



Hosted by The Royal Marsden NHS Foundation Trust

Investigation of a Raised Ferritin

Defined as >300mcg/l male & >200mcg/l female

Raised serum ferritin levels can be due to a number of aetiologies including:

- Iron overload- Hereditary Haemochromatosis
- Multiple transfusions (Thalassaemia/Sickle cell disease/MDS)
- Inflammatory disorders- HLH/SLE/IBD/RA/Stills disease
- Viral infections- EBV/CMV/SARS-CoV-2
- Liver or renal disease
- Malignancy
- Metabolic syndrome

Suggested approach to investigate isolated elevated serum ferritin in patients without known secondary iron overload:

- Check FBC/LFT/Transferrin saturation (T-sat)
 - Normal T-sat: Consider causes other than iron accumulation
 - Alcohol
 - Inflammatory disorder
 - Inflammatory Disorders
 - Metabolic syndrome
 - Tissue damage/cell turnover/malignancy
 - o Raised T-sat, FBC normal: Consider iron accumulation
 - Hereditary Haemochromatosis (HFE genotyping)
 - Iron loading anaemia eg Thalassaemia intermedia/HbH
 - o No cause found:
 - Ferritin <1000mcg/l→ repeat serum ferritin and Tsat in 3-6 months and if persistent refer to haematology
 - Ferritin >1000mcg/l→abnormal LFTs→ refer to hepatology
 - Ferritin >1000mcg/l→normal LFTs→refer to haematology
 - Consider assessment of liver iron stores (Ferriscan/T2*MRI or liver biopsy) and rare causes

Investigations: FBC/Film

- Ferritin/T-Sat
- LFTs/GGT
- Auto antibody screen
- Viral Serology: CMV/EBV/SARS-CoV-2
- Random glucose/HbA1c
- HFE genotype
- Haemoglobinopathy screen
- Tumour markers/protein electrophoresis
- US abdomen/CT/PET
- · Referral for genetic testing of rare diseases
- Bone marrow biopsy

Table 1: Causes of raised ferritin

Increased ferritin synthesis due to iron accumulation	Increase in ferritin synthesis not associated with significant iron accumulation	Increased ferritin as a result of cellular damage
Hereditary Haemochromatosis	Malignancies	Liver disease: necrosis/viral hepatitis/Alcoholic & non-alcoholic steatohepatitis
Hereditary Acaeruloplasminaemia	HLH	
Secondary due to blood transfusion/excess iron intake/administration	Hereditary hyperferritinaemia with and without cataracts	Chronic excess alcohol consumption
Ineffective erythropoiesis eg sideroblastic anaemia, MDS	Gaucher disease	
	Acute	
Thalassaemia	and chronic infections	
Atransferrinaemia	Chronic inflammatory disorders	
Ferroportin disease	Autoimmune disorders	

If ferritin remains elevated discuss

with haematology

Raised Ferritin >300mcg/l male >200mcg/l female **Normal T-Sat** Iron loading anaemia-FBC abnormal, Check FBC/LFTs/T-Sat Consider: T-sat raised Thalassaemia Alcohol **HbH Disease** Inflammatory disorders Metabolic FBC normal, No Ferritin <1000mcg/l syndrome T-sat raised Tissue damage Malignancy HFE genotype Yeş Abnormal LFTs: Refer to hepatology Repeat ferritin & T-sat 3-6 Manage as per diagnosis **Normal LFTs:**

Algorithm

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Refer to haematology

Consider assessment of iron stores:

Ferriscan/T2*MRI

Liver biopsy

And

Rare causes