0	50.0	215.0	60.0%	3.10	30.0%	3.4	84.5	33.1	35.7	14.0%	8.8	
1/1/19	9.0	6.0	15.0	45.0	320.0	40.0%	3.15	31.0%	4.4	92.0	32.4	37.3	12.4%	9.2	
5/1/19	5.5	4.2	12.1	36.0	220.0	28.0%	2.99	32.0%	4.2	81.0	33.0	34.0	13.2%	9.9	
5/1/20	11.0	7.0	18.0	50.0	245.0	60.0%	4.10	28.5%	5.0	84.2	31.2	33.4	14.5%	7.6	
7/1/20	9.0		15 0			40.0%	3 99	33 0%	<mark>5 1</mark>	82.0	29.4	33.0	12 2%	82	
HOM	E CBC (CBC Charts	Blood Ch	emistry	Immunoglol	oulins	+				:	•			



Audience Questions & Answers

This program was made possible by grant support from





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Thank You for Attending!

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Join us in August for our webinar on PI3K Inhibitors

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