Anemias

LECTURE IN INTERNAL MEDICINE FOR IV COURSE STUDENTS

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Plan of the lecture

- Definition
- Epidemiology
- Etiology
- Mechanisms
- Adaptation to anaemia
- Classification
- Clinical investigation
- Diagnosis
- Treatment
- Prognosis
- Prophylaxis
- Abbreviations
- Diagnostic guidelines

Definition

Anemia is a disease and/or a clinical syndrome that consist in lowered ability of the blood to carry oxygen (hypoxia) due to decrease quantity and functional capacity and/or structural disturbances of red blood cells (RBCs) or decrease hemoglobin concentration or hematocrit in the blood.

A severe form of anemia, in which the hematocrit is below 10%, is called the hyperanemia.

WHO criteria is $Hb < 13 \text{ g/dL in men and } Hb < 12 \text{ g/dL in women}$ (revised criteria for patients with malignancy $Hb < 14 \text{ g/dL in men and } Hb < 12\text{g/dL in women}$)
Epidemiology 1

Worldwide Prevalence of Anemia, by severity

Severity of Anemia
- Normal (<5.0%)
- Mild (5.0-19.9%)
- Moderate (20.0-39.9%)
- Severe (≥40.0%)
- No data

https://www.k4health.org/sites/default/files/anemia-map_updated.png
Epidemiology 2

[Bar chart showing the percentage of males and females with anemia by age group.]

Percent Who Have Anemia (%) by Age Group (years):
- 1-16: Male 6.0%, Female 1.5%
- 17-49: Male 8.7%, Female 4.4%
- 50-64: Male 12.2%, Female 6.8%
- 65-74: Male 7.8%, Female 8.5%
- 75-84: Male 15.7%, Female 10.3%
- 85+: Male 26.1%, Female 20.1%
Epidemiology 3

Prevalence of severity of anemia

- Normal: 140
- Mild: 151
- Moderate: 24
- Severe: 85
Etiology 1 (basic forms)

Basic forms
- Blood loss
- Deficient erythropoiesis
- Excessive hemolysis (RBC destruction)
- Fluid overload (hypervolemia)

https://en.wikipedia.org/wiki/Anemia
http://content.onlinejacc.org/data/JAC/23133/04044_gr2.jpeg
Etiology 2 (blood loss)

• Blood loss can be acute or chronic
• Anemia does not develop until several hours after acute blood loss, when interstitial fluid diffuses into the intravascular space and dilutes the remaining RBC mass
• During the first few hours, however, levels of polymorphonuclear granulocytes, platelets, and, in severe hemorrhage, immature WBCs and normoblasts may rise
• Chronic blood loss results in anemia if loss is more rapid than can be replaced or, more commonly, if accelerated erythropoiesis depletes body iron stores

Etiology 3a (deficient erythropoiesis)

- Deficient erythropoiesis has myriad causes
- Complete cessation of erythropoiesis results in a decline in RBCs of about 7 to 10%/week (1%/day)
- Impaired erythropoiesis, even if not sufficient to decrease the numbers of RBCs, often causes abnormal RBC size and shape

Etiology 3b (deficient erythropoiesis)

Deficient erythropoiesis has myriad causes
Etiology 4a (excessive hemolysis)

- Excessive hemolysis can be caused by intrinsic abnormalities of RBCs or by extrinsic factors, such as the presence of antibodies on their surface, that lead to their early destruction.
- An enlarged spleen sequesters and destroys RBCs more rapidly than normal.
- Some causes of hemolysis deform as well as destroy RBCs.
- Excessive hemolysis does not normally decrease reticulocyte production unless iron or other essential nutrients are depleted.
Etiology 4b (excessive hemolysis)
Etiology 5 (Fluid overload (hypervolemia))

• Fluid overload causes decreased hemoglobin concentration and apparent anemia:
• General causes of hypervolemia include excessive sodium or fluid intake, sodium or water retention and fluid shift into the intravascular space
• Anemia of pregnancy is induced by blood volume expansion

https://en.wikipedia.org/wiki/Anemia
Etiology 6
(cause of anemia)

- Blood loss
  - Acute
  - Chronic
- Deficient erythropoiesis
  - Microcytic
  - Normochromic-normocytic
  - Macrocytic
- Excessive hemolysis due to extrinsic RBC defects
- Reticuloendothelial hyperactivity with splenomegaly
- Immunologic abnormalities
- Mechanical injury
- Excessive hemolysis due to intrinsic RBC defects
- Membrane alterations, acquired
- Membrane alterations, congenital
- Metabolic disorders (inherited enzyme deficiencies)
- Hemoglobinopathies

Mechanisms 1a
(anemia due to blood loss)

• With anemia due to blood loss, a reduction in oxygen-carrying capacity occurs along with a decrease in intravascular volume, with resultant hypoxia and hypovolemia

• Hypovolemia leads to hypotension, which is detected by stretch receptors (in the carotid bulb, aortic arch, heart, and lungs) and transmitted by their impulses along afferent fibers of the vagal and glossopharyngeal nerves to the medulla oblongata, cerebral cortex, and pituitary gland

http://emedicine.medscape.com/article/198475-overview#a4
Mechanisms 1b
(anemia due to blood loss)

A person can have a low hematocrit and not be anemic

http://web2.airmail.net/uthman/anemia/anemia.html
Mechanisms 2a
(anemia due to blood loss)

• In the medulla, sympathetic outflow is enhanced, while parasympathetic activity is diminished

• Increased sympathetic outflow leads to norepinephrine release from sympathetic nerve endings and discharge of epinephrine and norepinephrine from the adrenal medulla

• Sympathetic connection to the hypothalamic nuclei increases antidiuretic hormone (ADH) secretion from the pituitary gland

http://emedicine.medscape.com/article/198475-overview#a4
Mechanisms 2b
(anemia due to blood loss)

Peptic ulcers extend beyond the lamina propria, whereas erosions are superficial.
Mechanisms 3
(anemia due to blood loss)

• Antidiuretic hormone (ADH) increases free water reabsorption in the distal collecting tubules

• In response to decreased renal perfusion, juxtaglomerular cells in the afferent arterioles release renin into the renal circulation, leading to increased angiotensin I, which is converted by angiotensin-converting enzyme (ACE) to angiotensin II

• Angiotensin II has a potent pressor effect on arteriolar smooth muscle and stimulates the zona glomerulosa of the adrenal cortex to produce aldosterone

http://emedicine.medscape.com/article/198475-overview#a4
Mechanisms 4
(anemia due to blood loss)

• Aldosterone increases sodium reabsorption from the proximal tubules of the kidney, thus increasing intravascular volume

• The primary effect of the sympathetic nervous system is to maintain perfusion to the tissues by increasing systemic vascular resistance (SVR)

• The augmented venous tone increases the preload

http://emedicine.medscape.com/article/198475-overview#a4
Mechanisms 5  
(anemia due to blood loss)

• In states of hypovolemic hypoxia, the increased venous tone due to sympathetic discharge is thought to dominate the vasodilator effects of hypoxia

• Counterregulatory hormones (e.g., glucagon, epinephrine, cortisol) are thought to shift intracellular water to the intravascular space, perhaps because of the resultant hyperglycemia
Mechanisms 6a
(anemia due to blood loss)

• Tissue oxygen delivery is the major controlling factor of erythropoiesis through the synthesis and release of erythropoietin (EPO) by the proximal tubular cells or the peritubular interstitial cells in the kidney.

• EPO synthesis is governed by the activation of hypoxia inducible factor-1 (HIF-1), which controls the metabolic responses of multiple gene products to hypoxia.

http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/
Mechanisms 6b
(anemia due to blood loss)

Pathophysiological mechanisms contributing to anemia in patients with Inflammatory Bowel Disease

http://www.haematologica.org/content/95/2/175
Mechanisms 7a
(anemia due to blood loss)

• HIF-1 binds and activates the hypoxia-responsive transcriptional enhancer in the EPO gene regulatory region that upregulates EPO expression

• EPO stimulates erythroid precursor cells, leading to increased proliferation and shortening of their maturation time

• The marrow responds to increased EPO maximally in 4 to 7 days if enough iron is available

http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/
Mechanisms 7b
(anemia due to blood loss)

A. The peripheral blood in severe megaloblastic anemia
B. The bone marrow in severe megaloblastic anemia

Mechanisms 8
(anemia due to acute blood loss)

• Erythropoiesis can be increased by as much as a factor of 8

• Typical of an endocrine loop feedback mechanism, there is an inverse relation between the hemoglobin and EPO levels measured in the blood

• This relation is somewhat distorted in the anemia associated with inflammation or chronic disease, in which there may be a blunted EPO response
Adaptation to anemia

- Modulation of oxygen affinity—largely mediated by an increase in red blood cell 2,3-biphosphoglycerate
- Increased production of erythropoietin—the main growth factor for red blood cell production
- Redistribution of flow to benefit the myocardium, brain, and muscle
- Increase in cardiac output
- Reduction of mixed venous oxygen tension to increase the arteriovenous oxygen difference

http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/
# Classification (morphological)

<table>
<thead>
<tr>
<th>Cell Size</th>
<th>Normal RDW</th>
<th>High RDW</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Microcytosis</strong>&lt;br&gt;MCV&lt;70 µm³</td>
<td>Thalassemia minor, anemia of chronic disease, some hemoglobinopathy traits</td>
<td>Iron deficiency, hemoglobin H disease, some anemia of chronic disease, some thalassemia minor, fragmentation hemolysis</td>
</tr>
<tr>
<td><strong>Normocytosis</strong>:&lt;br&gt;High reticulocyte count</td>
<td>Anemia of chronic disease, hereditary spherocytosis, some hemoglobinopathy traits, acute bleeding</td>
<td>Early or partially treated iron or vitamin deficiency, sickle cell disease</td>
</tr>
<tr>
<td>Low reticulocyte count</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Macrocytosis</strong>&lt;br&gt;MCV&gt;100 µm³</td>
<td>Aplastic anemia, some myelodysplasias</td>
<td>Vitamin B12 or folate deficiency, autoimmune hemolytic anemia, cold agglutinin disease, some myelodysplasias, liver disease, thyroid disease, alcohol</td>
</tr>
</tbody>
</table>

RDW = red cell distribution width; MCV = mean corpuscular volume
## Classification

*(kinetic: red blood cell loss (bleeding) or destruction (hemolysis)*)

### Acquired

<table>
<thead>
<tr>
<th>Mechanical (march hemoglobinuria, artificial heart valves)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Microangiopathic (disseminated intravascular hemolysis, thrombotic thrombocytopenic purpura, vasculitis)</td>
</tr>
<tr>
<td>Parasites and microorganisms (e.g., malaria, bartonellosis, babesiosis, clostridium perfringens)</td>
</tr>
</tbody>
</table>

### Antibody mediated

- Warm-type autoimmune hemolytic anemia
- Cryopathic syndromes (cold agglutinin disease, paroxysmal cold hemoglobinuria, cryoglobulinemia)
- Transfusion reactions

### Hypersplenism

### Red cell membrane disorders

- Spur cell hemolysis
- Acquired acanthocytosis and acquired stomatocytosis

### Chemical injury and complex chemicals (arsenic, copper, chlorate, spider, scorpion, and snake venoms)

### Physical injury (heat, oxygen, radiation)

[http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/#figure1](http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/#figure1)
# Classification

(kinetic: red blood cell loss (bleeding) or destruction (hemolysis))

## Hereditary

<table>
<thead>
<tr>
<th>Hemoglobinopathies (sickle cell disease, unstable hemoglobins)</th>
</tr>
</thead>
</table>

### Red cell membrane disorders

- Cytoskeletal membrane disorders (hereditary spherocytosis, elliptocytosis, pyropoikilocytosis)
- Lipid membrane disorders (hereditary abetalipoproteinemia, hereditary stomatocytosis)
- Membrane disorders associated with abnormalities of erythrocyte antigens (McLeod syndrome, Rh deficiency Syndromes)
- Membrane disorders associated with abnormal transport (hereditary xerocytosis)

### Red cell enzyme defects (pyruvate kinase, 5' nucleotidase, glucose-6-phosphate dehydrogenase deficiencies)

### Porphyrias (congenital erythropoietic and hepatoerythropoietic porphyrias, rarely congenital erythropoietic protoporphyria)

http://www.clevelandclinicmeded.com/medicalpubs/diseasemanagement/hematology-oncology/anemia/#figure1
**Classification**

(kinetic: red blood cell underproduction)

<table>
<thead>
<tr>
<th>Acquired</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Pluripotent stem cell failure</strong></td>
</tr>
<tr>
<td>- Aplastic anemia (radiation induced, drugs and chemicals, viruses, idiopathic)</td>
</tr>
<tr>
<td>- Anemia of leukemia and of myelodysplastic syndromes</td>
</tr>
<tr>
<td>- Anemia associated with marrow infiltration (multiple myeloma, myelofibrosis, carcinoma)</td>
</tr>
<tr>
<td><strong>Erythroid progenitor cell failure</strong></td>
</tr>
<tr>
<td>- Pure red cell aplasia (parvovirus B19 infection, drugs, associated with thymoma, autoantibodies)</td>
</tr>
<tr>
<td>- Endocrine disorders (thyroid, adrenal, pituitary hypofunction)</td>
</tr>
<tr>
<td>- Acquired sideroblastic anemia (drugs, copper deficiency, etc.)</td>
</tr>
<tr>
<td><strong>Functional impairment of erythroid and other progenitors due to nutritional and other causes</strong></td>
</tr>
<tr>
<td>- Nutritional deficiencies-iron, vitamin B$_{12}$, folic acid, pyridoxine</td>
</tr>
<tr>
<td>- Chronic renal disease</td>
</tr>
<tr>
<td>- Anemia of chronic disease and inflammation</td>
</tr>
</tbody>
</table>
**Classification**

*(kinetic: red blood cell underproduction)*

<table>
<thead>
<tr>
<th>Hereditary</th>
</tr>
</thead>
<tbody>
<tr>
<td>Pluripotent stem cell failure (Fanconi, Shwachman and dyskeratosis congenital syndromes)</td>
</tr>
<tr>
<td>Erythroid progenitor cell failure (Diamond-Blackfan, congenital dyserythropoietic syndromes)</td>
</tr>
<tr>
<td>Functional impairment of erythroid and other progenitors due to nutritional and other causes</td>
</tr>
<tr>
<td>- Megaloblastic anemias (Imerslund-Gräsbeck disease, intrinsic factor deficiency, transcobalamin II deficiency)</td>
</tr>
<tr>
<td>- Inborn purine and pyrimidine metabolic defects (Lesch-Nyhan syndrome, hereditary orotic aciduria)</td>
</tr>
<tr>
<td>- Disorders of iron metabolism (atransferrinemia, divalent metal transporter or DMT-1 mutation)</td>
</tr>
<tr>
<td>- Hereditary sideroblastic anemia</td>
</tr>
</tbody>
</table>
Iron-deficiency anemia is the most common form of anemia in the world
Clinical investigation
(symptoms)

- Easy fatigue and loss of energy
- Unusually rapid heart beat, particularly with exercise
- Shortness of breath, particularly with exercise
- Pale skin
- Leg cramps
- Coldness in the hands and feet
- Insomnia
- Light-headedness
- Faintness
- Signs of heart failure
Clinical investigation
(symptoms: anemia caused by iron deficiency)

- A hunger for strange substances such as paper, ice, or dirt (a condition called pica)
- Upward curvature of the nails, referred to as koilonychias
- Soreness of the mouth with cracks at the corners

http://www.webmd.com/a-to-z-guides/understanding-anemia-symptoms
http://www.nhs.uk/conditions/nail-abnormalities/Pages/Introduction.aspx
http://mystuf123.blogspot.com/p/p-i-c.html
Clinical investigation (symptoms: anemia caused by vitamin B12 deficiency)

• A tingling, "pins and needles" sensation in the hands or feet
• Lost sense of touch
• A wobbly gait and difficulty walking
• Clumsiness and stiffness of the arms and legs
• Dementia

Hand stiffness

Clinical investigation
(symptoms: anemia caused by chronic lead poisoning)

• A blue-black line on the gums referred to as a lead line
• Abdominal pain
• Constipation
• Vomiting

A blue-black line on the gums referred to as a lead line

http://www.webmd.com/a-to-z-guides/understanding-anemia-symptoms
http://www.medicaljournals.se/acta/content/?doi=10.2340/00015555-1201&html=1
Clinical investigation
(symptoms: anemia caused by chronic red blood cell destruction)

• Jaundice (yellow skin and eyes)
• Brown or red urine
• Leg ulcers
• Failure to thrive in infancy
• Symptoms of gallstones

Yellow around eyes anemia

Clinical investigation
(symptoms: sickle cell anemia)

• Fatigue
• Susceptibility to infection
• Delayed growth and development in children
• Episodes of severe pain, especially in the joints, abdomen, and limbs

Sickle cell anemia
Clinical investigation
(symptoms: anemia caused by sudden red blood cell destruction)

- Abdominal pain
- Brown or red urine
- Jaundice (yellow skin)
- Small bruises under the skin
- Seizures
- Symptoms of kidney failure

The bruise under the skin

http://www.webmd.com/a-to-z-guides/understanding-anemia-symptoms
http://www.wisegeek.org/what-are-the-most-common-causes-of-green-skin.htm
Clinical investigation
(accents on history 1)

• The duration of anemia can be established by obtaining a history of previous blood examinations and, if necessary, by acquiring those records

• Similarly, a history of rejection as a blood donor or prior prescription of hematinic provides clues that anemia was detected previously

Clinical investigation
(accents on history 2)

• Obtain a family history for anemia, jaundice, cholelithiasis, splenectomy, bleeding disorders, and abnormal Hbs

• Document the patient's occupation, hobbies, prior medical treatment, drugs (including over-the-counter medications and vitamins), and household exposures to potentially noxious agents (insecticides, paints, solvents, hair dyes)
Clinical investigation
(accents on history 3)

• In searching for blood loss, carefully document pregnancies, abortions, and menstrual loss

• Patients do not appreciate the significance of tarry stools, but changes in bowel habits can be useful in uncovering neoplasms of the colon

• Hemorrhoidal blood loss is difficult to quantify, and it may be overlooked or overestimated from one patient to another

Clinical investigation
(accurate on history 4)

• Seek a history of gastrointestinal (GI) complaints that may suggest gastritis, peptic ulcers, hiatal hernias, or diverticula

• Abnormal urine color can occur in renal and hepatic disease and in hemolytic anemia

• A dietary history must include foods that the patient eats and those that he/she avoids, as well as an estimate of their quantity
Clinical investigation
(accents on history 5)

• Changes in body weight are important with regard to dietary intake and can suggest the presence of malabsorption or an underlying wasting disease of infectious, metabolic, or neoplastic origin

• Obtain a history of fever or identify the presence of fever, because infections, neoplasms, and collagen vascular disease can cause anemia
Clinical investigation
(accents on history 6)

• The occurrence of purpura, ecchymoses, and petechiae suggest the occurrence of either thrombocytopenia or other bleeding disorders; this may be an indication either that more than 1 bone marrow lineage is involved or that coagulopathy is a cause of the anemia because of bleeding

• Cold intolerance can be an important symptom of hypothyroidism or lupus erythematosus, paroxysmal cold hemoglobinuria, and certain macroglobulinemias

Clinical investigation (accents on history 7)

- The relation of dark urine to either physical activity or time of day can be important in march hemoglobinuria and paroxysmal nocturnal hemoglobinuria
- Explore the presence or the absence of symptoms suggesting an underlying disease, such as cardiac, hepatic, and renal disease; chronic infection; endocrinopathy; or malignancy
- A geographic history can also be important in establishing an etiology

Clinical investigation
(accents on physical examination 1a)

• The skin and mucous membranes are often bypassed, so that pallor, abnormal pigmentation, icterus, spider nevi, petechiae, purpura, angiomas, ulcerations, palmar erythema, coarseness of hair, puffiness of the face, thinning of the lateral aspects of the eyebrows, nail defects, and a usually prominent venous pattern on the abdominal wall are missed in the rush to examine the heart and the lungs

Clinical investigation
(accurats on physical examination 1b)

Purpura
Spider nevi
Thinning of the lateral aspects of the eyebrows

https://en.wikipedia.org/wiki/Purpura
Clinical investigation
(accents on physical examination 2)

• Examine optic fundi carefully but not at the expense of the conjunctivae and the sclerae, which can show pallor, icterus, splinter hemorrhages, petechiae, comma signs in the conjunctival vessels, or telangiectasia that can be helpful in planning additional studies.

• Perform systematic examination for palpable enlargement of lymph nodes for evidence of infection or neoplasia.
Clinical investigation
(accents on physical examination 3a)

• Bilateral edema is useful in disclosing underlying cardiac, renal, or hepatic disease, whereas unilateral edema may portend lymphatic obstruction due to a malignancy that cannot be observed or palpated

• Carefully search for hepatomegaly and splenomegaly because in patients with chronic disorders, these organs are firm, nontender, and nonnodular, and in patients with carcinoma, they may be hard and nodular

Massive hepatosplenomegaly in a patient with severe malarial anemia due to Plasmodium falciparum infection
Clinical investigation
(accents on physical examination 4)

• A rectal and pelvic examination cannot be neglected, because tumor or infection of these organs can be the cause of anemia

• The neurologic examination should include tests of position sense and vibratory sense, examination of the cranial nerves, and testing for tendon reflexes
Clinical investigation
(accents on physical examination 5)

• The heart should not be ignored, because enlargement may provide evidence of the duration and the severity of the anemia, and murmurs may be the first evidence of infective endocarditis that could explain the etiology of the anemia

Diagnosis

initial assessment

• Patients may present in several ways
• The urgency with which anemia is evaluated depends on the severity at presentation
• Patients with an acute severe hemorrhage present with hypovolaemia and symptoms and signs of the underlying cause
• Many patients with no acute or active bleeding are asymptomatic, and the anaemia is only noted on an full blood count (FBC) taken as part of the assessment of an unrelated condition
• The first step in diagnosis is to identify the type of anemia that is present, using the results of the FBC

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis algorithm for the assessment of anemia

1. Examination of FBC and peripheral bloodsme
   - MCV < 80
     - Microcytic anaemia
       - Serum iron studies
         - Low iron and ferritin with high TIBC
           - Iron deficiency anaemia
             - Mentzer index (MCV/RBC) < 13
               - Thalassaemia
         - Low iron and ferritin with low TIBC
           - Anaemia of chronic disease
     - MCV 80-100
       - Normocytic anaemia
         - Reticulocyte count
           - < 2% (hypoproliferative)
             - Leukaemias
               - Aplastic anaemia
               - Pure red cell aplasia
           - > 2% (hyperproliferative)
             - Haemorrhage
               - Haemolytic anaemias
               - Vitamin B12 and/or Folate deficiency
               - Drug-induced
     - MCV > 100
       - Macrocytic anaemia
         - Megalocytes and segmented neutrophils on peripheral smear
           - Present: megaloblastic
             - Alcohol abuse
               - Myelodysplastic syndrome
               - Liver disease
               - Congenital bone marrow failure syndromes
           - Absent: non-megaloblastic
             - Drug-induced
               - Vitamin B12 and/or Folate deficiency
               - Liver disease
               - Congenital bone marrow failure syndromes

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

microcytic anemia: abnormal serum iron studies 1/4

• A low serum iron, a high total iron-binding capacity (TIBC), and a low ferritin indicate iron deficiency anaemia

• Iron deficiency produces an associated reactive thrombocytosis that provides an additional clue

• Iron deficiency is not a diagnosis and requires further investigation to elucidate the cause

http://bestpractice.bmj.com/best-practice monograph/93/diagnosis.html
Diagnosis

microcytic anemia: abnormal serum iron studies 2/4

- Diets low in meat
- Generalized malabsorption and malnutrition (combined vitamin B12 and/or folate deficiency)
- A history of bleeding (excessive menstrual losses in women, upper and lower GI bleeding, peptic ulcer disease, cirrhosis, idiopathic pulmonary hemosiderosis, excessive blood donation, runner's anaemia, etc.)

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

microcytic anemia: abnormal serum iron studies 3/4

- Signs of iron deficiency (koilonychia, angular cheilosis, glossitis, thinning hair)
- Investigations are guided by the history and examination, and include the fecal occult blood testing, upper GI endoscopy, immunoglobulin A-tissue transglutaminase (IgA-tTG) test (positive in coeliac disease), colonoscopy, flow cytometry (if there is a history of passing red urine), transvaginal ultrasound (causes of menorrhagia)

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
microcytic anemia: abnormal serum iron studies 4/4

• A low serum iron, a low total iron-binding capacity, and a low ferritin suggest anaemia of chronic disease
• A history of an underlying inflammation (infection, neoplasms, autoimmune reactions, and injury to tissue from trauma and surgery) is usually present
• A serum erythropoietin level is usually normal or mildly elevated
• Hypothyroidism and vitamin C deficiency may produce a falsely low ferritin level
Diagnosis
microcytic anemia: normal serum iron studies 1/5

- The most important cause to exclude is thalassemia
- A family history is usually present
- The disease is more common in individuals of Mediterranean, Middle Eastern, or Southeast Asian descent
- The severity ranges from asymptomatic to severe transfusion-dependent symptoms
Diagnosis
microcytic anemia: normal serum iron studies 2/5

• The examination findings may be normal, or reveal splenomegaly, jaundice, abdominal distension, and icterus

• Morphological changes including skeletal abnormalities, a large head, chipmunk facies, and misaligned teeth are seen in beta-thalassemia intermedia and major

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
microcytic anemia: normal serum iron studies 3/5

- Distinct features on the FBC that suggest the diagnosis include a marked decrease in MCV (usually close to 70 femtolitres [fL]) with a low mean corpuscular Hb, target cells on the peripheral smear, and an elevated reticulocyte count (>2%)

- A Mentzer's index (MCV/RBC) <13 is suggestive of thalassemia, and an index >14 suggests iron deficiency

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
microcytic anemia: normal serum iron studies 4/5

• Thalassemia is diagnosed using Hb electrophoresis
• The presence of Hb H, Hb Bart, and concomitant hemoglobinopathies (Hb E, Hb S, Hb C, Hb D) is diagnostic of alpha-thalassemia
• A high HbF with minimal or absent HbA and an elevated HbA2 is diagnostic of beta-thalassemia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

microcytic anemia: between IDA and Thalassemia 5/5

Diagnosis
normocytic anemia: hypoproliferative 1/11

• Normocytic anaemia include disorders that decrease RBC production
• Hematological malignancies and aplastic anaemia are the most important diagnoses to exclude, and are usually associated with multiple cytopenias
• An isolated anemia is usually due to pure red cell aplasia, which may be self-limiting or persistent

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hypoproliferative 2/11

• Chronic renal failure or hypothyroidism can cause an isolated anaemia
• Secondary hyperparathyroidism exacerbates the anaemia of chronic renal failure
• Symptoms of bleeding, easy bruising, night sweats, or weight loss suggest hematological malignancy or aplastic anaemia
Diagnosis

normocytic anemia: hypoproliferative 3/11

- Parvovirus infection, infectious mononucleosis, viral hepatitis, malaria, respiratory infections, gastroenteritis, primary atypical pneumonia, and mumps can result in a self-limiting pure red cell aplasia

- Antiepileptic medications (phenytoin, carbamazepine, valproate sodium), azathioprine, sulfonamides, isoniazid, and procainamide cause pure red cell aplasia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hypoproliferative 4/11

- Benzene, penicillamine, and gold can cause aplastic anaemia
- Chloramphenicol can cause either aplastic anaemia or pure red cell aplasia
- Chemotherapy causes pancytopenia.
- Radiotherapy, especially to pelvic or sternal areas, can cause pancytopenia
- A history of immunosuppression or chronic hepatitis suggests persistent pure red cell aplasia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hypoproliferative 5/11

• There may be a history or features of chronic renal failure or hypothyroidism

• Ecchymoses or petechiae due to thrombocytopenia suggest hematological malignancy, myelodysplastic syndrome, or aplastic anaemia

• Lymphadenopathy or fever suggest malignancy or infections (e.g., infectious mononucleosis)

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 6/11

• Splenomegaly may be seen in hematological malignancies

• Clinical features of systemic lupus erythematosus (SLE), rheumatoid arthritis, dermatomyositis, polyarteritis nodosa, or scleroderma resulting in persistent pure red cell aplasia may be present

• Abnormal lung examination (if lung cancer is the primary cancer) or a breast mass (if breast cancer is the primary) may be present

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 7/11

- A positive Trousseau's sign or Chvostek's sign in patients with chronic renal failure indicates hypocalcemia, probably due to associated secondary hyperparathyroidism
- FBC may show an associated cytopenia and characteristic changes specific to a hematological malignancy
- A pancytopenia suggests aplastic anaemia, or may be due to chemotherapy or radiotherapy

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 8/11

- An isolated anemia suggests pure red cell aplasia or anemia due to chronic renal failure
- Bone marrow aspiration provides a definitive diagnosis of aplastic anemia, acute leukemia, chronic myelogenous leukemia (CML), or bone marrow metastases
- Antiparvovirus antibodies are positive in parvovirus infection, the most common infectious cause of pure red cell aplasia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 9/11

Other tests to consider:

• Hepatitis serology, to exclude an active hepatitis
• Monospot test or Epstein-Barr virus (EBV) IgM, to exclude infectious mononucleosis
• Thick and thin peripheral smear, to exclude malaria if history and findings suggest it
• Thyroid function tests; TSH is elevated and free T4 reduced in hypothyroidism
• Antinuclear antibodies, which are positive in SLE or scleroderma

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 10/11

Other tests to consider:

- Rheumatoid factor, which is positive in rheumatoid arthritis
- Serum CK, which is elevated in dermatomyositis
- Chest x-ray, which may show infiltrates in atypical pneumonia or a smooth mass in thymoma
- Erythropoietin levels, which may be decreased in patients with chronic renal failure
- Serum calcium and parathyroid hormone levels should be considered if associated secondary hyperparathyroidism is suspected

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hypoproliferative 11/11
Diagnosis

normocytic anemia: hyperproliferative 1 /15

• Potential diagnoses include hemolytic anemias (microangiopathic hemolytic anemias, autoimmune hemolytic anemia, drugs, infections, inherited conditions, transfusion reactions, or burns)

• Drugs that can cause hemolysis include penicillin, methyldopa, levodopa, quinidines, cephalosporins, and some NSAIDs

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 2/15

• Cyclosporine, tacrolimus, clopidogrel, oral contraceptive pills, and some chemotherapy drugs may cause hemolytic uremic syndrome

• The triggers of disseminated intravascular coagulation (DIC), that include ongoing severe infection, sepsis, malignancy, obstetric emergency, trauma, burns, envenomation, drug overdose, or any cause of endothelial damage

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 3/15

• The presence of acute-onset neurological symptoms, including headache, confusion, focal weakness, seizures, or coma, should prompt suspicion of thrombotic thrombocytopenic purpura (TTP)

• Female patients may have associated menorrhagia
Diagnosis
normocytic anemia: hyperproliferative 4/15

• Sudden-onset dizziness, headache, mental status changes, loss of sensation or motor strength, chest pain or pressure, dyspnoea, or oedema in a patient with known hypertension should prompt suspicion of malignant hypertension; a history of renal failure or eclampsia may also be present

• An expanding vascular skin lesion in a young infant or child should prompt suspicion of a hemangioma
Diagnosis

normocytic anemia: hyperproliferative 5/15

- A history of prosthetic valve replacement may indicate hemolysis induced by the prosthesis
- Cutaneous burns affecting more than 10% of the body surface area can cause a hemolytic anaemia, or trigger DIC
- Infective causes include cytomegalovirus (CMV), infectious mononucleosis, toxoplasmosis, and leishmaniosis

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hyperproliferative 6/15

• Bloody diarrhoea should prompt suspicion of *Escherichia coli* infection and hemolytic uraemic syndrome

• Patients with inherited hemolytic anemias such as sickle cell anaemia, hereditary spherocytosis, or glucose-6-phosphate dehydrogenase (G6PD) deficiency may have a positive family history

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hyperproliferative 7/15

- Persistent pain in the skeleton, chest, or abdomen; priapism; lower-extremity skin ulcers; or an acute pneumonia-like syndrome suggest sickle cell anemia

- There may be a previous history of autoimmune disease (e.g., SLE, rheumatoid arthritis, scleroderma) or lymphoproliferative disorders (non-Hodgkin's lymphoma, chronic lymphocytic leukemia), which can lead to autoimmune hemolytic anemia

http://bestpractice.bmj.com/best-practice monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 8/15

• The autoimmune diseases may also cause pure red cell aplasia, in which case the reticulocyte count would be low, with normal lactate dehydrogenase, haptoglobin, and bilirubin levels

• Recent blood transfusion may indicate hemolysis due to a transfusion reaction

• Occupational or home exposure to lead should prompt suspicion of lead toxicity

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 9/15

• Features of microangiopathic disease: purpura or ecchymoses, malignant hypertension, edema, oliguria or polyuria, focal neurological signs, and hypertensive retinopathy
• Splenomegaly is seen in hereditary spherocytosis
• Lymphadenopathy may indicate infectious mononucleosis, leukemia, lymphoma, or autoimmune disease
Diagnosis

normocytic anemia: hyperproliferative 10/15

- A thrombocytopenia with schistocytes strongly suggests a microangiopathic hemolytic anemia
- Spherocytes suggest autoimmune hemolytic anaemia or hereditary spherocytosis
- Hereditary spherocytosis is associated with increased mean corpuscular Hb
- Sickling of RBCs is diagnostic of sickle cell anemia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 11/15

- Heinz bodies, eccentrocytes, or bite cells are seen in G6PD deficiency
- Elevated lactate dehydrogenase and bilirubin levels with a decreased haptoglobin are strongly suggestive of a hemolytic anemia
- Serum creatinine may be elevated in patients with hemolytic uraemic syndrome or malignant hypertension

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hyperproliferative 12/15

• Prothrombin time and activated partial prothrombin time, which are prolonged in DIC but normal in other microangiopathic hemolytic anemias

• DIC panel shows elevated D-dimers and fibrin degradation products with low fibrinogen in patients with DIC

• X-rays and MRI scanning of suspected regions reveal internal hemangiomas

http://bestpractice.bmj.com best-practice monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 13/15

- Direct antiglobulin (Coombs') test is positive in autoimmune hemolytic anemia
- Sickle cell anaemia is diagnosed on FBC
- Osmotic fragility test is positive in hereditary spherocytosis
- G6PD assays identify deficiencies of the enzyme

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

normocytic anemia: hyperproliferative 14/15

• Monospot test or EBV IgM is positive in infectious mononucleosis
• CMV IgM is positive in CMV infection
• Double-sandwich IgM ELISA or IgG avidity test is positive for IgM in acute toxoplasmosis
• Splenic or bone marrow aspirate shows amastigotes of the parasite in leishmanianosis
• Blood lead levels, which are elevated in lead toxicity

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
normocytic anemia: hyperproliferative 15/15

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Diagnosis

macrocytic anemia: megaloblastic 1/5

- The main causes to consider are vitamin B12 or folate deficiency, or drugs that interfere with DNA synthesis
- Discontinuation of causative medications leads to resolution of the anaemia
- Poor intake due to malnutrition, alcohol abuse, or strict vegan or low-protein diets can produce deficiency of vitamin B12 and/or folate

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

macrocytic anemia: megaloblastic 2/5

- A history of coeliac disease, tropical sprue, Crohn's disease, previous gastric or intestinal surgery, or bacterial overgrowth may indicate malabsorption
- A swollen, red, painful tongue; angular stomatitis; patchy hyperpigmentation of the skin and mucous membranes; and a persistent mild pyrexia are symptoms of folate deficiency

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: megaloblastic 3/5

• Known causative medications include purine analogues, pyrimidine analogues, reductase inhibitors, methotrexate, trimethoprim, anticonvulsants, oral contraceptives, cycloserine, p-aminosalicylic acid, metformin, colchicine, neomycin, and biguanides

• Hydroxyurea, in particular, is known to cause oval macrocytosis with MCV >110 femtolitres (fL)

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: megaloblastic 4/5

• Serum vitamin B12 levels are decreased and serum methylmalonic acid levels are elevated in vitamin B12 deficiency
• Normal serum homocysteine levels make folate deficiency unlikely
• Anti-intrinsic factor and parietal cell antibodies are positive in pernicious anemia
Diagnosis
macrocytic anemia: megaloblastic 5/5

http://www.medical-labs.net/wp-content/uploads/2014/03/Macrocytic-Anemia-Interpretation-Diagram.jpg
Diagnosis

macrocytic anemia: non-megaloblastic 1/6

• Causes to consider non-megaloblastic macrocytic anemia include alcohol abuse, myelodysplastic syndrome, chronic liver disease, and congenital bone marrow failure

• High alcohol intake indicates alcohol-induced anaemia, which usually persists for months after total abstinence

• A history of chronic liver disease indicates liver disease-induced anaemia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: non-megaloblastic 2/6

• History of prior exposure to petroleum distillates (especially benzene), chemotherapy, or radiotherapy should prompt suspicion of myelodysplastic syndrome

• A history of fever, chills, fatigue, weakness, recurrent infection, anorexia, night sweats, shortness of breath, and easy bruising should prompt suspicion of myelodysplastic syndrome

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: non-megaloblastic 3/6

• Recurrent infections in an infant should prompt suspicion of congenital bone marrow failure syndromes
• Examination may reveal stigmata of chronic alcoholism or chronic liver disease
• Dyskeratosis congenital is characterised by the triad of abnormal nails, reticulated skin rash, and leukoplakia

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: non-megaloblastic 4/6

• Skeletal abnormalities and growth retardation are seen in Shwachman-Diamond syndrome
• FBC shows associated neutropenia and thrombocytopenia with macro-ovalocytes in myelodysplastic syndrome
• Bone marrow aspiration and biopsy shows myeloblasts with immature precursors in myelodysplastic syndrome

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis
macrocytic anemia: non-megaloblastic 5/6

• Diagnostic features of congenital bone marrow failure syndromes are also identified
• Cytogenetics reveal chromosomal translocations in myelodysplastic syndrome
• Additional tests for congenital bone marrow syndromes: diepoxybutane or mitomycin-c fragility test is positive in Fanconi anaemia
• Genetic testing reveals underlying mutations

http://bestpractice.bmj.com/best-practice/monograph/93/diagnosis.html
Diagnosis

macrocytic anemia: megaloblastic 6/6

http://www.medical-labs.net/wp-content/uploads/2014/03/Macrocytic-Anemia-Interpretation-Diagram.jpg
Clinical case
(A Young Adult with Aplastic Anemia and Gray Hair 1)

- A man in his early twenties presented with hematuria and moderate pancytopenia [white blood cell count, 1.9 * 10^9 /L; hemoglobin, 13.3 g/dL; platelets, 61 * 10^9 /L]
- A bone marrow (BM) aspirate and biopsy demonstrated hypocellularity (20%) without dysplasia
- Results for BM cytogenetics, fluorescence in situ hybridization (FISH), and a diexpoybutane test for Fanconi anemia were normal
- Flow cytometry findings for peripheral blood showed no lymphoproliferative or myeloid-maturation disorder, nor any protein defect consistent with paroxysmal nocturnal hemoglobinuria
- Over the next 5 years, the patient remained asymptomatic and transfusion free with stable blood counts and an unchanged BM histology

http://www.clinchem.org/content/59/1/47.full.pdf
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 1)

• A 58-year-old African American male (AAM) was admitted from a nursing home (NH) with a chief complaint (CC) of being lethargic and not acting appropriately

• The patient stated that his legs hurt, and they had been hurting for a long time
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 2)

- Diabetes mellitus type II (DM), hypertension (HTN), venous stasis ulcers, anemia, hepatitis C, peripheral vascular disease (PVD), congestive heart failure (CH)
- Bilateral lower extremity (B) LE) stasis ulcers status post (S/P) debridement and multiple failed skin grafts 2 years ago, a right hallux amputation
- Tylenol (acetaminophen), Ambien (zolpidem), hydrocodone, FeSO4, clonidine, amitriptyline, Oxycontin (oxycodone), Lantus (insulin glargine), Lasix (furosemide), metformin, Actos (pioglitazone), metoprolol
- Remote history of heroin and cocaine abuse, former smoker and drinker.
Clinical case

(Anemia due to Iron Deficiency and Chronic Disease 3)

A diabetic patient with (B) infected stasis ulcers
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 4)

Physical examination

• VS 38-126-24-137/81
• Chest: CTA (B)
• CVS: tachycardic but regular with no murmurs
• Abdomen: Soft, NT, ND, +BS
  Extremities: severe venous stasis ulcers of the lower extremities (B)
• Neuro
  • AAO x 2
  • No focal neurological deficits apart from diminished sensation on (B) LE
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 5)

http://clinicalcases.org/2004/06/anemia-due-to-iron-deficiency-and.html
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 6)

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http://clinicalcases.org/2004/06/anemia-due-to-iron-deficiency-and.html
Clinical case (Anemia due to Iron Deficiency and Chronic Disease 7)

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</table>

http://clinicalcases.org/2004/06/anemia-due-to-iron-deficiency-and.html
Mainly hemodilution: the patient was with a 3-L positive balance.
Clinical case
(Anemia due to Iron Deficiency and Chronic Disease 4)

What did we learn from this case?

• A decrease in the hemoglobin can be due to hemodilution
• This is especially true in hypovolemic patients with preexisting anemia
• The rule of thumb is that one liter of IVF can decrease the hemoglobin by as much as one gram/dL

http://clinicalcases.org/2004/06/anemia-due-to-iron-deficiency-and.html
Treatment
patient education

• Inform patients of the etiology of their anemia, the significance of their medical condition, and the therapeutic options available for treatment

• If no effective specific treatment of the underlying disease exists, educate patients requiring periodic transfusions about the symptoms that herald the need for transfusion

• Likewise, they should be aware of the potential complications of transfusion

http://emedicine.medscape.com/article/198475-overview#a3
Treatment
iron deficiency anemia

• Iron deficiency anemia is treated with changes in diet and iron supplements
• If the underlying cause of iron deficiency is loss of blood — other than from menstruation — the source of the bleeding must be located and stopped

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment
vitamin deficiency anemias

• Folic acid and vitamin C deficiency anemias are treated with dietary supplements and increasing these nutrients in diet
• If patient’s digestive system has trouble absorbing vitamin B-12 from the food, he may receive vitamin B-12 injections

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment
anemia of chronic disease

• There's no specific treatment for anemia of chronic disease

• If symptoms become severe, a blood transfusion or injections of synthetic erythropoietin, a hormone normally produced by your kidneys, may help stimulate red blood cell production and ease fatigue

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment
aplastic anemia

• Treatment for aplastic anemia may include blood transfusions to boost levels of red blood cells
• Patient may need a bone marrow transplant if bone marrow can't make healthy blood cells

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment of anemias associated with bone marrow disease can include simple medication, chemotherapy or bone marrow transplantation.
Managing hemolytic anemias includes avoiding suspect medications, treating related infections and taking drugs that suppress immune system, which may be attacking your red blood cells. Depending on the severity of anemia, a blood transfusion or plasmapheresis may be necessary. Plasmapheresis is a type of blood-filtering procedure. In certain cases, removal of the spleen can be helpful. 

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment sickle cell anemia

• Treatment for sickle cell anemia may include the administration of oxygen, pain-relieving drugs, oral and intravenous fluids, blood transfusions, folic acid supplements and antibiotics

• A bone marrow transplant may be an effective treatment in some circumstances

• A cancer drug called hydroxyurea (Droxia, Hydrea) also is used to treat sickle cell anemia
Treatment thalassemia

- Thalassemia may be treated with blood transfusions, folic acid supplements, removal of the spleen (splenectomy), a bone marrow transplant or another drug

http://www.mayoclinic.org/diseases-conditions/anemia/basics/treatment/con-20026209
Treatment
oral iron b)

• Mild to moderate iron-deficiency anemia is treated by oral iron supplementation with ferrous sulfate, ferrous fumarate, or ferrous gluconate
• When taking iron supplements, stomach upset and/or darkening of the feces are commonly experienced
• The stomach upset can be alleviated by taking the iron with food; however, this decreases the amount of iron absorbed.
• Vitamin C aids in the body's ability to absorb iron, so taking oral iron supplements with orange juice is of benefit

https://en.wikipedia.org/wiki/Anemia
Treatment
oral iron b)

**ORAL IRON THERAPY**

- Oral iron treatment may require 3-6 months to replenish body stores.

**TABLE SHOWING SOME COMMON ORAL IRON MEDICATION.**

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<th>Preparation</th>
<th>Tablet size</th>
<th>Elemental iron per tablet</th>
<th>Usual Adult Dosage (per tab)</th>
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</thead>
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<td>65mg</td>
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<td>65mg</td>
<td>3 to 4</td>
</tr>
<tr>
<td>Ferrous gluconate</td>
<td>325mg</td>
<td>36mg</td>
<td>3 to 4</td>
</tr>
<tr>
<td>Ferrous fumarate</td>
<td>100mg</td>
<td>33mg</td>
<td>6 to 8</td>
</tr>
<tr>
<td></td>
<td>325mg</td>
<td>106mg</td>
<td>2 to 3</td>
</tr>
</tbody>
</table>

Treatment
injectable iron a)

• In cases where oral iron has either proven ineffective, would be too slow (for example, pre-operatively) or where absorption is impeded (for example in cases of inflammation), parenteral iron can be used

• The body can absorb up to 6 mg of iron daily from the gastrointestinal tract

• In many cases the patient has a deficit of over 1,000 mg of iron which would require several months to replace

• This can be given concurrently with erythropoietin to ensure sufficient iron for increased rate of erythropoiesis

https://en.wikipedia.org/wiki/Anemia
Treatment
injectable iron b)

<table>
<thead>
<tr>
<th>Iron Product</th>
<th>Iron-Repletion Dose in Patients on Chronic Hemodialysis</th>
<th>Iron-Repletion Dose in Patients Not on Hemodialysis</th>
</tr>
</thead>
<tbody>
<tr>
<td>Iron dextran</td>
<td>100-mg IV q dialysis treatment × 10 doses</td>
<td>100-mg IV or IM daily × 10 doses 250-1,000-mg slow IV infusion(^a)</td>
</tr>
<tr>
<td>Iron sucrose</td>
<td>100-mg IV q dialysis treatment × 10 doses</td>
<td>200-mg IV on 5 different occasions in 14 days</td>
</tr>
<tr>
<td></td>
<td>200-mg IV q dialysis treatment × 5 doses</td>
<td>500-mg slow IV infusion on Day 1 and Day 14</td>
</tr>
<tr>
<td>Sodium ferric gluconate</td>
<td>125-mg IV q dialysis treatment × 8 doses</td>
<td>300-mg, 300-mg, 400-mg IV infusion each 14 days apart</td>
</tr>
<tr>
<td>Ferumoxytol</td>
<td>510-mg IV × 2 doses 3 to 8 days apart</td>
<td>510-mg IV × 2 doses 3 to 8 days apart</td>
</tr>
</tbody>
</table>

\(^a\) Not FDA approved. IM: intramuscular.
Source: References 11-17.
Treatment
blood transfusions a)

• Blood transfusions in those without symptoms is not recommended until the hemoglobin is below 60 to 80 g/L (6 to 8 g/dL)

• In those with coronary artery disease who are not actively bleeding transfusions are only recommended when the hemoglobin is below 70 to 80 g/L (7 to 8 g/dL)

• Transfusing earlier does not improve survival

• Transfusions otherwise should only be undertaken in cases of cardiovascular instability

https://en.wikipedia.org/wiki/Anemia
Treatment
blood transfusions b)
Treatment
erthropoiesis-stimulating agent a)

• The motive for the administration of an erythropoiesis-stimulating agent (ESA) is to maintain hemoglobin at the lowest level that both minimizes transfusions and meets the individual persons needs

• They should not be used for mild or moderate anemia and are not recommended in people with chronic kidney disease unless hemoglobin levels are less than 10 g/dL or they have symptoms of anemia

• Their use should be along with parenteral iron

https://en.wikipedia.org/wiki/Anemia
Treatment
erthropoiesis-stimulating agent b)

Recombinant human erythropoietin injections

http://www.searchhomeremedy.com/drugs-and-medications-to-treat-anemia/
Treatment
hyperbaric oxygen a)

• Treatment of exceptional blood loss (anemia) is recognized as an indication for hyperbaric oxygen (HBO) by the Undersea and Hyperbaric Medical Society

• The use of HBO is indicated when oxygen delivery to tissue is not sufficient in patients who cannot be given blood transfusions for medical or religious reasons

• HBO may be used for medical reasons when threat of blood product incompatibility or concern for transmissible disease are factors

https://en.wikipedia.org/wiki/Anemia
Treatment

hyperbaric oxygen b)
Prognosis

- The prognosis depends on the underlying cause of the anemia
- The severity of the anemia, its etiology, and the rapidity with which it develops can each play a significant role in the prognosis
- Similarly, the age of the patient and the existence of other comorbid conditions influence outcome

http://emedicine.medscape.com/article/198475-overview#a4
Prophylaxis

• Many types of anemia can't be prevented
• Iron deficiency anemia and vitamin deficiency anemias can be prevented by a diet that includes:
  • Iron (meats, beans, lentils, iron-fortified cereals, dark green leafy vegetables, and dried fruit)
  • Folate (citrus fruits and juices, bananas, dark green leafy vegetables, legumes, and fortified breads, cereals and pasta)
  • Vitamin B-12 (meat and dairy products)
  • Vitamin C (citrus fruits, melons and berries)

http://emedicine.medscape.com/article/198475-overview#a4
Abbreviations

- ADH - antidiuretic hormone
- EPO - erythropoietin
- FBC - full blood count
- GI - gastrointestinal
- Hct - hematocrit
- HIF-1 - hypoxia inducible factor-1
- IVF - intravenous fluid
- MCV - mean corpuscular volume
- NSAID - non-steroidal anti-inflammatory drug
- TBV - total blood volume
Diagnostic guidelines

Europe
• Anaemia management in people with chronic kidney disease(external link)
• Guidelines for the diagnosis and management of adult aplastic anaemia(external link)
• Guideline for the laboratory diagnosis of functional iron deficiency(external link)
• British consensus guidelines on intravenous fluid therapy for adult surgical patients (GIFTASUP)(external link)
• Guidelines for the management of iron deficiency anaemia(external link)
• Significant haemoglobinopathies: guidelines for screening and diagnosis(external link)

North America
• Clinical practice guidelines for evaluation of anemia(external link)