

Top Ten RBC Morphologic Changes Helpful in Classifying Anemia

Dennis B. DeNicola, DVM, PhD, Diplomate ACVP Chief Veterinary Educator IDEXX Laboratories

Erythron – Top Ten List

Erythron – "Top Ten List"



Normal Erythrocyte Morphology

- Features similar in dogs, cats, horses and ruminants
- All lack nuclei
- Stain reddish to reddish-orange
- Generally biconcave discoidshaped cells
- Major differences are in size and central pallor







Normal Erythrocyte Morphology

- Features similar in dogs, cats, horses and ruminants
- All lack nuclei
- Stain reddish to reddish-orange
- Generally biconcave discoidshaped cells
- Major differences are in size and central pallor







Erythron – Top Ten List

Erythron – "Top Ten List"

1. Polychromasia



Classification of Anemia



Absolute Reticulocyte count most objective method of differentiating regenerative from non-regenerative anemia

Polychromasia correlates with Reticulocyte count



Polychromasia

- Variation in color among cells
- Bluish color in erythrocytes (RNA)
- Cells generally larger than mature RBCs
- Represent <u>reticulocytes</u>
- Few are normal in dogs and cats
- Good morphologic evidence of a bone marrow response to a peripheral demand





Morphologic Features of Regeneration

- Polychromasia
- Potential findings only if associated with polychromasia
 - Anisocytosis
 - Target cell formation
 - Howell-Jolly bodies
 - Basophilic stippling
 - Nucleated red blood cells



Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis



Spherocytosis

- Appear smaller than normal mature erythrocyte
- More dense staining than normal mature erythrocyte
- No central zone of palor
- As name suggests, they have lost their normal biconcave disc shape and are "spherical"
- Supportive of extravascular immune mediated destruction
 - Partial phagocytosis





Immune Mediated Hemolytic Anemia





Immune Mediated Hemolytic Anemia





Immune Mediated Hemolytic Anemia





Spherocytosis







Spherocytosis





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination



Autoagglutination





Agglutination

- Three dimensional clumps of erythrocytes
 - Unorganized in contrast to "stack of coins" as with Rouleaux
- Relatively strong binding between cells because of cross-linking of antibodies on surface
- Supportive of an immunemediated mechanism





Agglutination



Agglutination







Saline Agglutination Test



Autoagglutination





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis



Acanthocytosis

- Specific type of poikilocyte
- Spherical shape with 2-10 surface projections
- Projections are:
 - Blunt
 - Variably sized
 - "Finger-like"
- Cholesterol:phospholipid ratio in RBC membrane is increased
- Supportive of potential liver, splenic or metabolic disorder





Metabolic Disorders

• RBCs have phospholipid bilayer





Metabolic Disorders

- RBCs have phospholipid bilayer
- Insertion in either the outer or inner leaflets can cause shape changes in RBC





Acanthocytosis

- Specific type of poikilocyte
- Spherical shape with 2-10 surface projections
- Projections are:
 - Blunt
 - Variably sized
 - "Finger-like"
- Cholesterol:phospholipid ratio in RBC membrane is increased
- Supportive of potential liver, splenic or metabolic disorder





Acanthocytes





Acanthocytes





Acanthocytes – Feathered Edge





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis



Target Cells - Leptocytosis

- "Floppy" cells with excess cell membrane compared to normal erythrocyte
- When associated with "regenerative" blood picture
 - Typical for immature erythrocytes
- When not associated with "regenerative" blood picture
 - Indicative of "lipid loading" potentiall associated with liver, splenic or metabolic disorder





Target Cells







Target Cells - Leptocytosis

- "Floppy" cells with excess cell membrane compared to normal erythrocyte
- When associated with "regenerative" blood picture
 - Typical for immature erythrocytes
- When not associated with "regenerative" blood picture
 - Indicative of "lipid loading" potentially associated with liver, splenic or metabolic disorder





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis
- 6. Hypochromasia



Hypochromasia

- Pale staining cells compared to normal
- Increased zone of central palor
- Supportive of decreased mean cellular hemoglobin content
- In veterinary medicine primarily suggests chronic blood loss





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis
- 6. Hypochromasia
- 7. Schistocytosis



Schistocytosis

- Irregularly shaped erythrocyte fragments
- Indicate mechanical injury to erythrocytes
- Damage often in small blood vascular system
 - "microangiopathy"
- Could prove helpful in recognizing underlying Disseminated Intravascular Coagulopathy (DIC)





Schistocytes







Schistocytes





Schistocytosis

- Irregularly shaped erythrocyte fragments
- Indicate mechanical injury to erythrocytes
- Damage often in small blood vascular system
 - "microangiopathy"
- Could prove helpful in recognizing underlying Disseminated Intravascular Coagulopathy (DIC)





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis
- 6. Hypochromasia
- 7. Schistocytosis
- 8. Metarubricytosis



Metarubricytosis

- Metarubricytes are late stage nucleated red blood cells
- Typically restricted from movement into the blood
- Potentially present during:
 - Marked erythroid hyperplasia in "regenerative" marrow response
 - Marrow stromal damage without marked erythroid hyperplasia
 - Acute lead toxicity
 - Marrow infiltrative disease
 - Septicemia





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis
- 6. Hypochromasia
- 7. Schistocytosis
- 8. Metarubricytosis
- 9. Heinz Body Formation



Heinz Body Formation

- Short blunt projections from surface of cell
- Represent oxydized denatured hemoglobin
 - Onion toxicity dog
 - Acetominophen toxicity cat
 - Phenothiazide horse
- Identification enhanced with New Methylene Blue staining





Heinz Bodies







Heinz Bodies





Heinz Bodies and "Keratocytes"

- Previous "blister cell"
- One or two projections that form from a ruptured vesicle
- Usually seen with Heinz body anemias





Blister Cell





Keratinocyte





Eccentrocyte Formation

Fusion of opposed oxidized RBC membrane

- Potential accumulation of small portions of oxidized hemoglobin between membranes (velcro)
- Leaves a pale area on one side of RBC



Eccentrocyte Formation

Fusion of opposed oxidized RBC membrane

- Potential accumulation of small portions of oxidized hemoglobin between membranes (velcro)
- Leaves a pale area on one side of RBC





Side view

Top view



Heinz Bodies and Eccentrocytes





Erythron – "Top Ten List"

- 1. Polychromasia
- 2. Spherocytosis
- 3. Agglutination
- 4. Acanthocytosis
- 5. Target Cell Formation / Leptocytosis
- 6. Hypochromasia
- 7. Schistocytosis
- 8. Metarubricytosis
- 9. Heinz Body Formation
- **10.** Miscellaneous Inclusion Identification



Erythron – Top Ten List

Miscellaneous Inclusions



Classification of Anemia





Questions?





