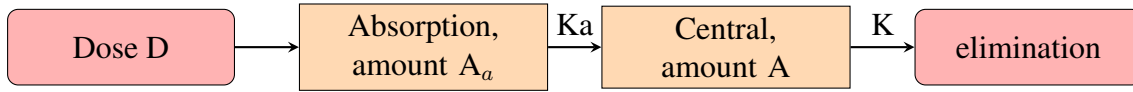


First order PO, ODE derivation



First order, thus ODE:

$$\begin{aligned} \frac{dA_a}{dt} &= -K_a A_a, & \text{IC : } A_a &= D \text{ when } t = 0; \\ \frac{dA}{dt} &= K_a A_a - K A, & \text{IC : } A &= 0 \text{ when } t = 0; \end{aligned}$$

$$\frac{d}{dt} \begin{bmatrix} A_a \\ A \end{bmatrix} = \begin{bmatrix} -K_a & 0 \\ K_a & -K \end{bmatrix} \begin{bmatrix} A_a \\ A \end{bmatrix}$$

Solve for eigenvalues and eigenvectors and we obtain

$$\begin{aligned} \lambda_1 &= -K_a, & \mathbf{v}_1 &= \begin{bmatrix} K - K_a \\ K_a \end{bmatrix} \\ \lambda_2 &= -K, & \mathbf{v}_2 &= \begin{bmatrix} 0 \\ 1 \end{bmatrix} \end{aligned}$$

Thus the equation is

$$\begin{bmatrix} A_a \\ A \end{bmatrix} = C_1 \mathbf{v}_1 \exp(-K_a t) + C_2 \mathbf{v}_2 \exp(-K t)$$

in which C_1 and C_2 are constants. Given the initial conditions, we could obtain $C_1 = \frac{D}{K - K_a}$ and $C_2 = \frac{-K_a D}{K - K_a}$, thus the final equation will be:

$$\begin{bmatrix} A_a \\ A \end{bmatrix} = \frac{D}{K - K_a} \begin{bmatrix} K - K_a \\ K_a \end{bmatrix} \exp(-K_a t) + \frac{-K_a D}{K - K_a} \begin{bmatrix} 0 \\ 1 \end{bmatrix} \exp(-K t)$$

We are interested in central compartment which could represent the plasma drug amount/concentration

$$\begin{aligned} A &= \frac{K_a D}{K_a - K} (e^{-K t} - e^{-K_a t}) \\ Conc &= \frac{A}{V_d} = \frac{K_a D}{(K_a - K) V_d} (e^{-K t} - e^{-K_a t}) \end{aligned}$$