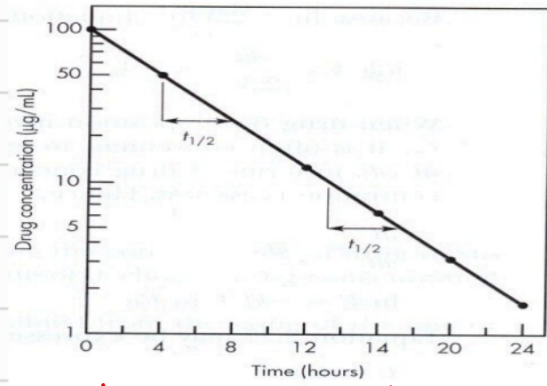


This color means it is my understanding (سُفْرِي، سَفْرِي)

* First order Drug Elimination

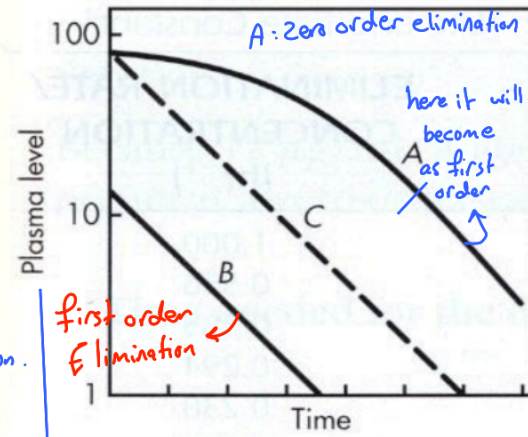
- it occurs when the rate of elimination is directly proportional to the amount of drug in the body.
- Occurs in many drugs at therapeutic concentrations
- Constant fraction of the drug is eliminated per time
- the elimination rate constant is designated as k and the unit is fraction per time ($1/\text{time}$)



* the y axis is logarithmic scale so it will never be zero

* Zero-Order Drug Elimination

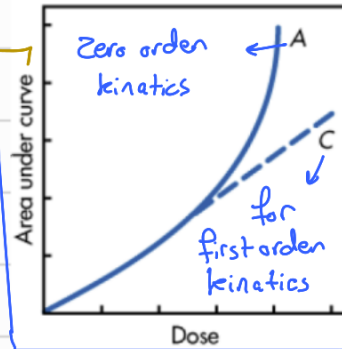
- Saturable elimination, occur with few drugs as aspirine, phenytoin, ethanol (as in alcoholic drinks)
- Elimination rate is not proportional to drug amount
- Constant Amount is removed per unit time because the saturation of elimination.



$$\text{Rate of elimination} = \frac{V_{\max} \cdot C}{K_m + C}$$

Drug conc where the elimination rate is 50% of V_{\max} ← $K_m + C$

Area under the curve represent Bio availability



* now we talk about first order kinetics

* Flow dependent Drug Elimination → some Drugs are cleared very rapidly by the organ of elimination (liver) so most of the drug perfusing the organ is eliminated by first pass effect, and the rate of elimination determined by the hepatic Blood flow. This drug property called high extraction ratio.

* Half life of elimination ($t_{1/2}$): is the time required for the amount of drug in the body or plasma conc. of drug to drop by 50%

* in this case it is constant and not related to the dose

it will never reach 100% elimination because you deal with percentage not an amount.

* After 4 Half lives, most of the drug will be eliminated from the body

* $k \times t_{1/2} = 0.693$ → for all first order elimination.

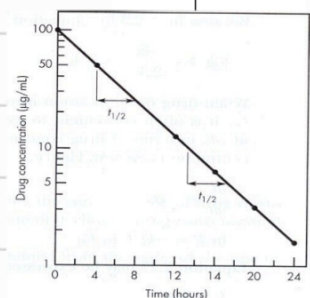
* half life is related to volume of distribution and drug clearance

$$CL = k \cdot V_d \Rightarrow t_{1/2} = \frac{0.693 \times V_d}{CL}$$

* there are 4 parameters of drugs

CL V_d $t_{1/2}$ k

| Half-lives | % of drug removed |
|------------|-------------------|
| 1 | 50 |
| 2 | 75 |
| 3 | 87.5 |
| 4 | 93.75 |

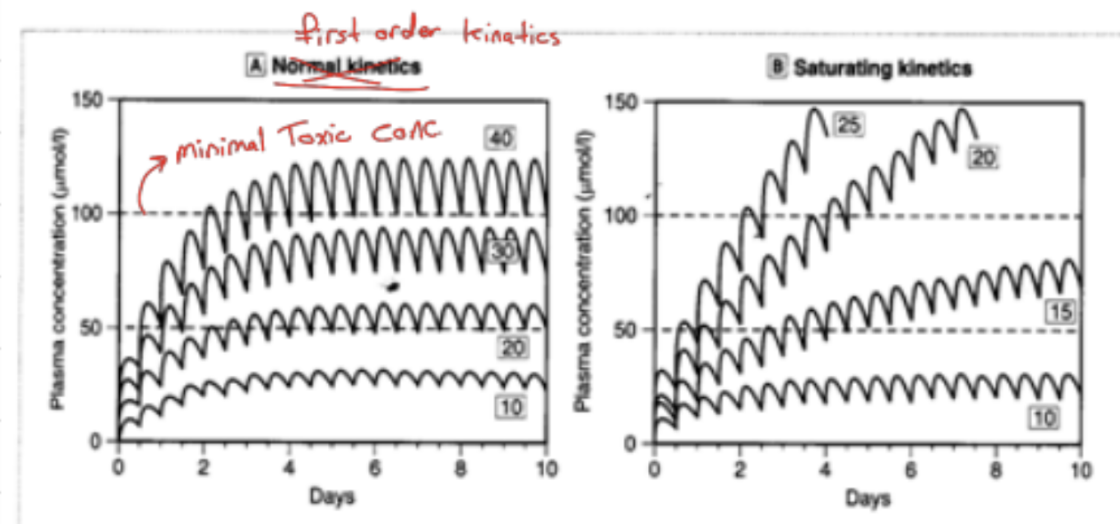
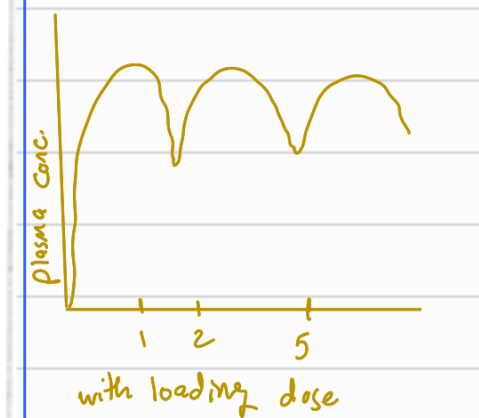
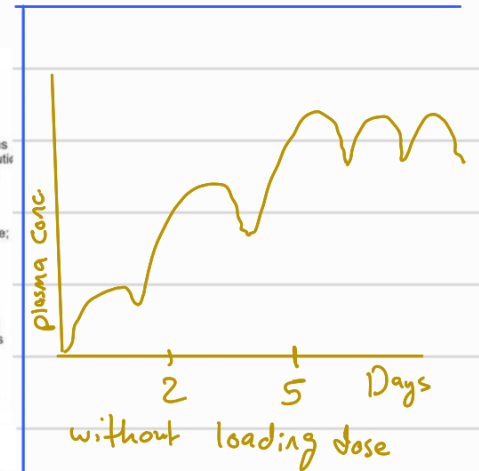
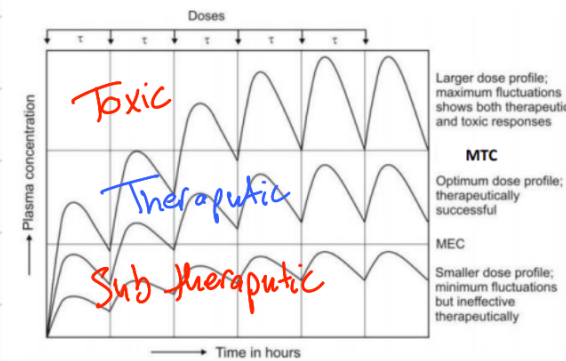
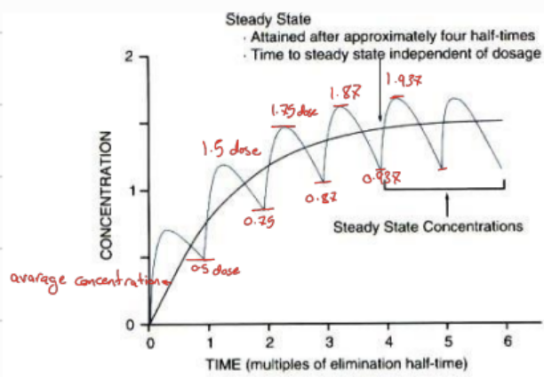


* in Zero-order elimination \Rightarrow Higher conc. \rightarrow longer Half life of elimination.
 dose and plasma conc. for drug under go zero order elimination \leftarrow

* Steady-State: Condition achieved following repeated drug administration

Simply it means state which drug is in static conc. in body

- * it is occur when Dosing rate = eliminating rate
- * in Steady-State a constant, trough and average drug conc. is achieved.
- * it is achieved after 4 repeated half lives of administration.
- * So our goal is to attain (C_{ss}) \Rightarrow steady-state conc. of drug in therapeutic range not below it (sub-therapeutic) or toxic one.



- * Loading dose
- when the half life is too long, steady state will take along time to be achieved.
- * So we may need to give a loading dose to achieve drug conc. within the therapeutic range sooner.

$$LD = V_d \cdot C_{ss}$$

* Maintenance Dose

\rightarrow to attain and maintain a desired C_{ss} of drug we need to adjust the dose so that, the drug administration is equal to the rate of drug elimination

$$MD = CL \cdot C_{ss}$$

\rightarrow also called target conc.

Done By: Abd Arrahman Dabbas.