

Laboratory assessment of nutritional status in children

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Malnutrition

CHO, proteins, lipids, vitamins, mineral and trace elements are nutrients that are crucial for human life, growth and well being

Malnutrition : Inadequate intake or use of these nutrients

It is associated with risk of:

- impaired physiologic function
- increased morbidity and mortality

>50% of deaths among children < 5 years old have malnutrition

Causes of child mortality with malnutrition due to:

- diarrhea
- Infections : acute respiratory infection, measles etc..

Malnutrition:

Risk factors:

- Low socio-economic status
- Malabsorption
 - Chronic diarrhea
 - Chronic liver disease
 - Pancreatic diseases
 - Parasites infestation
- Post operative state (ileus with long periods without oral intake)
- Chronic disease: AIDS ,TB , chronic renal disease
- Hypermetabolic conditions:
major burns; septicemia ;ARF ; multiple organic failure

Malnutrition

Early detection and treatment is needed to avoid morbidity and mortality :

- Increased rates of infections
- Impaired wound healing
- Extended lengths of hospital stay
- Higher mortality rates

Lab work up for malnutrition

Hematology:

- FBC and peripheral smear :anemia from nutritional deficiency such as Iron ,folate and vit B12

Biochemistry:

- S-Pre-albumin and transferrin :better short term indicators of protein malnutrition
- S-Albumin : measure of long term malnutrition
- RBP
- S-BUN
- S-electrolytes and creatinine
- LFT
- Blood glucose
- CMP

Microbiology/parasitology /Serology

- Septic screening
- Stool and urine for parasites and bacteria
- HIV

Lab work up for malnutrition

Other tests:

- Trace elements: Zn ; Iodine
- Creatinine /proline ratio
- Coeliac serology :
 - Screening test if family history or presence of other autoimmune diseases
 - IgA tissue transglutaminase Ab (IgA tTG)
 - IgA endomysial antibody (IgA EMA)

Complication of PEM:

- Hypoglycemia
- Hyponatremia
- Dehydration and shock
- Infections (bacterial ,viral and trush)
- Hypothermia

Ideal selection of serum proteins in malnutrition

- Proteins with a short biologic half-life reflecting changes in the serum
- Associated with an :
 - Severe infection
 - Stress injury
 - End stage liver disease
 - Renal disease

Ideal s-proteins:

- Albumin
- Transferrin
- Prealbumin
- Retinol binding protein (RBP)
- IGF-1
- Nitrogen balance

Albumin

- Long Biological $\frac{1}{2}$ life of 20 days
- Not a good indicator of short term PEM
- Good indicator in chronic protein deficiency under conditions of adequate non-protein-calorie intake such chronic liver disease ,NS , PLE etc...
- Low S-alb : predictor of high morbidity and mortality in hospitalised patients

Classification of malnutrition from pl. albumin levels:

- Levels >35 g/l : normal
- Levels of 28 to 35 g/l : mild malnutrition
- Levels of 23 to 30 g/l : moderate malnutrition
- Levels < 25 g/l : severe albumin depletion

Transferrin

- Biologic half-life : 9 days
- Synthesized in the liver and binds and transports ferric iron
- High levels in iron deficiency(in proportion to deficiency)
- An early indicator in iron deficiency
- Last analyte to return to normal when iron deficiency is corrected
- More sensitive indicator of protein depletion than s-Alb
- Low levels can also be seen :
 - Nephrotic syndrome
 - Liver disorders
 - Neoplastic disease

Prealbumin

- Biological half-life : 2 days
- Each prealbumin subunit contains one binding site for RBP
- Major transport protein for thyroxin
- Better indicator of visceral protein status and positive nitrogen balance than albumin and transferrin
- Has a superior indicator for monitoring short term effects of nutritional therapy
- In protein-energy malnutrition, there is a greatly decrease levels of prealbumin and RBP complex ; but levels return towards normal after nutritional replenishment

RBP

- Biologic half-life : 12 hrs
- Can be used to monitor short term changes in nutritional status
- Interacts strongly with plasma prealbumin and circulates in the plasma as a 1:1 mol/l prealbumin-RBP complex

IGF-1

- Important for stimulation of growth
- Regulated by GH and nutritional intake
- Used as a nutritional marker
- Circulates in plasma bound to IGF-BP3
- IGF-BP3 modulates the biologic effect of IGF-1 in the stress response

Malnutrition

- Protein-calorie malnutrition: Kwashiorkor
- Calorie malnutrition : Marasmus
- Protein losing enteropathy
- Stress hypermetabolism

Kwashiorkor

- Calorie-protein malnutrition
- Maladaptive response to starvation where the body utilizes proteins and conserve s/c fat
- Significant muscle and nitrogen losses from proteolysis

Clinical presentation :

- Edema
- Mental apathy
- Growth retardation
- Muscle wasting
- Desquamating patchy rash
- Hepatomegaly
- Dehydration (diarrhoea and vomiting)
- Signs of vitamin deficiencies
- Signs of infections



Lab findings:

- ↓TP ; ↓ Alb; ↓ pre-alb
- ↓BUN; ↓ Chol
- ↓ transferrin ; ↓ ferritin
- ↓ B12; ↓ folate ; ↓ lymphocyte count with severe anemia

Marasmus (wasting)

- Inadequate intake of proteins and calories
- An adaptive response to starvation, the body utilizes all fat stores before using muscle
- Seen most commonly in the 1st yr of life due to lack of breast feeding

Clinical presentation :

- Emaciation
- Severe loss of muscle tissue due to gluconeogenic activity
- Severe loss of adipose tissue due to increased lipolytic activity
- Severe growth retardation ;child looks older than his age
- No edema or hair changes
- Alert but miserable ; child appears hungry
- Dehydration (diarrhoea)
- No protein deficiency : normal serum transport proteins levels

Risk factors:

- Poverty and famine
- Diarrhea
- Ignorance and poor maternal nutrition



Protein losing enteropathy

Loss of s-protein into the lumen of GIT due to increased permeability as a result of cell damage or death

Causes:

Any condition causing serious inflammation, erosion, in the intestines such as :

- Intestinal parasite or bacteria infection
- Coeliac sprue
- Necrotizing enterocolitis
- Measles
- Whipple disease
- TB ; sarcoidosis
- Lymphatic obstruction

Symptoms :

- depend on underlying disease
- Diarrhea with or without bleeding
- Fever
- Abdominal pain and /or weight loss
- Edema secondary to decreased pl.oncotic pressure

Lab workup of Protein losing enteropathy

- ↓ s-alb (exclude NS, chronic liver disease ,malnutrition)
- ↓ Igs and lymphocytes
- Abnormal presence of α -1 antitrypsin in stool (important in diagnostic) :
 - α -1AT is an endogenous protein not present in diet , not normally actively secreted , absorbed or digested ; not secreted in urine
 - α -1AT is stable in feces at 37 ° C ; detected by IMX
- ↓ fat soluble vitamins (ADEK) in fat malabsorption if lymphatic obstruction

Imaging Tests :MRI or CT of abdomen

Histologic findings: Biopsies for definitive diagnosis of underlying disease

Stress hypermetabolism

- Catabolic state associated with stress injury (trauma ,burns and sepsis)
- Driven by cytokines release (IL-1, IL-6 and TNF- α); proportionate to the amount of injury
- Linked to the release of stress hormones such as catecholamine ,glucagon , GH,cortisol and thyroid hormones
- Catecholamine and glucagon stimulate hepatic glycogenolysis
- Adrenal cortisol opposes the effects of insulin (insulin resistance):
 - muscle proteolysis provides gluconeogenic precursors through alanine
 - Muscle weakness and wasting

Stress Hypermetabolism will result in:

- Hyperglycemia
- Release of FFA and associated ketosis
- Release of IGF-I
- Increase metabolic rate
- Negative nitrogen balance from gluconeogenesis due to massive proteolysis (conversion of lean body protein to a.a)

Stress hypermetabolism

- Patients may become marasmic when the gradual loss of fat tissue is associated with starvation
- Kwashiorkor is also present in stress injury when there is visceral protein loss