

PHA 5127
Answers Case Study 4
Fall 2007

1. For the following situations, indicate whether the drug is filtered, reabsorbed or actively secreted (Assume GFR is 130 mL min⁻¹, urine flow is 1.5 ml min⁻¹)

- A drug with $f_u = 0.02$ and a $Cl_{REN} = 20 \text{ mL min}^{-1}$ is _____
 (GFR* $f_u = 2.6 < Cl_{REN}$, Secreted)
- A drug with $f_u = 0.40$ and a $Cl_{REN} = 52 \text{ mL min}^{-1}$ is _____
 (GFR* $f_u = 52 = Cl_{REN}$, Filtered)
- A drug with $f_u = 0.30$ and a $Cl_{REN} = 0.45 \text{ mL min}^{-1}$ is _____
 (GFR* $f_u = 39 > Cl_{REN} = \text{Urine flow} * f_u = 1.5 * .3$, Complete Reabsorbed)

2. Calculate the k_e of a 56 year old, 79 kg male patient with a serum creatinine of 1.6 mg/dL who requires an aminoglycoside treatment. [Use $k_e = 0.00293 (CrCL) + 0.014 \text{ hr}^{-1}$].

Solution :

with Cockcroft-Gault Equation

$$CL_{Cr} = (140 - \text{age}) \cdot Wt / (72 \cdot \text{SerumCr}) = (140 - 56) \cdot 79 / (72 \cdot 1.6) = 57.6 \text{ ml / min}$$

$$k_e = 0.00293 (CrCL) + 0.014 \text{ h}^{-1} = 0.183 \text{ h}^{-1}$$

3. Mark whether the following statements for a **high extraction drug** are True or False

$$Cl = \frac{Q_H \cdot f_u \cdot Cl_{int}}{Q_H + f_u \cdot Cl_{int}} \approx Q_H \quad E = \frac{f_u \cdot Cl_{int}}{Q_H + f_u \cdot Cl_{int}} \approx 1 \quad F = 1 - E \approx \frac{Q_H}{f_u \cdot Cl_{int}}$$

- T F** The oral bioavailability (F) will be close to 1. (F)
- T F** Clearance will increase significantly after induction of the relevant enzyme. (F)
- T F** The hepatocyte membranes do not represent a barrier. (T)

4. Mark whether the following statements are True or False

- T F** a. Since creatinine is endogenous and predominantly eliminated by kidney, its clearance is a good estimation of renal active secretion. (F)
- T F** b. Creatine clearance can only be used to estimate the renal clearance of drugs that are similar to creatine, which does not show plasma albumin binding. (F)

T F c. “Linear pharmacokinetics” means that the plasma drug concentration versus time plots will result in a straight line. (F)