#### **CONCEPT:** INTRAMOLECULAR ACID/BASE CATALYSIS

• Intramolecular Catalyst: An acid/base or nucleophilic catalyst present within the reacting molecule.

### **Intramolecular General Acid Catalysis**

Acidic groups can catalyze reactions by intramolecular \_\_\_\_\_\_.

**EXAMPLE:** Write a plausible mechanism for the following esterification reaction.

## STEP 1: Protonation and Nucleophilic Attack: Carbonyl group is protonated through intramolecular protonation.

☐ Methanol molecule attacks the protonated carbonyl group.

STEP 2: Proton Transfer: An H+ is transferred from the \_\_\_\_\_oxy O to a \_\_\_\_oxy O.

STEP 3: Leaving Group and Proton Transfer: Carbonyl group is reformed, and \_\_\_\_\_ is kicked out.

☐ Final product is formed after an intramolecular proton transfer.

#### **CONCEPT:** INTRAMOLECULAR ACID/BASE CATALYSIS

### **Intramolecular General Base Catalysis**

• Basic groups can catalyze hydrolysis reactions by \_\_\_\_\_ of water.

Intramolecular General Base Catalysis							
			Reaction				Relative Rate
	+	H <sub>2</sub> O		OH	+	но	
	+	H <sub>2</sub> O	Weak base	OH	+	но	
O Na <sup>+</sup>	+	H <sub>2</sub> O	<b>→</b>	O O Na <sup>+</sup>	+	но	

**EXAMPLE:** Write a plausible mechanism for the following hydrolysis reaction.

# STEP 1: Deprotonation and Nucleophilic Attack: Carboxylate anion deprotonates H<sub>2</sub>O to produce \_\_\_\_\_.

□ OH⁻ ion attacks the carbonyl group to form a tetrahedral intermediate.

# STEP 2: Leaving Group and Intramolecular Proton Transfer: Alkoxy \_\_\_\_ kicks out the phenoxide ion.

□ An intramolecular H+ transfer takes place as the phenoxide is kicked out.

## **CONCEPT:** INTRAMOLECULAR ACID/BASE CATALYSIS

**PRACTICE:** Which of the following compounds should undergo hydrolysis faster at neutral pH? Write mechanism for the hydrolysis of that compound.

**PRACTICE:** The following compound, when heated in a solution, undergoes hydrolysis without an acid catalyst. Write a plausible mechanism for the hydrolysis reaction.