



	S <sub>N</sub> 2 and E2	S <sub>N</sub> 1/E1
<b>mechanism</b>	one step—this single step is the rate-determining step (RDS)	two steps—RDS is formation of carbocation
<b>big obstacle</b>	S <sub>N</sub> 2: steric hindrance blocking Nu (Nu is in RDS) E2: blocking B isn't a big obstacle (B doesn't join substrate)	stabilizing carbocation (Nu/B isn't in RDS, so blocking it isn't an obstacle)
<b>stereo-chemistry</b>	S <sub>N</sub> 2: inversion (backside attack, since LG blocks frontside) E2: cis vs. trans determined by anti-periplanar transition-state	S <sub>N</sub> 1: racemization (planar carbocation intermediate) E1: both cis and trans isomers will be produced
<b>regio-chemistry</b>	E2: possible products from deprotonation of any β-carbon major product w/ bulky base: less substituted (steric hindrance) major product with non-bulky base: more substituted	E1: possible products from deprotonation of any β-C major product: more substituted alkene (e <sup>-</sup> -donating alkyl substituents stabilize alkenes)
<b>rate expression</b>	Rate = $k$ [substrate] [Nu <sup>-</sup> or B <sup>-</sup> ], so [Nu <sup>-</sup> /B <sup>-</sup> ] ↑ → rate ↑ (substrate and Nu <sup>-</sup> /B <sup>-</sup> are in RDS)	Rate = $k$ [substrate], so [Nu <sup>-</sup> /B <sup>-</sup> ] ↑ → rate unchanged (only the substrate is in RDS)
<b>Nu quality</b>	requires good Nu/strong B (Nu/B is in RDS) bulky Nu/B favors E2 vs. S <sub>N</sub> 2 (blocking B isn't a big obstacle)	can work with a poor Nu/weak B (Nu/B isn't in RDS)
<b>LG quality</b>	requires good leaving group (because leaving group is in RDS)	requires good leaving group (because LG is in RDS)
<b>preferred solvent?</b>	polar aprotic (no O-H or N-H bonds) (for S <sub>N</sub> 2, hydrogen-bonds to solvent would block Nu) (for E2, protic solvent would protonate the base)	polar protic (at least one O-H or N-H bond) (hydrogen-bonds to solvent stabilize carbocation)
<b>substrate</b>	S <sub>N</sub> 2: methyl > 1° > 2°; 3° gives no S <sub>N</sub> 2 (substituents block Nu) E2: 1°, 2°, or 3° (blocking B is not a big obstacle)	3° > 2°; methyl and 1° give no S <sub>N</sub> 1/E1 (alkyl substituents stabilize the carbocation)

comparing the same element

	charge	resonance
<b>nucleophilicity</b>	negative charge → better Nu	resonance → worse Nu (charge is stabilized)
<b>basicity</b>	negative charge → stronger base	resonance → weaker base (charge is stabilized)
<b>leaving-group ability</b>	positive charge → better LG (more willing to accept electrons)	resonance → better LG (charge will be stabilized)

comparing different elements

	same row	same column
<b>big difference</b>	electronegativity	size
<b>nucleophilicity</b>	less electronegative → better Nu (willing to donate electrons)	bigger → better Nu (usually) (big Nu's are less hindered by solvent, more polarizable)
<b>basicity</b>	less electronegative → stronger base (willing to donate electrons)	bigger → weaker base (large base can spread out and stabilize electron density)
<b>leaving-group ability</b>	more electronegative → better LG (willing to accept electrons)	bigger → better leaving group (big LG's can spread out and stabilize electron density)

nucleophiles, leaving groups, bases

nucleophiles				leaving groups			
N	O	F	N <sup>-</sup> O <sup>-</sup> F <sup>-</sup>	N	O	F	N <sup>+</sup> O <sup>+</sup> good LG
P	S	Cl	P <sup>-</sup> S <sup>-</sup> Cl <sup>-</sup>	P	S	Cl	P <sup>+</sup> S <sup>+</sup> not a LG
	Se	Br	Se <sup>-</sup> Br <sup>-</sup>			Br	
	I		I <sup>-</sup>			I	
cyanide NC <sup>-</sup> (charge on the C) azide N <sub>3</sub> <sup>-</sup>				sulfonate $\text{R}-\text{S}(=\text{O})_2-\text{OR}$ The $\alpha$ carbon is attached to the oxygen, not to the sulfur.			
bases				Nucleophiles and bases shown with charges before attacking. Leaving groups shown with charges before leaving. The tables for individual atoms assume no resonance. Resonance makes atoms into worse nucleophiles and bases and into better leaving groups.			
N	O	F	N <sup>-</sup> O <sup>-</sup> strong base (E2)				
P	S	Cl	P <sup>-</sup> S <sup>-</sup> Cl <sup>-</sup> weak base (E1)				
		Br	Br <sup>-</sup> not a base				
		I	I <sup>-</sup>				

what happens in S<sub>N</sub>2, S<sub>N</sub>1, E2, and E1 mechanisms

	what happens	big obstacle
<b>S<sub>N</sub>2</b>	One step: Nucleophile joins $\alpha$ carbon and leaving group leaves $\alpha$ carbon	steric hindrance
<b>S<sub>N</sub>1</b>	Step one: Leaving group leaves $\alpha$ carbon Step two: Nucleophile joins $\alpha$ carbon	stabilizing the carbocation
<b>E2</b>	One step: Base takes $\beta$ hydrogen, $\pi$ bond forms between $\alpha$ and $\beta$ carbons, leaving group leaves $\alpha$ carbon.	none
<b>E1</b>	Step one: Leaving group leaves $\alpha$ carbon Step two: Base takes $\beta$ hydrogen, $\pi$ bond forms between $\alpha$ and $\beta$ carbons	stabilizing the carbocation

how to determine S<sub>N</sub>2 vs. E2 vs. S<sub>N</sub>1 vs. E1 for haloalkane and alkylsulfonate substrates

	poor Nu / weak base O with no formal charge	good Nu / weak base Cl <sup>-</sup> , Br <sup>-</sup> , I <sup>-</sup> , NC <sup>-</sup> , N <sub>3</sub> <sup>-</sup> , S <sup>-</sup> , Se <sup>-</sup> , or CH <sub>3</sub> COO <sup>-</sup> or N, S, or Se with no formal charge	good Nu / strong base N <sup>-</sup> , O <sup>-</sup>
<b>methyl <math>\alpha</math>-carbon</b> <b>1° <math>\alpha</math>-carbon</b>	no reaction	S <sub>N</sub> 2 <sup>1</sup>	E2 with <i>tert</i> -butyl-oxide (bulky base) <sup>2</sup> Otherwise, S <sub>N</sub> 2
<b>2° <math>\alpha</math>-carbon</b>	95% S <sub>N</sub> 1 5% E1 (usually not shown)	S <sub>N</sub> 2 <sup>1</sup>	E2
<b>3° <math>\alpha</math>-carbon</b>	95% S <sub>N</sub> 1 5% E1 (usually not shown)	95% S <sub>N</sub> 1 5% E1 (usually not shown)	E2

For cases with “95% S<sub>N</sub>1, 5% E1”, E1 products are generally not shown unless the problem specifies “all possible products”.

<sup>1</sup>No reaction if beta-carbon is 4°.

<sup>2</sup>S<sub>N</sub>2 for methyl  $\alpha$ -carbon.

The table displays the major reaction(s) for each case—in some cases there may be significant levels of other competing reactions.

This table may not give the correct answer in all real-world situations, but it will generally be accurate for the questions that are typical of exams.