

In the Name of GOD



Rickets and Vitamin D Deficiency

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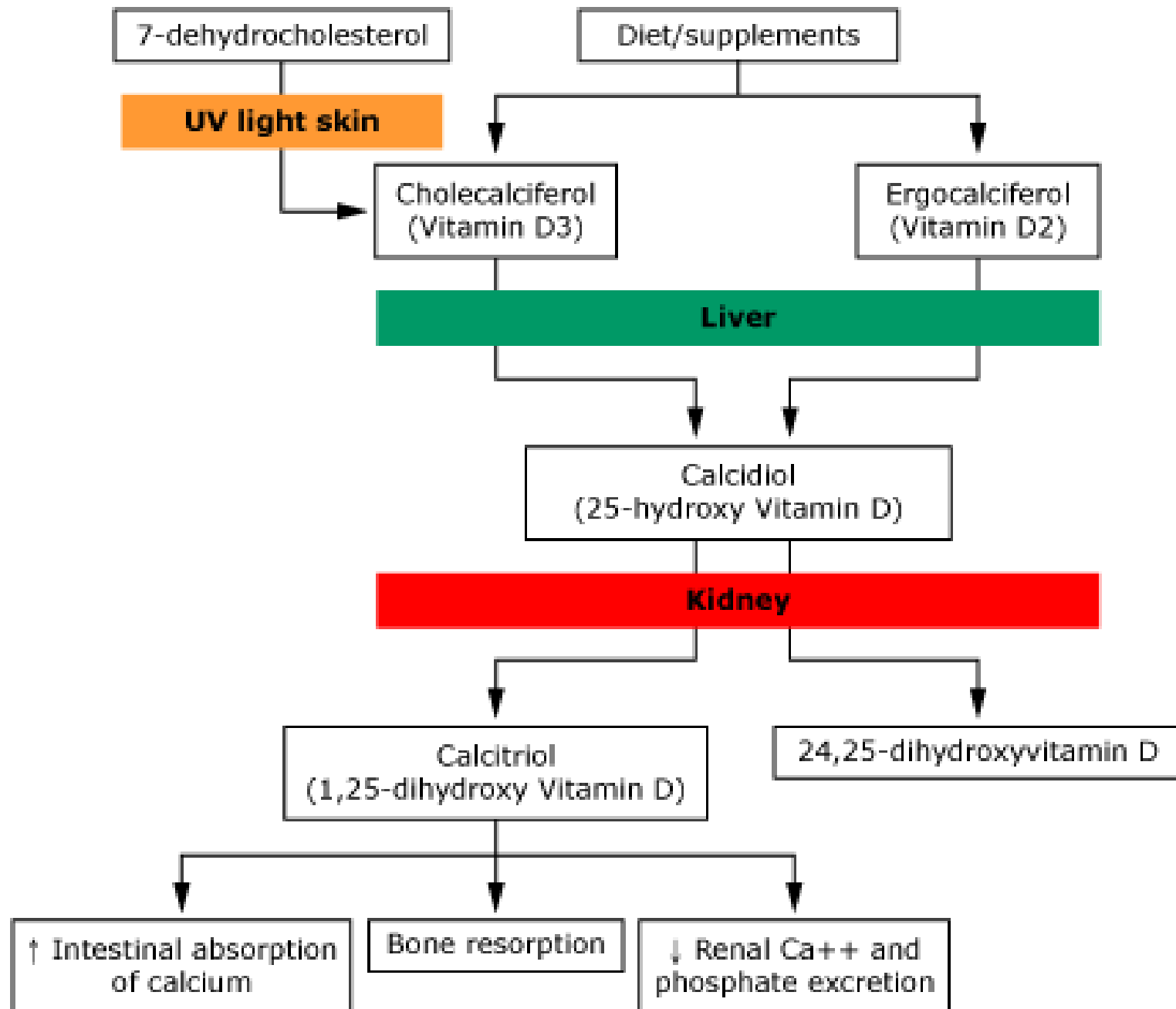
PEDIATRIC ENDOCRINOLOGY AND METABOLISM

SHIRAZ UNIVERSITY OF MEDICAL SCIENCES

Forms of Vitamin D

- ▶ Cholecalciferol (Vitamin D3)
 - ▶ Cutaneous synthesis
 - ▶ Animal products
 - ▶ Vitamin D supplements
- ▶ Calcidiol: $25(\text{OH})\text{D}$
 - ▶ **Storage form**
- ▶ Calcitriol: $1,25(\text{OH})_2\text{D}$
 - ▶ **Active form**
- ▶ Ergocalciferol (Vitamin D2)
 - ▶ Plant dietary sources
 - ▶ Vitamin D supplements

Metabolism of Vitamin D



Sources of Vitamin D

1. Exposure to Sunlight
2. Sardines, Salmon, Mackerel, Tuna
3. Cod liver oil
4. Raw milk
5. Caviar
6. Eggs
7. Mushrooms
8. Fortified food



Targets for Vitamin D Intake (RDA)

- ▶ Term Infants: 400 IU/day vitamin D3
- ▶ Healthy children 1-18 y/o: 600 IU/day
- ▶ Preterm infants:
 - ▶ Very low birth weight (<1500 g): 200-400 IU/day
 - ▶ >1500 g: 400 IU/day
- ▶ Higher doses in:
 - ▶ Obese children
 - ▶ Medical conditions (malabsorption ,...)
 - ▶ Children on mentioned medication

National Protocol for Vitamin D Supplementation

دستور العمل کشوری
مکمل یاری با مگادوز ویتامین د

- ▶ < 2 y/o: 400 IU/day
- ▶ 2-18 y/o: 50,000 units every 2 months
- ▶ 18-60 y/o: 50,000 units every 1 month
- ▶ > 60 y/o: 50,000 units every 1 month
- ▶ Pregnancy: 1000 IU/day
 - ? any previous vitamin consumptions ?
 - **No need** to check 25(OH)D
 - Sunlight exposure recommended
 - Signs of toxicity should be mentioned

Prevalence

Vitamin D deficiency prevalence is increasing globally.

- ▶ Increasing frequency since the mid 1980s
 - ▶ 15 % (overall), 1-2 % severe deficiency (United States)
 - ▶ 60% (even up to 80%) in different areas of Iran
 - ▶ Higher prevalence in females

Causes of Vitamin D Deficiency

- ▶ Decreased nutritional intake
- ▶ Decreased synthesis
 - Latitude
 - Sunlight
 - Midday
 - Season
 - Arms+ legs Or
 - Hands+ arms + face
 - No sunblock
 - Light skinned: 10-15 min
 - Dark skinned: 3-10 times more

Causes of Vitamin D Deficiency (cont'd)

- ▶ Obesity
- ▶ Liver & kidney diseases
- ▶ Perinatal factors
 - Maternal vit D deficiency
 - Prematurity
 - Exclusive breastfeeding
- ▶ Malabsorption
 - Celiac
 - CF
 - IBD
 - Surgery

Causes of Vitamin D Deficiency (cont'd)

▶ Genetic disorders

- 1-alpha-hydroxylase deficiency or **vitamin D-dependent rickets type I A**
- 25-hydroxylase deficiency or **vitamin D-dependent rickets type 1B**
- Hereditary resistance to vitamin D or **vitamin D-dependent rickets type II**
 - mutations in the vitamin D receptor gene

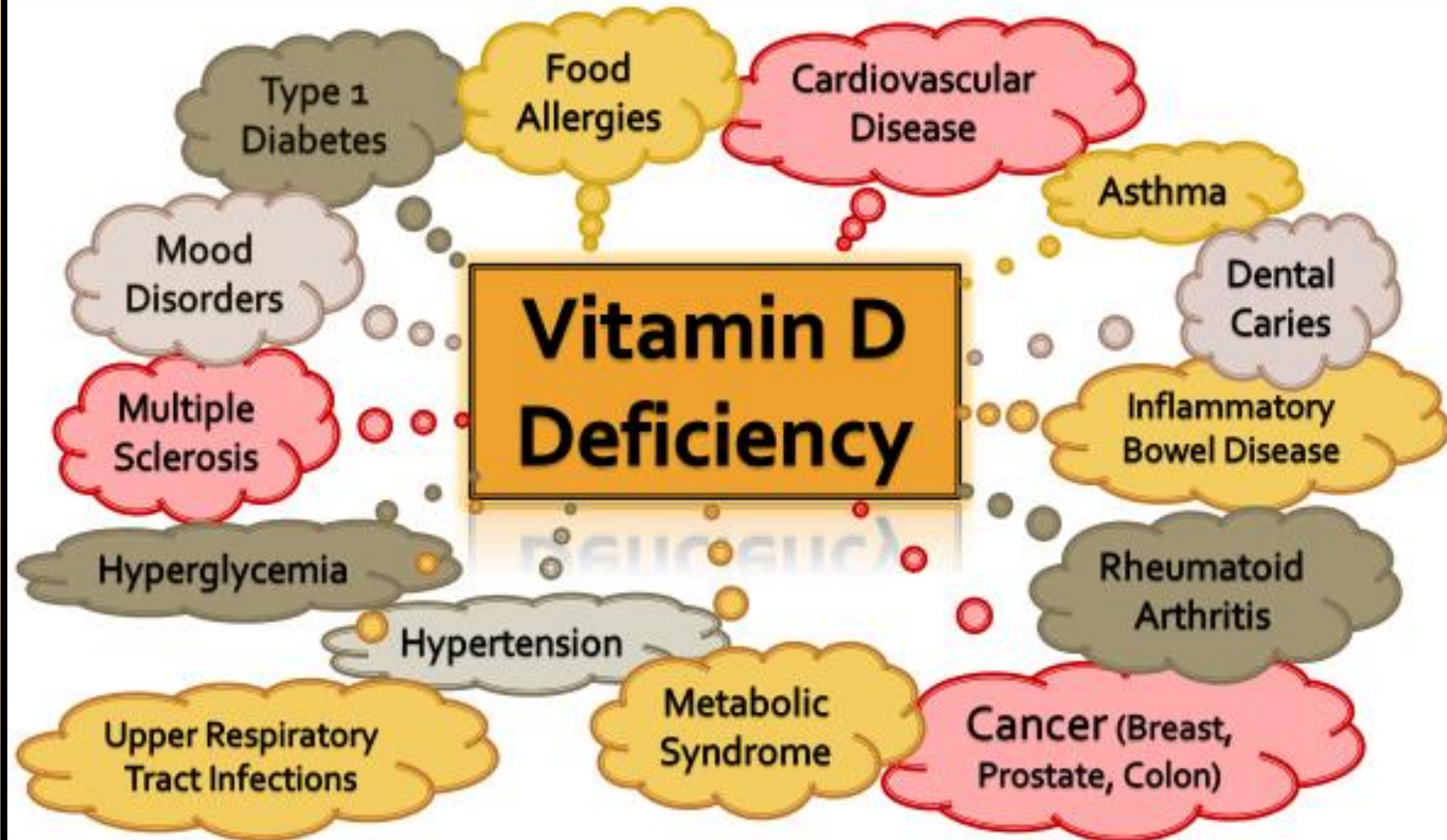
▶ Medication

- ▶ **Anti Epileptic Drugs (AEDs)**
- ▶ **Antiretrovirals**
- ▶ **Glucocorticoids**
- ▶ **Antifungals**

Why is Vitamin D Important?

- ▶ An essential nutrient that plays an important role in calcium homeostasis and bone health.
- ▶ Can **reduce cancer cell growth**, help control infections and reduce inflammation.
- ▶ Covid-19 infection & Vitamin D:
 - ▶ Reduced vitamin D values resulted in a **higher infection risk, mortality** and **severity COVID-19 infection**. Supplementation may be considered as **preventive and therapeutic measure**.

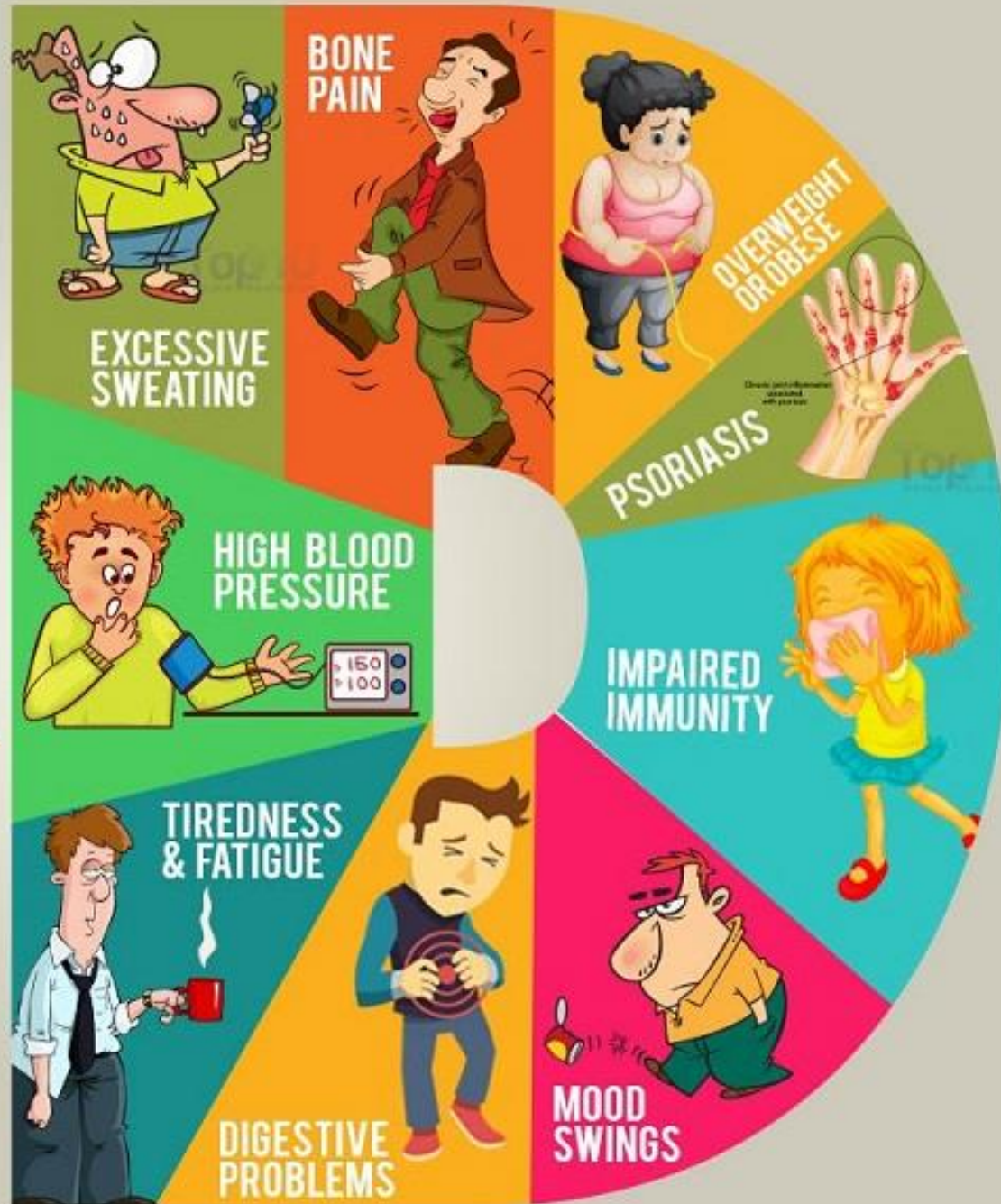
Associations



SIGNS

You May Have a

VITAMIN D DEFICIENCY



Rickets & Vitamin D Deficiency

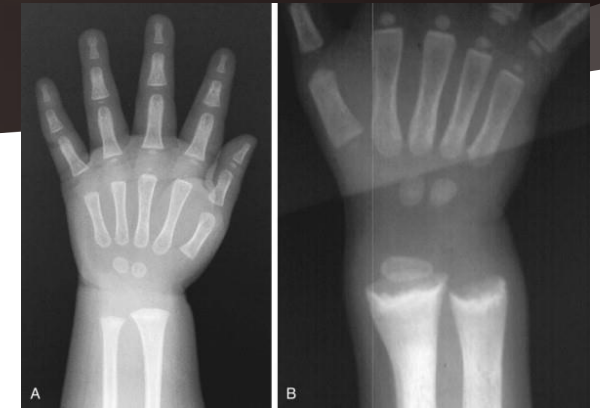
- ▶ Rickets is the **principle manifestation** of vitamin D deficiency in **infants and young children**
 - ▶ Asymptomatic, pain and irritability, motor delays, and poor growth
 - ▶ Younger children: delayed closure of fontanelles, craniotables, frontal bossing, prominence of costochondral junctions, widening of wrists and ankles, and bow legs or knock knees
 - ▶ Muscle weakness and discomfort, difficulty standing or walking



Rickets & Vitamin D Deficiency (cont'd)

- ▶ Advanced vitamin D-deficient rickets:
 - ▶ seizures, tetany or apneic spells, stridor, wheezing, hypotonia, and hyperreflexia, particularly in very young children.
- ▶ These are a consequence of **severe hypocalcemia**, more likely during periods of very rapid growth (infancy and adolescence)
- ▶ Osteomalacia
 - ▶ **Principle** manifestation of vitamin D deficiency in **older adolescents and adults**
 - ▶ Asymptomatic
 - ▶ Isolated or generalized muscle & bone pain

What to test?



- **25(OH)D level**
 - **Gold standard: HPLC & LC-MS**
 - Radioimmunoassay (?)
 - Significant controversy in determining standards
- Suspicious to rickets if:
 - Growing child with 25(OH)D < 20 ng/mL
 - Very young age (< 3 y/o)
 - Physical signs of rickets



Check :

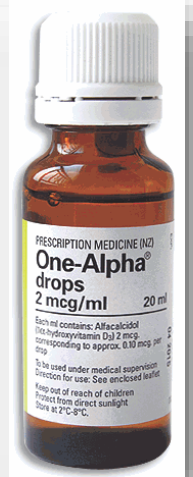
- Ca
- P
- Alk ph
- PTH
- Radiographs

Definition (children & adolescents)

- ▶ Vitamin D **sufficiency**:
 - ▶ 20-100 ng/mL
- ▶ Vitamin D **insufficiency**:
 - ▶ 12-20 ng/mL
- ▶ Vitamin D **deficiency**:
 - ▶ < 12 ng/mL
- ▶ Vitamin D **intoxication**:
 - ▶ > 150 ng/mL



Treatment



Treatment



- ▶ If 25(OH)D: 20-30 ng/mL and NO symptoms & NO risk factors:
 - ▶ Review the diet
 - ▶ Vitamin D supplements
 - ▶ Monitor 25(OH)D periodically and treat if < 20 ng/mL

Treatment

- ▶ If $25(\text{OH})\text{D} < 20 \text{ ng/mL}$ or Rickets → Treat
 - ▶ Infants < 12 m/o:
 - ▶ 2000 IU/day for 6-12 wks
 - ▶ Followed by maintenance: 400 IU/day
 - ▶ Children > 12 m/o:
 - ▶ 2000 IU/day for 6-12 wks or 50,000 IU/wk for 6 wks
 - ▶ Followed by maintenance: 600-1000 IU/day
 - ▶ Need for higher doses in:
 - ▶ Obesity, malabsorption, on medication
 - ▶ 6000 IU/day for 6-12 wks & higher maintenance

Treatment of Rickets

- ▶ 2000-5000 IU/day for 4-6wks

OR

- ▶ Stoss therapy:

- ▶ <12 m/o: 300,000 IU in 5cc olive oil / 6 hrs

- ▶ >12 m/o: 600,000 IU in 5cc olive oil / 6 hrs

AED-Induced Vitamin D Deficiency

- ▶ Preventive Dose: 400-2000 IU/day
- ▶ Monitor 25(OH)D yearly
- ▶ If 25(OH)D < 20 ng/mL :
 - ▶ Treat
 - ▶ 2000 IU/day for 6-12 wks or 50,000 IU/wk for 6 wks
 - ▶ Monthly measurement of 25(OH)D
 - ▶ Increase the dose accordingly (upto 5,000-15,000 IU/day)

Treatment of Vitamin D Deficiency in Adults

- ▶ **25(OH)D < 10 ng/mL:**
 - ▶ 50,000 IU/wk for 6-8 wks then 800 IU/day
- ▶ **25(OH)D 10-20 ng/mL:**
 - ▶ 800-1000 IU/day then repeat test 3-4 m later
- ▶ **25(OH)D 20-30 ng/mL:**
 - ▶ 600-800 IU/day
- ▶ **Vit D deficiency in pregnancy:**
 - ▶ 600-800 (1000-2000) IU/day is safe
 - ▶ Maintain 25(OH)D > 30 ng/mL
 - ▶ Urinary Ca excretion should be monitored

Follow-Up

- ▶ Individual variation in response can be seen → therapy might need to be repeated.
- ▶ Low 25(OH)D:
 - ▶ Check 25(OH)D levels after 2-3 m
- ▶ Low 25(OH)D & biochemical changes:
 - ▶ Check 25(OH)D & other chemistries 6-8 wks later, then 6 m later, then annually
- ▶ Rickets:
 - ▶ Close follow up after 1-2 wks, then every month until Alk Ph normalizes



Thank You For Your Attention