

PHA 5127 Homework 1 solution:

1. (50)

a.  $k_e = -(\ln C_2 - \ln C_1) / (t_2 - t_1) = -(\ln 0.01 - \ln 0.0736) / (3 - 1) = 1 \text{ hr}^{-1}$

$t_{1/2} = 0.693 / k_e = 0.693 \text{ hr}$

b. Since  $C = C_0 e^{-k_e t}$ ,  $C_0 = C / e^{-k_e t} = 0.0736 / e^{-1 \cdot 1} = 0.2 \text{ mg/L}$

(Take 1 hr data point here. Any point is fine).

c.  $V = \text{Dose} / C_0 = 30 / 0.2 = 150 \text{ L}$

d. At  $t = 4 \text{ hr}$ ,  $C = C_0 e^{-k_e t} = 0.2 * e^{-4} = 0.00366 \text{ mg/L}$ .

Hence, the amount at 4 hr  $A = C * V = 0.00366 * 150 = 0.549 \text{ mg}$ .

e.

Time interval	AUC
0-0.5	0.0803
0.5-1	0.0487
1-2	0.0503
2-3	0.0185
3-∞	0.010
<b>0-∞</b>	<b>0.208</b>

2. (10)

It is a zero-order process and there is no half-life. Equal AMOUNT of drug is eliminated at equal time interval. At different times, half-lives will be different.

3. (20)

a.  $V_p(\leftrightarrow) + V_i(\leftrightarrow) \frac{F_u(\uparrow)}{F_{u,T}(\leftrightarrow)} = V(\uparrow)$

b.  $V_p(\leftrightarrow) + V_i(\leftrightarrow) \frac{F_u(\leftrightarrow)}{F_{u,T}(\downarrow)} = V(\uparrow)$

c.  $V_p(\leftrightarrow) + V_i(\leftrightarrow) \frac{F_u(\uparrow)}{F_{u,T}(\uparrow)} = V(\leftrightarrow)$

d.  $V_p(\leftrightarrow) + V_i(\leftrightarrow) \frac{F_u(\uparrow)}{F_{u,T}(\downarrow)} = V(\uparrow)$

4. (20)

X is permeability limited and Y is perfusion limited. Under physiological condition, most of Y will exist in unionized form and the permeability is not a limiting factor for Y due to its high partition coefficient. Drug X is exactly the opposite.