# Investigation and Management of the Adult Patient with Anaemia

**Microcytic Anaemia** 



STEP 2       Se ferritin < 30 ug/L when CRP < 30 ug/L       Ferritin > 70 ug/L, CRP normal or increased and TSAT > 20%         I ron studies, including ferritin and transferring schurching (TSAT)       Se ferritin < 100 ug/L when CRP > 30 ug/L       Ferritin > 70 ug/L, CRP normal or increased and TSAT > 20%         STEP 3       IRON DEFICIENCY ANAEMIA       Discuss management of anoemia with an Obstetrician if pregnant       Microcytic AnAEMIA UNLIKELY TO BE DUE IN the pregnant         Investigate cause if unknown (not in parenteral iron therapy fin unknown investigation and in preenteral iron therapy fin unknown (not in parenteral iron therapy fin unknown investigation and in preenteral iron therapy fin unknown investigation and in preenteral iron therapy fit unknown investigation and in preenteral iron therapy fit unknown investigation and iron indefence or poor compliance       Acte or chronic inflammatory disease         Intense tube       Intense tube       Sectoration is source of chronic bleeding       Acte or chronic inflammatory disease         Intense tube       Permenopousol fenale       Permenopousol fenale       Permenopousol fenale         Permenopousol fenale       Permenopousol fenale       Permenopousol fenale       Permenopousol fenale         Permenopousol fenale       Permenopousol fenale       Permenopousol fenale       Permenopousol fenale         Permenopousol fenale       Permenopousol fenale       Permenopousol fenale       Permenopousol is unknown         STEP 4       IRON DEFICIENCY ANAEMIA       Check f	STEP 1 <ul> <li>Full Blood Picture Test</li> </ul>	* MCV < 76fl or MCH < 27pg Microcytic anaemia	* Normal range values may differ between hospital laboratories
<ul> <li>Correct cancernia</li> <li>Start oral iron therapy to normalise Hb and replenish iron stores</li> <li>Investigate cause if unknown (unless further investigation not in patient's best interests)</li> <li>Review history &amp; examination for source of chronic bleeding Refer to Gastroenterologist if - Adult male - Postmenopausal female - Premenopausal female - Premenopausal female - Premenopausal female - Premenopausal female - Postmenopausal female - Postmenopausal female - Postmenopausal female - Premenopausal female - Premenopausal female - Premenopausal female - Premenopausal female - Postmenopausal female - Postmenopausal female - Postmenopausal female - Premenopausal female - Premenopausal female - Postmenopausal fem</li></ul>	<ul> <li>Iron studies, including ferritin and transferrin saturation (TSAT)</li> </ul>	Se ferritin < 100 ug/L when CRP > 30 mg/l or TSAT <20% Manage as IRON DEFICIENCY ANAEMIA	Ferritin > 70 ug/L, CRP normal or increased and TSAT >20%
<ul> <li>Monitor response to replacement therapy</li> <li>Treat disease causing the anaemia</li> <li>Check full blood picture (FBP) after 2-4 weeks of iron therapy</li> <li>If Hb has improved (10-20g/L increase):</li> <li>Check if Hb normalised after 2-4 months iron therapy</li> <li>Continue iron therapy for another 3 months to replenish iron stores</li> <li>If no improvement, consider:</li> </ul>	<ul> <li>Correct anaemia</li> <li>Investigate cause if unknown         (unless further investigation not in patient's best</li> </ul>	<ul> <li>a) Start oral iron therapy to normalise Hb and replenish iron stores Start with parenteral iron therapy if: <ul> <li>History of oral iron intolerance or poor compliance</li> <li>Impaired gastrointestinal absorption</li> <li>Haemodialysis</li> <li>Major surgery must take place in &lt; 3 weeks</li> </ul> </li> <li>b) Review history &amp; examination for source of chronic bleeding Refer to Gastroenterologist if <ul> <li>Adult male</li> <li>Postmenopausal female with gastro-intestinal symptoms or bleeding</li> </ul> </li> <li>Refer to Gynaecologist if <ul> <li>Post menopausal bleeding</li> </ul> </li> </ul>	IRON DEFICIENCY         Assess for:         • Acute or chronic inflammatory disease         • Chronic infection         • Malignancy         • Liver disease         Check differential white cell count (WCC), liver function tests (LFTS)         Refer to a Haematologist if:         • Thalassaemia or sideroblastic anaemia suspected or
• Switch to initiavenous iron merapy	<ul> <li>Monitor response to replacement therapy</li> <li>Treat disease causing</li> </ul>	<ul> <li>Check full blood picture (FBP) after 2-4 weeks of iron therapy</li> <li>If Hb has improved (10-20g/L increase):</li> <li>Check if Hb normalised after 2-4 months iron therapy</li> <li>Continue iron therapy for another 3 months to replenish iron stores</li> </ul>	A diagnosis of exclusion Unresponsive to parenteral iron unless iron deficiency also present

# Investigation and Management of the Adult Patient with Anaemia



#### GUIDANCE ON THE INVESTIGATION AND MANAGEMENT OF THE ADULT PATIENT WITH ANAEMIA

**Iron deficiency anaemia** is the most prevalent cause of anaemia. It can be due to blood loss, impaired iron absorption or failure to utilise iron stores.

- 1. Have a high index of suspicion for iron deficiency anaemia in patients with:
- Chronic gastro-intestinal symptoms or evidence of chronic bleeding from gastrointestinal tract
- Acute or chronic inflammatory bowel disease or malabsorption
- Malignancy
- Menorrhagia
- Recent pregnancy

- If nausea, abdominal pain, constipation or diarrhoea occur, advise ferrous sulphate be taken with food, reduce daily dose of ferrous sulphate or change to ferrous gluconate 600 mg twice a day
- Haemoglobin (Hb) should rise by 20 g/L every 3-4 weeks and be within normal range after 2-4 months of iron therapy
- Continue oral iron therapy for 3 months after Hb has returned to normal range, to replenish iron stores.
- Measure Hb every 3 months for a year, then monitor annually, prescribing additional iron therapy as required

#### Intravenous iron therapy is indicated if:

Iron should be administered intravenously for patients on haemodialysis, unless this route is contraindicated.

Consult with patient's Nephrologist about addition of Erythropoiesis-Stimulating Agent (ESA) if inadequate response in Hb with iron therapy alone.

**Macrocytic anaemia** is most commonly due to excessive alcohol intake, folate and / or vitamin B12 deficiency but may also occur with myelodysplasia, myeloma, cytotoxic drug therapy and following gastrectomy or terminal ileum resection

• Do not delay replacement therapy with

Goddard AF, James MW, McIntyre AS, Scott BB, on behalf of the British Society of Gastroenterology, Guidelines for the management of iron deficiency anaemia. GUT 2011; 60: 1309 - 1316

Goodnough LT, Maniatis A, Earnshaw P et al. Detection, evaluation, and management of preoperative anaemia in the elective orthopaedic surgical patient: NATA guidelines BJA 2011; 106: 13-22

Hoffbrand AV, Moss PAH, Pettit JE. Essential Haematology 2006, Blackwell Publishing Ed. Musallam et al. Preoperative anaemia and postoperative outcomes in non-cardiac surgery: a retrospective cohort study. www.thelancet.com October 2011.

2. Investigate the cause of anaemia promptly unless the aetiology is already known or further investigations are not in the patient's best interests (e.g. palliative care patient).

Anaemia should be investigated more urgently in men with a haemoglobin concentration < 120 g/L and in women with a haemoglobin concentration < 100 g/L, because such low levels of Hb may be indicative of serious disease.

- 3. Once iron deficiency has been detected treatment should not be delayed
- Oral iron therapy is the treatment of choice in most cases, even if temporary discontinuation for 1 week is indicated to facilitate colonoscopy
- Prescribe ferrous sulphate 200mg twice or three times a day (at least 6hr apart), preferably between meals, if tolerated

- Oral iron therapy has failed or was not tolerated by patient
  - History of malabsorption or chronic inflammatory bowel disease
- Functional iron deficiency e.g. chronic kidney disease (CKD) on haemodialysis
- Major surgery must proceed in less than 3 weeks time

**Normocytic anaemia** can be due to CKD or anaemia of chronic disease but may also be a sign of early iron deficiency anaemia.

- \*\* In CKD treat as iron deficiency anaemia if transferrin saturation (TSAT) is less than 20% AND ferritin is less than 100 ug/L. Measure **both** of these parameters to assess iron deficiency and response to replacement therapy.
- folic acid and / or vitamin B12. Investigate cause, if unknown.
- Ensure that vitamin B12 therapy is commenced with folic acid when both nutritional deficiencies exist, to avoid neurological complications.

#### Anaemia prior to scheduled major

**surgery** whether mild or severe, is an independent risk factor for postoperative major morbidity and mortality. Major surgery should therefore be delayed until anaemia is corrected, if it is in the patient's best interests.

#### References

World Health Organisation. Worldwide Prevalence of Anaemia 1993-2005. WHO 2008.

Cook J. Diagnosis and Management of Iron Deficiency Anaemia. Best Practice Research and Clinical Haematology 2005; 18: 319-32. \*\*National Institute for Clinical Excellence, Anaemia management in people with chronic kidney disease. June 2015 cks.nice.org.uk/anaemia-iron-deficiency

NHS Clinical Knowledge Summaries Anaemia - iron deficiency, February 2013 cks.nice.org. uk/anaemia-iron -deficiency.

\*\*\*NHS Clinical Knowledge Summaries, Anaemia - vitamin B12 and folate deficiency, July 2015 cks.nice.org.uk/anaemia-b12-andfolate-deficiency

S Pavord, B Myers, S Robinson et al, on behalf of the British Committee for Standards in Haematology, UK Guidelines on the management of iron deficiency in pregnancy July 2011. **www.bcshguidelines.com.** 

## **Northern Ireland Transfusion Committee**

# **Investigation and Management of the Adult Patient with Anaemia**

**Normocytic Anaemia** 



STEP 1 <ul> <li>Full Blood Picture Test</li> </ul> STEP 2	Ferritin < 30 ug/L when CRP < 30 mg/L	* MCV 76 - 100 fl MCH 27-32pg Normocytic anaemia Ferritin < 100 ug/L and TSAT < 20% **	* Normal range values may differ between hospital laboratories Ferritin > 30 ug/L and TSAT > 20%	
<ul> <li>Iron studies, including ferritin and TSAT</li> <li>Check folate and vitamin B12 levels</li> <li>Urea, creatinine, eGFR</li> </ul>	Ferritin < 100 ug/L when CRP > 30 mg/L TSAT < 20% Normal eGFR and blood creatinine Treat as IRON DEFICIENCY ANAEMIA See STEP 3 for microcytic anaemia	Normal vitamin B12 level Low or normal folate level Elevated blood creatinine, eGFR < 60 ml/min Consider CHRONIC KIDNEY DISEASE	Normal folate and vitamin B12 levels Normal renal function (eGFR > 60 ml/min) Go to STEP 3	
<ul><li>STEP 3</li><li>Correct anaemia</li><li>Investigate cause if unknown</li></ul>		CHRONIC KIDNEY DISEASE Refer to a Nephrologist if new diagnosis or deterioration of renal function Treat as IRON DEFICIENCY ANAEMIA See STEP 3 for microcytic anaemia Seek advice from a Nephrologist or a Haematologist	NORMOCYTIC ANAEMIA OF DIALACTIC ANAEMIA OF Assess for: • Acute or chronic inflammatory disease • Chronic infection • Malignancy • Liver disease Check differential WCC & LFTs If cause still unknown refer to a Haematologist	
<ul> <li>STEP 4</li> <li>Monitor response to replacement therapy</li> <li>Treat disease causing the anaemia (if appropriate)</li> </ul>		<ul> <li>CHRONIC KIDNEY DISEASE</li> <li>Continue oral iron if Hb has improved by 10-20 g/L after 2-4 weeks. Aim for: <ul> <li>Hb &gt; 110 g/L</li> <li>TSAT &gt; 20%</li> </ul> </li> <li>Ferritin greater than 100 ug/L but less than 800 ug/L**</li> <li>Offer intravenous iron If: <ul> <li>No improvement in Hb</li> <li>Intolerant of oral iron</li> <li>On regular haemodialysis</li> </ul> </li> <li>Discuss addition of § ESA with patient's Nephrologist Treat concurrent folate deficiency</li> </ul>	ANAEMIA OF CHRONIC DISEASE A diagnosis of exclusion, unresponsive to parenteral iron Iron deficiency may also be present Monitor and treat the underlying cause	
Northern Ireland Transfusion Committee       § Erythropoiesis-Stimulating Agent         Investigation and Management of the       Adult Patient with Anaemia         Macrocytic anaemia       GANNO				
STEP 1	Refer to a Haematologist if haemolysis detected	* MCV > 100 fl or	* Normal range values may differ between hospital laboratories	

**STEP 2** 

 Check folate and vitamin B12 levels Urea, creatinine, eGFR

Full Blood Picture Test

Liver function tests

**VITAMIN B12 DEFICIENCY** 

Low folate and / or low vitamin B12 level

FOLATE AND / OR VITAMIN B12 DEFICIENCY

\*\*\* Seek urgent advice from a Haematologist if

neurological symptoms secondary to folate or

vitamin B12 deficiency or if patient is pregnant

**MACROCYTIC ANAEMIA OF** 

Go to STEP 3

levels

### **STEP 3**

- **Correct anaemia**
- Investigate cause if unknown
- a. Start oral folic acid 5 mg daily If co-existing vitamin B12 deficiency start vitamin B12 injections at same time to avoid neurological complications

in FBP (confirmed on repeat FBP)

- b. Assess for cause poor diet, liver disease, alcohol misuse, gastro-intestinal surgery, recent pregnancy, chronic inflammatory disease (e.g. Crohn's disease or TB), malignancy and drug therapy (e.g. anticonvulsants)
- a. Hydroxocobalamin intramuscular injections: 1 mg alternate days for 2 weeks, then 1 mg every 3 months for life

MCH > 32 pg

- b. Investigate for possible cause, e.g.
  - Malabsorption
  - Post gastrectomy
  - Terminal ileum disease or resection

#### **UNKNOWN CAUSE**

Investigate for possible cause:

- Liver disease
- Alcohol misuse
- Hypothyroidism ٠
- Drug causes e.g. cytotoxics

If myelodysplasia or myeloma suspected or if cause of anaemia still unknown refer to a Haematologist

Normal renal function, folate and vitamin B12

### **STEP 4**

- Monitor response to replacement therapy
- **Treat disease causing** the anaemia (if appropriate)

### FOLATE DEFICIENCY

FOLATE DEFICIENCY

Monitor Hb and reticulocyte count

- after10 days for response
- after 8 weeks to check if Hb has returned to normal range
- after 4 months course completed

### **VITAMIN B12 DEFICIENCY**

Monitor Hb and reticulocyte count

- after 10 days for response
- after 8 weeks to check if Hb has returned to ٠ normal range

#### **MACROCYTIC ANAEMIA not** due to folate or vitamin B12 deficiency

Monitor Hb

Treat and monitor cause if identified

## **Northern Ireland Transfusion Committee**

August 2015