





قبل ما نبلش المحاضرة... عشان أنا كتير منيحة الله يرضى عني الهاقد الموادة الموادة الله يرضى عني الهاقد الموادة الموادة

بتستخدمه).



[7] Drug dependence:

ومنية هندية • Habitation: - psychic craving of the drug.

- No physical disturbance
- If sudden stoppage...> emotional distress. e.g. coffee and tea habits.

[8] Genetic a bnormalties (idiosyncrasy):

 It is abnormal response to drugs due to genetic abnormality in drug metabolism. These genetic abnormalities are revealed only by the effect of drugs.

. Genetics Il ables - H lie Cos *

* وخلاب بالكم انه كل حالة نقتص على أدوية معينة.

1. Acetylation Polymorphism:

• People can be classified according their rate of acetylation reaction in liver into **Rapid** and **Slow** acetylators

Examples, in slow acetylators:

a. Isoniazid → peripheral neuropathy (due to interference with

pyridoxine (vit B6) metabolism).

مرجن الذئبة الحراد b. Hydralazine → <u>SLE-like</u> (systemic lupus erythematosus-like).

u ay line will is in this disease, the immune starts · Examples in rapid acetylators: attacking body tissues

 a. Isoniazid → hepatocellular necrosis (due to accumulation of toxic metabolites)

In Isoniazid Slow acetylation -> slow metabolism in the liver -> accumulation of peripheral Competition with Vitamin B6 neuropathy — which is needed by the brain to —
have a healthy nervous system

> المريف بيجب عند الطبيب دايخ ، حاسس بتنفيل ، أعصابه منعينة. بهيك يفض لا عابة [Vit B 6] فيهدل المرض مع isoniazid .

> rapid acetylation _ excessive isoniazid toxic metabolites accumulates in liver

> Hepato cellular necrosis

فبجي المربين عند الدلتورعدة Jaundice منده اسيال ، امساك.

2. Hemolytic Anemia due to G6PD Deficiency

- Glucose-6-phosphate dehydrogenase (G6PD) is an important source of reduced glutathione which protects RBCs from hydrolysis by oxidizing drugs.
- Congenital (G6PD) deficiency → acute hemolysis in presence of some oxidant drugs as antimalarials, sulfonamides and fava beans (favism).

الى NADPH الى NADP الى مهمته يعول مهمته بيعنها المجتها الله بساعد RBCs المتحاقف على المجتها تبعتها المجتها المجتها المجتها المجتها بيعني عدم وجود مفاص لعمليات Oxidation الى معكن تجبير عالخلية المهك لو المريف أخد Which approximates المهك لو المريف أخد Which approximates وعوير عنا Hydralysisis عيمير عنا كالمهك لو المريف أخد كالمتعالم المهلك لو المريف أخد كالمتعالم المهلك لو المريف أخد كالمتعالم المتعالم ال

3. Porphyrias

- Normally, porphyrins precursors delta-aminolevulinic acid porphyrins second heme (ALA) synthase enzyme heme
 - Genetic deficiency of second enzyme → ↑ level of porphyrins with some drugs stimulating (ALA) synthase → cyanosis, severe CNS disturbances & may cause death. Porphyrin معناها عناها الأعرافا خطيرة لهك بدنا ننته للمرضان الي عناهم الأعرافا خطيرة لهك بدنا ننته للمرضان الي عناهم
 - Barbiturates and sulfonamides precipitate porphyria.

مرحى للعضلات

4. Succinylcholine Apnea توقف التنفس

• Pseudocholine esterase enzyme is responsible for breakdown of neuromuscular blocker (succinylcholine). In genetic defect of the enzyme,

Contraction العقلات مو قادق قعل Succinylcholine → respiratory muscle paralysis with apnea.

Succinylcholine is a neuromuscular blocker metabolized by pseudocholinestrase enzyme. Some individuals with deficient PsChE, when they take succinylcholine, severe muscle paralysis occurs due to lack of succinylcholine metabolism, and may lead to death from respiratory paralysis (succinylcholine apnea).

* مشكلة برجوع SR و Ca+2 بعد ما استخدمناها بالContraction *

5. Malignant Hyperthermia

• Genetic disorder in which skeletal muscles fail to sequester Ca ⁺⁺ in sarcoplasmic reticulum following administration of succinylcholine and halothane → marked muscle rigidity & sever hyperthermia. 42 محارة أعلى من 42

Drug hypersensitivity
[9] Drug allergy: جهان المناعة يسِر أ بمهاجمة الدواء

- It is Abnormal response to drug mediated by immunogenic mechanisms.

أشهر مثال مجموعة penicillin ومجموعة cephalosporin هدول مجموعتين شبه بعض بالتركيب (عندهم بتركيبهم beta-lactam ring) لو في مريض وعنده allergy من ال penicillin وبدك تعطيه دواء غيره... ما بينفع تعطيه cephalosporin لأنهم نفس التركيب فممكن يكون عنده allergy من cephalosporin برضو

Type	Mechanism	Examples
Type I Reaction (immediate type; anaphylactic)	Antigen/IgE reaction on mast cell → degranulation → release of allergotoxins e.g. histamine → fever, rash, urtecaria, angioedema & even anaphylactic shock	Penicillins

Type II Reaction	Antigen + IgG or IgM antibodies +	Methyldopa
(cytotoxic)	complement are fixed to a cell → cell lysis e.g. hemolytic anemia in RBCs	to treat of blood pressure
		anemia.

Type III Reaction Antigen + IgG antibodies complement are fixed to e → vasculitis, glomerulone	ndothelium Penicillin
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Type IV Reactions			
(Delayed type;	allergic contact dermatitis		
cell-mediated)	Pumpers Condibility 11 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1		

(التقضيحات بالصنحة الناليت

Type I: the antigen such as allergen dry binds the lyte antibodies attached to the surface of mast cells [Basophils] this binding will induce a cascade of IC reactions resulting in the release of large amounts of Histomine from the storage vesicles that will induces various allergic reactions, <u>Such as</u>: rash, fever

Type II: the antigen isn't an external allergen floating in the blood

but rather certain molecules attached to external surface of the pateint

cells. Binding of IgGer 19M antibodies stimulates a cascade of

immune reactions _____ resulting in the activation

of _____ the complement system _____ which induce further damage _____ matural killers _____ which induce further damage _____ that affect tissues.

Type III: the small antigen antibody complexes float through the blood and accumulate in various tissues such as Joints
in their dendsthelium

Blood vessels

this accumulation triggers the complement's system to attack the hot tissues.

Type IV: isn't antibody mediated but mainly cell mediated immunity.

In this type T-cells recognize the intracellular antigens.

the stimulated T-cells release cytokines or lymphokinase

which activates various immune system components

Such as phagocytes

engulf affected body cells

Diagnosis of Drug Allergy

1. History and type of reaction.

و فحوصات للتأكد من حساسين دراد معين.

2. Intradermal and conjunctival tests. بتنعمل مع الأدوية المشهور فيها الحساسية



Treatment of anaphylactic shock

Epinephrine- hydrocortisone - antihistamines.

Physiological antagonist of histamine

Quiz Time

- 1) Failure of the patient to breath after surgical operation may be due to:
- A. Pseudocholin estrase deficiency
- B. Methemoglobin reductase deficiency
- C. G-6-PD deficiency
- D. VitaminK epoxide reductase deficiency
- E. Monoamine oxidase deficincy
- 2) Failure of some children with rickets to respond to the rapeutic doses of vitamin D is most likely to be due to:
- A. Differences in sex
- B. Differences in body weight
- C. Genetic variation
- D. Tolerance
- E. Intolerance

- 3) Which of the following best describes what the term "tachyphylaxis" means?
- A. An increase in the rate of the response, for example, an increase of the rate of muscle contraction
- B. Immediate hypersensitivity reactions (i.e., anaphylaxis)
- C. Prompt conformational changes of the receptor such that agonists, but not antagonists, are able to bind and cause a response
- D. Quick and progressive rises in the intensity of drug response, with repeated administration, even when the doses are unchanged
- E. Rapid development of tolerance to the drug's effects
- 4) A 44-year-old black male is brought to the emergency department with 6 h of worsening lethargy and confusion. Past medical history is significant for easy bruising, 3 months of bone pain, and frequent pneumococcal infections. Labs were ordered, revealing serum cal- cium of 17 mg/dL (normal: 9.0 to 10.5 mg/dL). To rapidly lower his serum calcium, you administer calcitonin. However, calcitonin alone is insufficient be- cause it is known to rapidly and suddenly lose its effectiveness within 2 to 3 days of repeated dosing. For this reason, a bisphosphonate, which take 2 to 3 days to become effective, is added simultaneously. What is the term for the rapid decrease in response to calcitonin?
- (A) Anaphylaxis
- (B) Prophylaxis
- (C) Tachyphylaxis
 - (D) Tolerance
- 5)Regarding the use of a daily baby aspirin, oral fiber supplements, and a daily "water" pill in an 89-year-old man with hypertension and coronary artery disease, which of the following statements is true regarding pharmacology in the elderly patient?
- (A) Coexisting disease states are unlikely to produce additive impairment
- (B) Elderly patients are less sensitive to drug effects
- (C) Elderly patients are less sensitive to drug side effects
- (D) Elimination of drugs becomes impaired with age
- (E) Responses to compensate for drug accumulation are satisfactory

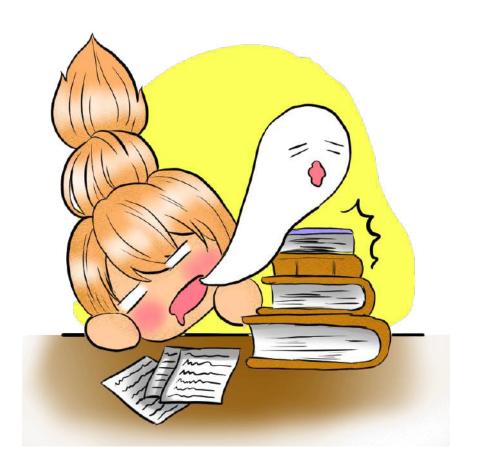
(D) DrugAB = I	DrugB > DrugA			
	yould be an exan	ual magnitude. If	Drug A and Drug B are he following?	combined
(C) Potentiatio				
(D) Synergism				
8)Tolerance de	velops because	of:		
a) Diminished a	absorption			
b) Rapid excret	ion of a drug			
c) Both of the a	above			
d) None of the	above			
combination is: a. Less the b. Equal to c. Greater		vidual actions of the two dual actions ndividual actions	on to two drugs given in o drugs	
receive:		120	surface area of 1 m ² should	
a. 100 mg	b. 87 mg	c. 120 mg	d. 95 mg	
	قت لتكون داعمًا		ت للخيبة ، هناك حلم ح يء داخلك ،لم تخلق نف صنع المستح	,

6) Which of the following drug equations exemplifies the concepts of potentiation?

(A) DrugAB > DrugA + DrugB

(B) DrugAB = DrugA = DrugB

(C) DrugAB < DrugA < DrugB



معلش ما قدرت ما أحطها وهي معبرة عن وضعي وانا مريضة وبكمل تدقيق هالتفريغ ١٠٠٠

وهيك خلصنا مادة الدكتور شريف الفخم الله يعطيه العافية ويجزيه كل خير وان شاء الله الأحد رح نبلش مع دكتورة ياسمين ويارب تكون بداية خير معها

أسأل الله لكم التوفيق والفلاح في الدنيا والأخرة 🎔 😽