

Summary

- 1) Here's an important general rule to be followed in writing all reaction mechanisms. If a reaction is executed under acidic conditions, no strong bases can be present. Any negatively charged species must be a weak base (e.g., Cl^-). If a reaction is executed under basic conditions, no strong acids can be present. Any positively charged species must be a weak acid.
- 2) Under basic conditions, a hydroxyl group cannot be substituted directly because HO^- is too poor of a leaving group. The hydroxyl group must first be transformed into a derivative that is a suitable leaving group.
- 3) Under acidic conditions, the $\text{R}_3\text{C}-\text{X}^+$ bond can ionize spontaneously to give $\text{X}:$ and R_3C^+ . The likelihood of ionization depends on the stability of the carbocation product, the leaving group and the solvent.
- 4) Under acidic conditions, a lone pair nucleophile bearing a proton (e.g., H_2O , ROH , RCO_2H , RNH_2) is always deprotonated after it adds to the electrophile (e.g., carbocation).
- 5) Under acidic conditions, epoxides open by attack of the nucleophile onto the protonated ring. The nucleophile attacks the most highly substituted carbon.
- 6) Under basic conditions, epoxides open by nucleophilic attack onto the least substituted carbon (the reaction resembles an $\text{S}_{\text{N}}2$ reaction; the leaving group is a poor one, but the process is driven by the release of ring strain).
- 7) If carbocations are intermediates, rearrangement is possible.