A Sample Lecture from an Organic Chemistry Course Designed for Pre-Med, Nursing, and Biology Students

Jason Dinsmore, Ph.D Facebook.com/OrganicChemistOnCall

# Design of Course

#### A Few Common Observations...

Organic chemistry is typically taught from a chemical point of view Organic chemistry is the one course *EVERYONE* has heard of and many <u>DREAD</u> taking Organic chemistry is usually a "make it or break it" class for pre-med students

### Does it have to be this way?

### **ABSOLUTELY NOT!!!**

You can learn the same material that has been taught semester after semester at universities around the world but from a medical/biological point of view

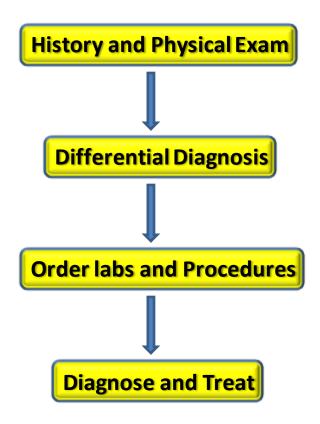
Or·gan·ic \or-'ga-nik\ adj: 1. of, relating to, or arising in a bodily organ; 2. of, relating to, or containing carbon compounds; 3. relating to, being, or dealt with by a branch of chemistry concerned with the carbon compounds of living beings and most other carbon compounds

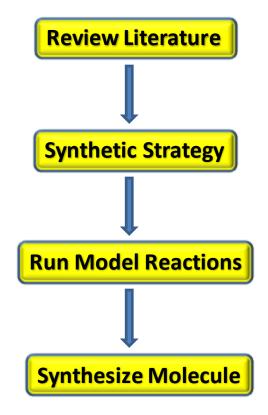
**Chem·is·try** \'ke-mə-str $\bar{e}$ \ n: a science that deals with the composition, structure, and properties of substances and with the transformations that they undergo

**Med·i·cine** \'me-də-sən\ n: **1.** a substance or preparation used in treating disease; **2.** the science and art dealing with the maintenance of health and the prevention, alleviation, or cure of disease

### MEDICINE

### ORGANIC CHEMISTRY





### The format of the lectures will be as follows:

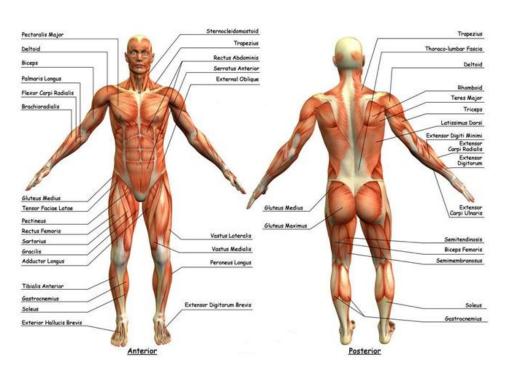
### Presentation of Case Study

- Ex. 24 year old female presents to the ED with...

### Discuss Functional Group(s)/Reaction(s)

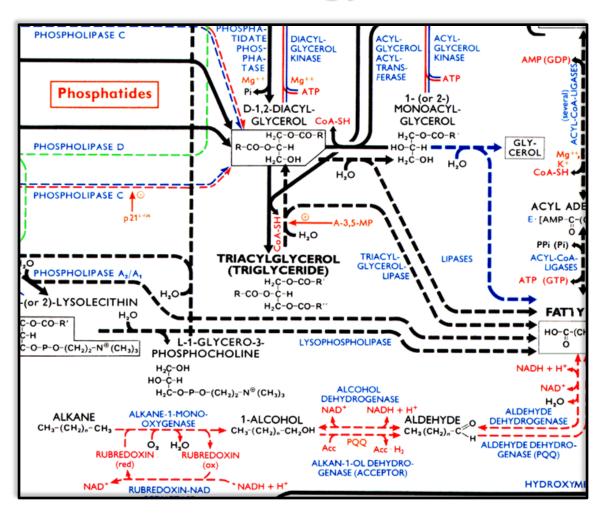
- How this functional group/reaction(s) relates to the disease

### Medicine



Blood, Plasma, Serum	Reference Range	SI Reference Intervals
<ul> <li>Alanine aminotransferase (ALT, GPT at 30°C)</li> </ul>	8-20 U/L	8-20 U/L
Amylase, serum	25-125 U/L	25-125 U/L
* Aspartate aminotransferase (AST, GOT at 30°C)	8-20 U/L	8-20 U/L
Bilirubin, serum (adult)		
Total // Direct	0.1-1.0 mg/dL // 0.0-0.3 mg/dL	2-17 µmol/L // 0-5 µmo
* Calcium, serum (Total)	8.4-10.2 mg/dL	2.1-2.8 mmol/L
* Chalesteral, serum	140-250 mg/dL	3.6-6.5 mmol/L
* Creatinine, serum (Total)	0.6-1.2 mg/dL	53-106 µmol/L
Electrolytes, serum		
Sodium	135-147 mEq/L	135-147 mmol/L
Chloride	95-105 mEq/L	95-105 mmol/L
* Potassium	3.5-5.0 mEq/L	3.5-5.0 mmol/L
Bicarbonate	22-28 mEq/L	22-28 mmol/L
Gases, arterial blood (room air)	an and Mar	
P <sub>O<sub>2</sub></sub>	75-105 mmHg	10.0-14.0 kPa
P <sub>CO<sub>2</sub></sub>	33-44 mmHg	4.4-5.9 kPa
pH	7.35-7.45	[H+] 36-44 nmol/L
* Glucose, serum	Fasting: 70-110 mg/dL	3.8-6.1 mmol/L
	2-h postprandial: < 120 mg/dL	< 6.6 mmol/L
Growth hormone - arginine stimulation	Fasting: < 5 ng/mL	< 5 µg/L
	provocative stimuli: > 7 ng/mL	> 7 µg/L
Osmolality, serum	275-295 mOsm/kg	275-295 mOsm/kg
Phosphatase (alkaline), serum (p-NPP at 30°C)	20-70 U/L	20-70 U/L
Phosphorus (inorganic), serum	3.0-4.5 mg/dL	1.0-1.5 mmol/L
* Proteins, serum		
Total (recumbent)	6.0-7.8 g/dL	60-78 g/L
Albumin	3.5-5.5 g/dL	35-55 g/L
Globulins	2.3-3.5 g/dL	23-35 g/L
Urea nitrogen, serum (BUN)	7-18 mg/dL	1.2-3.0 mmol urea/L
* Uric acid, serum	3.0-8.2 mg/dL	0.18-0.48 mmol/L
Cerebrospinal Fluid		
Glucose	40-70 mg/dL	2.2-3.9 mmol/L
Hematologic		
Erythrocyte count	Male: 4.3-5.9 million/mm3	4.3-5.9 × 1012/L
	Female: 3.5-5.5 million/mm <sup>3</sup>	3.5-5.5 × 1012/L
Hematocrit	Male: 41-53%	0.41-0.53
	Female: 36-46%	0.36-0.46
Hemoglobin, blood	Male: 13.5-17.5 g/dL	2.09-2.71 mmol/L
	Female: 12.0-16.0 g/dL	1.86-2.48 mmol/L

### **Biology**



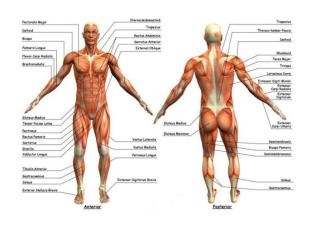
### **Organic Chemistry**

$$\begin{array}{c|c} OH & O \\ \hline CH_3-CH-CH_3 & \frac{K_2Cr_2O_7}{H_2SO_4} \rightarrow \begin{array}{c} CH_3-C-CH \\ \hline propanone \end{array}$$

$$\begin{array}{c} \text{OH} & \text{O} \\ | \\ \text{CH}_3 - \text{CH}_2 - \text{CH} - \text{CH}_3 & \xrightarrow{\text{KMnO}_4} & \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH}_3 \\ \text{2-butanol} & \text{80}^{\circ} & \text{2-butanone} \end{array}$$

$$\begin{array}{c} \text{OH} & \text{O} \\ | \\ \text{CH}_3 - \text{CH}_2 - \text{CH} - \text{CH}_3 & \hline{\text{CrO}_3} \\ \text{2-butanol} & \text{CH}_3\text{COOH} \end{array} \\ \begin{array}{c} \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH} \\ \text{2-butanone} \end{array}$$

#### Medicine



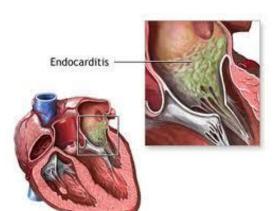


#### **Organic Chemistry**

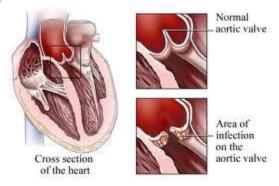
$$\begin{array}{c} \text{OH} & \text{OH} \\ -1 & \text{CH}_3 - \text{CH} - \text{CH}_3 \\ 2\text{-propanol} & \frac{\text{K}_2\text{Cr}_2\text{O}_7}{\text{H}_2\text{SO}_4} \\ \text{Propanone} & \text{CH}_3 - \text{C} - \text{CH} \\ -1 & \text{CH}_3 - \text{CH}_2 - \text{CH} - \text{CH}_3 \\ 2\text{-butanol} & \frac{\text{KMnO}_4}{\text{H}_2\text{SO}_4} \\ \text{SO}^\circ & \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH}_3 \\ -1 & \text{CH}_3 - \text{CH}_2 - \text{C} + \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH}_3 \\ -1 & \text{CH}_3 - \text{CH}_2 - \text{C} + \text{CH}_3 - \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH}_3 \\ -1 & \text{CH}_3 - \text{CH}_2 - \text{C} + \text{CH}_3 - \text{CH}_3 - \text{CH}_2 - \text{C} - \text{CH}_3 \\ -1 & \text{CH}_3 - \text{CH}_2 - \text{C} + \text{$$

### Case 1

A 30-year-old man is evaluated in the emergency department for complications of difficulty breathing, chills, and chest pain for the past 24 hours. He denies any previous history of medical problems. On physical examination, he appears ill. His temperature is 40°C (104°F), his blood pressure is 90/50 mm Hg, and his heart rate is 110/min. Cardiac examination reveals a 3/6 diastolic murmur; however, the patient denies any history of a murmur. ECG results are normal.



### **Diagnosis: Acute Endocarditis**



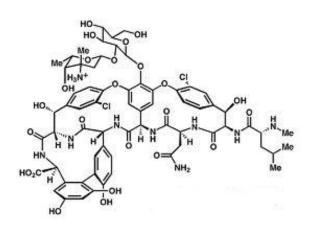
How does this case relate to organic chemistry?

## Treatment of Endocarditis

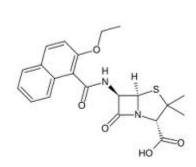
#### **CEFTRIAXONE**

$$R_1$$
  $H_2N$   $H_2N$   $H_2N$   $H_2N$   $H_3C$   $H$ 

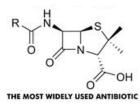
**GENTAMICIN** 



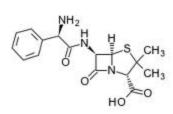
**VANCOMYCIN** 



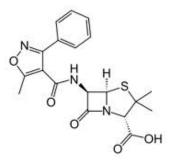
**NAFCILLIN** 



#### **PENICILLIN**



#### **AMPICILLIN**

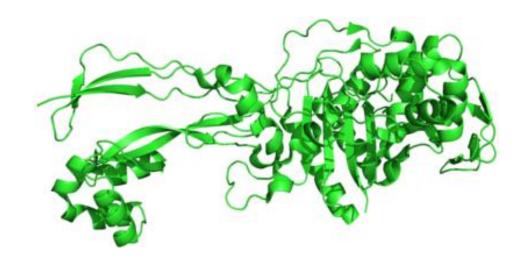


#### **OXACILLIN**

### Penicillin

 $\beta$ -lactam structure is involved in the mechanism of action

### Penicillin Mechanism of Action



#### **Penicillin Binding Protein**

- PBPs are all involved in the final stages of the synthesis of peptidoglycan, which is the major component of bacterial cell walls.
- Bacterial cell wall synthesis is essential to growth, cell division (thus reproduction) and maintaining the cellular structure in bacteria.
- Inhibition of PBPs leads to irregularities in cell wall structure such as elongation, lesions, loss of selective permeability, and eventual cell death and lysis.

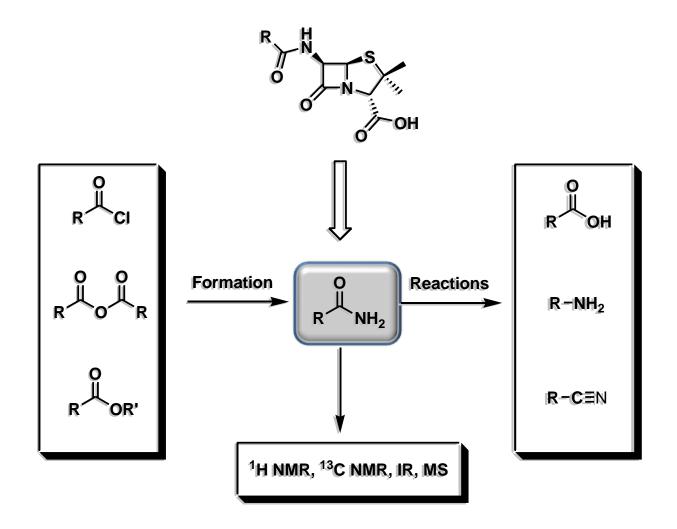
## Penicillin Mechanism of Action

**Nucleophilic Attack on the Cyclic Amide Carbonyl Carbon** 

# The Chemistry of Amides

Biology sets the reaction up perfectly; however, this is not the case in a chemistry laboratory....

# The Chemistry of Amides



## The Formation of Amides

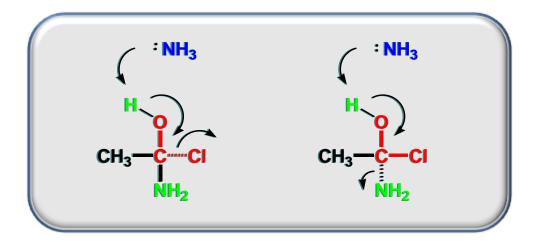
$$CH_3$$
  $-C$   $-CI$  + 2  $NH_3$   $-C$   $-NH_2$  +  $NH_4$   $+ CI$  Acid Chloride

$$CH_3-\overset{O}{C}-O-\overset{O}{C}-CH_3 + 2 NH_3 \longrightarrow CH_3-\overset{O}{C}-NH_2 + CH_3-\overset{O}{C}-O^- NH_4^+$$
 Anhydride

$$CH_3-C-O-CH_2CH_3 + NH_3 \longrightarrow CH_3-C-NH_2 + HO-CH_2CH_3$$
Ester

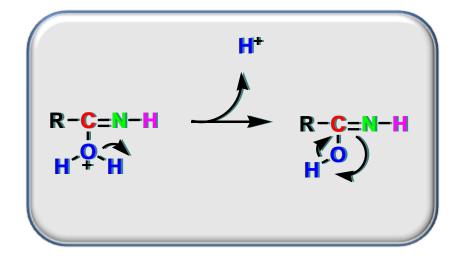
### Mechanisms





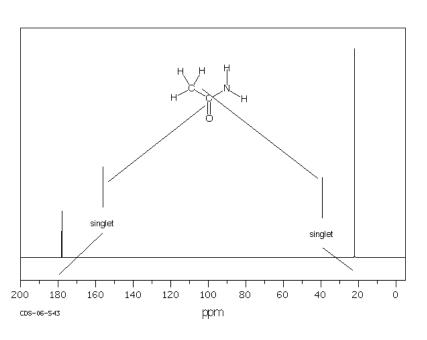
Halides are great leaving groups

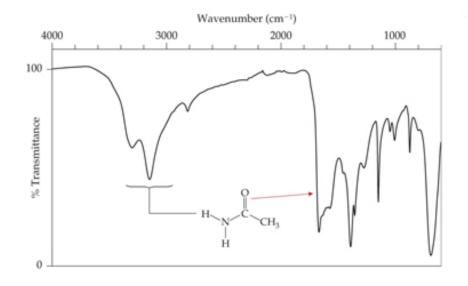
### Mechanisms



Regenerated acid catalyst

### 13 CNMR and IR of Amide





#### 13C NMR spectra

Carbonyl carbon: 180 ppm

Alkyl carbon: 20 ppm

#### IR spectra

Carbonyl group: 1700 cm<sup>-1</sup>

N-H group: 3200 cm<sup>-1</sup>

- Physicians obtain laboratory values and perform diagnostic tests
- Organic chemists obtain spectroscopic data and perform model reactions

# The Chemistry of Amides

What compounds can we make from the amide functional group?

# The Chemistry of Amides

O  

$$CH_3 - \overset{\circ}{C} - NH_2 + Br_2 + 2NaOH \longrightarrow CH_3 - \frac{NH_2}{Amine} + CO_2$$

# Stability of Amides

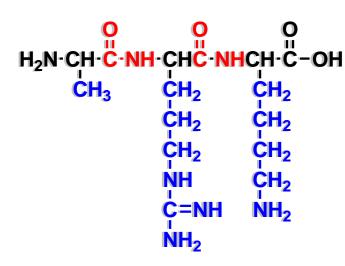
Less electrophilic that other carbonyl functional group

### Amides and Amino Acids

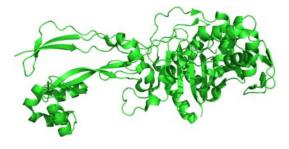
**Alanine** 

**Arginine** 

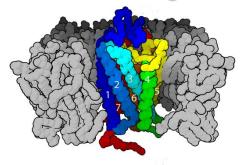
## Amides and Amino Acids







**Penicillin Binding Protein** 



**G-coupled Protein Receptor** 



Na/K Channel