

- 1) synthetic utility: alkyl halide \rightarrow alcohol
ether
thiol

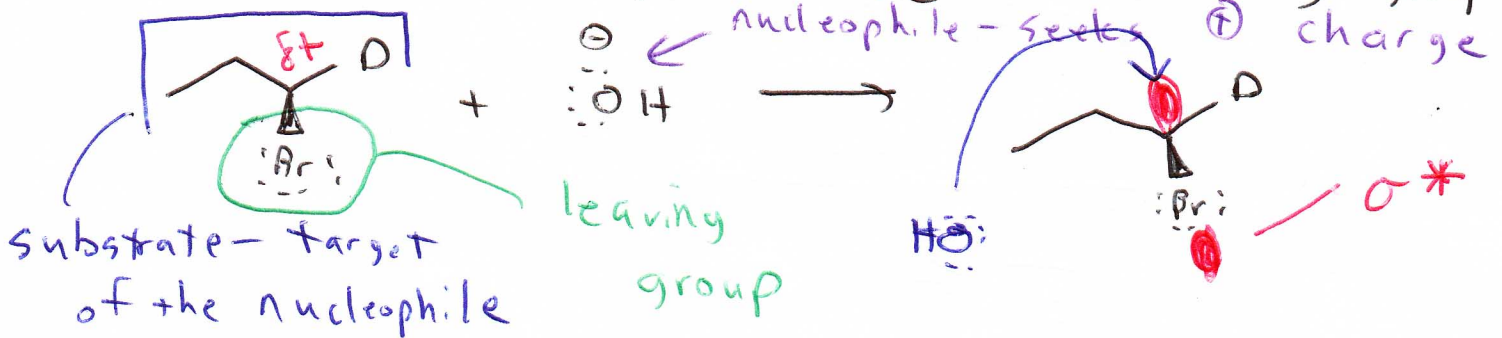
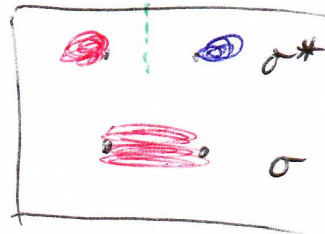
2) reagents: basic nucleophile
ex: NaOH, NaOCH₃, NaCN

3) conditions: polar aprotic solvent

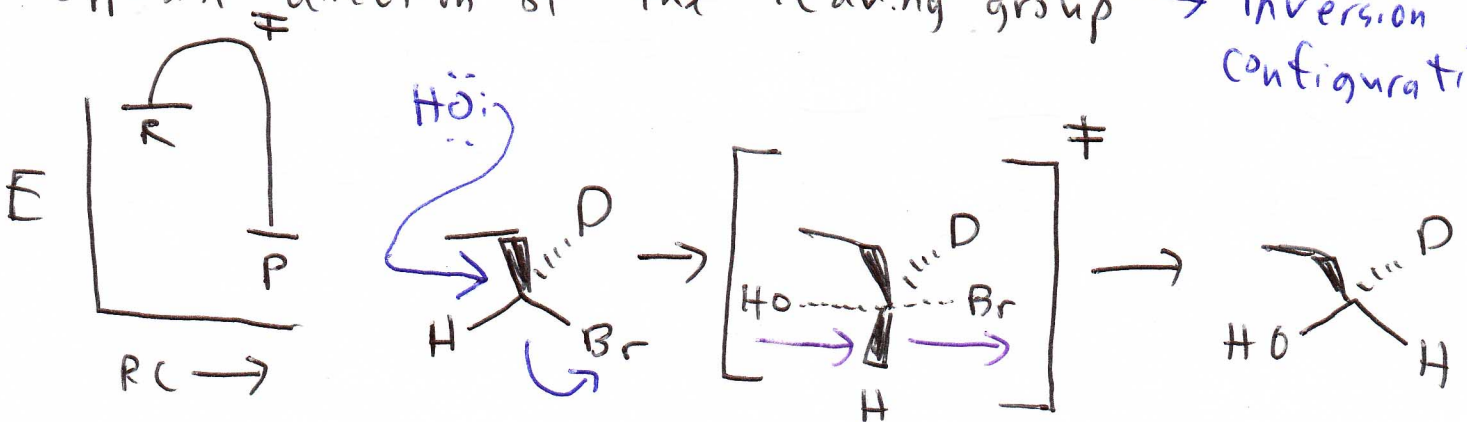
4) mechanism: anionic, concerted

5) stereochemistry: inversion of configuration

6) regiochemistry: specific (@ site of leaving group)

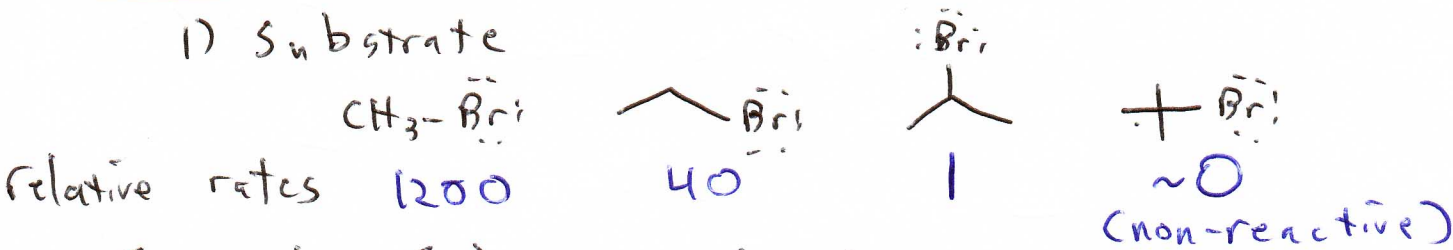


If a S_N2 rxn, the nucleophile attacks by adding into the σ^* bond of the leaving group. Since e^- are added into the antibonding orbital, the bond with the leaving group breaks, this allows a new bond with the nucleophile to form, but in the exact opposite direction of the leaving group \rightarrow inversion of configuration

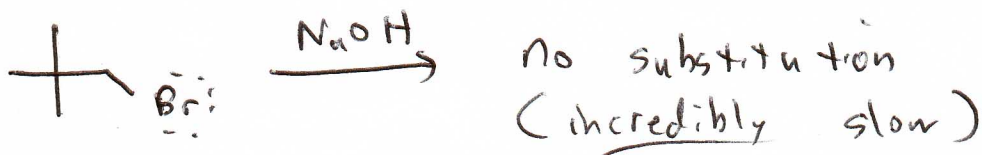


Factors that affect an S_N2 rxn

1) Substrate



Since the S_N2 rxn depends on the nucleophile directly attacking the carbon bearing the leaving group, steric hindrance around that position will slow the rate of rxn (or prevent it)



Nu: phenyl \equiv benzene substituent
phenyl halides do not undergo S_N2 because of too much steric hindrance (attack would have to occur through the ring),

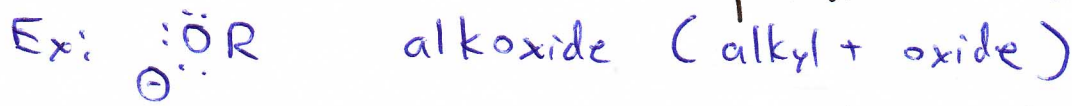
2) Nucleophiles

Nucleophilicity - how strongly does a compound seek \oplus charge,



hydroxide is much more nucleophilic because it has a \ominus charge on it,

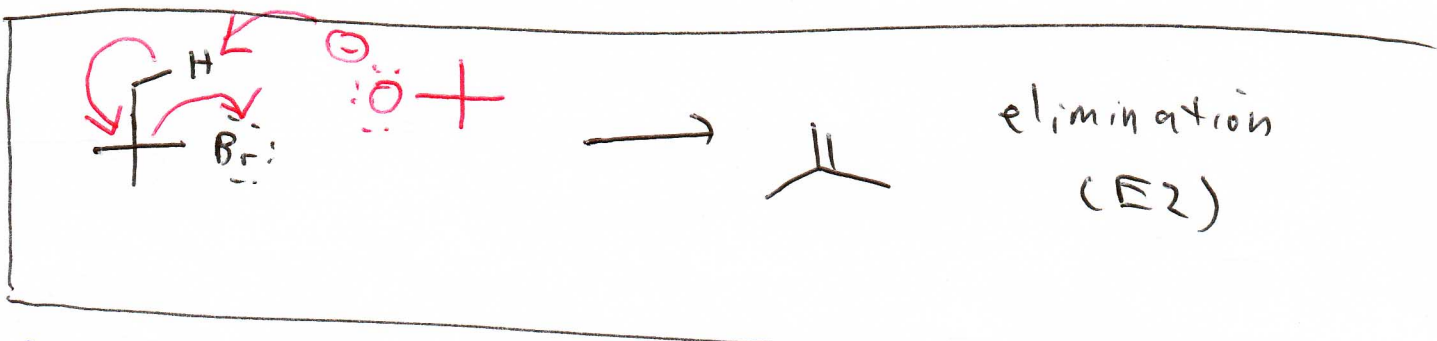
basicity - how able is a compound to remove H^+



OH^-	vs	SH^-		H_2O	$\text{pK}_a = 15.74$
Stronger base		stronger nucleophile		H_2S	$\text{pK}_a = 7.00$

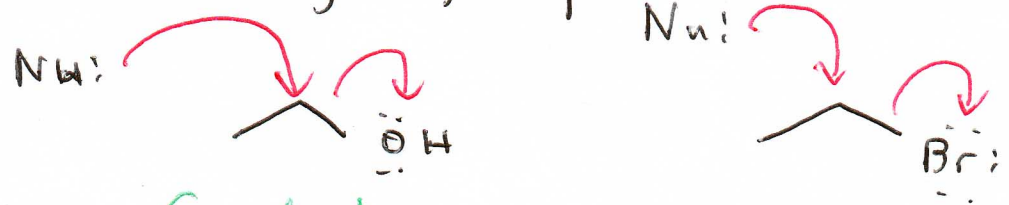
polarizability - able to become polarized -
larger ions are "squishier," meaning that electron density can be more easily displaced.

Since sulfur is more polarizable than oxygen, the process of forming a bond occurs more easily - and therefore more quickly - for sulfur versus oxygen → sulfur is more nucleophilic than oxygen (in similar molecules).



S_N2 rxns require strong nucleophiles, which are often, but not always, basic. * Extremely basic nucleophiles, especially sterically hindered ones, can undergo elimination versus substitution.

3) leaving group



Good leaving groups are conjugate bases of strong acids.