Name:	
OR UFID #:	
PHA 5127	
First Exam	
Fall 2004	
On my honor, I have neither given nor received unauthorized aid in doing this assignment.  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	

Name:	
	OR
UFID #:	

# **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

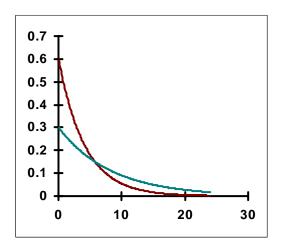
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# **Question Set II**

(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:

Parameter

- A.
- Dose

B.

Vd

C.

Clearance

Name:	
	OR
UFID #:	

# **Question Set III (Matching)**

(20 points)

For the physiological changes listed below, select the induced changes on the pharmacokinetic parameters for a lipophilic, unionizable (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

Physiol	logical	change
1 11 9 510	ogicar	change

- 7: Increase in metabolic enzymes\_\_C\_\_\_
- 8: Decrease in plasma protein binding \_\_C or D\_\_
- 9: Increase in liver blood flow\_**B**\_
- 10: Decrease in creatinine clearance\_A\_\_\_

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#### **Question Set IV (Matching)**

(20 points)

(Assume GFR is 130 mL min<sup>-1</sup>, urine flow is 1.5 ml min<sup>-1</sup>) 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  $Cl_{REN} = 40$  mL min<sup>-1</sup> is  $\underline{C}$
- 12: A drug with  $f_u = 0.20$  and a  $Cl_{REN} = 26$  mL min<sup>-1</sup> is  $\_A\_\_$
- 13: A drug with  $f_u = 0.30$  and a  $Cl_{REN} = 0.45$  mL min<sup>-1</sup> is \_B\_\_
- 14: A drug with  $f_u = 1.0$  and a  $Cl_{REN} = 0.15$  mL min<sup>-1</sup> is **\_D**\_\_\_

Name:	
	OR
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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 h<sup>-1</sup>
  - B: 0.0260 h<sup>-1</sup>
  - C: 0.1560 h<sup>-1</sup>
  - D: 1.600 h<sup>-1</sup>
  - E: 0.390 h<sup>-1</sup>

Name:	
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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  $0.150 \ h^{-1}$ .

What would be the concentration at 8 pm?

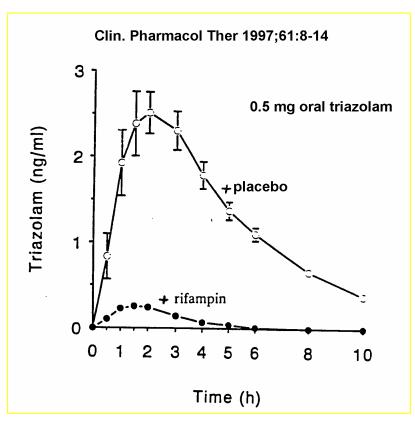
- A 1.15 mg/L
- B 0.8 mg/L
- C 1.0 mg/L
- D 0.1 mg/L
- E 0.2 mg/L

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

## **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: **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.