

MIRACLE Academy

سموم
زميلتكم نهى حسن



لجان الرفعات

قال تعالى (يَرْفَعُ اللَّهُ الَّذِينَ آمَنُوا مِنْكُمْ وَالَّذِينَ أُوتُوا الْعِلْمَ دَرَجَاتٍ)

اكثر الناس عرضه اله
الي يشتغلو بمحطات
البنزين

METALSHEAVY

هسا يا جماعه في
أشياء قراتهم الدكتور ه
قراءه فانا حاولت
اشرحهم



Lead toxicity

HEAVY METALS

LEAD ✓

✓IRON

MERCURY

ARSENIC

NICKEL

CADMIUM

THALLIUM

ALUMINUM

GOLD

يعتبر من العادن الي شديد
السميه لذلك جسمي ما
بحتاجه كثير

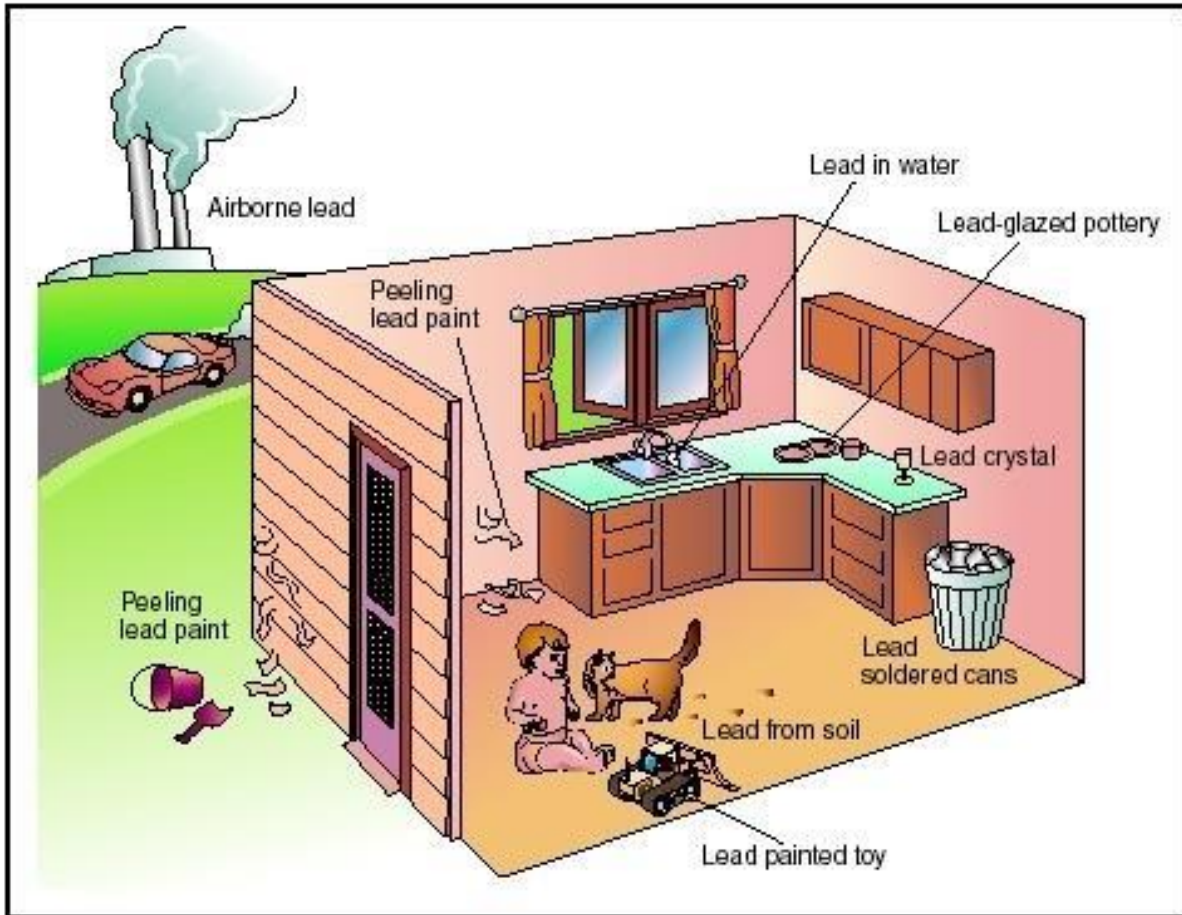
- Some metals needed in trace amounts
- Body lacks any major system to remove excess metals

بالغالب بتخلص منهم عن
طريق renal excretion

LEAD

- Lead poisoning is one of the oldest occupational and environmental diseases in the world
- Exposure from: environment (water, air, soil, food), fuels, paints, production of storage batteries, glass polishing, shooting
- Environmental lead exposure has declined considerably in the last three decades.
- elimination of lead as an additive in gasoline, as well as diminished contact with lead-based paint and other lead-containing consumer products.

هون يحكولنا صح يدخل بصناعه
الأشياء الي فوق ولكن حاولو يقللوا
نظرا لانه سميته عاليه



LEAD

هادر نادر اءوءءه Acut
هادر اءطفر Chronic
وفعلم مشاكل اءطفره

- Lead is a cumulative poison that causes **both chronic (plumbism) and acute intoxication**
- Acute** poisoning is **rare** but **chronic** one is a **serious problem** (low-level lead exposure)

معناها انو فكون معرض له لفءره قلفله ولكن لمة طوله فعنف مءلا كل ما فروح للمصنع ساعه بءعرض اله ولمءه طوفله

- The **intestinal tract** is the primary route of entry in **non-industrial exposure....from food & water**

اول اشف ممكن فءءل اللفء ففه للجسم هو عن طرق اكله فعنف اشف فوصل للجهاز الهضمف

- Lead-containing paint** is a **1ry** source of lead exposure in **children (pica)**

ءكفنا الءهان فءءوف على اللفء وهاء ممكن فءعرض الاءفال وءعرفو بفءبو فمسكو وفلعبو بكل اشف

- Lead exist in both **inorganic** and **organic** form

اعلى
absorption



معنف pica هف condition بءكفها عن الاءفال لانهم فوكل الاشفاء الففر صالءه للاكل مثل ءءراب ممكن وفكون فءءوف على اللفء او بطارفاء ءعرفوهم الصغار الءائرفاء

Toxicokinetics

عنا اكثر من طريقه ليصير
امتصاص لليد ويدخل للجسم

- **Absorption:**

امتصاصه بالنسبه للصغار اعلى
مقارنه مع الكبار

- **Oral exposure:**

- adult diet (**10% absorbed, children absorb 50%**)

- **Dietary deficiencies** of calcium, iron, zinc enhance lead absorption as well as its tissue storage

اذا واحد عندو نقص بالكالسيوم او الزنك
وكان معه انيميا رح يزيد هاد امتصاص
الليد وتخزينه محل الكالسيوم

- **Inhalation:** absorption is greater and more rapid by pulmonary route....is **the major route of industrial exposure** (lead fumes, fine particles)

تعتبر الطريقه الرئيسيه لامتناسه داخل
المصانع وسريع جدا

- **Dermal absorption** is poor, Cutaneous absorption of lead is limited (typically far less than 1%), except in case of organic lead

امتصاصه بكون قليل من خلال الجلد الا اذا
كان organic

Toxicokinetics

- After absorption lead circulate through the blood associated **99% with erythrocytes** and **1% present in plasma**

هسا صار امتصاصا للأسف ف روح يروح على circulation ومنها على bone cns
الخ...

ف يمكنوب انه 99% منه بيروح على erythrocyte

- Distributed first to soft tissues** (renal tubule and liver) and then **incorporates into bone, hair and teeth** for storage

يخل على الكبد والكليه وبعدها
على العظم والشعر والاسنان
ويتخزن هناك

- Crosses the placenta and the BBB**

- High affinity for bone and other calcified tissue.....90% deposited in bone "lead lines" (tertiary lead phosphate)

اعطل اشي انو يجل من خلال المشيمه والbbb
و90% يصيرله ترسيب بالعظم بسبب ارتباطه مع
الفوسفات

LEAD

شايقين هاد
اللون الأبيض
المشع هاد
للليد

رح يوقف النمو
للعظام ومشاكل
خطيره بالعظم



LEAD LINES



Toxicokinetics

إذا كان بالدم أو الكليه أو الكبد مده
التخلص منه بتكون من شهر لشهرين
أما إذا كان بالعظم وترسب هناك لسنين

- **Clearance:** half life in the blood and soft tissues is 1–2 months; while in bone is years to decades
- ~70% of lead excretion occurs via the urine
- Less amounts are eliminated via the feces and exfoliation of epithelial tissue, sweat, and breast milk
- **A dose of 0.5g of absorbed lead is estimated to represent a fatal dose**

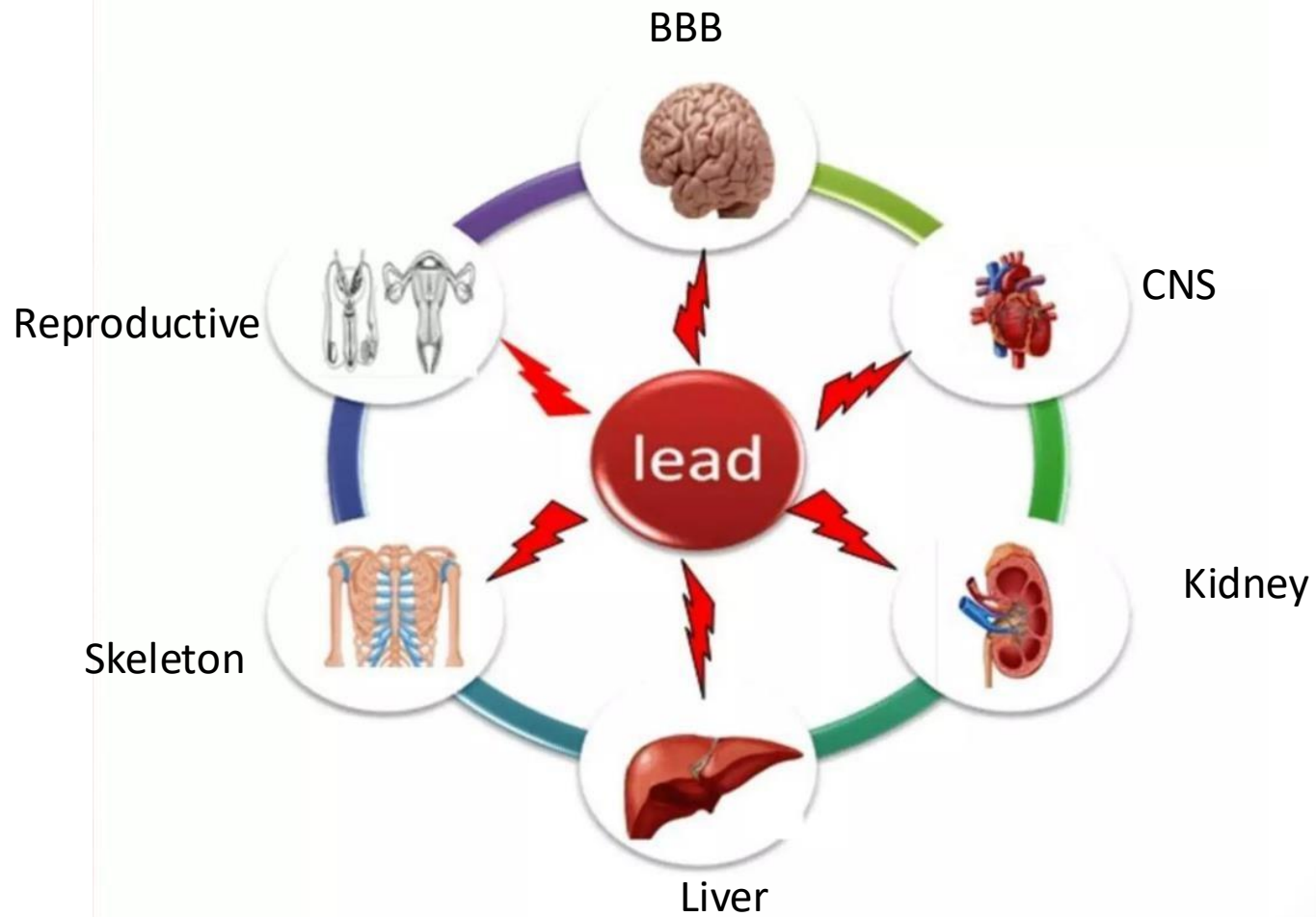
إذا اخذ
500mg تكون
جرعه قاتله

Toxic dose

	Severe toxic	Non toxic
بالوضع الممتاز المفروض مياه الشرب ما تحتوي على الليد ولكن للأسف فيه ولكن ضمن رينج محدد الا وهو 15ppb	Children:600 micro g(3) Adult :800 micro g(4)	<15microg /L(1mmol)

- Whole blood lead concentrations are non toxic **if < 150 µg/L (1 mmol/L)**
- Concentrations over **600 µg/L [3 mmol/L]** (children) or **800 µg/L [4 mmol/L]** (adults) are usually associated with severe toxicity.
- level for lead in drinking water **is 15 ppb**.....however, the maximum contaminant level **goal 0 ppb**

Lead toxicity



Toxicity

لما يصير تسمم عندي رح يتم
تنشيط عده انزايمز مسؤولين عن
عده عمليات حيويه بالجسم منهم
sulfhydryl group يرتبطو مع
ويثبطو عمل الانزيم

- The toxic **effects** range from
- inhibition of enzymes to the **production of severe pathology or death**
- Lead exerts **multisystemic toxic effects** that are mediated by multiple modes of action:
- Primarily by binding to sulfhydryl group of protein molecules....cause inactivation of several enzyme systems
- Lead affect the nervous system, the GI, hematopoietic, reproductive & CV systems

Hematologic Effects

- Decreased heme synthesis.....increase
production of RBCs by bone marrow
(compensatory mech.)
- These cells are released as immature
reticulocytes and **stippled cells (basophilic**
stippling)
- When blood smear is stained erythrocytes display
dots.....**accumulation of mRNA**
- Normocytic or Microcytic and hypochromic
anemia

الليد بقلل الهيم تمام فيحاول العظم بزيد من
صناعه RBCs ولكن الي تكون رح تكون
immature
وتكون زي مرقطه فلما نشوفها تحت
المايكروسكوب تكون dots



أنواع الانيميا
الي يسببها

Renal Toxicity

- **Chronic lead nephrotoxicity** consists of interstitial fibrosis, progressive nephron loss, azotemia & renal failure
- **Acute lead nephrotoxicity** consists of proximal tubular dysfunction and azotemia.....can be reversed by Tx with chelating agents
- Impairs the renal synthesis of heme-containing enzymes involved in vitamin D metabolism....affect bone
- Hyperuricemia with gout

لما يصير عندي فشل بالكليه رح تبطل تقدر تحول
vit D الى شكله النشط وهاد باثر على العظم لانه نص
الفيتامين هاد بقلل امتصاصا Ca+

لانه يتم التخلص من اليوريك اسيد
عن طريق الكليه وهون صار فشل
فيها ف صار يتراكم بالدم

الدكتوراه روان حكت بنركز على
الي بالاحمر بالاضافه انها هي الي
كاتبه الاسئله

Neurologic, Neurobehavioral, and Developmental Effects in Children

- Manifestations range from **impaired concentration, headache, diminished visual-motor coordination**, & tremor to overt **encephalopathy**: lethargy or delirium, vomiting, irritability, loss of appetite, dizziness, and **convulsions** مشاكل ب cns
- May progress to obvious ataxia, and reduced level of consciousness....may progress to coma and death
- Lead affects virtually **every neurotransmitter system in the brain** (glutamatergic, dopaminergic, and cholinergic systems).... الدكتوراه قراتهم قراءه
- Recovery is often accompanied by sequelae including retardation....in mentalepilepsy, some cases, optic neuropathy and blindness

Effects on Cardiovascular System

- The pathogenesis of lead-induced **hypertension** is multifactorial including:
 1. **Inactivation of endogenous nitric oxide and cGMP**, possibly through lead-induced reactive oxygen species;
 2. Changes in the **RAAS** and increases in sympathetic activity.....important humoral components of HTN;
 3. possible rise in **endothelin & thromboxane===vasoconstrictors**

الليد بقلل افراز الناييترو أكسيد
وهو مسؤول عن vasodilator

Other Toxic Effects

- Lead decreases immunoglobulins, peripheral B lymphocytes, and other components of the immunologic system.....**immunosuppressive agent**
- Retention and mobilization of lead in bone occur by the same mechanisms involved in calcium regulation.....**competes with Ca for GI absorption**
- Lead affects osteoblasts, and osteoclasts.....has been associated with osteoporosis and delays fracture repair

Other Toxic Effects

يعمل عقم

- Lead toxicity has long been associated also with **sterility** and **spontaneous abortion and low birth weight**
- **GI effects:**
 - Abdominal cramp
 - Constipation, Nausea
 - Less common Diarrhea

Diagnosis

- Skeletal x-ray's fluorescence measurement of lead
- Blood levels of lead
- Anemia microcytic, hypochromic (with basophilic stippling)
- Azotemia, Gout

-)

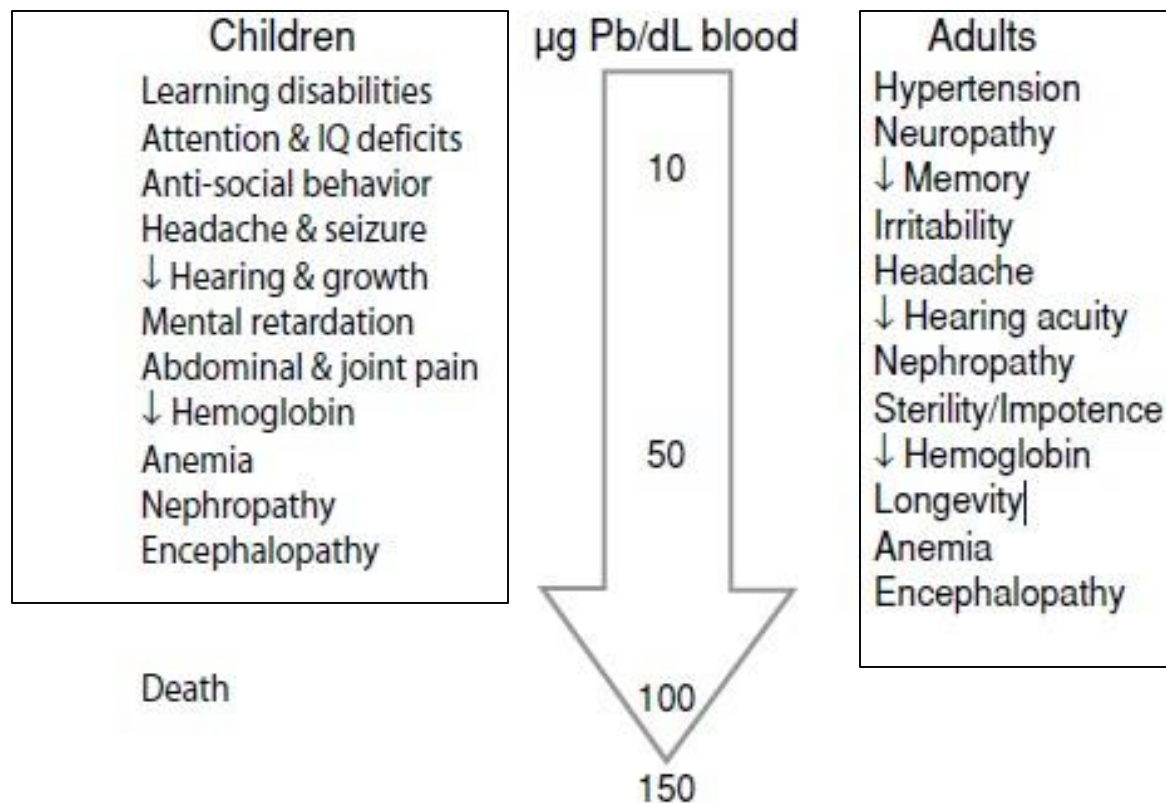
هاي حكينا
تدل على انو
acute في
renal

Diagnosis

- N.B: consider lead poisoning in any patient with **multisystem findings** with abdominal pain, headache, anemia, and, less commonly, motor neuropathy, gout, and renal insufficiency.
- Consider lead encephalopathy in any child or adult with delirium or convulsions (especially with coexistent anemia)

S.E and
does
مش مطلوب

FIGURE 1. Effects of lead poisoning on human health^a



^a Adapted from Gurer and Ercal (49).

Dose
dependant

LEAD TREATMENT

TREATMENT:

- REMOVAL OF THE SOURCE & STABILIZE THE PATIENT
- CHELATING THERAPY:
 - **BAL**
 - **Calcium EDTA**
 - **SUCCIMER**
 - **D - PENICILLAMINE**
- **SUPPORT**

هسا هدول بعطيهم
لناس الي تسممو
باليد عشان يعملو
complex مع
الليد ويقللو من
نسبه امتصاصه
وتراكمه

Treatment

بالاضافه الى
CHELATING THERAPY
رج يحتاج الى علاج
مساعد في حاله حدوث الاتي

- Treat seizures and coma if they occur
- Provide adequate fluids to maintain urine flow but avoid overhydration.....may aggravate cerebral edema

بعطيه كميه منيحه من السوائل حتى احسن
لانه يتم التخلص من الليد من خلاله ولكن بكميه
محدد حتى اتجنب حدوث edema

- Patients with increased intracranial pressure may benefit from corticosteroids or mannitol

اذا عنده ضغط على الجمجمه بعطيه هدول

- Decontamination by activated charcoal and whole bowel irrigation

CHELATING AGENTS

WHAT MAKES A GOOD CHELATING AGENT?

- NONTOXIC & FORMS NONTOXIC COMPOUNDS
- HIGH WATER SOLUBILITY
- SIMILAR DISTRIBUTION TO THE METAL
- LOW AFFINITY FOR CALCIUM and other ions
- EASILY REMOVED FROM THE BODY
- GREATER AFFINITY FOR THE METAL THAN ENDOGENOUS LIGANDS
- *Treatment with chelating agents decreases blood lead concentrations and increases urinary excretion*

– **DIMERCAPROL (BAL): British AntiLewisite** comp. (I.M)

برضو استخدمه في حال اتسمم بهدول الاشياء

ليش بستخدهه حتى
يربط بدل الليد مع
sulfhydryl

- Forms complexes with sulfhydryl groups

Used for inorganic mercury, arsenic and in lead poisoning

- **Chelate lead in serum and cerebral spinal fluid**

- Usually used in combination with calcium EDTA

- The complex is rapidly excreted in the urine

- May cause **hemolysis in patient with G6PD deficiency**

الي يعملهم الدواء
الاثار

- **ADE:** transient hypertension, tachycardia, N/V, fever

CALCIUM DISODIUM EDETATE (CaNa_2EDTA)

(im/iv)

- Mobilize lead from soft tissue and bone
- Forms a stable, nonionizable, water soluble compound with lead
- Complex rapidly excreted in urine
- ADE: fever, headache, N/V, anorexia, myalgia, hypotension
- **ADEs: nephrotoxicity minimized by adequate hydration**
- May deplete manganese, zinc & iron

SUCCIMER (DMSA)....p.o

- DIMERCAPTOSUCCINIC ACID....water soluble analog of BAL
- Enhances the urinary excretion of **lead** and **mercury** without affecting the elimination of the endogenous minerals as Ca, Fe, and Mn
- ADEs: GI disturbances, mild reversible increase in transaminase enzymes, allergic reaction

PENICILLAMINE.....p.o

- Penicillin derivative without antimicrobial activity...allergy!
- Widely replaced by succimer because of its poor safety profile

Symptomatic	Tx. regimen
	EDTA for 5 days

Asymptomatic	Tx. regimen
Blood lead 10-24 $\mu\text{g}/\text{dl}$	Chelation no recommended
Blood lead 25-44 $\mu\text{g}/\text{dl}$	Succimer for 2-4weeks OR EDTA for 5 days
Blood lead 45-69 $\mu\text{g}/\text{dl}$	EDTA for 2 weeks
Blood lead >70 $\mu\text{g}/\text{dl}$	BAL for five days + EDTA for 5 days

LEAD

زي ما حكيينا قبل شوي لازم اضبط
كميه الماء الي يوخذاها

• SUPPORT:

- Establish adequate urine output before administering chelating agent (fluid bolus but monitor coz may aggravate cerebral edema)

- Dialysis for patients with severe renal insufficiency

ممکن يكون بحاجه الها اذا عنده مشكله بالرينل
لانه ما رح يقدر يتخلص من الليد

- Blood lead levels: stop chelation if level <30µg/dl

اذا اصبح مستوى الليد في الدم هلقد لازم أوقف
chelation

- Recurrent blood level assessment before and after treatment with chelating agents at regular interval

لازم يضل يعمل
فحوصات لدم
على فترات
منتظمة حتى
اشوف مستويات
chelating اذا
استخدمتهم
بالعلاج