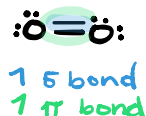
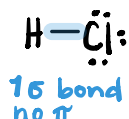
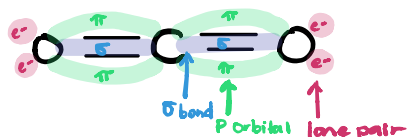
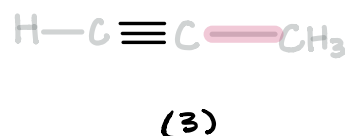
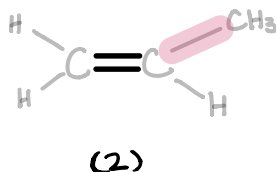
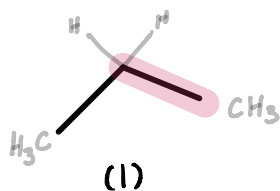


BONDING



Q: Which of the following bonds are strongest?



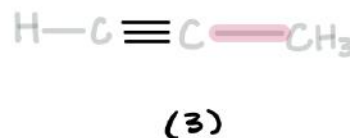
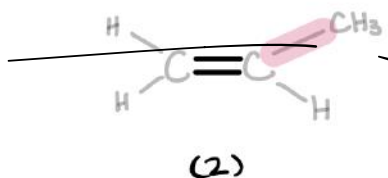
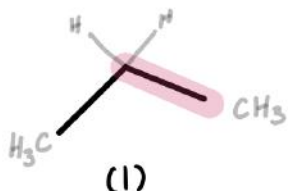
A: Propyne (3) is shorter and stronger than the others.

This is because its orbitals sp and sp^3 have the most overlap. Propene (2) is the next strongest followed by propane (1).

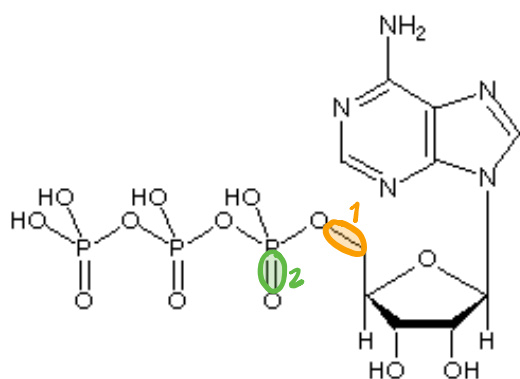
Q: Are "pi" bonds or "sigma" bonds stronger?

A: $\pi < \sigma$ **HOWEVER** $1\sigma + 1\pi$ is stronger than a single σ bond.

Q: Which bond is the most reactive?



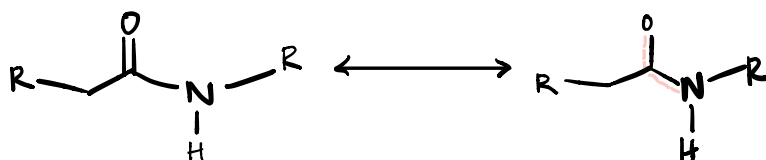
A: 3! triple bonds have an extra π bond which tends to react.



Q: Why in this ATP molecule is bond 1 more likely to dissociate than bond 2?

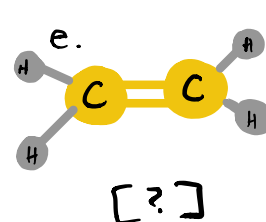
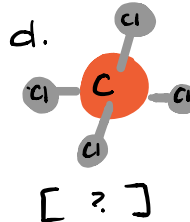
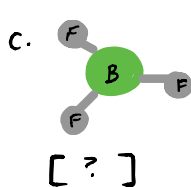
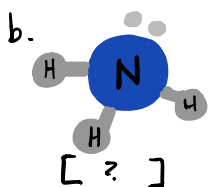
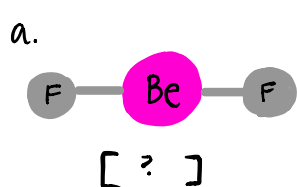
A: There are several reasons but the bond energy of bond 2 is much higher making it harder to break than bond 1.

Q: Do peptide bonds rotate?



A: No! Resonance gives double bond character to both the C=O and C-N bonds.

HYBRIDIZATION



a. [sp]

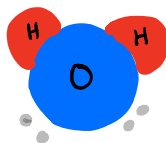
b. [sp³]

c. [sp²]

d. [sp³]

e. [sp²]

* This phenomenon occurs due to high and low energy orbitals mixing.



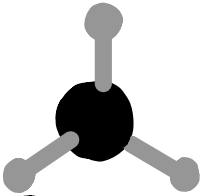
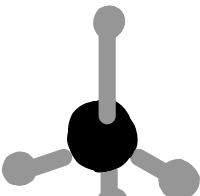
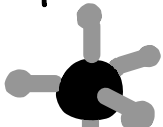
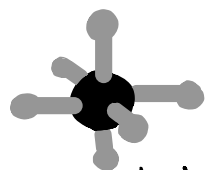
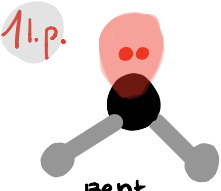
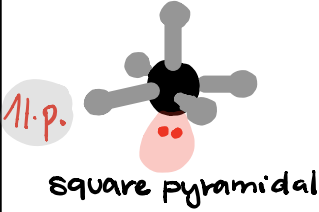
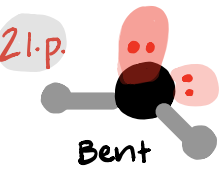
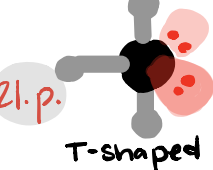
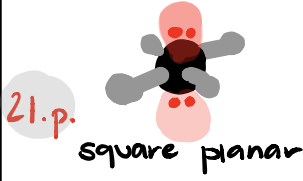
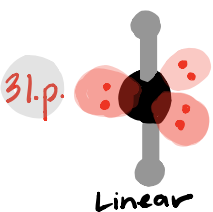
Sp³ hybridized
Q: What is the "s" character?

A: 25% "s" character
 +
 75% "p" character

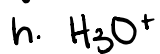
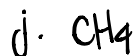
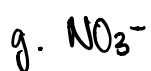
$$[\underset{\uparrow 1}{s} + \underset{\uparrow 1}{p} + \underset{\uparrow 2}{p} + \underset{\uparrow 3}{p} = sp^3]$$

$= 1/4 = 25\% s$

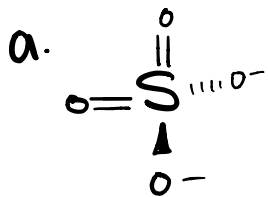
Shapes

sp^2	sp^3	sp^3d	sp^3d^2
<u>TRIGONAL PLANAR</u>	<u>TETRAHEDRAL</u>		<u>OCTAHEDRAL</u>
 Trigonal planar	 Tetrahedral	 Trigonal bipyramidal	 Octahedral
 Bent	 Trigonal pyramidal	 Seesaw	 Square pyramidal
	 Bent	 T-shaped	 Square planar
		 Linear	

Q: What shape do the following molecules take:



Answers



Tetrahedral



sp^3d^2 - Square pyramidal



$\text{Br} = 7 \text{ VE}$

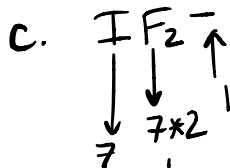
$\text{F} = 7 \text{ VE}$

$$7(1) + 7(5) =$$

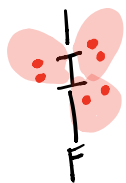
$$35 + 7 = 42 - 40 = 2e^- = 1 \text{ l.p.}$$

lone pairs = 1.p.

(multiple of)
8-octet



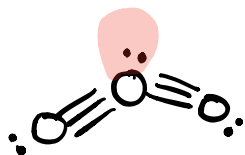
$$1 + 1 + 7 = 22 - 16 = 6e^- = 3 \text{ l.p.}!$$



sp^3d
linear



$$6 \times 3 = 18 - 16 = 2e^- = 1 \text{ l.p.}$$

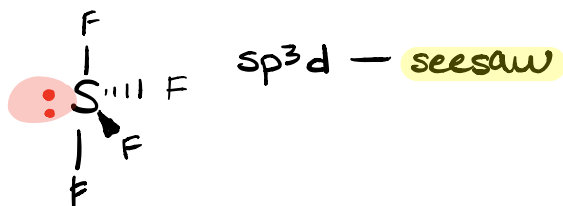


sp^2 - Bent

e. SF_4



$$6 + 4(7) = 28 + 6 = 34 - 32 = 2e^- = 1 \text{ p.}$$



f. SOF_4

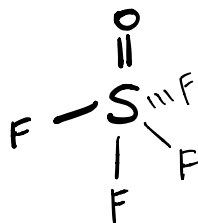


$$6 + 6$$

$$+ 7 \times 4$$

$$12$$

$$+ 28 = 40 - 40 = 0e^- = 0 \text{ p.}$$



sp^3d - trigonal bipyramidal

g. NO_3^-

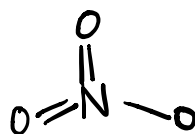


$$5 + (6 \times 3) + 1$$

$$5 + 18 + 1 = 24 - 24 = 0e^- = 0 \text{ p.}$$

$$\uparrow$$

(8×3)



sp^2 - trigonal planar

h. H_3O^+

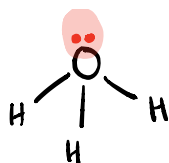


$$(1 \times 3)$$

$$+ 6$$

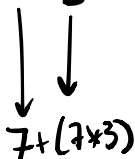
$$+ 1$$

$$1 + 3 + 6 = 10 - 8 = 2e^- = 1 \text{ p.}$$

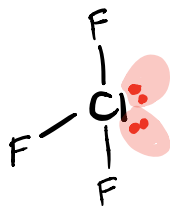


sp^3 — trigonal pyramidal

i. ClF_3

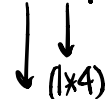


$$7 + 21 = 28 - 24 = 4e^- = 2 \text{ l.p.}$$

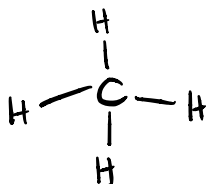


sp^3d — T-shaped

j. CH_4

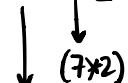


$$4 + 4 = 8 - 8 = 0e^- = 0 \text{ l.p.}$$



sp^3 — tetrahedral

k. $SnBr_2$



$$4 + 14 = 18 - 16 = 2e^- = 1 \text{ l.p.}$$



sp^2 — Bent

Single bonds

Double bonds

Triple bonds



STABILITY



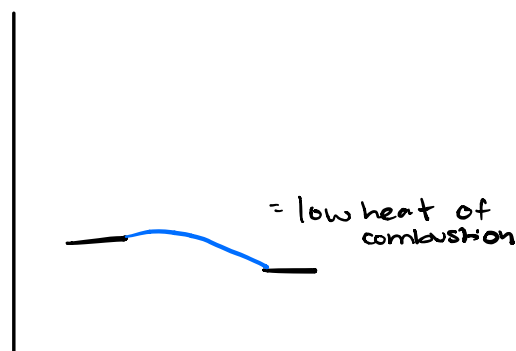
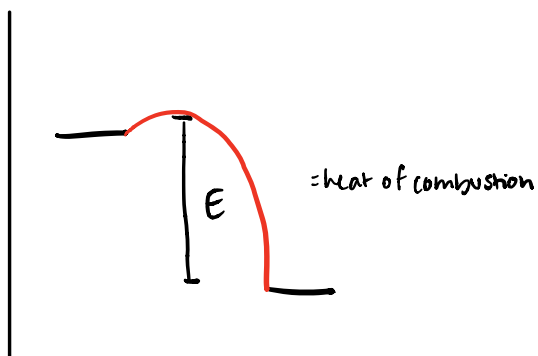
REACTIVITY



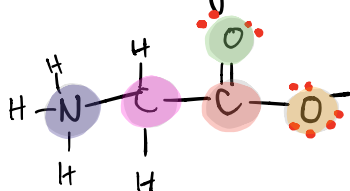
STRENGTH



Bond strength = harder to break (shorter)



Formal Charge of glycine



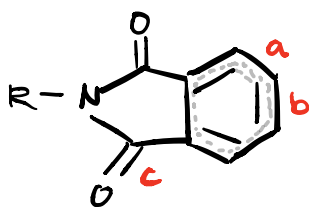
$$N = 5 - 4 = \text{FC of } 1+$$

$$C = 4 - 4 = \text{FC of } 0$$

$$C = 4 - 4 = \text{FC of } 0$$

$$O = 6 - 6 = \text{FC of } 0$$

$$O = 6 - 7 = \text{FC of } -1$$



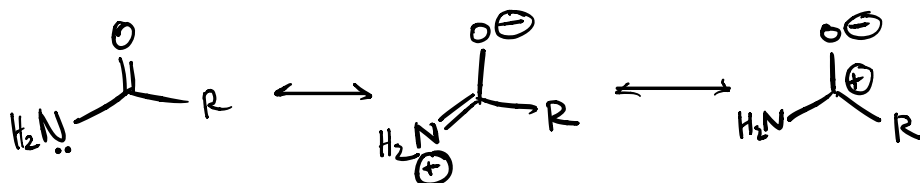
Q: Order the bonds from longest to shortest:

Longest $\longrightarrow$ shortest

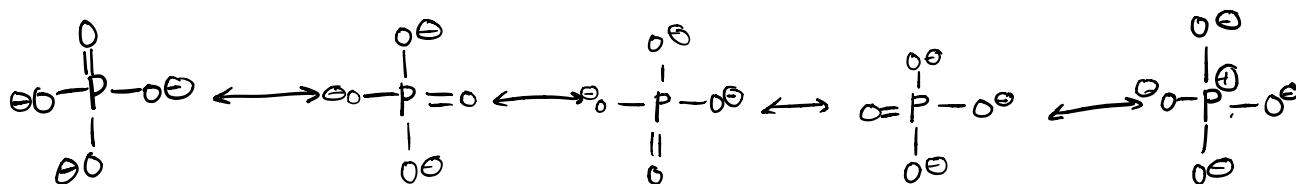
A: $a = b > c$
 $\uparrow$
 due to resonance

Q: Draw all resonance forms for:

a. $R-\text{CONH}_2$ [Amide]



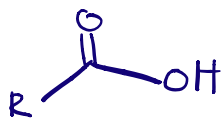
b. PO_4^{3-} [Phosphate]



NOMENCLATURE

priority (IUPAC)

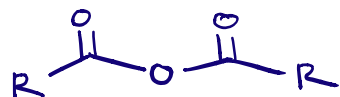
(1) Carboxylic acid



Prefix: carboxy

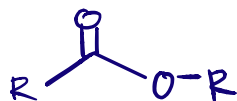
Suffix: oic acid

(2) acid anhydride



Suffix: oic anhydride

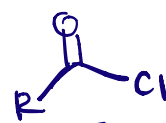
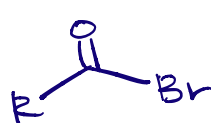
(3) Esters



Prefix: alkoxy carbonyl

Suffix: oate

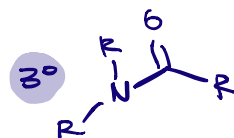
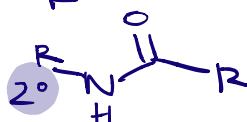
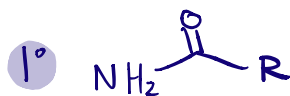
(4) Acyl Halides



Suffix: *oyl chloride*

(5) Amides

Prefix: *Carbamoyl*
Suffix: *amide*



(6) Nitriles



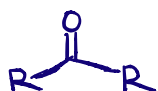
Prefix: *Cyano*
Suffix: *Nitrile*

(7) Aldehydes



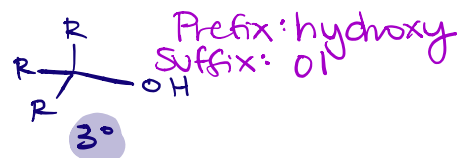
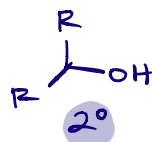
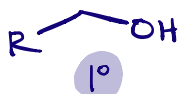
Prefix: *oxo*
Suffix: *al*

(8) Ketones



Prefix: *oxo*
Suffix: *al*

(9) Alcohols



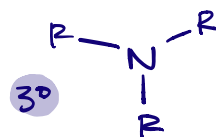
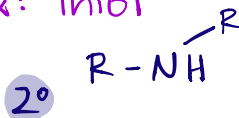
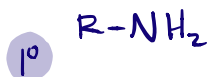
Prefix: *hydroxy*
Suffix: *ol*

(10) Thiol



Prefix: *mercapto*
Suffix: *thiol*

(11) Amines



Prefix: *amino*
Suffix: *amine*

(12) Sulfides



(13) Alkenes



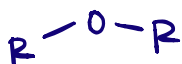
Prefix: *enyl*
Suffix: *ene*

(14) Alkynes



Prefix: *ynyl*
Suffix: *yne*

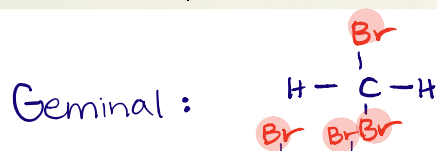
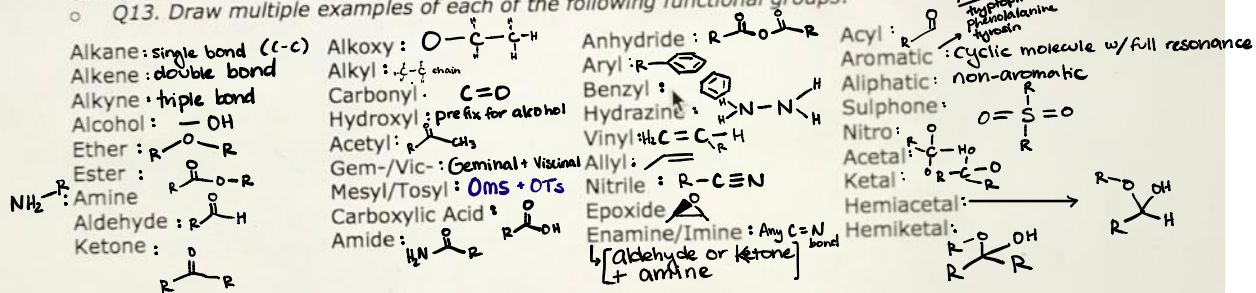
(15) Alkanes



Prefix: *yl*
Suffix: *ane*

Organic Nomenclature

- > **Functional Groups: You must memorize all of these!** The MCAT will refer to functional groups by either their names or their structures and expect you to understand.
- o Q13. Draw multiple examples of each of the following functional groups:



Mesylate: Add a methyl group ($-\text{OSO}_2\text{CH}_3$) also known as Oms
 ↳ excellent leaving group. (+ no acidic proton)

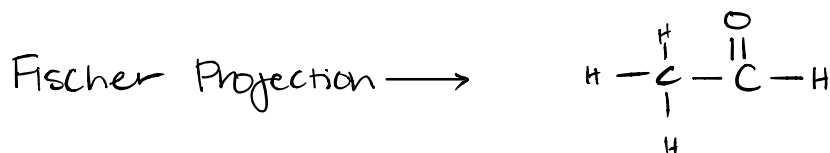
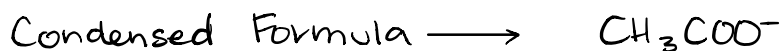
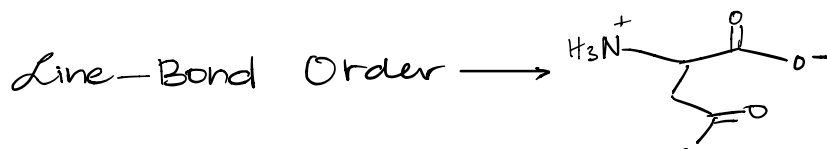
Tosylate: OTs (similar to Oms)

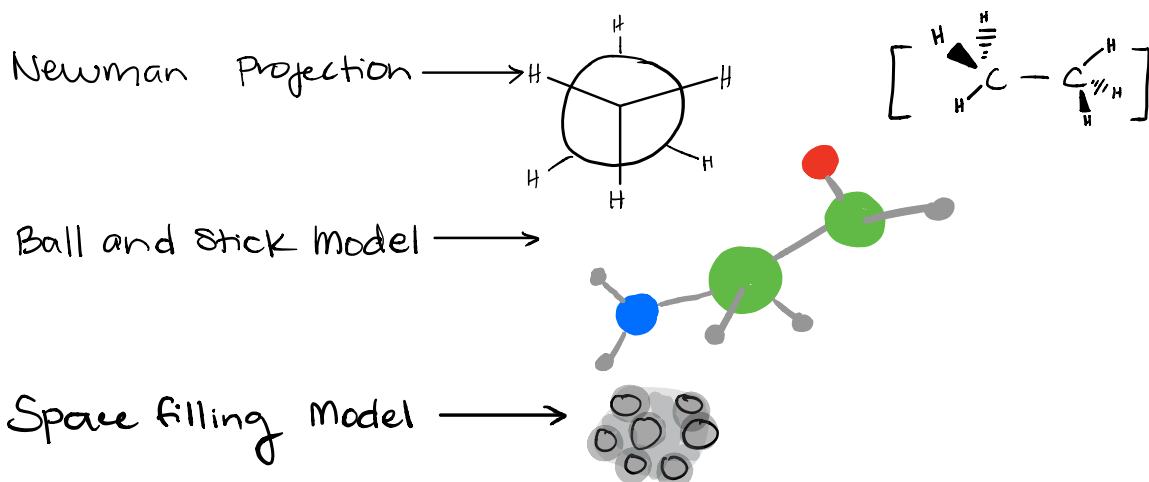
Aryl: Any functional group derived from an aromatic ring
 $\text{R}-\text{C}_6\text{H}_5$

Vinyl: $=$ group

Allyl: $=$ group

Q: Lewis Dot Structure $\longrightarrow$ $\text{H}-\ddot{\text{O}}-\text{H}$



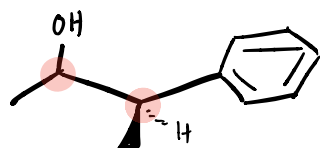


Isomers:

* Same molecular formula, different compounds

Q: How do we determine how many isomers something has?

A: 2^n (Where n = # chiral centers)



How many stereoisomers?

$$2^n = 2^2 = \boxed{4}$$

#1 Conformational isomers
(NOT true isomers)

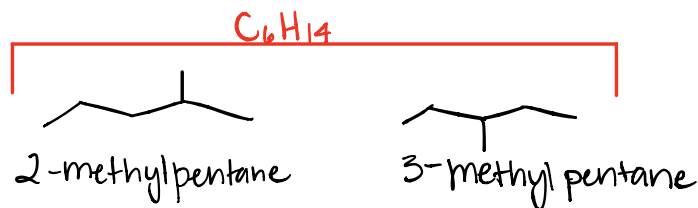
Conformational isomers are the exact same molecule.

Butane



#2 Structural Isomers

Same formula but different connectivity



#3 Stereoisomers

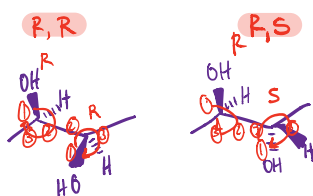
Same formula, same connectivity, different 3D arrangements

→ Enantiomers: same molecular formula + same connectivity but are non-identical.
Non-super imposable mirror images

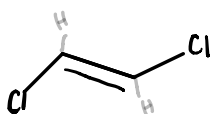
(R+S)



→ Diastereomers



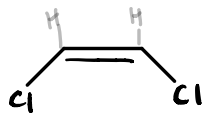
Geometric Isomers: (cis/trans)



trans
[E]

↑
no dipole moment

↑
no steric hindrance

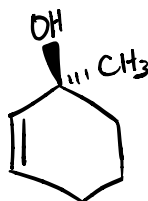


cis
[Z]

↑
often have a dipole moment

↑
often experience "steric hindrance" (↑ energy)

practice assigning



Reaction Types

E1: Elimination Rxn ~ Unimolecular

1st order (one species is concentrated)

2 steps

① Dissociation of leaving group, form carbocation

② Abstract a proton + form double bond

slow

fast

E2: Elimination Rxn ~ Bimolecular

2nd order (two species are concentrated)

1 step

① Abstraction of a proton + electron collapse to form a double bond and ejection of the leaving group.

SN1: Substitution Rxn ~ Unimolecular

1st order

2 steps

① Dissociation of leaving group and formation of carbocation

② Nucleophilic attack of the carbocation

SN2: Substitution Rxn ~ Bimolecular

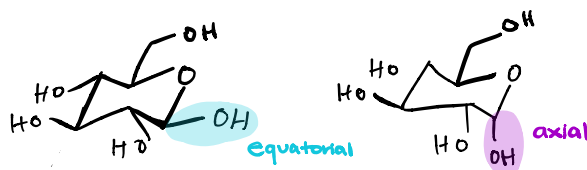
2nd order (two involved concentrated species)

1 step

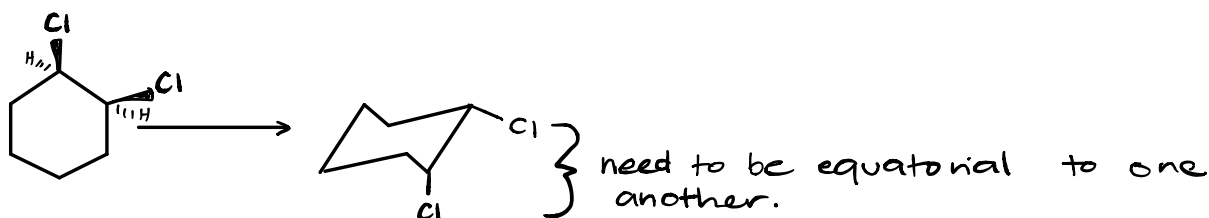
① Single step back side attack of electrophile + ejection of leaving group.

	#Steps	Rxn order	Carbocation?	methyl or hydride shifts?	Product Stereochemistry	Favored by:
E1	2	1 st	yes	yes	planar	weak bases 3° carbons polar protic
SN1	2	1 st	yes	yes	racemic mix	poor nucleophiles 3° carbons polar protic solvents
E2	1	2 nd	no	no	planar	Strong and/ or bulky bases
SN2	1	2 nd	no	no	inversion of configuration	good nucleophiles methyl 1° or 2° carbons

Cyclic Compounds :



a large substitute would be most comfortable in the axial position as it eliminates repulsion.



COMBUSTION OF ALKANES



- ↑ energy of activation
- exothermic

Q: Will N_2 have a high heat of combustion?

- Stable molecule = low heat of combustion
- N_2 = stable
- N_2 = low heat of combustion



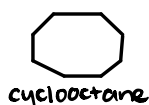
(1)



(2)



(4)



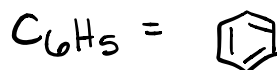
(3)

Highest heat of combustion = 1
Lowest = 4

Radicals = single unpaired electron (NOT ON MCAT)

→ Free radicals may be a consequence of cell damage

Donating groups donate their electron density into a conjugated π system



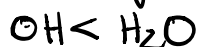
Unsaturated

BP ↑ with ↑ MW
BP ↓ with ↑ branches

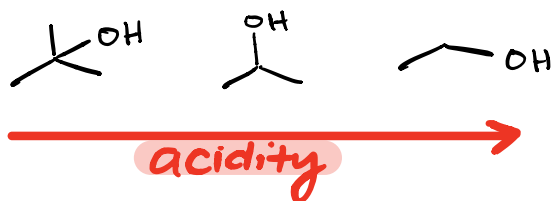
Q: Why is alcohol highly soluble in water?

A: Polarity, hydrogen bonding

Acidity



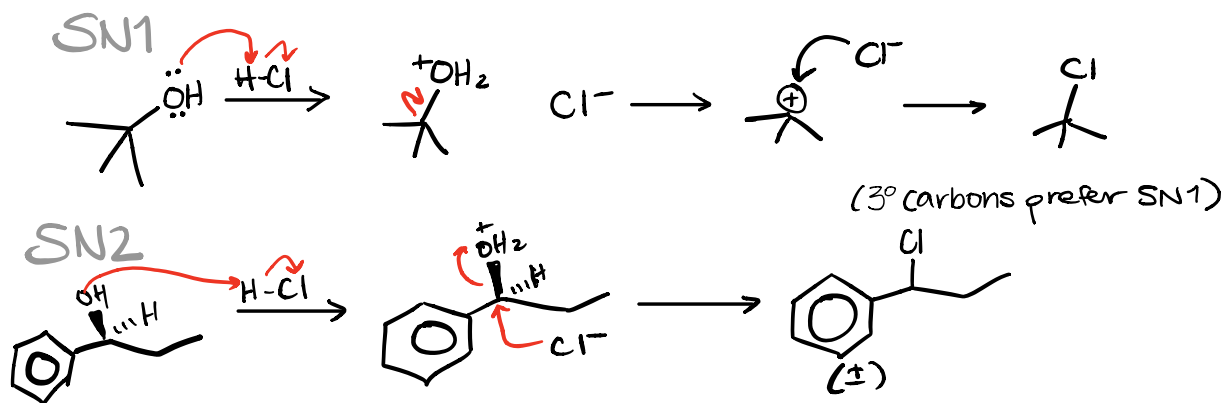
H^+ = acidic so H_2O has more proton than OH^- and OH^- causes an increase in pH.



This is because at less substituted areas the proton can be more easily donated (leaving O^-) whereas the $(-)$ charge on a tertiary molecule would be crowded and repulsive and less stable. This is due to the alkyl groups being weak electron donating group

REACTIONS [Galore]

Formation of an alkyl Halide from an alcohol



Oxidation of alcohols

1° alcohols $\rightarrow$ aldehydes $\rightarrow$ carboxylic acids

2° alcohols $\rightarrow$ ketones

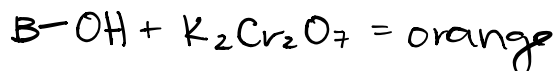
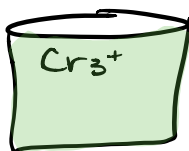
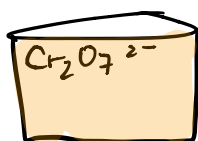
3° alcohols $\rightarrow$ cannot be further oxidized

Common Oxidizing Agents:

- O_3	- Jones
- Cr_2O_7	- Collins
- CrO_4	- PCC
- $KMnO_4$	- PDC

Q: Q28. Potassium dichromate is often used as an oxidizing agent. A solution of dichromate ions ($Cr_2O_7^{2-}$) is orange, but a solution of chromium ions (Cr^{3+}) is green. If alcohol A is reacted with potassium dichromate and produces a green solution, and alcohol B is reacted with the same reagent and produces an orange solution, what can be inferred about the two alcohols, A and B?

A:



dichromate ions are becoming chromium ions by oxidation. so unreacted dichromate is orange alcohol B is likely tertiary as it did not oxidize (react). Alcohol A is likely 1° or 2°.

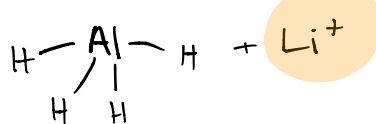
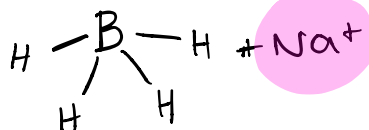
Reduction synthesis of an alcohol

*Reducing agents do the opposite of oxidizing agents

Reducing agents:

- $NaBH_4$ (only can reduce aldehydes + ketones)
 - $LiAlH_4$
 - H_2 + pressure
- } can reduce aldehydes, ketones, carboxylic acids and esters.

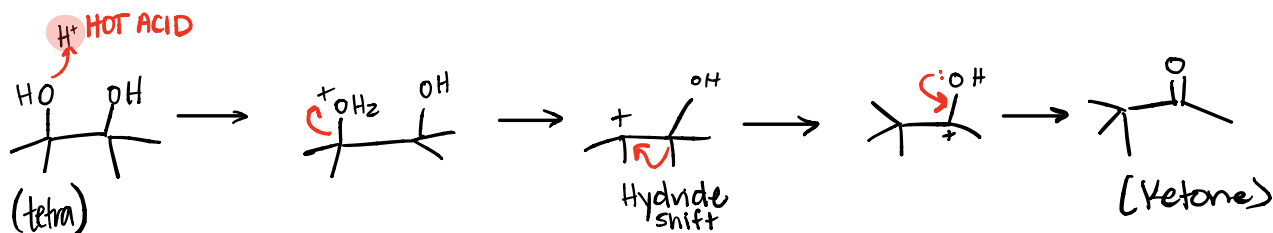
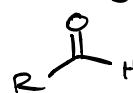
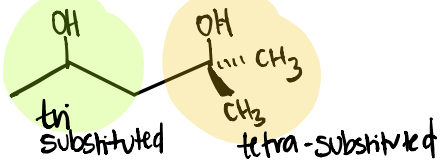
sp^3d



Boron is more electronegative, keeping the electron density inwards making the hydrides less nucleophilic. LiAlH_4 is a stronger reducing agent therefore as its hydrides can reduce "2x".

Pinacol Rearrangement

Vic-diol + hot acid $\longrightarrow$ Ketone or aldehyde



Alcohol Protection

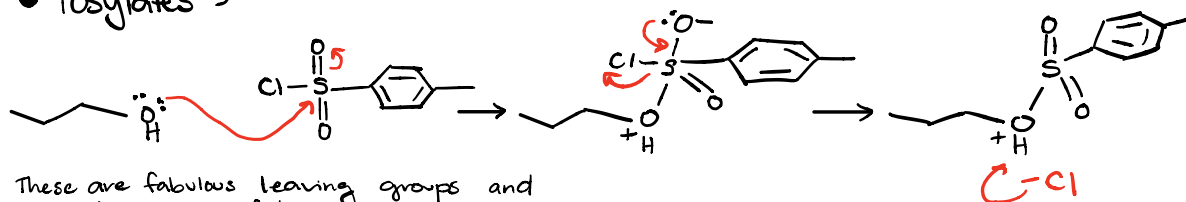
* used to protect alcohols from oxidation or reaction but allowing other groups on the molecule to react.

#1) TMS

#2) MOM

& acidification removes protecting group and restores alcohol.

- Mesylates
 - Tosylates
- } $\text{S}_{\text{N}}2$



These are fabulous leaving groups and are used because of this characteristic

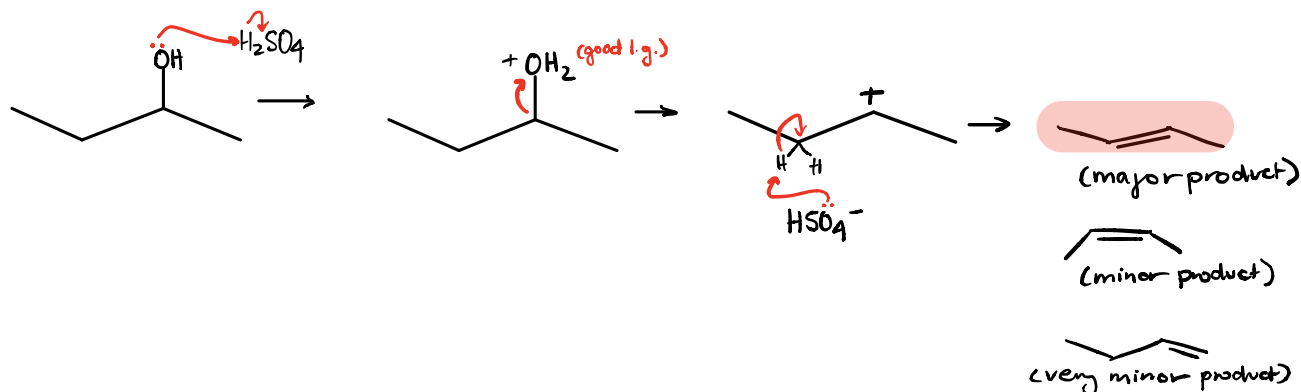
Dehydration Rxn

* Equilibrium reaction!!

alcohol favored by cold dilute acid

alkene favored by hot concentrated acid

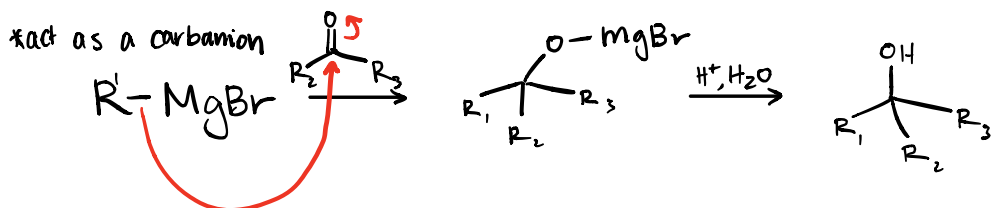
* Major product = most stable alkene



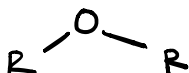
Grignard Synthesis

* increases # of carbons

* produces an alcohol



Ethers



- non reactive
- weakly polar
- low boiling point

Organic solvents

pI = isoelectric point

* like dissolves like

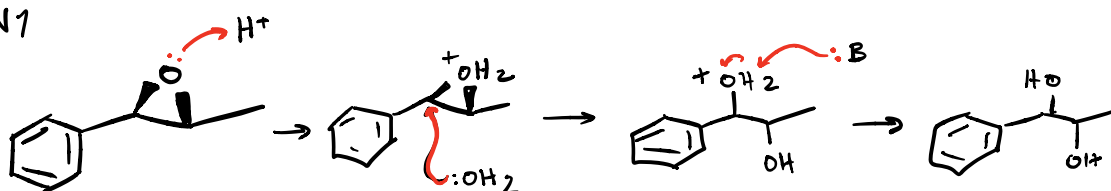
* unreactive

Epoxides

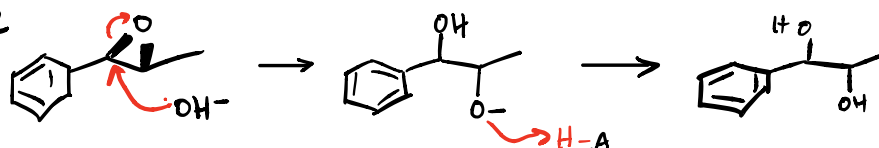


- Very reactive
- O gets protonated + becomes a better leaving group.

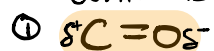
SN1



SN2



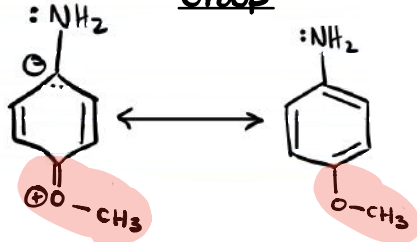
Which is more reactive?



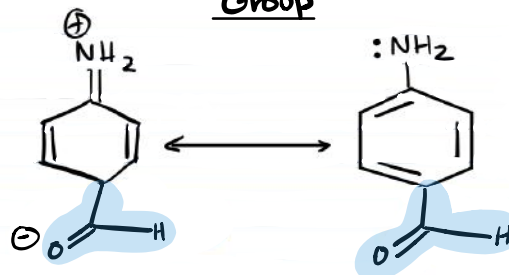
Electrophiles:
 δ^+ or $+$

Nucleophiles:
 δ^- or $-$

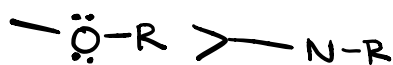
Electron Donating Group



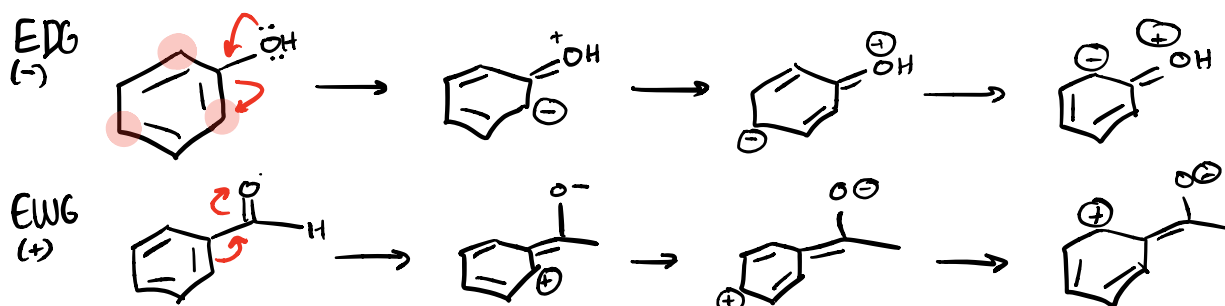
Electron Withdrawing Group



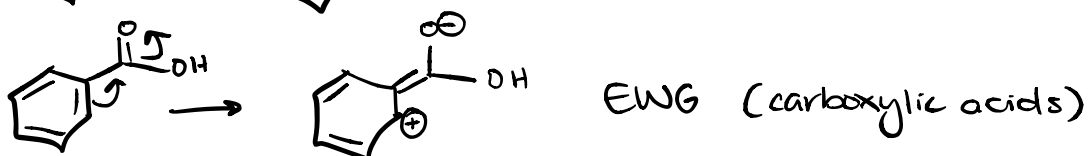
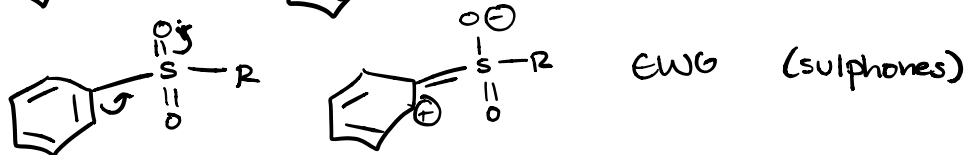
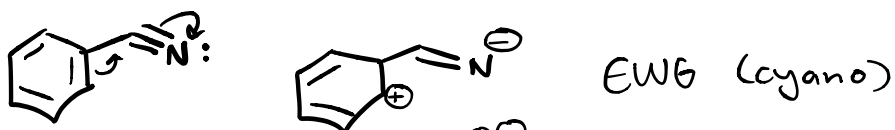
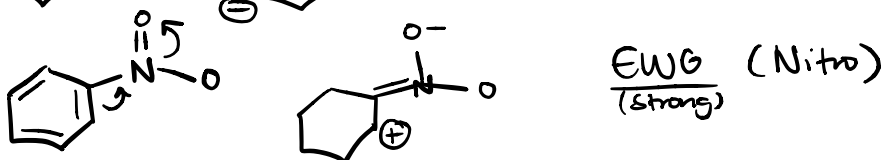
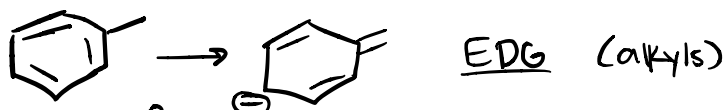
* lone pair

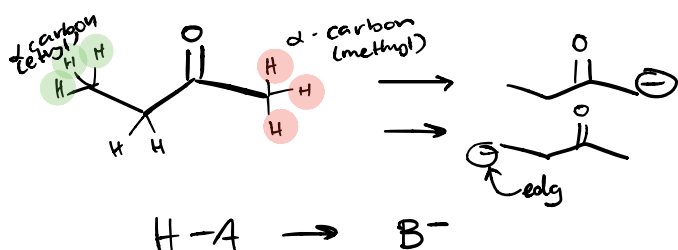
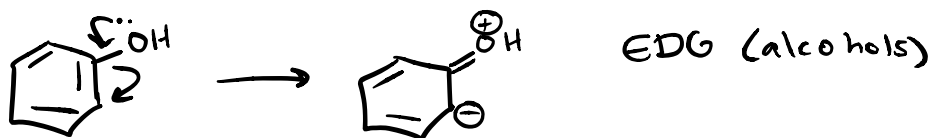
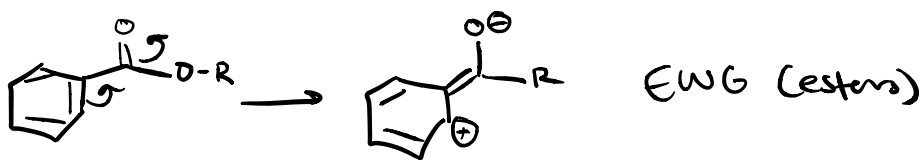


Nitrogen is less electronegative making it more available to donate its electrons



EWG or EDG??

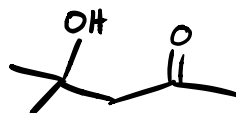




3° carbon must react via $\text{S}_{\text{N}}1$ or $\text{E}1$

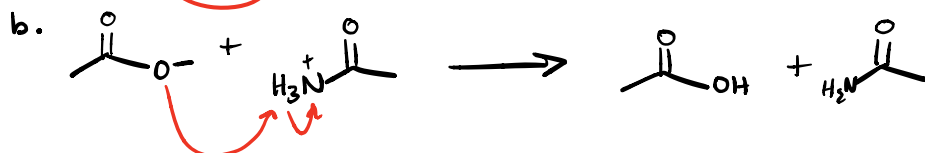
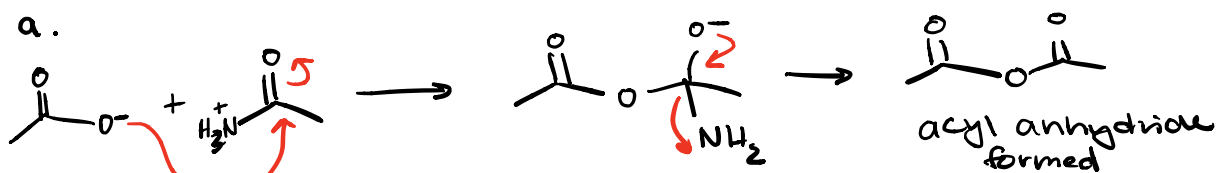
less stable cb = less acidic

$\uparrow$ cb stability = $\uparrow$ acidity



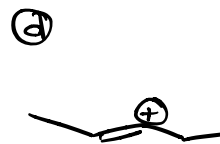
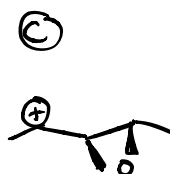
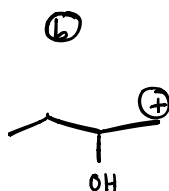
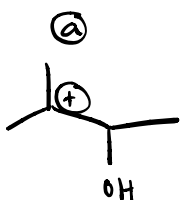
Resonance Stabilizes ♥

Q: which is more likely to occur?

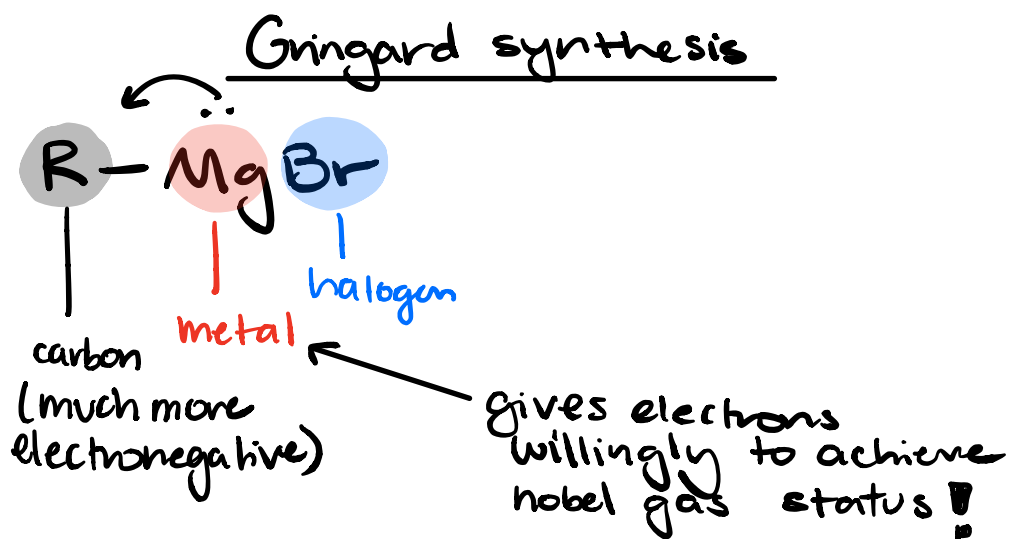


A: B! First and foremost a base!

Q: Which carbocation is most stable?

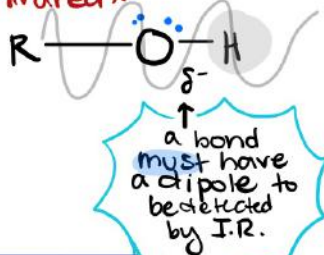


A: A!
tertiary substituted!



IR Spectroscopy

infrared



Spectra to know

- carbonyl : 1700cm^{-1} (C=O)
Sharp + deep
- alcohol : 3300cm^{-1} (O-H)
broad, separate from C-H
- Saturated alkane : 2800cm^{-1} (C-H)
Sharp + deep
- carboxylic acid : 3000cm^{-1} (O-H)
broad, overlaps CH
- amine : 3300cm^{-1} (N-H)
broad + shallow
- amide : 3300cm^{-1} (N-H)
broad + deep
- nitriles : 2250cm^{-1} (C≡N)
Sharp + deep

when...

IR radiation =

frequency of vibration in a bond

the bond is in resonance so it will absorb some IR energy.

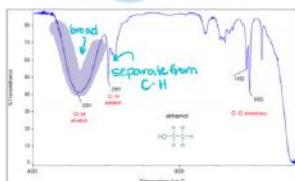
read by the detector

Bond frequency of vibration

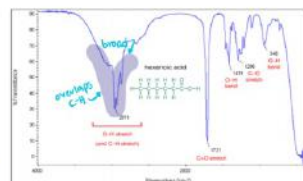
determined by...

- ① bond strength
- ② molecular weight of bonded atoms

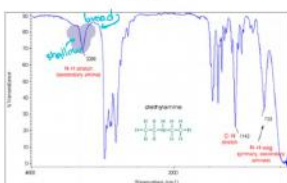
↑ bond strength = ↑ freq
↑ mass = ↓ freq



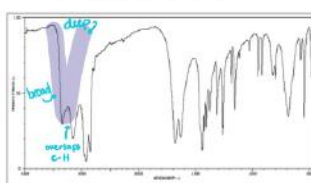
Alcohol



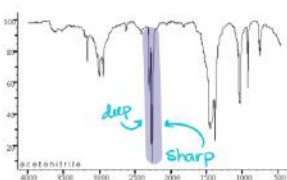
Carboxylic Acid (Note O-H and C=O)



Amine (notice how shallow it is)



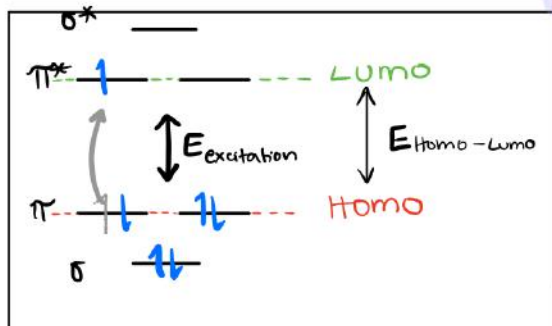
Amide (slightly overlapping C-H stretch)



Nitrile

UV Spectroscopy

ultraviolet



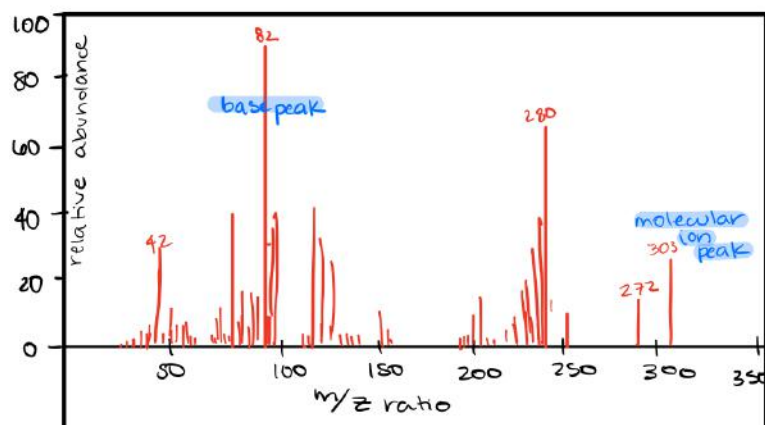
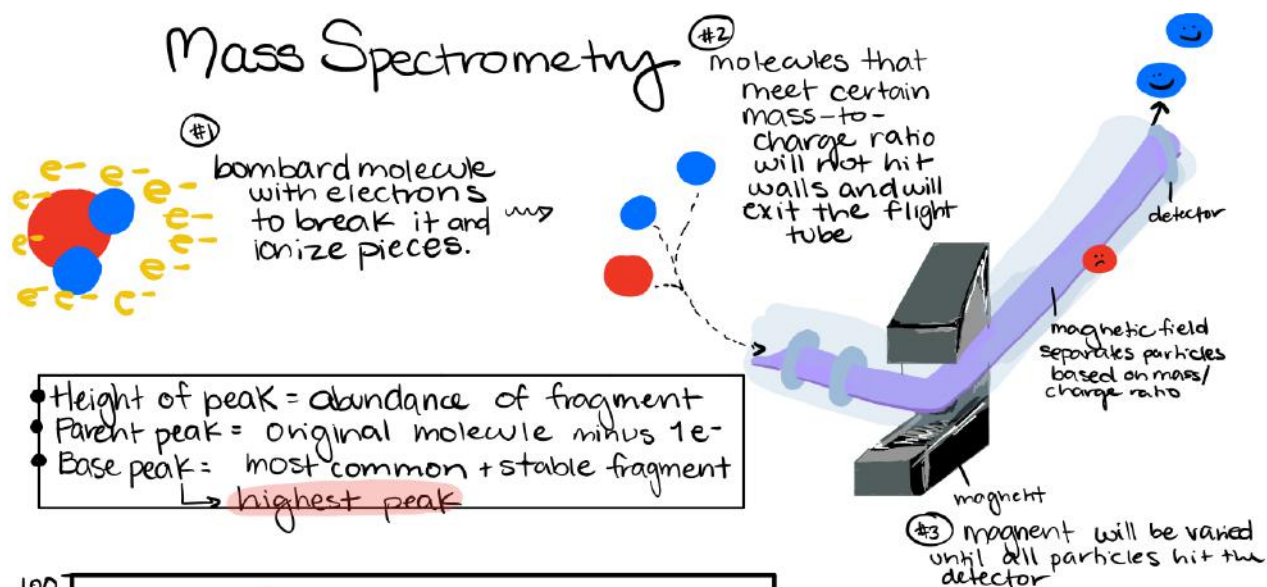
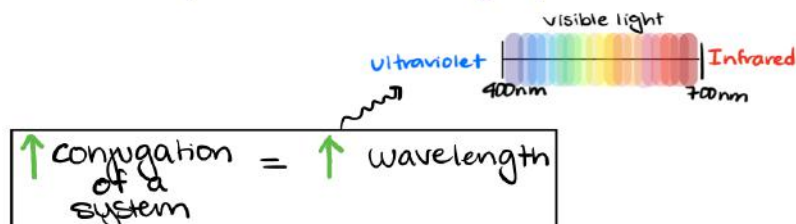
UV radiation

energy between molecular orbitals $\approx$ energy created by electromagnetic radiation in UV spectrum.

so...

when exposed to UV radiation electrons excite to next energy level. The absorbance is recorded.

- * single bonded molecules have low or no UV absorbance *
- * double bonds < triple bonds. Both show strong absorbance *
- * conjugated systems absorb even more light than others *
- * UV spectrum is a graph of absorbance vs wavelength *



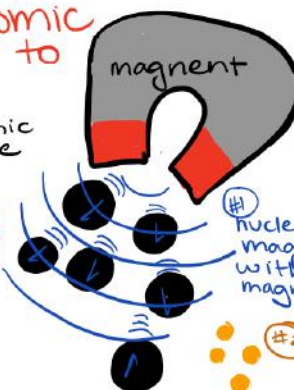
NMR Spectroscopy → differentiate molecules based on chemical environment of their (H) or (C)

nuclear magnetic resonance

atoms must have an odd atomic number or odd mass number to register on NMR.

all nuclei with odd atomic mass or mass # have a nuclear spin.

nuclear spin causes small changes in the electrical field which creates a magnetic field.

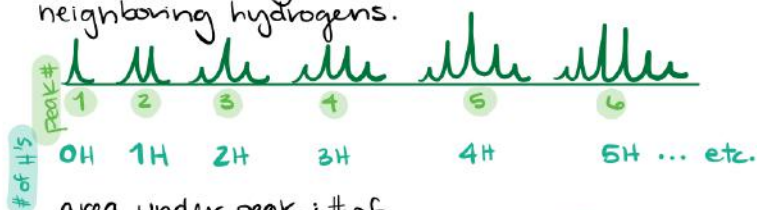


90% of NMR on MCAT

H-NMR

Peaks: all hydrogens in a molecule with a distinct chemical environment.

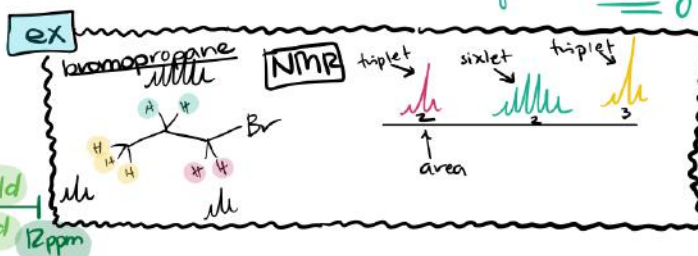
Spin-spin splitting: # of neighboring hydrogens.



#1 nuclei align their magnetic fields with the external magnetic field.

#2 when nuclei are exposed to photons some can absorb the energy and flip their field against magnetic field which is detected by the NMR.

#3 difference in energy needed to flip nuclei is based on degree of shielding



C¹³-NMR

- no spin-spin splitting - singlet peaks
- no integration - area under curve not important

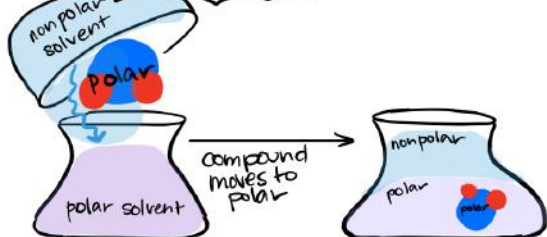
0-220 ppm



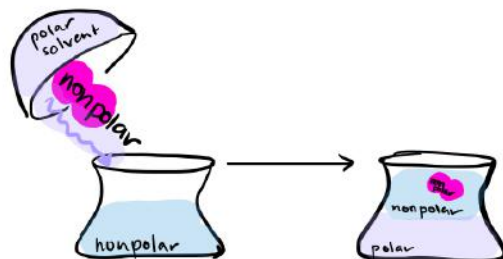
!absorptions to know! (still rare... just know in case)

- C-C $\rightsquigarrow$ 0-50 ppm
- C-O $\rightsquigarrow$ 50-100 ppm
- C=C $\rightsquigarrow$ 100-150 ppm
- C=O $\rightsquigarrow$ 150-200 ppm

Extraction



flow through contains product



polar layer drained solvent layer evaporated to obtain product.

(experimental design)

How to improve Separation

- #1 Repetition
- #2 Fractional extraction (use a 5mL x 10 over a 50mL)
- #3 Add an acid to protonate the product
- #4 add a base to deprotonate the product

*acids protonate bases to increase solubility in aqueous (polar) layer.

*bases deprotonate bases to increase solubility in polar layer

Chromatography

TLC

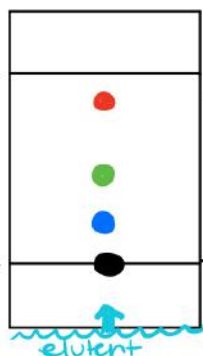
*thin layer *

paper = stationary phase

$$R_f = \frac{\text{distance traveled by component}}{\text{distance traveled by solvent}}$$

} closer to 1 means polarity of compound + solvent are similar

max distance eluent travels "B"



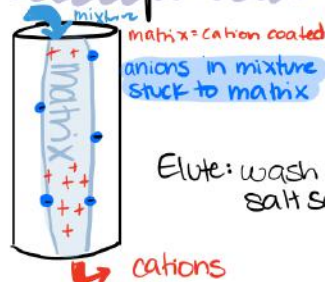
distance spot traveled "a"

$$\frac{a}{b} = R_f = 0.9 \rightarrow \text{non-polar compound}$$

Column Chromatography

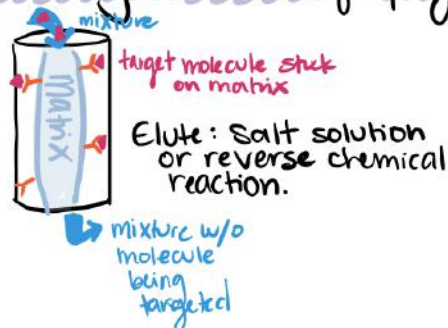


Ion-exchange Chromatography

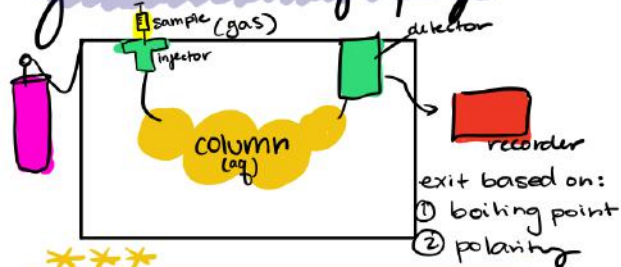


Elute: wash with salt solution

affinity Chromatography



gas chromatography



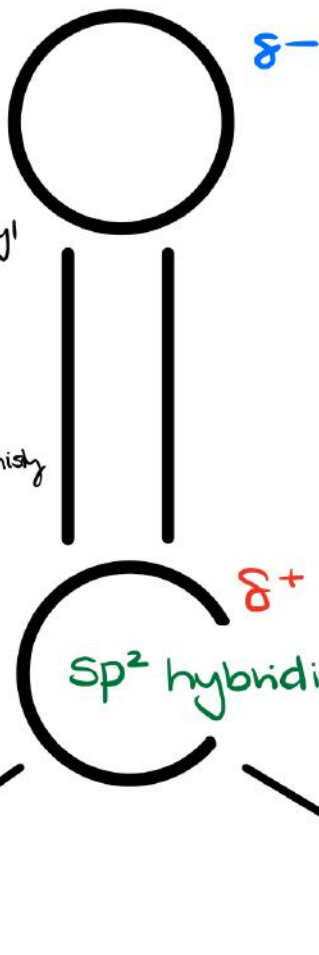
Peaks in gas chromatography are based on is equal to abundance of the compound

Lil Carbonyls 😊

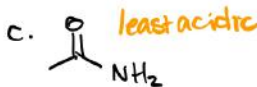
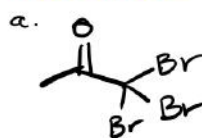
- Carbonyls > alkenes because oxygen is smaller and has better orbital overlap.

properties

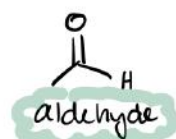
- 1- carbon is a good electrophile
- 2- α carbons are acidic
- 3- **EDG**: decrease reactivity of carbonyl carbon.
EWG: increase reactivity
- 4- Bulky substituents decrease carbonyl reactivity
- 5- planar stereochemistry



most acidic



α -carbon!
acidic hydrogens



-al

- H-bond acceptors
H₂O H-bond donor
- electrophiles
- function as Lewis acid



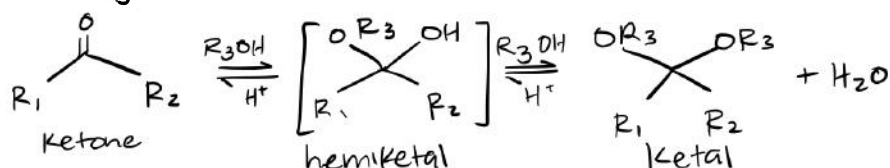
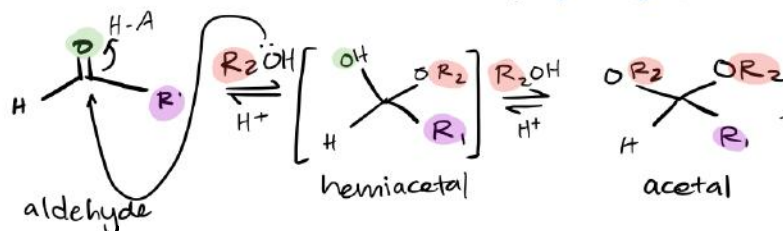
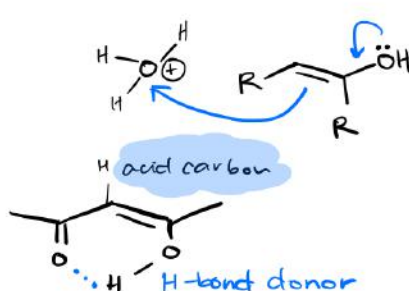
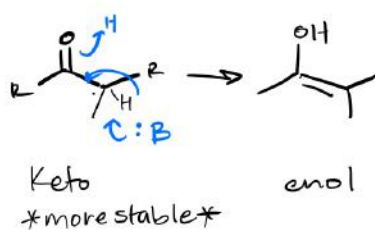
-one

- H-bond acceptors
H₂O H-bond donor
- electrophiles
- function as Lewis acid

solubility trends
alkanes < aldehydes + ketones < alcohols
no H-bond H-bond acceptor H-bond donor and acceptor
increasing →

boiling point
alkanes < aldehydes + ketones < alcohols
increasing →

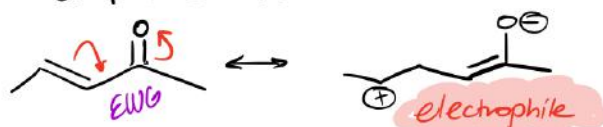
keto-enol tautomerization



* Done to protect aldehydes and ketones from reaction with nucleophiles and bases. *

* acidic conditions return an acetal and ketal to aldehydes and ketones *

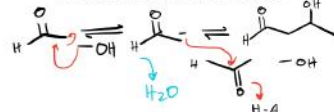
... α-β unsaturated carbons ...

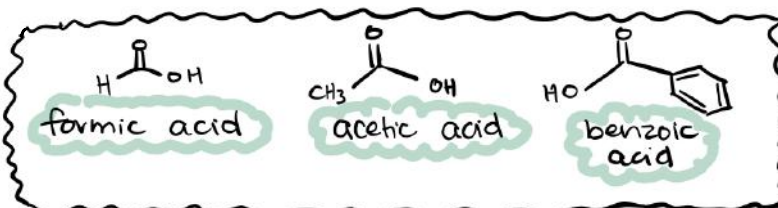
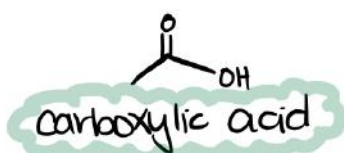


aldol condensation

condensation of an aldehyde with an aldehyde or a ketone with a ketone

* carbanion formed *

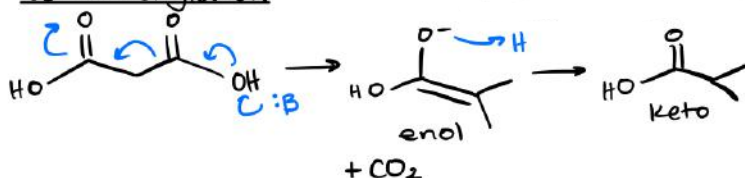




-oic

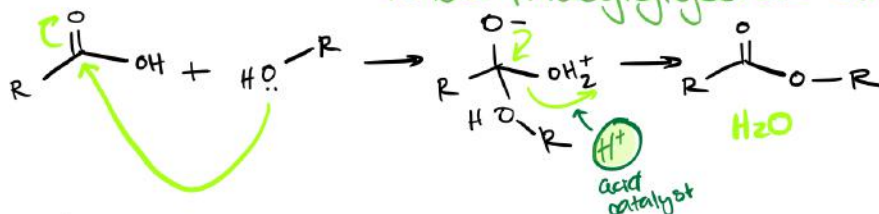
- carbonyl + hydroxyl substitute on alpha carbon
- high boiling points
- resonance stabilized

decarboxylation



esterification

How triacylglycerols are formed

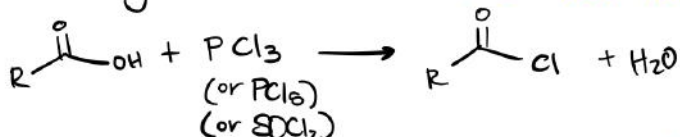


acid chlorides

-oyl chloride

- carbonyl + chloride

Cl⁻ will not form an acid chloride



anhydrides



- oic anhydride
- electrophilic

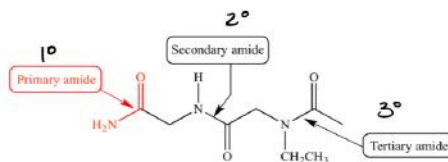
amides



-amide

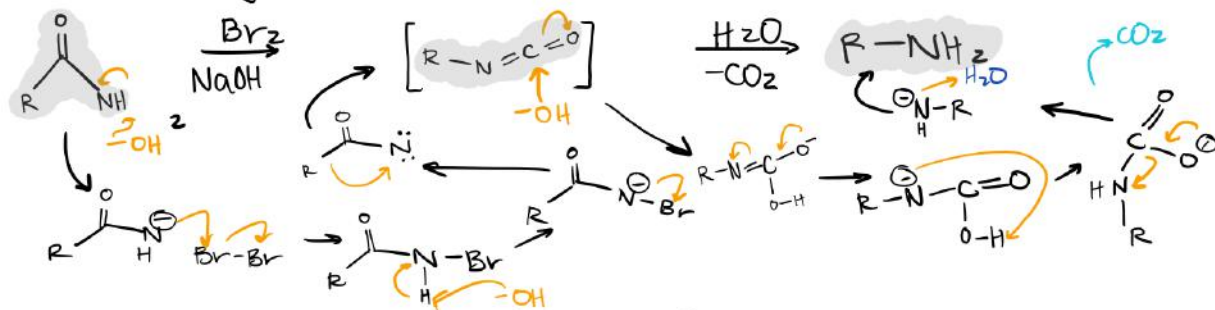
ie: benzoic acid → benzamide

- most stable acid derivative
- carbonyl carbons unreactive (formic acid)
- 1° and 2° amides can H-bond and are water soluble
- 3° amides cannot H-bond.

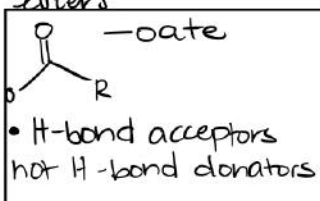


Hoffman Degradation

1° amides react in strong basic solutions of Cl_2 or Br_2 to form 1° amines $-\text{NH}_2$
 * Decarboxylation *



esters



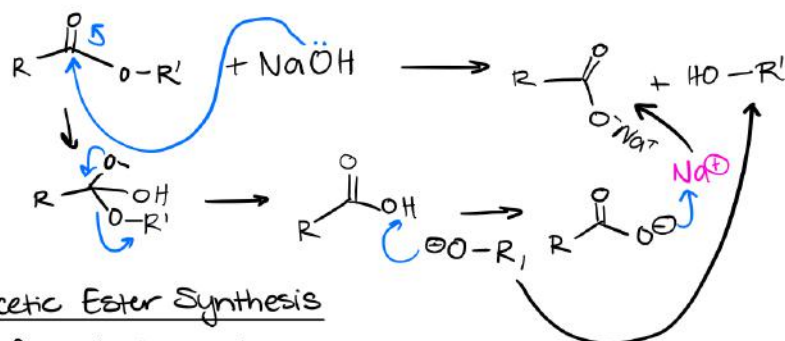
transesterification

ester + alcohol * acid catalysis *



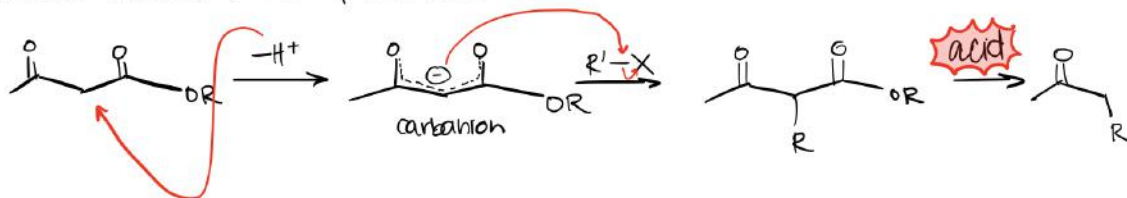
Saponification

ester hydrolysis to an alcohol and salt of carboxylic acid.



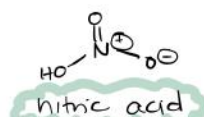
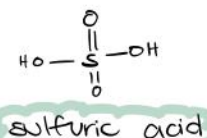
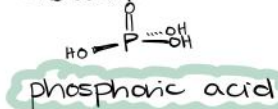
acetoacetic Ester Synthesis

ketone formed from a β -keto ester



Inorganic Esters

-oxoacids:



common inorganic ester examples?

- ATP, GTP, UTP (inorganic triphosphate esters)
- FADH_2 , NADH (inorganic diphosphate esters)
- FMN, DNA, RNA (monophosphate esters)

acid derivative
Leaving Groups

* Goodliest ~~~~~ ~~~~~ Crapdiest

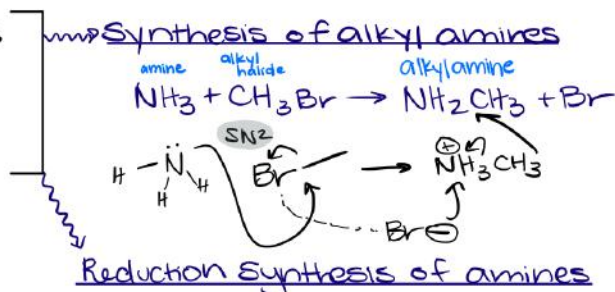
$-\text{Cl} > -\text{OCOR} > -\text{OH} > -\text{OR} > -\text{NH}_2$

* Stablest ~~~~~ ~~~~~ Not so stablest

amide > ester > carboxylic acid > anhydride > acid chloride

Amines

- bases or nucleophiles
- 1° or 2° = nucleophiles
- 3° = base
- NH_4^+ electrophiles



Reduction of amides, imines, nitriles, and nitro groups:

- **nitro groups:** reduced to 1° amine by any reducing agent (LiAlH_4 , NaBH_4 , H_2/cat)
- **nitrile groups:** reduced to 1° amine by any reducing agent
- **imines:** reduced to 1° amine by any reducing agent
- **Amides:** 1° amine only by LiAlH_4

adding amines to carbonyls

amines + ketones/aldehydes = imine or enamine

1° amines $\rightarrow$ imines

2° amines $\rightarrow$ enamines

3° amine $\rightarrow$ no rxn