Tutorial - Blood Cell Morphology

A Clinical Pathology 201 Study Module

by

Carolyn Sue Walters, MHS, MT(ASCP) Department of Pathology School of Medicine Louisiana State University Health Sciences Center New Orleans, Louisiana

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C. Sue Walters, MHS, MT(ASCP)

Associate Professor Department of Pathology LSU Health Sciences Center New Orleans, LA

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Special Acknowledgment

A special thanks and acknowledgment for the generosity of Mrs. Angela Foley, MS, MT(ASCP), Department of Clinical Laboratory Science, School of Allied Health, LSU Health Sciences Center New Orleans for the use of many of the blood cell images used in this presentation.



Feedback

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:

C. Sue Walters, MHS, MT(ASCP) LSU Health Sciences Center Department of Pathology 1901 Perdido Street New Orleans, LA 70112





The directions for navigating through the exercise are given on the next 2 pages. They are the same as those routinely used in Clinical Pathology 202 study modules. Please click on:



to visit the directions before continuing with the exercise.





to go directly to the first page of the exercise.



Directions, continued

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 to return to
the Main Menu selection.



in the lower right corner of the page to continue.

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





Directions, continued

"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 "hot point", 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.



Hematologic Cells Found in Peripheral Blood and Bone Marrow





MAIN MENU

Introduction

Leukocytes

Erythrocytes

Abnormal erythrocytes - terminology

Platelets

Disorders – characteristic morphology





Introduction





What is the purpose of this study module?

This study module is designed for LSUHSC L2 students enrolled in Clinical Pathology 201. It is intended as a reference for blood cell and bone marrow morphology.

The presentation of illustrative cells in this module is by no means a comprehensive study of blood cells. It is limited to the material covered in the lectures and laboratory sessions.

Unfortunately, a few cell illustrations are not available at this time but will be added later.





Leukocytes





How are the WBC identified and classified?

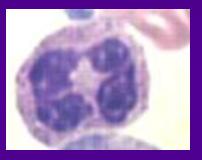
Typical nuclear and cytoplasmic morphologic features provide a means by which WBC can be identified and classified as to cell line (i.e.,):

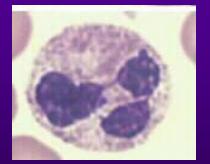
- granulocytes [neutrophils, eosinophils, or basophils]
- monocytes
- Iymphocytes



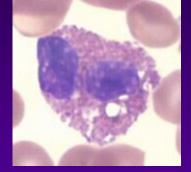
i.e., classified as granulocytes:

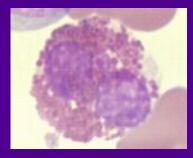


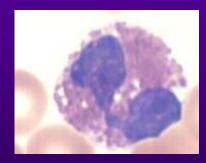




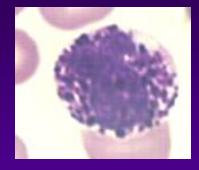
neutrophils

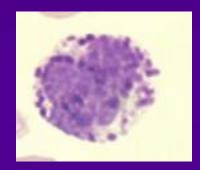


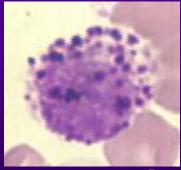




eosinophils



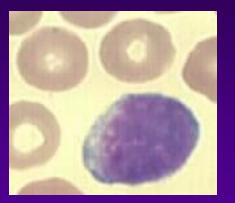


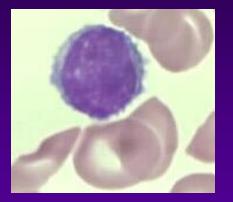


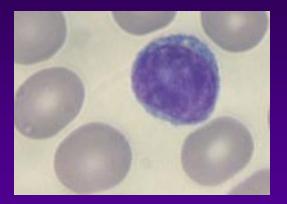
basophils



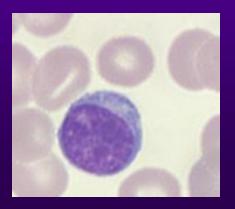
i.e., classified as lymphocytes:



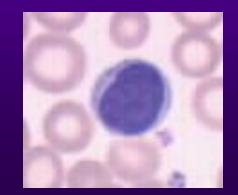






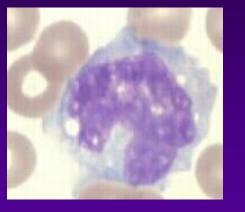


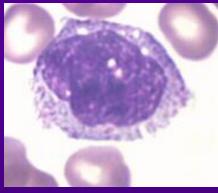


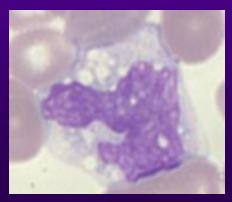


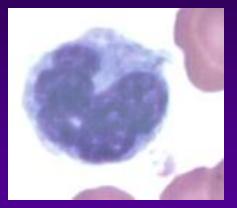


i.e., classified as monocytes:

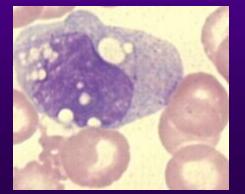


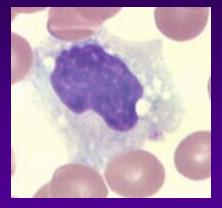


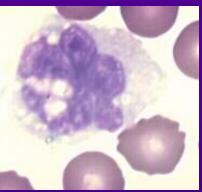


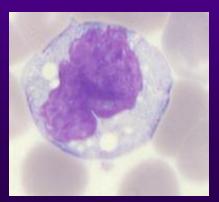
















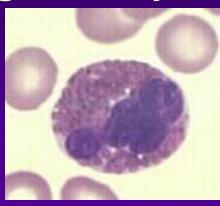


i.e., Identified as to cell lines

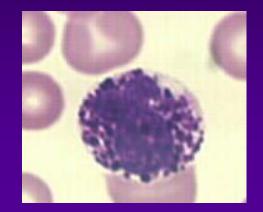
granulocytes



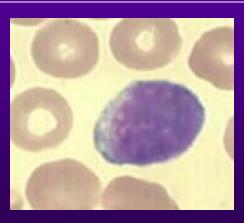
neutrophils



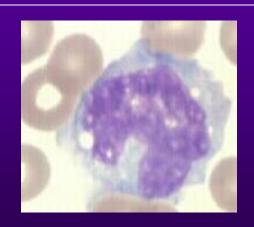
eosinophils



basophils



lymphocytes



monocytes





WBC can also be identified and classified as to...

maturity (i.e., mature cell or immature stage of development).

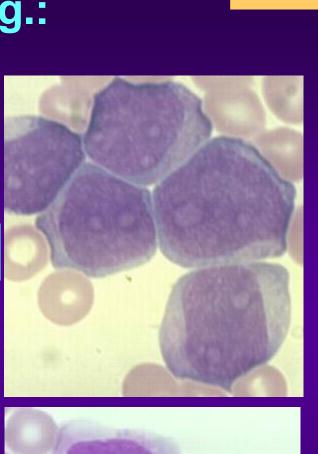


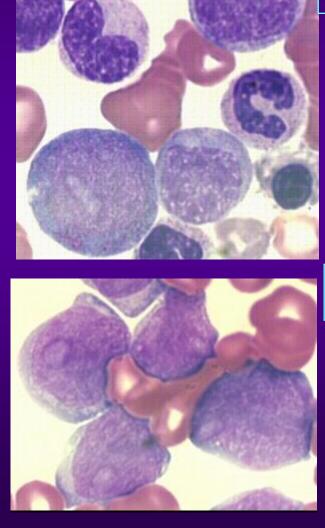
Immature WBC, e.g.:

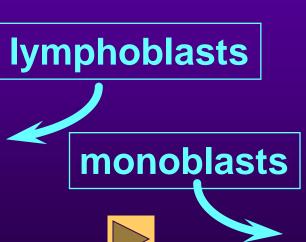
(various stages)

myeloblasts

granulocytes



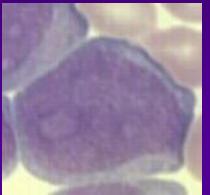




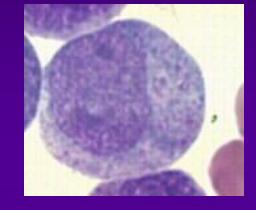


e.g., Neutrophils in various stages of maturation...

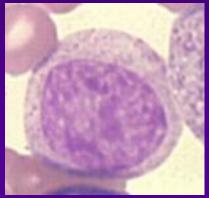
myeloblast



promyelocyte



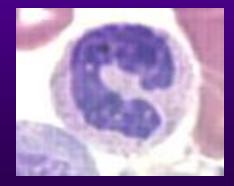
myelocyte



metamyelocyte



band



PMN (mature)



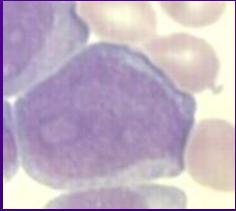






Neutrophilic Maturation

from immature blast to mature PMN



Nautrania abandés na "

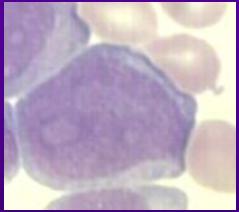






Neutrophilic Maturation

From mature PMN to myeloblast



Nautrania abandés na "





WBC can also be identified and classified as to...

menu

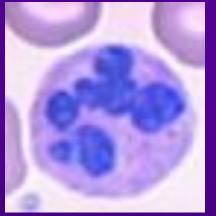
 abnormal morphology (i.e., nuclear or cytoplasmic alterations)





e.g., WBC with acquired nonneoplastic alterations...

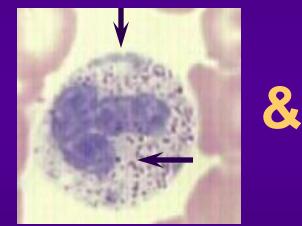
hypersegmented neutrophils in megaloblastic anemias



8

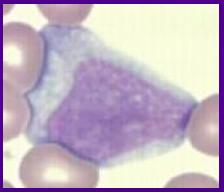


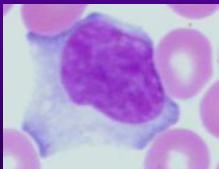
neutrophils In bacterial infections



with Döhle bodies and/or toxic granulation

reactive/atypical lymphocytes (ATL) In viral infections





inherited disorders

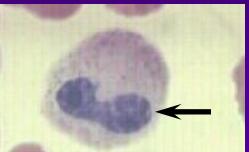




e.g., WBC with inherited nonneoplastic alterations...

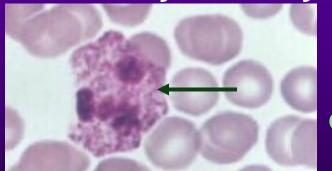
&

Pelger-Huet Anomaly



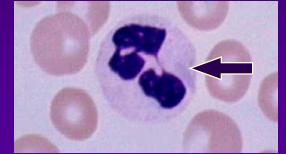
hyposegmented nuclei

Alder-Reilly Anomaly



cytoplasmic black granules &

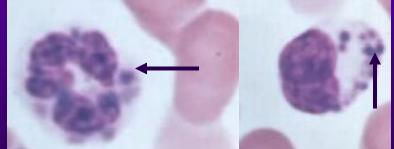
May-Hegglin Anomaly



&

cytoplasmic blue bodies

Chediak-Higashi Syndrome



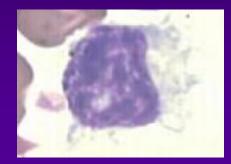
cytoplasmic large black granules





WBC with neoplastic alterations, e.g....

hairy cell lymphocytes



plasma cells



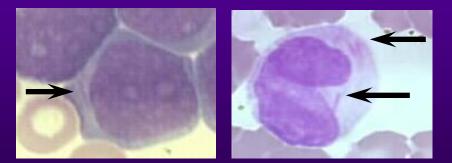
hairy cell leukemia

8

in multiple myeloma

myeloblasts w/ Auer rod(s)

&



in acute myelocytic leukemias





Leukocytic Maturation



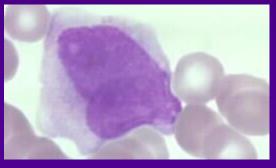




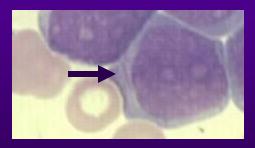
Blasts are the earliest leukocytic precursor that can be seen in peripheral blood.



myeloblast



monoblast



myeloblast w/ Auer rod



lymphoblast

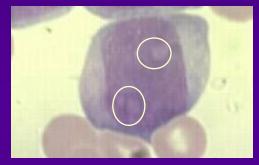








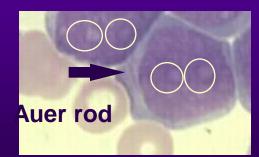
All blasts have nucleoli. <u>and</u> Cell lines are difficult to differentiate on Wright's stain without a distinguishing feature (eg, Auer rod in AML).



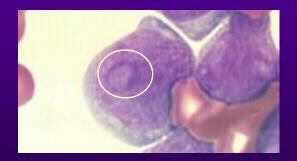
myeloblast



monoblast



myeloblast



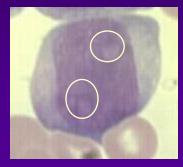
lymphoblast



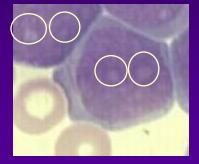


Blasts

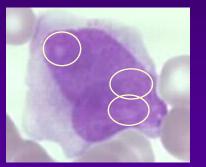
While the presence of nucleoli differentiates blasts from more mature forms,



myeloblast



myeloblast



monoblast



menu

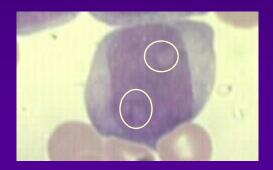
lymphoblast

special stains are usually needed for definitive identification of leukoblasts.

Leukoblasts must also be differentiated from proerythroblasts.





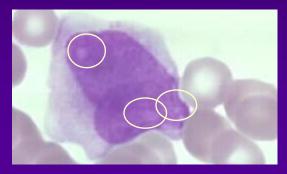


myeloblast

8

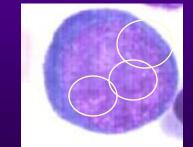


lymphoblast



monoblast

Proerythroblasts



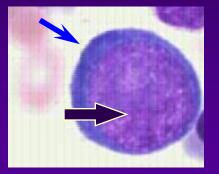
have nucleoli.





How do they differ morphologically?

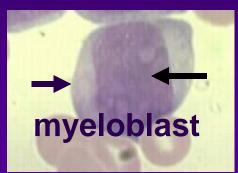
In the proerythroblast,

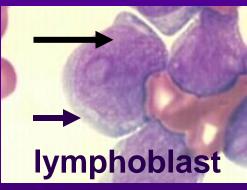


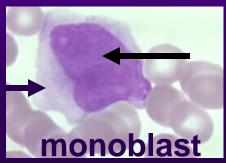
the cytoplasm is usually darker and bluer and the nuclear cor chromatin leu strands are wh linear and del distinct interval

compared to leukoblasts, which are more delicate and interlaced.

than the leukoblasts











Myelocytic (or Granulocytic) Series



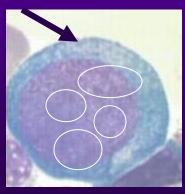


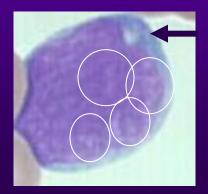
Myelocytic Maturation

There are three types of cells in the myelocytic (or granulocytic) series: neutrophils, eosinophils, and basophils.

Myeloblasts (ie, the earliest precursor) originate in the marrow from a stem cell common to erythroid, megakaryocytic, and granulocytic cells. Prominent nucleoli are seen in the nucleus and the cytoplasm is agranular. Morphologically, they are difficult to differentiate from lymphoblasts or monoblasts.

Examples of myeloblasts:





nucleoli

& agranular cytoplasm



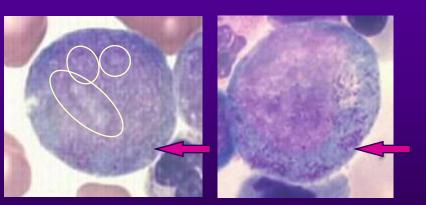


Myelocytic Maturation, continued

Nucleoli are also seen in the **promyelocyte** and the appearance of <u>large azurophilic (nonspecific) cytoplasmic granules</u> in the early stage of its transition is an indication that the cell is a granulocyte. However, morphologic determination as to neutrophilic, eosinophilic, or basophilic cannot yet be made.

Examples of promyelocytes:

prominent nucleoli and cytoplasmic non-specific granules begin to be visible



as the cell matures, the nucleoli begin to fade and the granules become more numerous and prominent

early promyelocyte

late promyelocyte

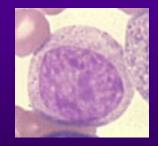




Myelocytic Maturation, continued

As the promyelocyte matures and reaches the myelocytic stage, it has definitely and visually differentiated into one of the three granulocytic types with characteristic cytoplasmic "specific" or secondary granules. Nucleoli are indistinct or not seen and this is the last mitotic stage.

Examples of myelocytes with specific granules:



neutrophilic



basophilic



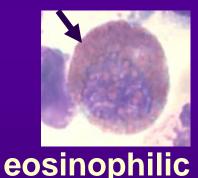
csw Isuhsc 2002

Myelocytic Maturation, continued Specific cytoplasmic granules:



ill-defined reddish granules within the bluish cytoplasm resulting in a lilac or pinkish color

neutrophilic



relatively large, spherical, orange granules



unevenly distributed large, blue-black granules, which are usually also visible on top of the nucleus





Neutrophilic Maturation

From this stage on, as the cell matures, there is little change in the cytoplasm but the nuclear chromatin becomes progressively more condensed and the characteristic nuclear shapes of the metamyelocyte, band (stab form), and mature segmented cell are noted.

Illustrated below in the images of neutrophilic cells:



metamyelocyte



band (or "stab")



mature segmented (or PMN)





Eosinophilic and Basophilic Maturation

menu

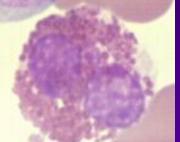
There is usually no differentiation made as to whether eosinophils and basophils are myelocytes, metamyelocytes, band, or mature cells. Regardless of the stage of maturation, they are still referred to only as eosinophils or basophils.

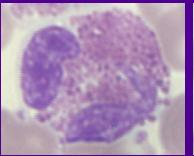
The granules observed in both cell lines are rather large, frequently dense and, in many cases, obscure the nucleus thus making it difficult to see the nuclear shape as illustrated in the images of eosinophils and basophils.

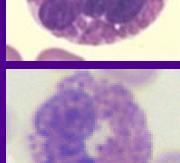


Various Stages Maturation

<-----> Eosinophils----->





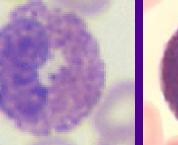


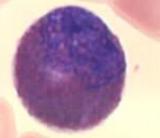


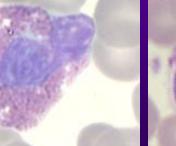




<----Basophils---->

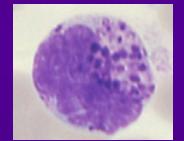
















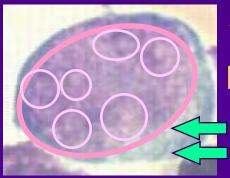


Morphologic Features of Granulocytes





The myeloblast is morphologically undifferentiated as to granulocytic cell line (ie, neutrophilic, eosinophilic, or basophilic).



Size: variable, usually 15 to 20 μm diameter

Nucleus: relatively round and large; predominantly red-stained, delicate, interlaced, well defined and evenly stained chromatin

Nucleoli: large and usually 2 or more

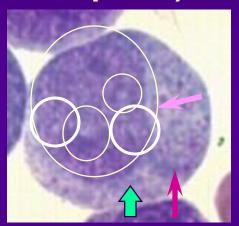
Cytoplasm: agranular, bluish and stains unevenly usually lighter next to the nucleus than at the periphery and may have cytoplasmic tags)





Promyelocyte

The **promyelocyte** is still undifferentiated as to a specific granulocytic cell line (ie, neutrophilic, eosinophilic, or basophilic).



Size: usually larger than blasts but variable depending on the stage in the mitotic cycle Nucleus: round and relatively large with predominantly red-stained chromatin Nucleoli: usually demonstrable

Cytoplasm:

- Stains blue with a relatively light area adjacent to the nucleus
- Granules nonspecific (or primary) granules and absence of secondary granules (ie, neutrophilic, eosinophilic, or basophilic)





Myeloblast vs. Promyelocyte

Myeloblast

∕ csw ∖ Isuhsc

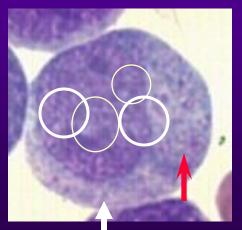
2002

Nucleoli: two or more, large, prominent

Cytoplasm: bluish, unevenly stained

Granules: none visible

Promyelocyte



Nucleoli: usually demonstrable

Cytoplasm: bluish, unevenly stained

Granules: distinct non-specific (or , primary), predominantly dark blue or reddish blue





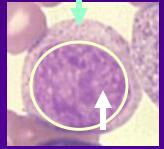
Granulocytes

Granulocytes are neutrophilic, eosinophilic, or basophilic. The cells cannot be morphologically differentiated on Wright's stain until they reach the myelocyte stage and develop specific granules.





This is the last stage in which nucleoli can be seen and mitosis can occur. However, if present, the nucleoli are usual y indistinct. Cell line differentiation is seen.



- Size: usually smaller than promyelocytes (10-18μm in diameter).
- **Nucleus:** round, oval, or flattened on one side.

Chromatin: fine, dispersed pattern in early cells which becomes more condensed as the cell matures.

Nucleoli: usually not demonstrable

Cytoplasm: more than promyelocytes (N/C ratio about 2:1).

- Stains blue but becomes less basophilic as it matures.
- Granules distinct specific (or secondary) granules that are neutrophilic, eosinophilic, or basophilic.



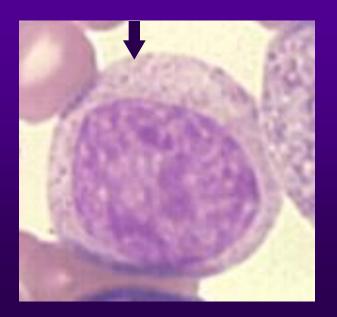


Myelocytes



The first morphologic evidence of specific or secondary granules that provide a means of identifying the cell as a neutrophil, eosinophil or basophil is seen in the myelocyte.

Neutrophils - neutrophilic granules give the cytoplasm a lilac or pinkish appearance



NOTE: an enlargement of the cell shown on the previous slide to better illustrate the granules.





Myelocytes

menu

The maturation sequence of the eosinophils and basophils is the same as the neutrophils. The nuclear features are identical but the color and/or size of the cytoplasmic granules differentiates these cells from the neutrophils.

Eosinophils - relatively large and spherical purplish-red granules give the cytoplasm a reddish-orange color.

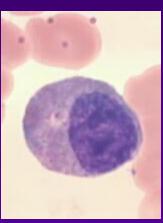
Basophils - large dark blue to black unevenly distributed granules may fill the cytoplasm and, when present in large numbers, may obscure the nucleus.



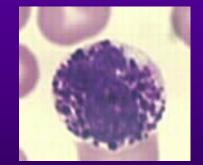
Neutrophilic, Eosinophilic, & Basophilic Myelocytes



eosinophilic



neutrophilic



basophilic



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Metamyelocyte (Neutrophilic)

As soon as the nucleus of the myelocyte (which may be neutrophilic, eosinophilic, or basophilic) becomes indented, the cell is classified as a metamyelocyte. The cell is no longer capable of mitosis.



Size: usually slightly smaller than myelocytes

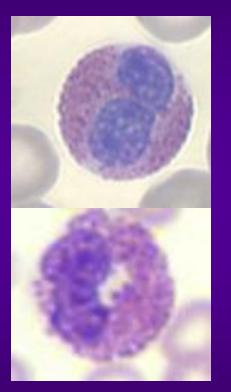
Nucleus: relatively smaller than myelocyte and, as the cell matures, indentation increases.

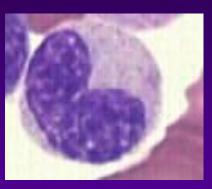
Chromatin: less well defined and becomes more condensed clumped, and darkly stained as the cell matures. Nucleoli: not demonstrable

Cytoplasm: progressively less basophilic than myelocytes and distinct specific (or secondary) granules that are neutrophilic, eosinophilic, or basophilic predominate.



Metamyelocytes





neutrophilic specific granules





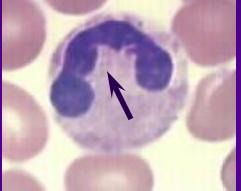
basophilic specific granules





Neutrophilic Band

As soon as the indention in the nucleus of the metamyelocyte becomes greater than 1/2 the diameter, the cell is classified as a band (neutrophil, eosinophil, or basophil).



Size: slightly smaller than metamyelocytes

Nucleus: indented greater than 1/2 diameter and opposite edges of the nucleus become approximately parallel (horse-shoe shape).

Chromatin: dense and clumped, usually with a pyknotic mass at each pole where the lobe will be.

Nucleoli: none present

Cytoplasm: no basophilia & may be slightly eosinophilic; distinct specific (or secondary) granules that are neutrophilic, eosinophilic, or basophilic predominate.



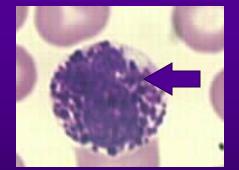




neutrophilic (lilac or pinkish) specific granules





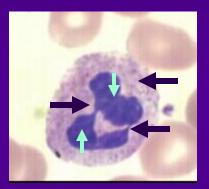


basophilic (blue-black) specific granules





Mature neutrophils have nuclei that are separated into definite lobes. They are frequently referred to as PMN (polymorphonuclear neutrophils).



Size: $10-15 \mu m$ (about twice the size of RBC). Nucleus: separated into definite lobes (usually 2 or 3 with occasional 4 or 5) which are connected by a very narrow filament or strand.

Chromatin: dense and clumped

Nucleoli: not present

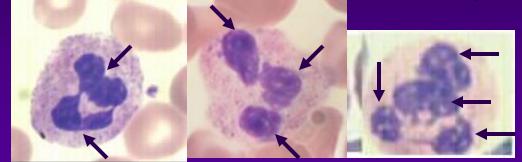
Cytoplasm: slightly eosinophilic or light pink with numerous pink to bluish-black evenly distributed small granules.





Normal Mature Neutrophils, Eosinophils, and Basophils

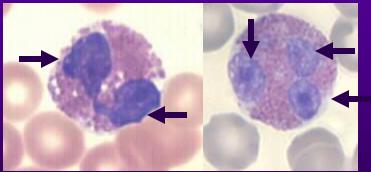
segmented meutrophils (PMN





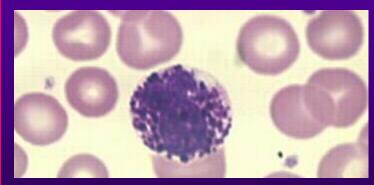
usually 2-4 lobed nucleus but may have a few w/ 5 lobes

eosinophils



usually bilobed nucleus but may have 3 or more lobes

basophil



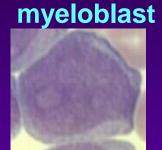
granules usually obscure nucleus



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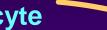
Review: Neutrophilic Maturation

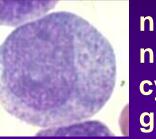




nucleoli & <u>ag</u>ranular cytoplasm







nucleoli & non-specific cytoplasmic granules cells become smaller, nucleus becomes smaller, loss of nucleoli, specific granules appear

neutrophilic metamyelocyte

neutrophilic myelocyte

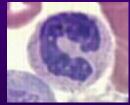




nucleus indented but less than half diameter & specific granules myelocyt

round nucleus & specific^{*} cytoplasmic granules

neutrophilic band



nucleus indented more than half diameter & specific granules





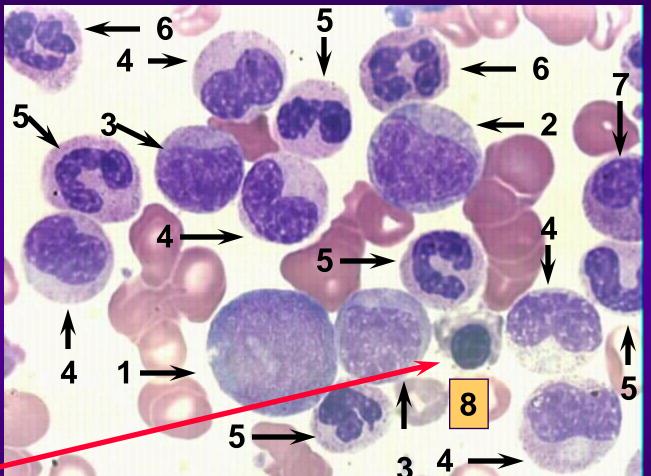
nucleus with separate lobes & specific granules





Examples of granulocytes in various stages of maturation:

- **1** early promyelocyte
- 2 late promyelocyte or early myelocyte
- 3 myelocyte
- 4 metamyelocyte
- **5** band neutrophil
- 6 <u>mature</u> segmented neutrophil (PMN)
- 7 eosinophil



8 Whoa! That's not a WBC. It's a nucleated RBC but will also be included in the total WBC count.



Now, Can you Identify the stages of granulocytes just illustrated?

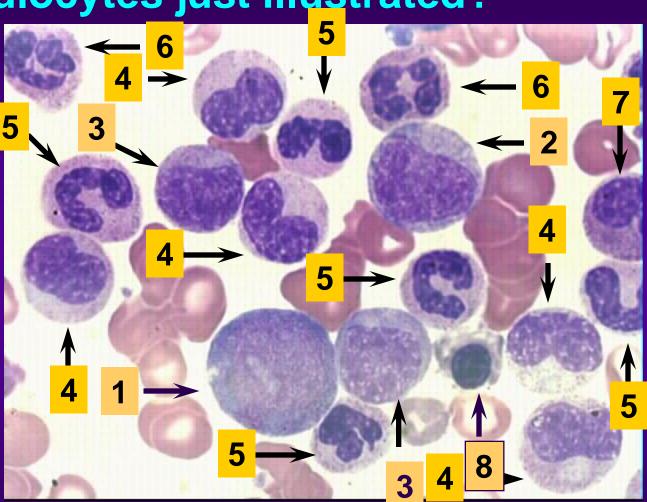
- arly promyelocyte
- 2 late promyelocyte or early myelocyte
- 3 myelocyte

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- 4 metamyelocyte
- **5** band neutrophil
- 6 <u>mature</u> segmented neutrophil (PMN)
- 7 eosinophil



8 Remember, it's a NRBC!





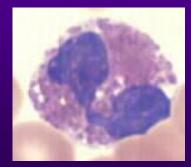
Review: Eosinophilic & Basophilic Maturation

The maturation sequence for eosinophils and basophils parallels that of neutrophils. Blast and promyelocyte stages are morphologically undifferentiated as to neutrophils, eosinophils, or basophils.

The cells can be differentiated in the myelocyte stage with the appearance of specific cytoplasmic granules (ie, neutrophilic pink, eosinophilic orange, or basophilic dark blue-black) that remain through maturity.



neutrophil



eosinophil



basophil



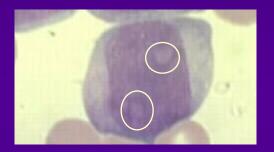


Lymphocytic Series





Blasts are the earliest leukocytic precursor that can be seen in peripheral blood and have nucleoli which help to differentiate them from mature cells.







menu

myeloblast

lymphoblast

monoblast

Special stains and/or flow cytometry are usually needed for definitive differentiation of the various leukoblast cell lines (i.e., myeloblasts, lymphoblasts, and monoblasts).

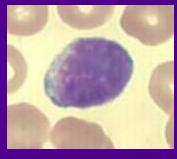
The least mature lymphoid cell seen in peripheral blood is the lymphoblast.



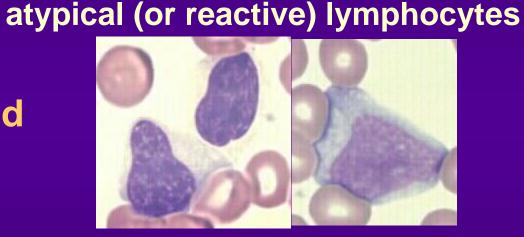


For Clinical Pathology 201, you will be expected to be able to identify and differentiate:

mature lymphocytes

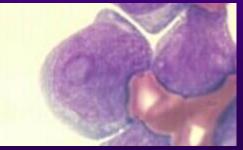


and



lympho<u>blasts</u>









Monocytic Series

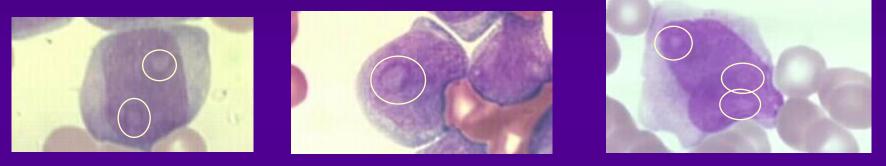




Blasts



Blasts are the earliest leukocytic precursor that can be seen in peripheral blood and have nucleoli which help to differentiate them from mature cells.



myeloblast

lymphoblast

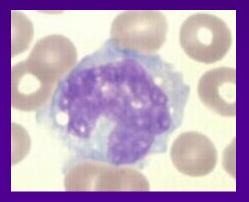
monoblast

The earliest monocytic cell seen in peripheral blood is the monoblast. Special stains are usually needed for definitive identification of blast cell lines. Refer to the Course Manual.





For Clinical Pathology 201, you will be expected to be able to differentiate mature monocytes



i.e., recognize the <u>mature</u> cell and differentiate

from monoblasts

with nucleoli



Special stains and/or flow cytometry are usually needed for definitive identification of the blast cell line.





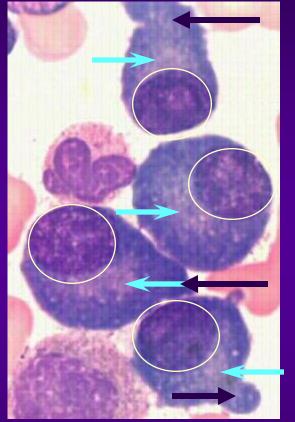
Plasma Cells





Plasma Cells





Size: mature cells vary greatly

Shape: usually oval shape with relatively smooth cytoplasmic margins, but, like the lymphocyte, the plasmocyte is easily traumatized and often has frayed or nebulous margins and pointed or filamentous cytoplasmic projections

Nucleus: relatively small and round and eccentrically located

Cytoplasm: abundant

The cytoplasm adjacent to the nucleus stains more lightly than the periphery of the cell which has a high saturation of red and blue dyes. The area is called a "golgi".



Plasma cells are never present in normal peripheral blood. They constitute about 1% of the nucleated cells in normal bone marrow where they tend to be grouped in small islands around blood vessels. They may be present in small numbers in chronic infections, in granulomatous and allergic diseases and in plasma cell myeloma.

Plasmoblasts (not shown) are cells with relatively large nuclei, nucleoli and delicate chromatin which takes a predominantly red color. Plasmoblasts are not recognizable except in malignancies of the plasmocytoid type.



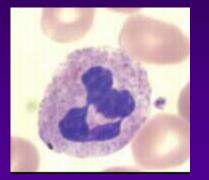


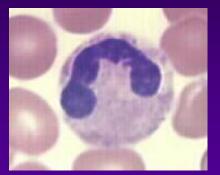
Leukocytes in Normal Perpheral Blood

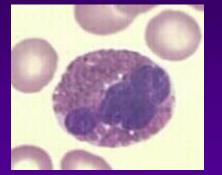


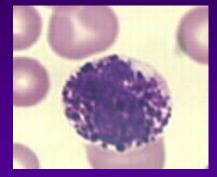
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Review: WBC found in normal peripheral blood:









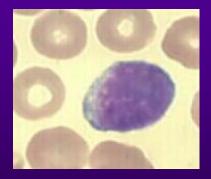
menu

mature neutrophils

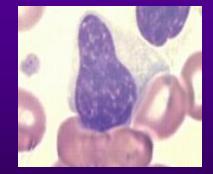
band neutrophils

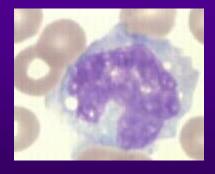
eosinophils

basophils



lymphocytes





atypical lymphocytes (<6% of lymphocytes) monocytes



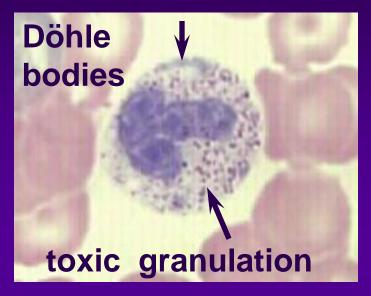


Leukocytes with Acquired Non-neoplastic Alterations





Toxic Granulation & Dohle Bodies in Neutrophils



Toxic granulation - dark blue to purple cytoplasmic granules and/or Döhle bodies - small blue cytoplasmic inclusions

Toxic granules may be seen in severe bacterial infections, burns, aplastic anemia, and following administration of toxic agents.

Frequently, Döhle bodies will also be seen concomitantly with toxic granulation.



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Hypersegmented Neutrophils



Hypersegmented neutrophil - (ie, > 5 lobes) which are presumably the result of abnormal nuclear maturation. Five lobes in more than 5% of the neutrophils constitute hypersegmentation, as do any neutrophils with 6 or more lobes. In this case, there are 7.

Hypersegmented neutrophils are characteristic features of megaloblastic anemias that are due to vitamin B_{12} or folate deficiency.

Refractory anemias that are megaloblastic usually do not include these granulocytic changes (ie, hypersegmentation).



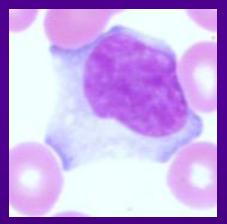


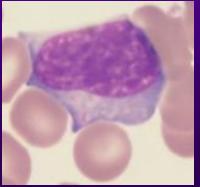


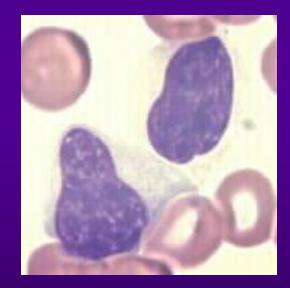
Atypical/Reactive Lymphocytes

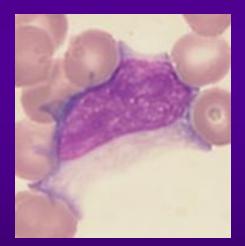
Atypical/reactive lymphocytes may be seen most typically in viral disorders.

Atypical lymphocytes may have abundant cytoplasm with scalloped or indented rims ...











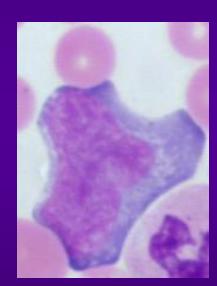




Atypical/Reactive Lymphocytes

...or have darker cytoplasm and more monocytoid nuclear or plasmacytoid features ..













Leukocytes with Inherited Non-neoplastic Alterations



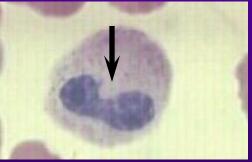




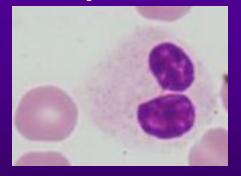
Pelger-Huet Anomaly

An inherited autosomal dominant condition in which there is a failure of normal segmentation of granulocytic nuclei (i.e., hyposegmented nuclei). The nuclei may be...

band shaped



bi-lobed or "pince-nez" shaped



or in very rare cases, round shaped (like a myelocyte).







Pelger-Huet Anomaly, continued

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Absence of symptoms of infection or other cause of a "left shift", history of persistent blood morphology, and/or similar blood morphology of other family members suggests the anomaly.

The cell morphology persists through life and the cells are functional.







Pseudo-Pelger-Huet Anomaly

An acquired disorder similar in appearance to Pelger-Huet anomaly may occasionally be found in cases of granulocytic leukemia, myeloproliferative disorders, some infections, and after exposure to certain drugs.

Band forms, neutrophils with only two segments or "pince-nez" appearance (not shown), and/or neutrophils with round non-segmented nuclei are seen. Neutrophils with > 2 segments (lobes) will <u>not</u> be seen in this disorder.

There is asynchronism between the shape of the nucleus and the maturity of the nucleus and cytoplasm.



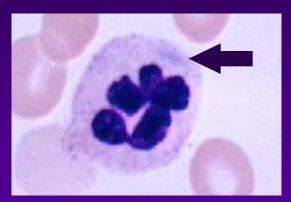








May-Hegglin Anomaly



Pale blue cytoplasmic inclusions that resemble Dohle bodies but are larger and more prominent. They may be found in neutrophils, eosinophils, basophils, and monocytes.

Bluish aggregations (RNA) particles can be seen in the cytoplasm of neutrophils. Giant platelets can also be seen.

This is a rare autosomal dominant condition.

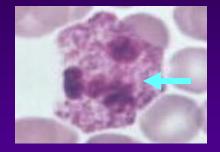
The cells are functional and the cytoplasmic inclusions in May-Hegglin persist through life. Acquired Dohle bodies are transient.





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Alder-Reilly Anomaly



This disorder is characterized by neutrophils with large azurophilic and basophilic granules in the cytoplasm that resemble toxic granulation.

This is an autosomal recessive trait. There is no apparent interference with leukocyte function.

However, these granules may also be seen in association with some but not all patients with <u>gargoylism (the Hurler</u> <u>syndrome)</u>, or more generally, the genetic mucopoly-<u>saccharidoses</u>).

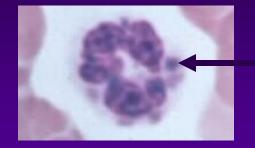
Alder-Reilly granules are persistent through life whereas acquired toxic granulation is transient.





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Chediak-Higashi Syndrome



Abnormally large cytoplasmic black granules which appear to be abnormal lysosomes may be seen in granulocytes, monocytes, and lymphocytes.

This is a rare sutosomal recessive disorder characterized by <u>paratial albinism</u>, <u>photophobia</u>, <u>and frequent pyogenic</u> <u>infections</u>. An accelerated lymphoma-like phase occurs, with lymphadenopathy, hepatosplenomegaly, and pancytopenia. Lymphoid infiltrates are widespread and <u>death</u> <u>ensues at an early age</u>. Leukocyte functional abnormalities exist.

The abnormal morphologic features are persistent throughout life.





Leukocytes with Neoplastic Alterations



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Hairy Cell Lymphocytes



The hallmark of hairy cell leukemia is the presence of lymphocytes with irregular long, delicate cytoplasmic projections which give them a hairy appearance.

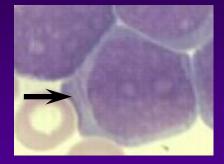
This is an uncommon chronic, low grade lymphoproliferative disease (or CLL) that occurs about 5 times more frequently in males than females.

Onset of disease is insidious; weakness and lethargy; or may be asymptomatic (10-15% of patients). May be bleeding and bruising.

Normocytic, normochromic anemia related to the neoplastic cell mass, marrow hypoplasia, and hypersplenism is also seen. Thrombocytopenia in about 75% of patients and Coomb's test may be positive.



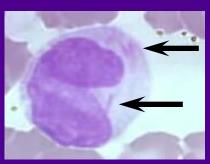
Myeloblasts with Auer Rods



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Auer rods in the cytoplasm of myeloblasts are associated with acute leukemias having a myeloid component. They appear as cytoplasmic reddish rods with Wright's or Wright's-Giemsa stains.

They may be seen in some, but not all, myeloblasts in some, but not all, of the variants of <u>acute</u> myelocytic leukemia. They are not seen in blasts in chronic myelocytic leukemia.

The presence of Auer rods in the cytoplasm of blasts effectively rules out a lymphoid disorder.





End of Leukocytes

This ends the section on leukocytes. Click on:

Erythrocytes to go to the next section of this study module as designed.

or

Menu to go back to the menu.

or

Quit to end the exercise.





Erythrocytes





How are the RBC identified?

Typical morphologic nuclear and/or cytoplasmic features provide a means by which RBC can be identified. For example:

Maturity

- Size
- Shape
- Color
- Hemoglobin content
- Inclusions (if any)





How are RBC classified as to maturity?

Characteristic nuclear and/or cytoplasmic morphologic features allow red blood cells to be classified as:

- pronormoblast (or rubriblast), the earliest form seen in peripheral blood (ie, least mature)
- basophilic normoblast (or prorubricyte)
- polychromatophilic normoblast (or rubricyte)
- orthochromatic normoblast (or metarubricyte)
- polychromatophilic erythrocyte (or diffusely basophilic erythrocyte
- mature erythrocytes



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Summary of the key features of erythrocyte development:

Cell	Cytoplasm	Nucleus	Nucleoli	chromatin
Pronormoblast	Scanty, basophilic	Large	Prominent	Dispersed, finely granular
Basophilic normoblast	Increased, still basophilic	Moderate	Indistinct	Dispersed but more condensed
Polychromatophilic normoblast	Mixed basophilic & eosinophilic	Smaller	None	Chromatin & parachromatin
Orthochromatic normoblast	More eosinophilic	Pyknotic	None	Condensed chromatin, no parachromatin
Polychromatophilic erythrocyte*	Eosinophilic	None	None	None
erythrocyte	Eosinophilic	None	None	None

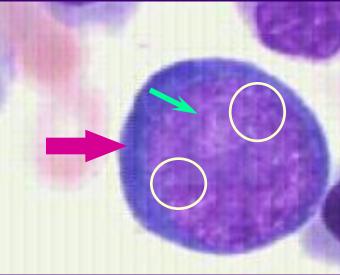
*called a reticulocyte after staining with a supravital stain







What are the characteristic features of pronormoblasts?



Nucleoli: usually visible

Nuclear chromatin: linear and distinct

Cytoplasm: in earliest form stains light blue; in later and more frequently occuring forms has a dark royal-blue color similar to that seen in some plasmacytes.







&

What are characteristic features of basophilic normoblasts (prorubricytes)?



Nucleoli: ill-defined or absent Nuclear chromatin: coarsening of the chromatin pattern

Cytoplasm: deeply basophilic due to the abundance of RNA with a reddish tinge produced by varying amounts of hemoglobin present

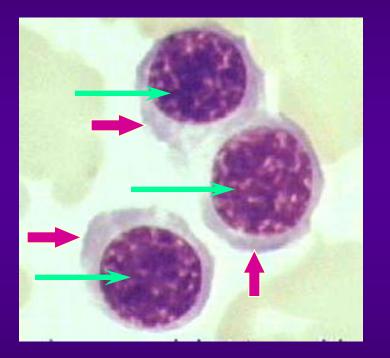
some cells may have a Golgi (clear) area adjacent to the nucleus (not visible in this cell).







What are characteristic features of polychromatophilic normoblasts (rubricytes)?



Nucleoli: no longer visible

Nuclear chromatin: thickened and irregularly coarsened

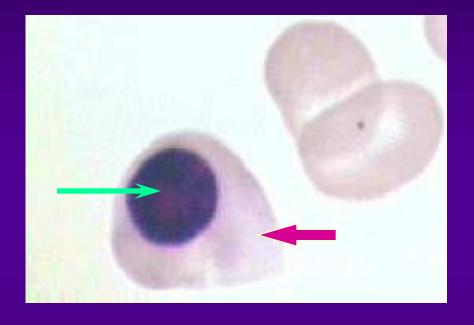
Cytoplasm: relatively more cytoplasm than the basophilic normoblast and takes varying mixtures of red and blue stain







What are characteristic features of orthochromatic normoblasts (metarubricytes)?



Nucleoli: none

Nuclear chromatin: nonlinear clumped structure or, as shown in this field, a solid reddishblue-black degenerated nucleus

Cytoplasm: predominantly red cytoplasm with minimal amounts of residual blue



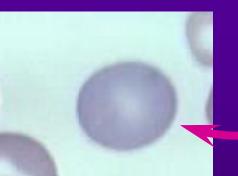




What are characteristic features of polychromatophilic erythrocytes (diffusely basophilic erythrocytes)?



orthochromatic erythro<u>blast</u> (erythro<u>blasts</u> have a nucleus)



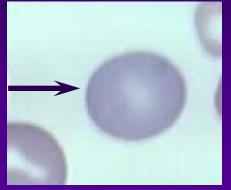
As the cell matures, the nucleus is **extruded** and the cell becomes a

polychromatophilic erythrocyte (erythrocytes do not have a nucleus)

Cytoplasm: predominantly red but may have a bluish tinge due to the reticulum strands (RNA) still present.



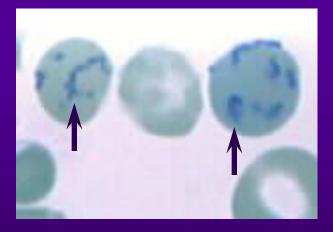
What is the correct name for this cell on a Wright's stained blood smear?



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On a Wright's stained blood smear, the cell is called a polychromatophilic erythrocyte.



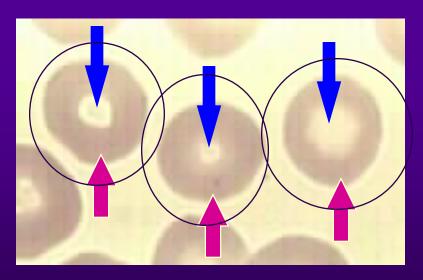
When these cells are stained with a supravital stain (e.g., new methylene blue), the residual RNA strands are precipitated,

and the cell is then called a reticulocyte.





Normal mature erythrocytes are anucleated biconcave discs that stain a reddish buff color with Wright's (or Wright's-Giemsa) stain and have a small (about 1/3 of the cell) central pallor.



The intensity of the stain in the center of the cell (i.e., the thin portion)

is less than at the outer rim of the cell (i.e., the thicker portion).



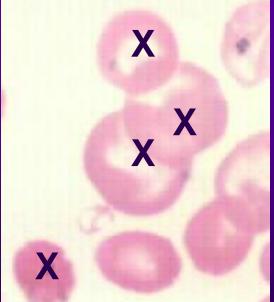


 Normocytic is the term used to indicate RBC that are normal size (6-8 μ in diameter) and normal shape.



Anisocytosis is a "generic" term used to indicate a variation in cell size, eg.,

normocytic microcytic macrocytic







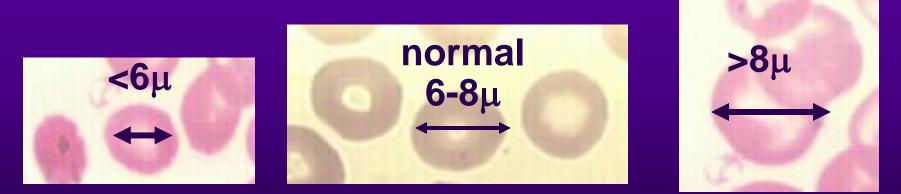
Normocytic (RBC 6-8 μ in diameter).

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- **Microcytic (RBC < 6** μ in diameter).
- **Macrocytic (RBC > 8** μ in diameter).







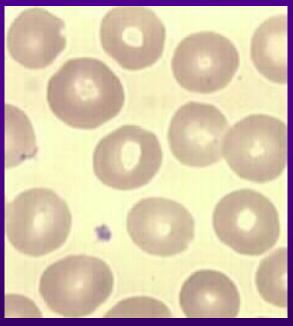


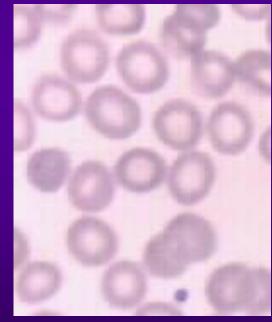
Depending upon the <u>predominant</u> cell size, an RBC population can be classified as...

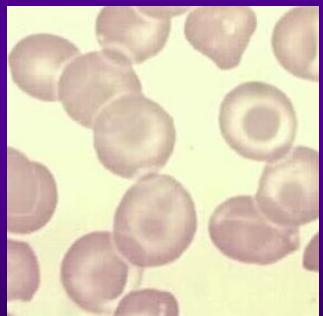
Normocytic

Microcytic

Macrocytic



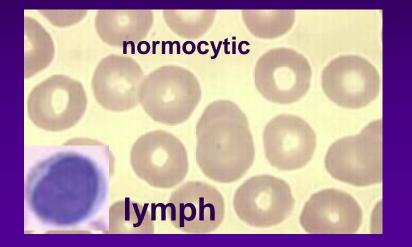




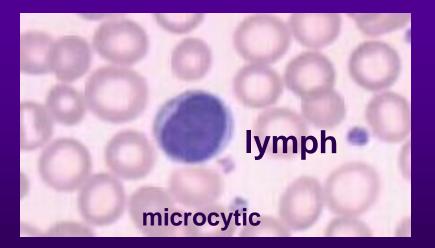


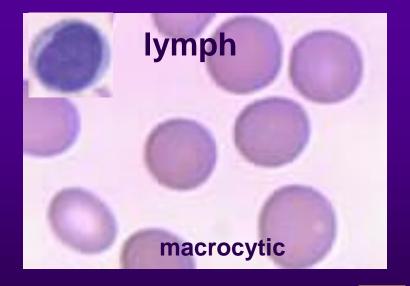


Illustrative Mature Erythrocytes



Comparison of erythrocytes with the normal small mature lymphocyte (which is about $6-10\mu$ in diameter) is helpful in determining whether cells are normocytic, microcytic, or macrocytic.



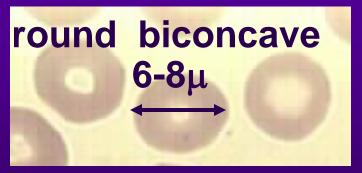






How are RBC classified as to shape?

Normocytic is the term used to indicate RBC that are normal size (6-8 μ in diameter) and normal shape (i.e., round, biconcave).

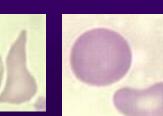


Poikilocytosis is the "generic" term used to indicate variation in shape.



Individual red cells can have numerous abnormal shapes, eg:















What are some of the RBC shape classifications?

Individual red cells can be classified as:

- > ovalocytes (elliptocytes)
- spherocytes
- target cells (leptocytes)
- Schistocytes (RBC fragments)

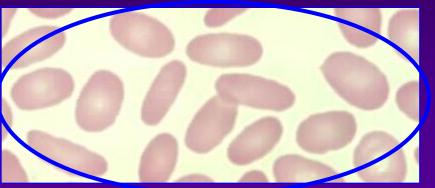


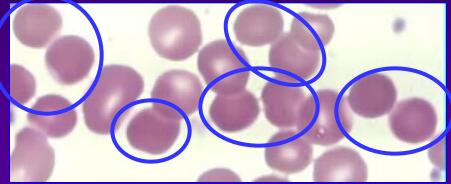


Illustrative RBC Shapes

ovalocytes (elliptocytes)

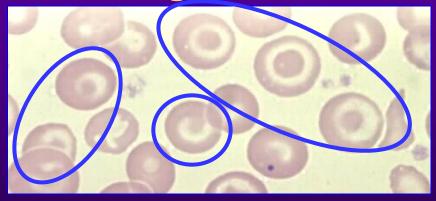
spherocytes

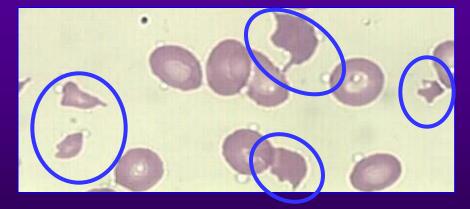




target cells

schistocytes



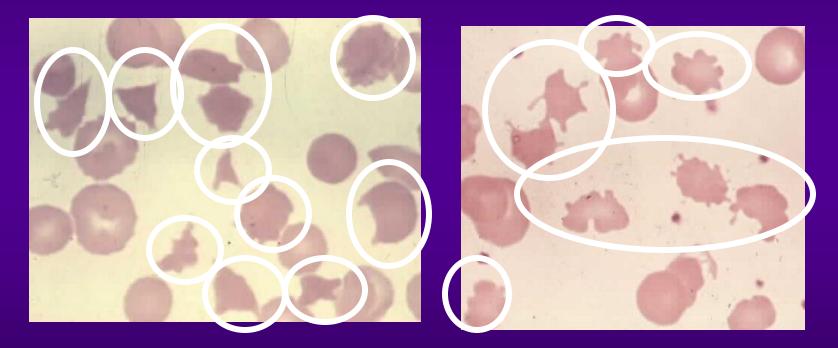






Schistocytes

Schistocytes are RBC fragments and may have a variety of shapes.









What other erythroid shapes can be seen?

Individual red cells can also be classified as:

- sickle cells (trepanocytes or meniscocytes)
- bitocytes (keratocytes)
- echinocytes or crenated
- acanthocytes



menu

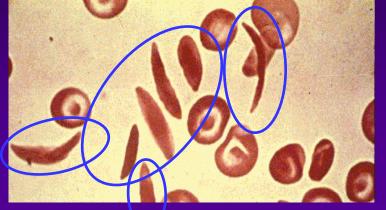
Illustrative RBC Shapes

sickle cells (drepanocytes)

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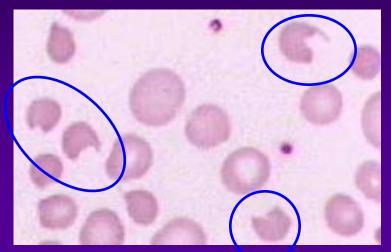
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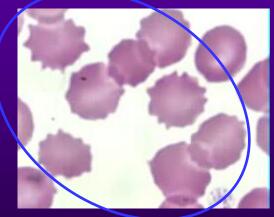
spiculated (acanthocytes)



bitocytes (keratocytes)



crenated (echinocytes)









What about groups of RBC?

Groups or clumps of red cells can also be classified as:

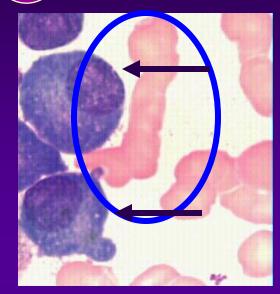
rouleaux

agglutination



Illustrative rouleau RBC formation:





CSW

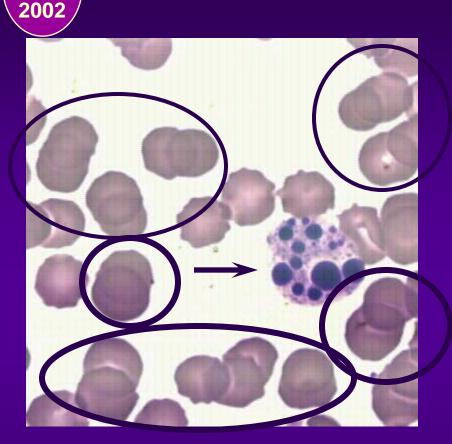
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> **Rouleau** is an aggreation of RBC that is aligned one upon the other resembling stacks of coins and is caused by elevated plasma fibrinogen or globulins.

This phenomenon causes an increased erythrocyte sedimentation rate (ESR) and interferes with the hemogram parameters. Rouleau is especially characteristic of paraproteinemia (monoclonal gammopathy), in which case plasma cells may also be seen.



Illustrative agglutinated RBC:



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Agglutination of red cells, caused by cold agglutinins, resembles rouleau but is more irregular and may appear in round clumps rather than linear rouleau.

The large cell in the field is a degenerated neutrophil with pyknotic nuclei and nuclear fragments.

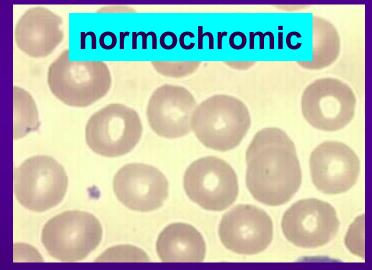




How are mature RBC classified as to hemoglobin content?

Depending upon the hemoglobin content, mature RBC may be classified as :

anucleated, pinkish cytoplasm with a small central pallor (about 1/3 of the cell diameter).



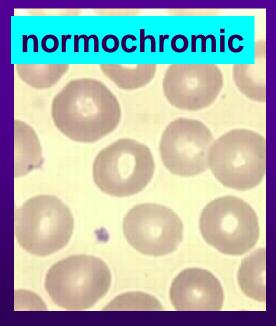


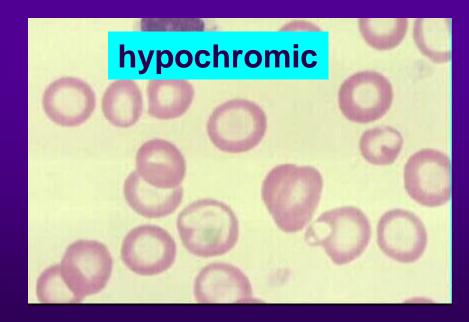




or hypochromic...

Anucleated, pinkish cytoplasm with a more pronounced central pallor (i.e., greater than 1/3 the diameter of the cell).



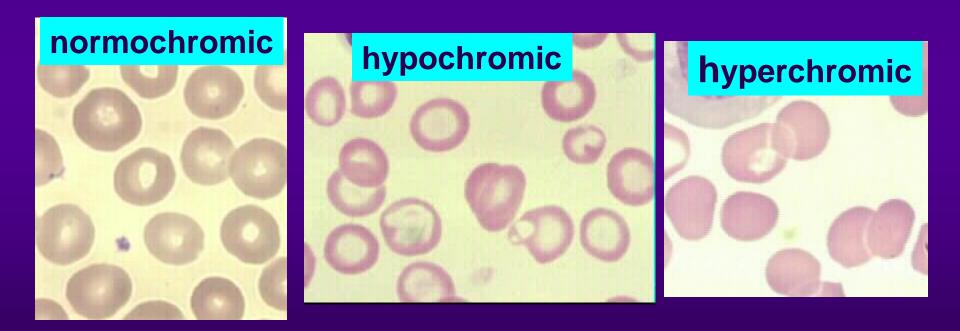








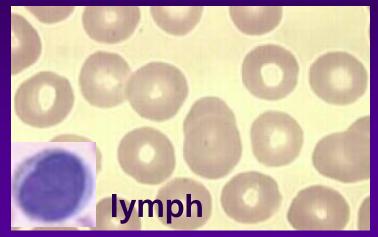
Anucleated, pinkish cytoplasm without central pallor (generally associated with megaloblastic anemias).







Illustrative Mature Erythrocytes Normochromic (Normocyte)

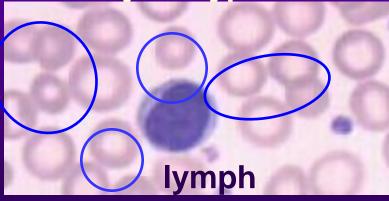


Microcytic Hypochromic

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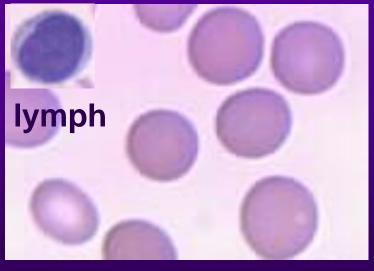
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Remember, comparison of the RBC with normal small mature lymphocytes is helpful in classifying them as normocytic, microcytic, or macrocytic.

Macrocytic Hyperchromic







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How are RBC inclusions classified?

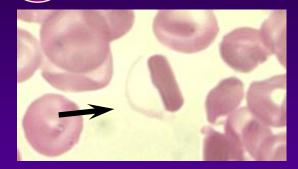
Cellular inclusions that may be found in erythrocytes may include:

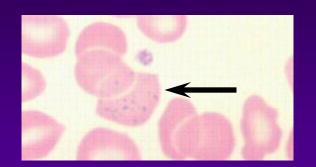
- Hemoglobin C crystals
- Basophilic stippling
- Howell-Jolly bodies
- Pappenheimer bodies
- Heinz bodies
- Cabot rings



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Illustrative RBC with inclusions:



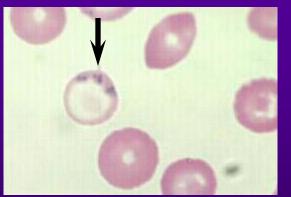




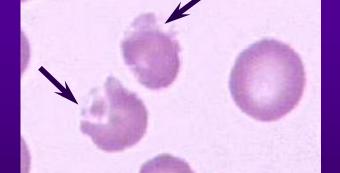
Howell-Jolly body



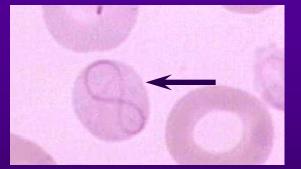




Pappenheimer bodies



(supravital stain) Heinz bodies

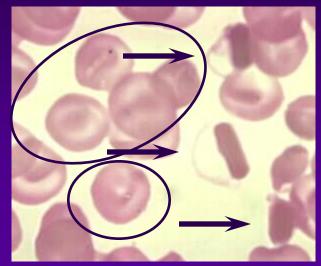


Cabot rings





Hemoglobin C Crystals



Target cells are characteristically seen in HbC disease and syndromes and may be the only abnormality in heterozygous HbC.

Hexaganol shaped HbC crystals may be seen in homozygous HbC disease but are not seen in heterozygous HbC trait. The crystals may be

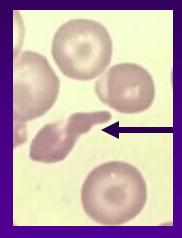
intracellular or extracellular.





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Hemoglobin C Crystals, cont'd



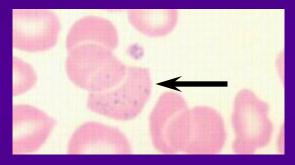
Other shaped forms of the crystals (e.g., glove shaped) are seen in HbSC disease.





Basophilic Stippling

Irregular basophilic granules, which may be coarse or fine, dispersed throughout an erythrocyte is called **basophilic stippling**. This finding is attributed to abnormal instability of the residual RNA in the cell.



Fine stippling is commonly seen when there is increased polychromatophilia, and, therefore, with increased production of red cells.

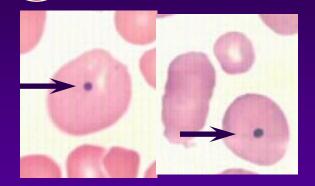
Coarse stippling may be seen in;

- lead poisoning or other diseases with impaired hemoglobin synthesis
- megaloblastic anemia
- other forms of severe anemia (eg, thalassemia major, sickle cell disease

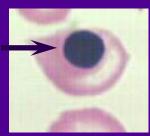


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Howell-Jolly Bodies



Howell-Jolly bodies are smooth, round, intracellular remnants of nuclear chromatin (DNA) that may be found in erythrocytes.



Their color may vary with the stain but are usually the same color as the nuclei of polychromatophilic erythroblasts.

Single Howell-Jolly bodies may be seen in megaloblastic anemia, hemolytic anemia, hemoglobinopathies, thalassemia major, and after splenectomy.

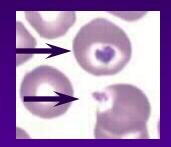
Multiple Howell-Jolly bodies in a single cell is usually indicative of abnormal erythropoiesis (e.g., megalo-blastic anemia).





Looks Like Howell-Jolly Bodies

Don't confuse Howell-Jolly bodies with platelets on top of a red cell.



Characteristically, platelets will appear to be surrounded by a clear "halo" where the hemoglobin has been displaced.



Howell-Jolly bodies usually have no halo.





Pappenheimer bodies:

Pappenheimer bodies appear as dark blue intracellular inorganic iron-containing granules when seen on Wright-Giemsa stained blood smears.



Pappenheimer bodies are few in a given red cell and are usually clustered at the edge of the cell membrane.

Wright-Giemsa stain

These cells are called <u>siderocytes</u> when observed after staining with an iron stain (e.g., Prussian blue).





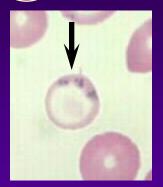
Prussian Blue stain





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Pappenheimer bodies, cont'd.

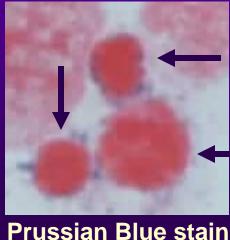


Pappenheimer bodies are associated with iron-loading disorders.

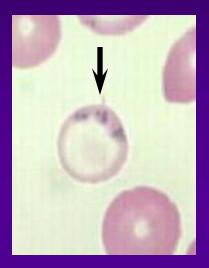
Wright-Giemsa stain

When Pappenheimer bodies are seen in peripheral blood, there may be a concomitant increase of siderocytes and sideroblasts in the bone marrow.

When the siderotic granules surround at least 2/3 of the circumference of the nucleus, the cell is called a "ringed sideroblast".



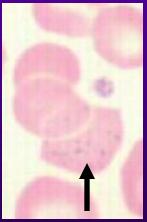
Don't confuse Pappenheimer bodies with basophilic stippling...



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While Pappenheimer bodies usually appear as one or several small round particles clustered together (usually near the rim of the cell),

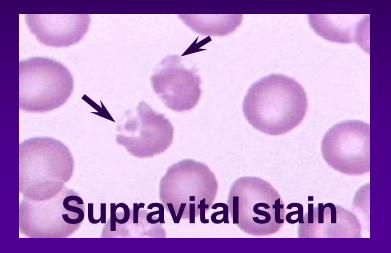


basophilic stippling particles are numerous and dispersed throughout the cell.





Heinz Bodies



Heinz bodies are not visualized on Wright's stained blood smears but are seen only after staining with supravital dyes. Even with these stains, exposure to an oxidizing drug is often required before they are detected.

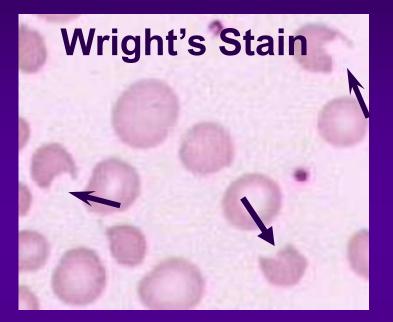






Heinz Bodies,cont'd

With removal of the Heinz body by the spleen, the cells observed on Wright-Giemsa stained blood smears appear to have had a bite taken out of the cell membrane and are called keratocytes (or "bitocytes").

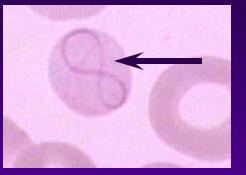


Heinz bodies are most frequently associated with G6PD and may be seen in hemolytic anemias and drugs such as phenacetin. They may also be associated with thalassemia major and hemoglobinopathies.





Cabot Rings



Cabot rings appear in erythrocytes as red or reddish purple intracellular structures. Their shape is usually in the form of a ring, figure-of-eight, or

loop with no internal structure. They are thought to be microtubules remaining from a mitotic spindle.

They are observed in erythrocytes in rare cases of pernicious anemia, lead poisoning, and certain other disorders of erythropoiesis. They are interpreted as evidence of abnormal erythropoiesis.

In this course, you will see them only in photos or computer images.





RBC Inclusions – Composition & Stains

Inclusions	Composition	Stain for ID
Basophilic Stippling	Unstable RNA	Wright-Giemsa
Cabot Rings	Mitotic remnant	Wright-Giemsa
Heinz Bodies	denatured hemoglobin	Supravital*
Hemoglobin C Crystals*	Hemoglobin C	Wright-Giemsa
Howell-Jolly Bodies*	DNA nuclear remnant	Wright-Giemsa
Normoblasts (NRBC)*	DNA	Wright-Giemsa
Pappenheimer Bodies	Iron particles	Wright-Giemsa
Reticulocytes	Precipitated RNA	Supravital*
Sideroblasts	Iron particles	Prussian Blue
Siderocytes	Iron particles	Prussian Blue

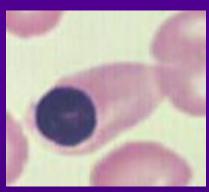
*may be observed with supravital but are identified with Wright-Giemsa ** e.g., new methylene blue, crystal violet





Nucleated Red Blood Cells (NRBC)

NRBC are not normally present in peripheral blood of adults. They may be seen normally in the peripheral blood of newborns and in some diseases in adults. The NRBC most commonly seen is the orthochromatophilic erythroblast.



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However, less mature stages may also be seen.









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How are immature RBC classified?

Immature RBC precursors are classified as:

- Normocytic when they are of normal size.
- Microcytic when they are smaller than normal.
- Macrocytic Non-megaloblastic when they are larger than normal with synchronized nucleus and cytoplasm maturation (i.e., normoblastic bone marrow).
- Megaloblastic when they are larger than normal (i.e., macrocytic) and have asynchronized nucleus and cytoplasm maturation (I.e., megaloblastic bone marrow).

Characteristic nuclear/cytoplasmic fea-Isuhsc tures at various stages of RBC maturation.

proerythroblast (earliest form w/ nucleoli)

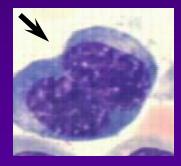
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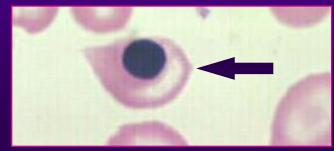
basophilic erythroblast (ill-defined or absent nucleoli)



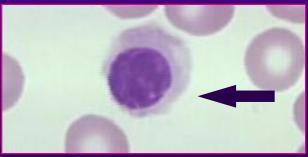


menu

orthochromatophilic erythroblast (last stage before extrusion of nucleus)



polychromatophilic erythroblast (cytoplasmic evidence of HGB)







Characteristic nuclear/cytoplasmic fea-Isuhsc tures of megaloblastic RBC maturation.

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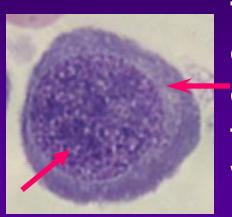
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The megaloblastic precursors are much larger than normal with asynchrony between the nucleus and cytoplasm maturation. The cytoplasm develops at the normal rate while the nucleus lags behind.

Therefore, it is difficult to assign a specific stage of development for an individual cell. For example, the nuclear features may be consistent with a basophilic megaloblast while the cytoplasm may be more mature and be consistent with a later stage.



Examples of nuclear/cytoplasmic asyn-Isuhsc 2002 chrony in megaloblastic precursors:



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The very large early megaloblastic precursor has a nucleus consistent with a pronormoblast, but the cytoplasm is consistent with the more mature basophilic erythroblast with visible evidence of hemoglobin (i.e., pink tinges).

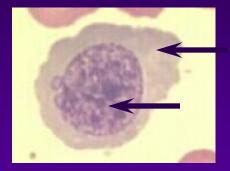
menu



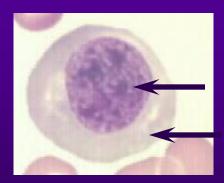
The nucleus of this megaloblast is consistent with a basophilic erythroblast but the cytoplasm is more consistent with a polychromatophilic erythroblast with varying mixtures of red and blood stain.

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Other examples of megaloblastic precursors:



This nucleus is consistent with the thickened and irregularly coarsened chromatin of the polychromatophilic erythroblast ... while the cytoplasm is predominantly red with minimal amounts of residual blue that is more consistent with the orthochromatic erythroblast.



Another megaloblast in which the nucleus looks consistent with a polychromatophilic erythroblast

while the cytoplasm is already as mature looking as the anucleated polychromatophilic erythrocyte.





More mature megaloblastic cells:

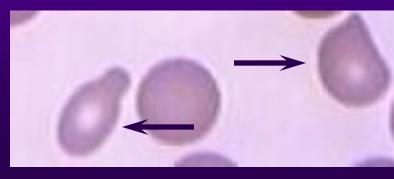
Polychromatophilic erythrocytes and mature RBC are larger than normal (i.e., macrocytic) and typically seen are:



macrocytes without central pallor



macroovalocytes



macro tear-drops







Abnormal Erythrocytes Terminology (Definitions)





Abnormal RBC are differentiated and identified as part of the "diff".

Changes in size, shape, hemoglobin content, and/or appearance of cellular inclusions may occur as a result of a disease process. Such changes are noted as part of the "diff".

What terminology is used to indicate the presence of abnormal red cells?

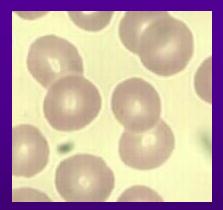




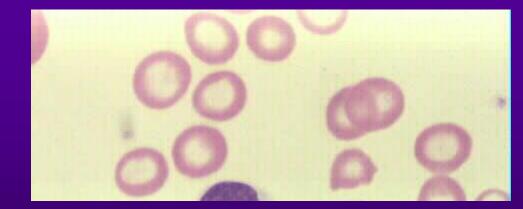
Definitions:



erythrocytes that demonstrate a central pale area that becomes larger and paler as the hemoglobin content diminishes (less than 1/3 of cell diameter.



normochromic



hypochromic

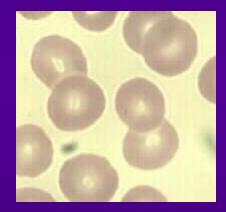




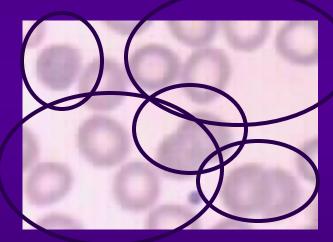
Definitions:

Anisochromic or dimorphic

indicates the presence of both normochromic and hypochromic cells in the same blood film.



normochromic



anisochromic or dimorphic normochromic

hypochromic



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Definitions:

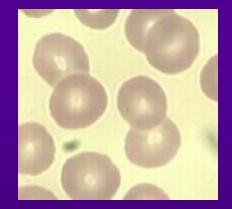
Polychromasia and polychromatophilia are interchangeable terms used to indicate the increased presence of non-nucleated immature erythrocytes (polychromatophilic erythrocytes) that contain residual RNA which gives a blue-gray tint to the red cells. These cells, which remain after ejection of the nucleus from the orthochromatic erythroblast, are slightly larger than mature erythrocytes. After exposure to a supravital stain, the cytoplasmic organelles of these cells clump into an easily recognized blue-staining reticulum and the cell is called a reticulocyte.

view cells

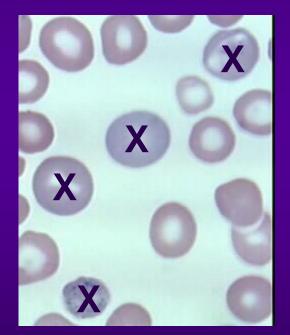


continued:

Polychromasia (polychromatophilia)



normochromic



polychromatophilic erythrocytes



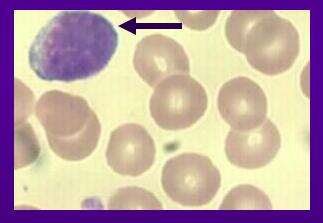


Definitions:



are abnormally small erythrocytes (i.e., less than 6 μ in diameter).

Compare with lymphocyte nuclei (\approx 8-10 μ in diameter).



normocytic RBC ≈ 6-8µ diameter

microcytic (predominant)



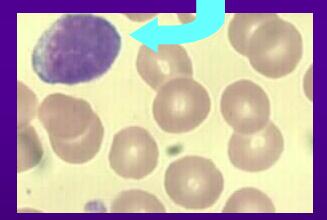


Definitions:



are abnormally large erythrocytes (i.e., greater than 8 μ in diameter.

lymphocyte (with nuclei about 8-10 μ in diameter)



normocytic RBC β 6-8μ diameter



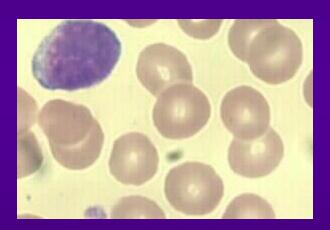
macrocytic





<u>Anisocytosis</u>

is a "generic" term used to indicate an abnormal variation in size of erythrocytes.



normocytic RBC 6-8µ diameter

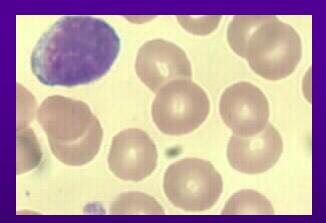


normocytic microcytic macrocytic





Poikilocytosis is a "generic" term used to indicate variation in shape of erythrocytes (e.g., oval, pearshaped, teardrop-shaped, saddle-shaped, helmetshaped, sickle-shaped, and irregularly shaped), eg:







RBC variable shapes



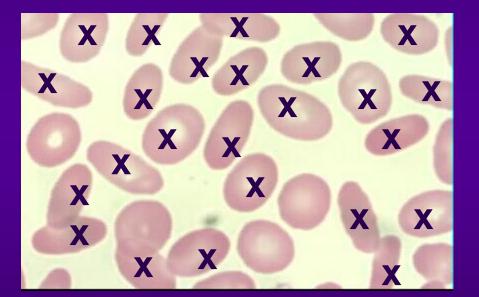


Elliptocytes and Ovalocytes

are interchangeable terms used to indicate ovalshaped erythrocytes.



normocytic RBC round biconcave



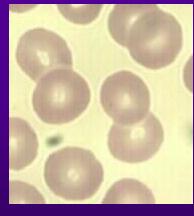
RBC predominantly ovalocytes



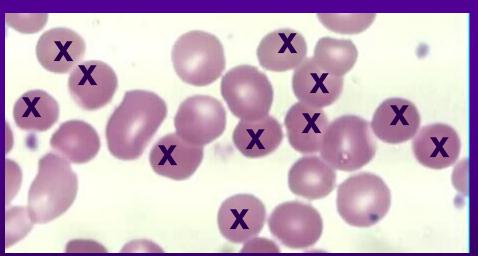


Spherocytes are nearly spherical erythrocytes which usually have a diameter smaller than normal. They lack the central pale area due to their **spherical** shape.

round, biconcave RBC



normocytic



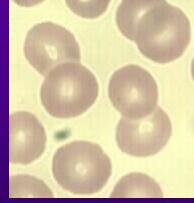
spherocytes



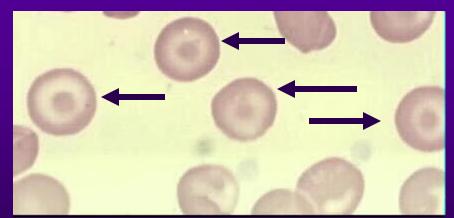


Target cells (leptocytes) are erythrocytes that are thinner than normal which show a peripheral rim of hemoglobin with a dark central hemoglobin-containing area. A pale unstained ring containing less hemoglobin separates the central and peripheral zones and gives the cell a target appearance.

round, biconcave RBC



normocytic



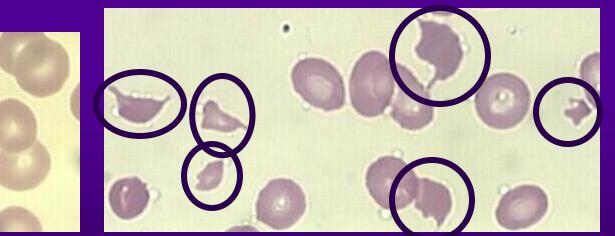






Schistocytes are fragmented red cell segments that are the result of some hemolytic process. The segments can be a variety of shapes but helmet cells and triangularly-shaped cells are particularly characteristic.

round, biconcave RBC



normocytic

schistocytes

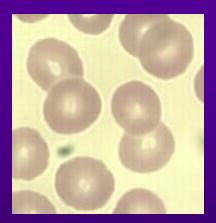




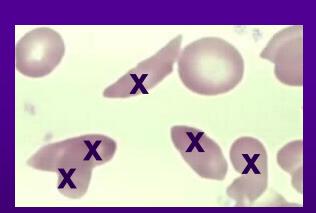
CSW

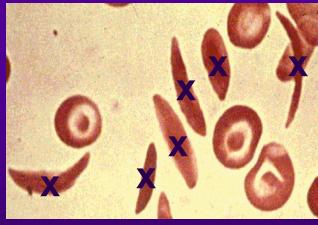
lsuhsc _2002 /

Sickle cells (drepanocytes, meniscocytes) are interchangeable terms used to indicate sickle-like forms of erythrocytes (crescent-shaped, irregular spines, filaments, holly-leaf appearance) noted when RBC containing HbS are subjected to reduction in oxygen tension or pH.



normocytic: round, biconcave RBC





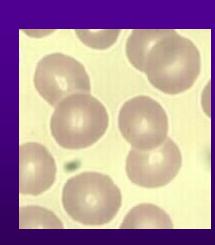
sickle cells

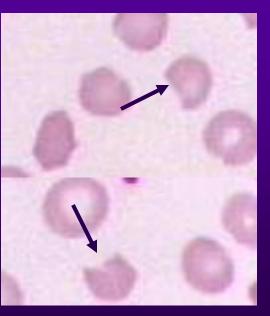




Keratocytes or "Bitocytes" interchangeable terms used to indicate irregularly contracted erythrocytes which stain densely and have contraction of hemoglobin from a part of the cell membrane, thereby giving the appearance that a "bite" has been taken out of the cell. These cells are thought to be cells from which Heinz bodies have been removed by the spleen.

normocytic RBC, round, biconcave





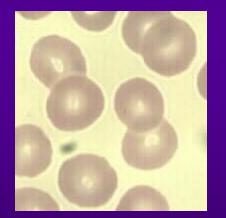
keratocytes



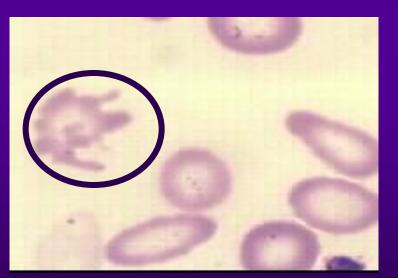




Acanthocytes are irregularly shaped red cells with spiny or thorny projections and dark centers which may be found in severe liver disease, infantile pyknocytosis (with underlying hemolytic process), abetalipoproteinemia, or anorexia nervosa.



normocytic RBC round biconcave



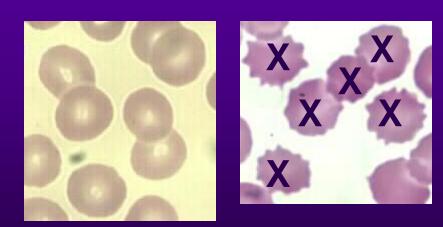
acanthocyte





Crenated red blood cells are uniformly shrunken red cells with uniform irregular, wrinkled cell membranes. Their presence is frequently an artifact of storage and all red cells in the field are usually affected. (By contrast, ecinocytes are intermixed with normal red cells.)

normocytic RBC round biconcave

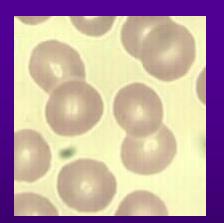


crenated RBC

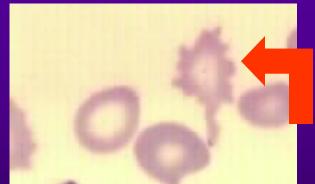




Echinocytes are irregularly shaped red cells with spiny projections and preserved central pallor. While their presence may be an artifactual phenomenon, they may be seen in liver and renal disease, hyperlipidemia, and red blood cell enzymopathies.



normocytic RBC round biconcave



echinocyte



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Definitions:

Rouleaux formation describes an aggregation of erythrocytes that are aligned one upon the other, resembling stacks of coins, caused by elevated plasma fibrinogen or globulins. This phenomenon causes an increased erythrocyte sedimentation rate. This finding is especially characteristic of paraproteinemia (monoclonal gammopathy).

normal





rouleau



Agglutination of red cells is caused by agglutinins and resembles rouleaux but is more irregular with round clumps rather than linear rouleaux.



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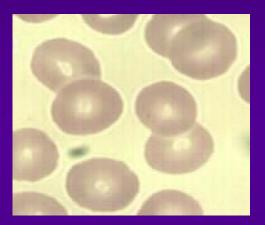




normal

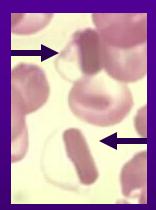


Hemoglobin C crystals are hexagonal crystals that may be found in individuals with HbC syndromes. The crystals may be intracellular or extracellular.



Normocytic RBC (round, biconcave, without inclusions)

intracellular HbC crystals



extracellular HbC crystal

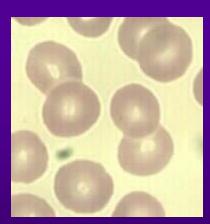






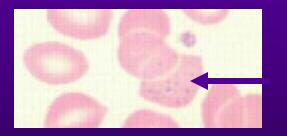


Basophilic stippling is the term used to indicate the presence of irregular basophilic granules in the cytoplasm of erythrocytes. The granules are composed of unstable RNA and may be fine or coarse.

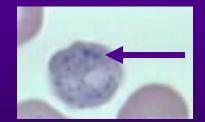


normocytic RBC round biconcave

basophilic stippling (fine)



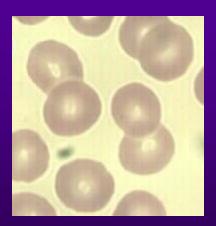
basophilic stippling (coarse)





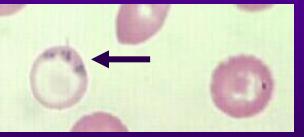


Pappenheimer bodies are intracellular inorganic iron-containing granules that may be observed on Wright's stained peripheral blood smears in ironloading disorders. When the inclusion bodies are demonstrated by stains for iron (e.g., Prussian Blue), the cells are called siderocytes.



normocytic

Pappenheimer bodies



(Wright stain)

siderocytes



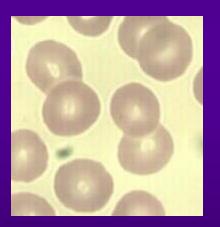
Prussian Blue stain (Prussian Blue stain)





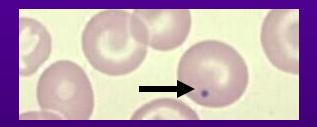


Howell-Jolly bodies are intracellular particles which are smooth, round remnants of nuculear chromatin (DNA). Usually, only one per cell is seen but, occasionally, there may be more than one.



normocytic

Howell-Jolly body (single)



Howell-Jolly body (multiple)



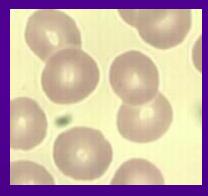






Nucleated red blood cells (NRBC)

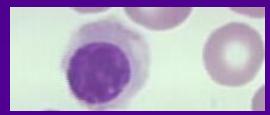
are precursors of the non-nucleated mature red cells, usually orthochromatic erythroblasts when noted in peripheral blood in disease states but earlier forms may also be seen,eg:



mature **RBC**



orthochromatic erythroblast



polychromatophilic erythroblast



proerythroblast

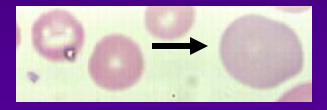


basophilic erythroblast



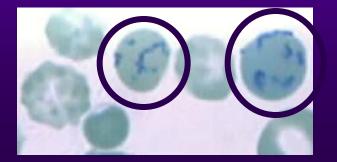


Reticulocytes are anucleated slightly immature erythrocytes, identified as polychromatophilic erythrocytes on Wright stained smears.



polychromatophilic erythrocyte (Wright's stain)

The cells are identified as reticulocytes only after exposure to a supravital stain which causes the cytoplasmic organelles of the cells to clump into an easily recognized blue-staining reticulum.



reticulocyte (supravital stain)

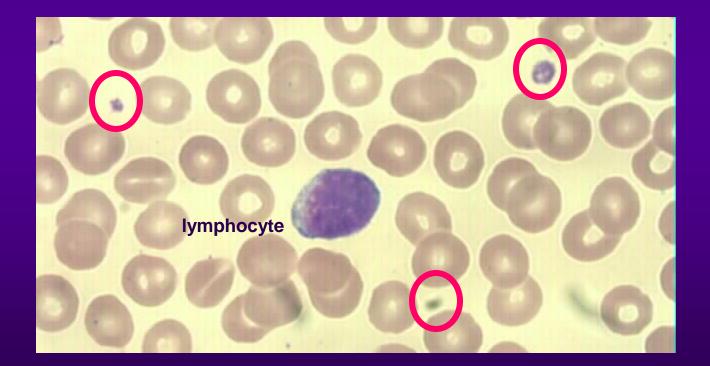








Normally, platelets are round or oval, 2 - 4 μ diameter, contain small fine granules that usually fill the cytoplasm, and are separated from one another.









Estimated Platelet Count

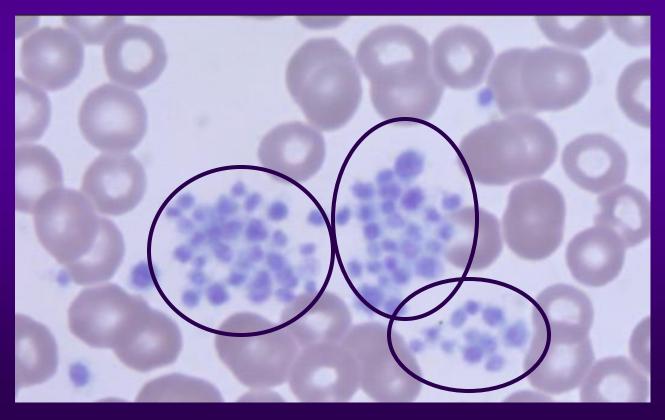
An estimated platelet count can be made on a peripheral blood smear. If the platelet count is normal, an average of about one platelet per 10 to 30 red blood cells. Using the oil immersion lens at 1000x magnification, that is about 5 to 25 platelets per field.

In Clinical Pathology 201, < 5 platelets/oil immersion field will be considered decreased and > 25 platelets/oil immersion field will be considered increased.





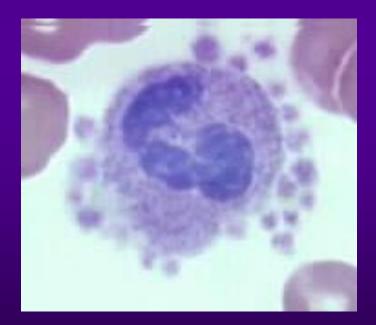
Platelet clumps may be found on blood smears that have been improperly prepared. They may also be seen in clotting disorders. Platelet estimates cannot be made from blood smears with platelet clumps.







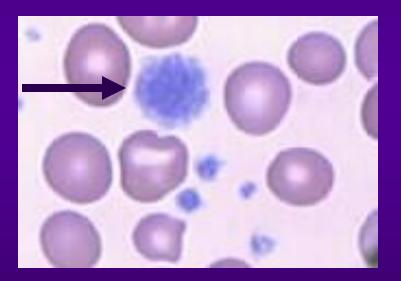
Occasionally, platelet satellites may also be seen. Platelets adhere to the outer surface of neutrophils. When platelet satellites are present, platelet estimates cannot be made from blood smears.







In some disorders, platelets may be larger than normal (i.e., giant platelets).







Disorders Characteristic Morphology



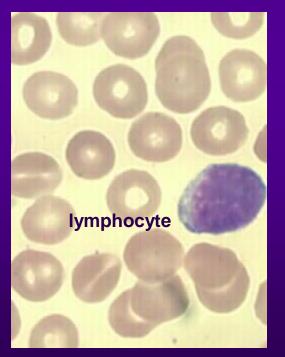


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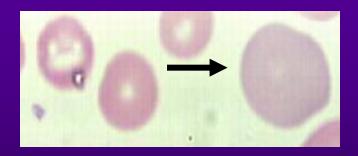
Macrocytic Non-Megaloblastic Anemias

Characteristic abnormalities associated with macrocytic non-megaloblastic anemias in diseases associated with reticulocytosis.

normal



numerous polychromatophilic erythrocytes like the one indicated by the arrow



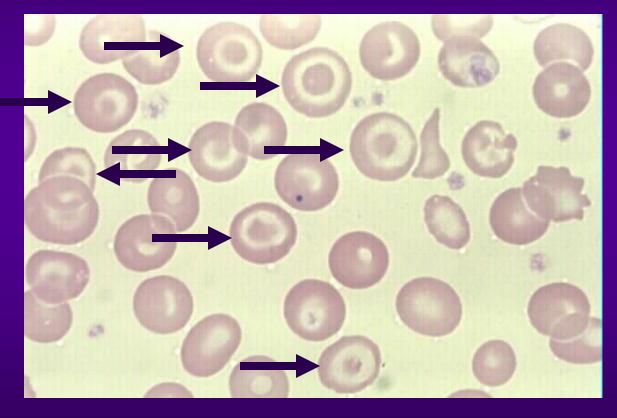




Macrocytic Non-megaloblastic Anemia

Characteristic abnormalities associated with macrocytic non-megaloblastic anemia in liver disease.

macrocytes and target cells



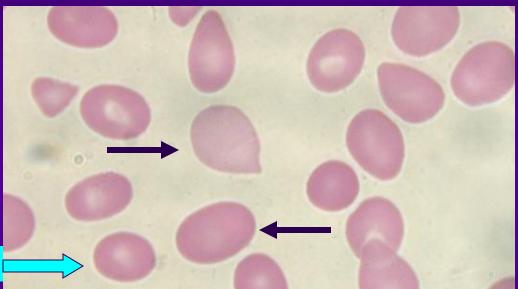




Macrocytic Megaloblastic Anemias

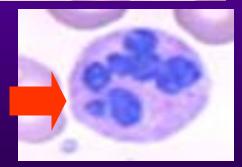
Examples of characteristic abnormalities associated with megaloblastic macrocytic anemias

macrocytes and macro-tear drops and macro-ovalocytes



Normocytic RBC





and erythrocytic precursors with asynchrony in nuclear and cytyplasm maturation

and hypersegmented PMN



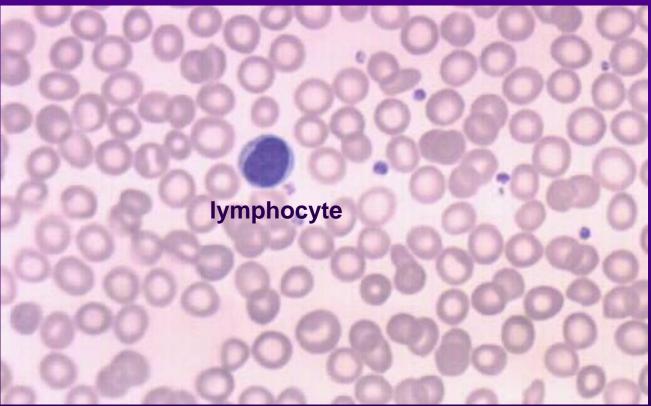


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Microcytic Hypochromic Anemias

Characteristic abnormalities associated with microcytic hypochromic anemias (eg, iron deficiency, chronic disease).

predominant cells are microcytic hypochromic erythrocytes







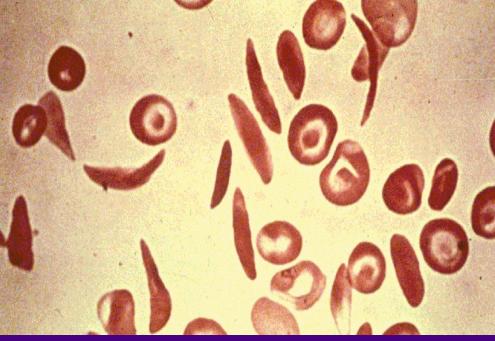
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Anemia in Sickle Cell Disease

Characteristic abnormalities associated with anemia in sickle cell disease

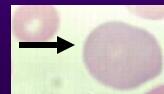
Sickled cells (may be crescent-shaped, irregular spines, filaments, holly-leaf appearance)

and target cells

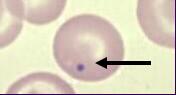


and also

NRBC polychromasia







basophilic stippling



Pappenheimer bodies







