

Evaluation of Anemia in Hospitalized Patients

Andrew Herber, PA-C Assistant Professor of Medicine Mayo College of Medicine Division of Hospital Medicine

No disclosures

Objectives

- Recognize common laboratory findings associated with different types of anemia
- Describe workup of anemia
- Review evidence-based guidelines on transfusions
- Describe impact of hospital acquired anemia on patient outcomes

First a refresher on the basics...

Definition

According to the World Health Organization anemia is defined as a hemoglobin level of less than 13 g/dL in men and less than 12 g/dL in women.

Symptom not a disease

HEMOGLOBIN

13.5g/dl-16.0g/dl

HIGH

-Hypoxia (Smoking, OSA, Lung Dx, High Altitude, CO)
-Dehydration
-Polycythemia Vera (JAK2V617F)
-EPO producing tumors (Liver, Renal, Hemangioblastoma, Pheo, Uterine)

LOW

-Nutritional Deficiency (Iron, B12, Folate) -Blood Loss (Trauma, GI Tract, Hematoma)

- -Hemodilution
- -Hemolysis
- -Renal Failure
- -Chronic Disease

Evaluation

Vital signs Reticulocyte Count MCV A Few Lab Geek Secrets

Vital Signs

Blood Pressure Heart Rate Oxygen saturation Respiratory Rate

RPI > 2.0

Acute Blood Loss Hemolytic Anemia Response to Therapy (5-7days)

But what if RPI <2.0???



Evaluation of anemia in the adult according to the mean corpuscular volume

CBC: complete blood count; MCV: mean corpuscular volume; RBCs: red blood cells; Fe: iron; TIBC: total iron-binding capacity (transferrin); LDH: lactate dehydrogenase

pToDate®

Patients don't always read the books...

CENSUS

Mrs. Salty Mr. Weakness Mister Carwash Mrs. Red Mrs. Organic Mrs. Carrot Mrs. Pale

Mrs. Salty

<u>PMH</u> Coronary Artery Disease

<u>PSH</u>: Cataract Removal

<u>SOCIAL HISTORY:</u> Single. Nonsmoker. No alcohol.

MEDS: Aspirin 325mg daily.

ROS:

Nausea. Vomit x 1. Assisted Living Traveling Cribbage Team all have the "GI Bug"

****Vomited x1 this morning. "Kinda dark colored" Came to ER. ****

Lab	Physical 1 week ago	ER Labs
Hemoglobin	13.3	6.8
Platelets	296	151
Sodium	138	146
Chloride	100	119
Potassium	4.8	3.1
Creatinine	1.0	0.9
BUN	20	16

********Vital signs stable. No current complaints****

"Saltines, Sierra Mist, and 2units of PRBCs and sending her up"

Lab	Physical 3 months ago	ER Lab	Floor Lab
Hemoglobin	13.3	6.1	14.6
Platelets	296	171	111
Sodium	138	146	140
Chloride	100	119	101
Potassium	4.8	3.1	5.1
Creatinine	1.0	0.9	1.1
BUN	20	16	21

Most important next step?

- A. Check Reticulocytes
- B. Octreotide infusion
- C. EGD
- D. Obtain H. Pylori Serology
- E. Fire the phlebotomist

Drip Arm



Speaking of IV fluids and anemia lets check on our next patient...

Mr. Weakness

<u>PMH</u>

PSH: None.

Hypertension Chronic Kidney Disease Osteoarthritis

SOCIAL HISTORY:

Married. Neversmoker. No ETOH.

MEDS:

Metoprolol, ASA, Simvastatin

ROS:

Progressive weakness, GERD, Joint Pain

LABS

Lab	Admission	Day 2	Day 3
Hemoglobin	9.8 (Baseline 10)	8.4	7.8
MCV	87	88	91
Platelets	206	259	214
Sodium	141	138	140
Potassium	4.8	5.2	5.2
Bicarbonate	20	21	23
Creatinine	1.4 (Baseline 1.5)	1.5	1.5
BUN	28	35	48
AST	52	-	-
ALT	55	-	-
UA	Negative	-	-
TSH	1.8	-	-

Your review of vitals...

Normal saline running 100cc/hr since admission Weight is up 3kg Fluid balance is +2.7L

LABS

Lab	Admission	Day 2	Day 3
Hemoglobin	9.8 (Baseline 10)	8.4	7.8
MCV	87	88	91
Platelets	206	259	214
Sodium	141	138	140
Potassium	4.8	5.2	5.2
Bicarbonate	20	21	23
Creatinine	1.4 (Baseline 1.5)	1.5	1.5
BUN	28	35	48
AST	52	-	-
ALT	55	-	-
UA	Negative	-	-
TSH	1.8	-	-

Now what?

- A. EGD
- B. Colonoscopy
- C. Peripheral Smear
- D. CT Abdomen
- E. Stop IVF and give Furosemide
- F. Transfer to SNF for continued PT/OT

BLOOD UREA NITROGEN 6-21mmol/L

AZOTEMIA

-Dehydration

-Rapid protein catabolism

-CHF

-Shock

-MI

-High protein diet

-Anabolic effect of systemic corticosteroids

LOW

-Liver failure

-Malnutrition

-Nephrotic syndrome

EGD = Duodenal Ulcer Hgb 7.8. What next?

- A. Transfuse 1 unit of PRBCs
- B. Transfuse 2 units of PRBCs
- C. Iron supplementation
- D. Monitor closely
- E. Toradol, Dexamethasone, and a Bang[™]

Caffeine in Bang™?

- A. 50mg
- B. 100mg
- C. 200mg
- D. 300mg
- E. Wait...what's Bang? Red Bull is my jam!

Transfuse?

Transfusion Strategies for Acute Upper Gastrointestinal Bleeding

Villanueva, MD et. al. The New England Journal of Medicine, 2013

Liberal or Restrictive Transfusion in High Risk Patients after Hip Surgery Jeffrey Carson, MD et al The New England Journal of Medicine, 2011

Lower versus Higher Hemoglobin Threshold for Transfusion in Septic Shock The New England Journal of Medicine , 2014

Transfuse?

Impact of More Restrictive Blood Transfusion Strategies on Clinical Outcomes: A Meta-analysis of Systematic Review Salpeter, MD et al The American Journal of Medicine

Restrictive versus liberal transfusion strategy for red blood cell transfusions: systematic review of randomized trials with meta-analysis and trial sequence analysis

Holst, et al BMJ 2015

Outcomes Using Lower vs Higher Hemoglobin Thresholds for Red Blood Cell Transfusion

Carson, MD et al Journal of American Medical Association 2013

Red Blood Cell Transfusion: A Clinical Practice Guideline From the AABB Carson, MD et al Annuals of Internal Medicine, 2012

Coronary Artery Disease?

Liberal versus restrictive transfusion thresholds for patients with symptomatic coronary artery disease *American Heart Journal, 2013*

Say you did transfuse...

Which of the following theoretically would not increase after transfusion?

- A. Potassium
- B. Bilirubin
- C. White Blood Cells
- D. Platelets
- E. Risk of TRALI/TACO

What about this next guy though?

Mr. Carwash

PMH None.

None.

PSH:

SOCIAL HISTORY:

Single. Positive THC and ETOH use on weekends only.

MEDS:

None.

ROS:

Fell asleep at work.

LABS

Hgb	6.8
Creatinine	6.8
Kidney Bx	FSGS
Transfuse?

- A. Yes
- B. No

Moving along...

Mrs. Red

PMH None.

PSH: None.

SOCIAL HISTORY:

Married. Beet farmer. Neversmoker. No alcohol.

MEDS:

Aspirin 325mg daily.

ROS:

Tired. Red urine. Weight loss.

LABS

Lab	Result
Hemoglobin	8.4
MCV	74
WBC	7000
Platelets	116,000
Sodium	136
Potassium	4.6
Glucose	133
Bicarbonate	19
Creatinine	0.9
BUN	19

Next test?

- A. Iron studies
- B. INR
- C. UA
- D. All of the above

Lab	Result
Hemoglobin	8.4
MCV	74
WBC	7000
Platelets	116,000
Sodium	136
Potassium	4.6
Glucose	133
Creatinine	0.9
BUN	19
UA	Negative
Iron	24
TIBC	578
Ferritin	9
INR	1.5

What next?

- A. Hold Aspirin. Give FFP and Vitamin K
- B. Urology consult for Cystoscopy
- C. Colonoscopy
- D. Hemoglobin electrophoresis
- E. Ferrous Sulfate BID. Recheck 1 month
- F. Avoid beets. Recheck 1 month

Mrs. Red

Red urine = Beeturia

Elevated Protime and Thrombocytopenia = Chronic DIC

Colonoscopy = Colon Cancer

Deferred on further treatment.

Discharged home.

What else causes microcytic anemia?

Microcytic Anemia

* Iron Deficiency
* Thalassemia
* Chronic Disease
* Lead Poisoning
* Sideroblastic Anemia
* Aluminum Toxicity
* Copper Deficiency
* Zinc Poisoning

Work Up:

*Serum Iron *TIBC *Ferritin *RDW *Peripheral Smear

LAB	Iron Deficiency Anemia	Anemia of Chronic Disease	
Iron			
TIBC			
Ferritin			

Cause of IDA?

Acute Blood Loss Decreased dietary intake Impaired absorption Increased Requirements

Our next patient awaits...

Mrs. Carrot

<u>PMH</u>

St. Jude Aortic Valve OSA Atrial Fibrillation HTN **<u>PSH</u>**: Right Total Hip Arthroplasty Aortic Valve Replacement

SOCIAL HISTORY:

Married. Neversmoker. No ETOH.

MEDS:

Warfarin, Metoprolol, HCTZ, and Melatonin

<u>ROS:</u>

Admitted from ER for weakness and dyspnea

LABS

Lab	On discharge from Cardiac Surgery	Admit Labs
Hemoglobin	10.7	7.7
MCV	87	88
Platelets	206	259
Sodium	141	138
Potassium	4.8	5.9
Bicarbonate	25	28
Creatinine	1.0	0.9
BUN	20	21
AST	79	251
ALT	86	
Bilirubin	1.2	3.8
INR	2.6	3.3

Now What?

- A. Abdominal CT
- **B. Call GI Bleed Team**
- C. FFP and Vitamin K
- **D. Peripheral Smear**
- E. Right Upper Quadrant US
- F. Plasma Potassium

Lab	Discharge from CV Surgery	Admit Labs
Hemoglobin	10.7	7.7
MCV	87	88
Platelets	206	259
Sodium	141	138
Potassium	4.8	5.9
Bicarbonate	25	28
Creatinine	1.0	0.9
BUN	20	21
AST	79	251
ALT	86	
Bilirubin	1.2	3.8
INR	2.6	3.3
Haptoglobin		3
LDH		980
Peripheral Smear		Schistocytes, Helmet Cells

Hemolysis

Down

Up

- Potassium
- AST
- LDH
- Bilirubin (Indirect)
- Reticulocytes

- Hemoglobin
- Haptoglobin

BILIRUBIN

- 0.1-1.0mg/dL -Choledocholithiasis (most common)
- -Liver Disease
- -Hemolysis (Indirect)
- -Recent transfusion
- -Gram Negative Sepsis
- -TPN
- -Obstruction (Tumor, Mass, Stone)
- -Gilberts Disease

LDH 122-222U/L

-Heart Disease (MI)

-Tissue Infarction (Renal, Pulmonary)

-Hemolysis

- -Liver Disease (Hepatitis, Cirrhosis, Cholangitis)
- -Malignancy (Lymphoma, Myeloma, Leukemia)

Present in liver, heart, kidney, RBC, WBC, Lungs, Platelets, skeletal muscle, prostate *Any cellular damage causes elevation*

Haptoglobin 30-200mg/dL

Increased:

- -Inflammation
- -Infection
- -Malignancy
- -Surgery
- -Trauma
- -Corticosteroids

Decreased:

- -Hemolysis
- -Liver disease
- -Malnutrition
- -Estrogens
- -Pregnancy

Reticulocytes 0.5-2.0%

Elevated:

Hemolytic Anemia Acute Blood Loss Response to Therapy (5-7days)

Low or Normal:

All other forms of anemia

Coombs?

- A. Positive
- B. Negative
- C. Dude... Come on!

Coombs

Immune vs. Nonimmune?

(+) Alloimmune, Autoimmune, Drug Induced (-) HS, G6PD, PNH, HUS, DIC, Mechanical, Infection

Moving along...

Mrs. Organic PSH: None.

PMH None.

SOCIAL HISTORY:

Married to a Hospitalist PA. 3 boys. (6 month old twins and 5 year old). Nonsmoker. No recent alcohol use because of nursing.

MEDS:

None.

ROS:

Dyspnea and fatigue.

LABS

Lab	
Hemoglobin	10.7
MCV	115
Platelets	206
Sodium	141
Potassium	4.6
Creatinine	1.0
BUN	20
AST	42
ALT	39
Bilirubin	1.2
TSH	1.0

What should we order for her?

- A. B12 and Folate
- B. Peripheral smear
- C. Iron studies
- D. Diapers and a New Minivan

LABS

Lab	
Hemoglobin	10.7
MCV	115
Platelets	206
Sodium	141
Potassium	4.6
Creatinine	1.0
BUN	20
AST	42
ALT	39
Bilirubin	1.2
TSH	1.0
B12	168
Folate	8

What did she eat for dinner last night?

- A. Steak
- B. Tofu
- C. Chicken
- D. Sushi
- E. White Castle Sliders (Organic ones of course)

Macrocytosis

- * **B12 Deficiency** (Pernicious Anemia, Surgical Resection of ileum, sprue, fish tapeworm, bacterial overgrowth, vegans)
- * Folate Deficiency (ETOH, Pregnancy, Medications)
- * Hypothyroidism
- * Drugs (AZT, MTX, Hydroxyurea, Bactrim, Valacyclovir, Triamterene, Phenytoin)
- * Liver disease
- * Myelodysplastic Syndromes
- * Reticulocytosis

B12 and Folate Pearls

- Higher the MCV, more likely the etiology
- Folate heavily influenced by diet *Fasting*
- RBC Folate?
- MMA and Homocysteine
- Low folate can falsely lower B12 (33% of time)
- Hypersegmented neutrophil
- Intrinsic Factor Antibody (70% Pernicious)

Last patient awaits...

Mrs. Pale

PMH Osteoporosis PSH: None.

SOCIAL HISTORY:

Married. Nonsmoker. Retired. Likes to knit hats.

MEDS:

Calcium and Vitamin D

ROS:

Frequent falls.

LABS

Lab	Admission	Day 2	Day 3	Day 4	Day 5
Hgb	13.0	12.3	11.7	11.1	10.4
MCV	87	88	91	91	91
Platelets	206	259	214	200	205
Sodium	141	138	140	138	142
Potassium	4.8	4.7	4.8	4.3	4.6
Creatinine	1.0	0.9	1.1	1.0	0.9
BUN	19	20	20	18	20
AST	52	50	51	58	54
ALT	55	48	44	46	49
Calcium	9.0	10.1	9.8	9.5	9.4
Albumin	4.0	3.9	4.2	4.4	3.8
Next best treatment?

- A. Stop drawing her blood
- B. Stop drawing her blood
- C. Stop drawing her blood
- D. Stop drawing her blood



Roughly, how much blood does your marrow make in one day?

- A. 10ml
- B. 50ml
- C. 100ml
- D. 350ml

Roughly, how much blood does it take to run CBC, CMP, and INR?

- A. 1ml
- B. 5ml
- C. 10ml
- D. 30ml

Hospital Acquired Anemia

Do Blood Tests Cause Anemia in Hospitalized Patients? Paaladinesh Thavendiranathan, MD, et al J GEN INTERN MED 2005; 20:520–524.

Hospital-Acquired Anemia: Prevalence, Outcomes, and Healthcare Implications

Colleen G. Koch et al Journal of Hospital Medicine September 2013 Volume 8

Hospital Acquired Anemia and in-hospital mortality in patients with acute myocardial infarction Salisbury et al

American Heart Journal, 2011

Causes

Procedural Blood Loss Recurrent Phlebotomy Impaired Erythropoiesis Hemodilution

Effects

Increased in hospital mortality Increased length of stay Increased hospital charges

What can you do?

Daily labs? Do you really need them? Microdraws Stored serum

"Lets Run The List"

Mrs. Salty Mr. Weakness Mister Carwash Mrs. Red Mrs. Carrot Mrs. Organic Mrs. Pale

References:

Do Blood Tests Cause Anemia in Hospitalized Patients? Paaladinesh Thavendiranathan, MD, et al J GEN INTERN MED 2005; 20:520–524.

Hospital-Acquired Anemia: Prevalence, Outcomes, and Healthcare Implications Colleen G. Koch et al Journal of Hospital Medicine September 2013 Volume 8

Hospital Acquired Anemia and in-hospital mortality in patients with acute myocardial infarction Salisbury et al American Heart Journal, 2011

Impact of More Restrictive Blood Transfusion Strategies on Clinical Outcomes: A Meta-analysis of Systematic Review Salpeter, MD et al The American Journal of Medicine

Restrictive versus liberal transfusion strategy for red blood cell transfusions: systematic review of randomized trials with meta-analysis and trial sequence analysis Holst, et al BMJ 2015

> Outcomes Using Lower vs Higher Hemoglobin Thresholds for Red Blood Cell Transfusion Carson, MD et al Journal of American Medical Association 2013

Red Blood Cell Transfusion: A Clinical Practice Guideline From the AABB Carson, MD et al Annuals of Internal Medicine, 2012

Questions?

Herber.Andrew@mayo.edu