

Name: \_\_\_\_\_

OR

UFID #: \_\_\_\_\_

**PHA 5127**

**First Exam**

**Fall 2004**

On my honor, I have neither given nor received unauthorized aid in doing this assignment.

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Name

Put all answers on the bubble sheet. **If you need to comment or question a problem please note this on the front page.**

TOTAL \_\_\_\_\_/160 pts

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**Question Set I (True or False)**

(25 points)

**True (A) or False (B). On the bubble sheet mark A for true or B for false**

For a high extraction drug

- 1: T F Hepatic clearance will be larger than that of a low extraction drug
- 2: T F Hepatic clearance will depend on liver blood flow
- 3: T F Hepatic clearance will depend on plasma protein binding
- 4: T F Oral bioavailability will be low
- 5: T F Hepatic clearance will be determined by the GFR

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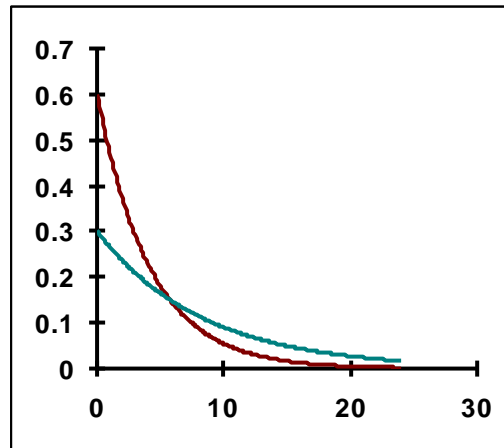
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**Question Set II**

(15 points)

Compare the following two concentration time profiles after a single bolus injection . The two lines differ in only one of the subsequent parameters. Please identify which parameter is different.



6: The 2 lines differ in:

- |    |                          |
|----|--------------------------|
| A. | <i>Parameter</i><br>Dose |
| B. | Vd                       |
| C. | Clearance                |

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**Question Set III (Matching)**  
(20 points)

For the physiological changes listed below, select the induced changes on the pharmacokinetic parameters for a lipophilic, uionizable (no acid or basic group in the molecule), protein bound **high extraction** drug that is also eliminated by renal elimination (only filtration, no reabsorption).

*Select the effect on kinetics*

(A)  $Cl_{REN} \downarrow$  (B)  $Cl_{HEP} \uparrow$  (C) oral bioavailability  $\downarrow$  (D)  $V_D \uparrow$  E. none of the listed answers

Physiological change

7: Increase in metabolic enzymes\_\_\_\_\_

8: Decrease in plasma protein binding \_\_\_\_\_

9: Increase in liver blood flow\_\_\_\_\_

10: Decrease in creatinine clearance\_\_\_\_\_

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**Question Set IV (Matching)**

(20 points)

(Assume GFR is  $130 \text{ mL min}^{-1}$ , urine flow is  $1.5 \text{ mL min}^{-1}$ ) For the following situations, indicate whether the drug is:

Select from the following choices:

(A) *only filtered*

(B) *filtered and reabsorbed through passive diffusion*

(C) *filtered and actively secreted*

(D) *filtered and reabsorbed through transporters*

11: A drug with  $f_u = 0.04$  and a  $Cl_{REN} = 40 \text{ mL min}^{-1}$  is \_\_\_\_

12: A drug with  $f_u = 0.20$  and a  $Cl_{REN} = 26 \text{ mL min}^{-1}$  is \_\_\_\_

13: A drug with  $f_u = 0.30$  and a  $Cl_{REN} = 0.45 \text{ mL min}^{-1}$  is \_\_\_\_

14: A drug with  $f_u = 1.0$  and a  $Cl_{REN} = 0.15 \text{ mL min}^{-1}$  is \_\_\_\_

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**Question Set V**

(20 points)

A drug is eliminated through glomerular filtration (no other clearance mechanisms is observed). **It does not bind to plasma proteins.** Glomerular filtration rate is normal (**130 ml/min**). No active renal secretion and passive or active reabsorption after renal filtration is observed. The volume of distribution is **50 L**.

15: What is the clearance? (10 points)

- A: 1.3 L/h
- B: 2.2 L/h
- C: 7.8 L/h
- D: 80 L/h

16: What is the  $k_e$  of the drug? (10 points)

- A:  $0.044 \text{ h}^{-1}$
- B:  $0.0260 \text{ h}^{-1}$
- C:  $0.1560 \text{ h}^{-1}$
- D:  $1.600 \text{ h}^{-1}$
- E:  $0.390 \text{ h}^{-1}$

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**Question Set VI**

(10 points)

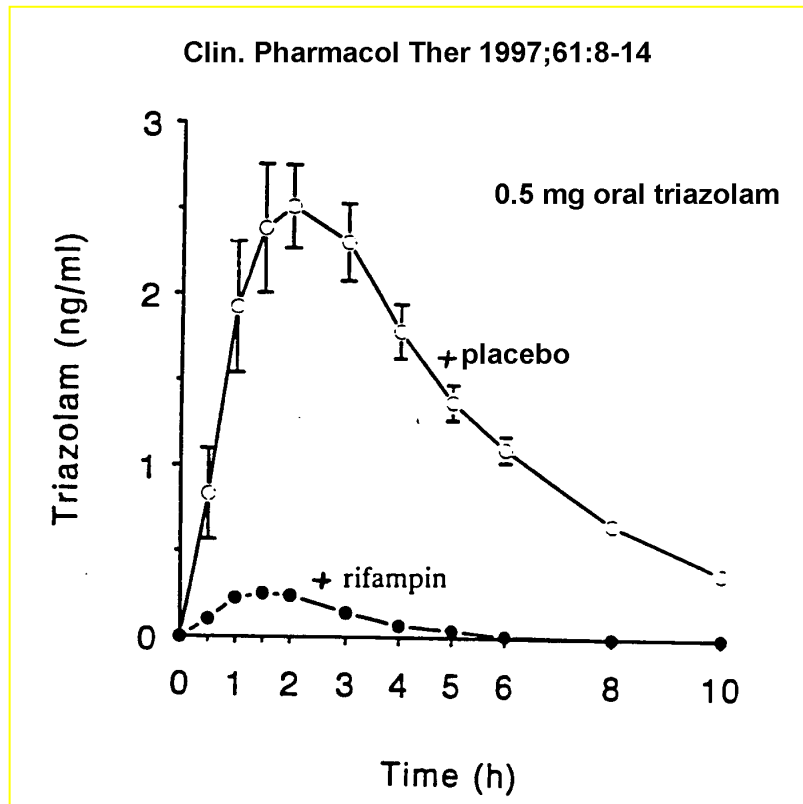
- 17: The nurses gave an iv bolus injection of an unknown drug at 7 a.m. They also did not record the dose. One hour after injection (8 a.m.) the concentration was found to be 6mg/L of plasma. Assume a  $k_e$  of  $0.150 \text{ h}^{-1}$ .

What would be the concentration at 8 pm?

- A 1.15 mg/L
- B 0.8 mg/L
- C 1.0 mg/L
- D 0.1 mg/L
- E 0.2 mg/L

**Question Set VII**

The same dose of triazolam was given either alone or with rifampin. Explain what is going on. (5 points)



Please choose the correct answers.

- 1: The clearance of triazolam is decreased in the presence of rifampin.
- 2: Triazolam is likely to be a high extraction drug.
- 3: Rifampin is an enzyme inducer.
- 4: Rifampin increases the volume of distribution of Triazolam.

18. Select the correct answer

- A: 1
- B: 1, 2
- C: 3
- D: 2, 3
- E: 4



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**Question Set VIII (True or False)**

(25 points)

**True (A) or False (B). On the bubble sheet mark A for true or B for false**

For a lipophilic unionized drug (no acid, or base)

- 19:    T     F    The renal clearance will depend on the tissue binding of the drug.
- 21:    T     F    The renal clearance will depend on plasma protein binding.
- 22:    T     F    Drinking a lot of water will increase the renal clearance.
- 23:    T     F    Involvement of renal transporters in the renal elimination of the drug is likely.
- 24:    T     F    The renal clearance will be smaller than the GFR.

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**Question Set IX (True or False)**

(20 points)

**True (A) or False (B). On the bubble sheet mark A for true or B for false**

- 25: T F The tissue uptake of a lipophilic unionized drug is more likely to be perfusion controlled.
- 26: T F The degree of plasma protein binding affects the metabolic clearance of all drugs that are metabolized in the liver.
- 27: T F Increase in plasma protein binding will decrease the volume of distribution of a lipophilic drug.
- 28: T F The renal clearance of a highly ionized drug is more likely to be affected by drug/drug interactions.