## Pharmacokinetics: 1<sup>st</sup> order elim

In biologic systems, most drugs are eliminated via zero- or first- order kinetics. With zero order kinetics, a constant, certain mass of drugs are eliminated per unit time. This means the half-life of a substance is proportional to the concentration of the substance. Substances eliminated via zero-order kinetics include alcohol, phenytoin, and salicylates.

Most substances are eliminated via first-order kinetics in which the rate of clearance/elimination is proportional to the concentration of the substance. This means the half-life is a constant. It is possible for the concentration of some substances to be high enough that its clearance method becomes saturated and for a time, the substance follows zero-order kinetics.

Mathematically these kinetics can be described as:

Zero-order kinetics:

d[A]/dt = -k $[A]_t = [A]_0 - kt$ 

First-order kinetics:

$$\label{eq:alpha} \begin{split} d[A]/dt &= -k[A]\\ ln([A]_t) &= ln([A]_0)\text{-}kt\\ \text{Which can be rearranged to}\\ [A]_t &= [A]_0 e^{\text{-}kt} \end{split}$$

Where  $[A]_t$  is the concentration of a substance at time t,  $[A]_0$  is the starting concentration of a substance, and k is system-specific elimination constant.

Second and higher order reactions exist but are rare enough in anesthetic practice (and not an ABA keyword) that I wouldn't worry about them.

Further reading: Kim, TK. Shinju, O. Johnson, KB. <u>Miller's Anesthesia</u>, 9th ed. "Basic Principles of Pharmacology". 18: 462-482. Elsevier: 2020

Wikipedia: https://en.wikipedia.org/wiki/Rate\_equation