

FACULTÉ DE PHARMACIE

TU07 – Refresher training in organic chemsitry

Year 2024-2025

Course outline

- Atoms and chemical bonds
 - Number of bonds around an atom
 - Types of chemical bonds
 - Covalent vs ionic
 - Single vs multiple
 - Impact of chemical bonds on structure
 - Functional groups
- Chemical reactivity
 - Nucleophile and electrophile
 - Bond polarisation
 - Inductive and mesomeric effects
 - Principal mechanisms





Atoms and chemical bonds



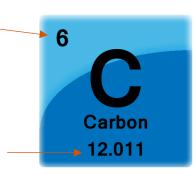
Atomic number 6 **Organic chemistry is... Carbon Chemistry!** Carbon 12.011 Atomic mass 18 2 PERIODIC TABLE OF THE ELEMENTS н He Helium 4.003 Hydrogen 1.008 14 15 16 17 2 4 10 Be B N 0 F Ne С Beryllium Boror Carbon 12.011 Oxygen 15,999 Neon 20.180 Nitrogen 18.998 5.941 9.012 10.811 12 13 Mg AI Si Na Ρ S CI Ar Sulfur Aluminum 26.982 Silicon Argon 39.948 Magnesius 24.305 hosphoru 30.974 Chlorine 35,453 12 10 11 25 27 29 22 23 24 26 28 30 20 21 31 Cu Sc Ti V Cr Fe Co Ni Zn Ga Ca Mn As Se Br Kr K Ge Cobalt Gallium Arsenic 74.922 Krypton 83,798 Calcium Scandium Titanium Vanadium Chromiun Manganese Iron Nickel Copper Zinc Germaniu 72.631 Bromine 79.904 40.078 44.956 47.88 50.942 51.996 54.938 55.845 58.933 58.693 63.546 65.38 69.723 78.971 39 40 41 42 43 44 45 46 47 48 50 38 49 54 Sb Rb Sr Zr Nb Rh Ag Silver 107.868 Cd Sn Xe Y Mo Tc Ru Pd In Te Vttrium Technetium Palladium Cadmium Antimony 121.760 Telluriur 127.6 lodine 126.904 Xenon 131.294 Strontium Zirconiun Niobium Molybdenum Rutheniu Rhodium Indium Tin 118.711 87.62 88.906 91.224 92.906 95.95 98.907 101.07 102.906 106.42 112.414 114.818 79 72 73 74 75 76 77 78 80 81 82 83 56 84 86 Ba Hf Та W Re Os Pt Au Hg TI Pb Bi Po Rn Cs Ir At Barium Hafnium Tantalum Tungsten Rhenium Osmium Iridium Platinum Gold Mercury 200.59 Thallium Lead Bismuth Astatine 209.987 Radon 222.018 137.328 178.49 183.85 186.207 190.23 192.22 195.08 196.967 204.383 207.2 208.980 208.982 180 948 88 89-103 104 105 106 107 108 109 110 111 112 113 114 115 118 116 Fr Ra Rf Db Sg Bh Hs Mt Rg Cn Nh FI Mc Og Ts Ds Lv Radium Rutherfordiun Dubnium Seaborg Bohrium Hassium Meitnerium Darmstadtiu oenta Copernicium Nihonium Flerovium [289] Moscovium Oganesso [294] [266] [264] [280] [285] [293] 226.02 [261] [262] [269] [278] [281] [286] [289] Pr Nd Pm Sm Eu Gd Тb Dv Ho Er Yb La Ce Lu Tm Cerium 140.116 Samariun Gadoliniu Terbium Erbium Thulium 168.934 Ytterbium Lutetium 174.967 Europiun ysprosiu 144.913 167.259 38 905 40 908 64 930 89 Th Pa Pu Cm Bk Cf Es Ac U Np Fm Md No Am Lr Actinium Thorium rotactiniu Uranium Curium Berkelium Californium Fermiun Aendelevium Nobelium Lawrencium Americiun insteiniur 227.028 232.038 231.036 237.048 244.064 243.061 247.070 258.1 259.101 [262] 247.070 251 080 257.095 Noble Alkaline Transition Basic Actinide Metalloid Ionmeta Halogen anthanide Metal Gas Earth Metal **FACULTÉ DE** universite

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Identity card of atoms

• Atomic number:



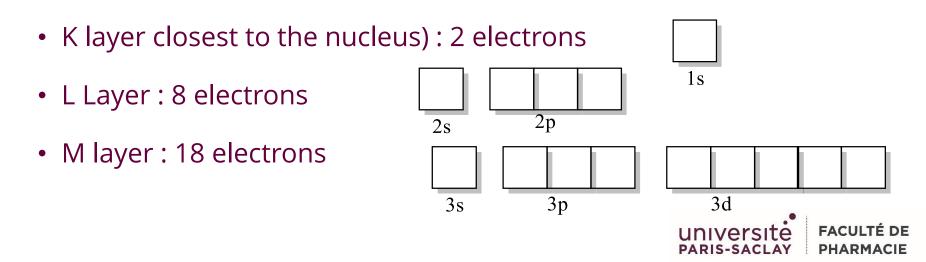
Atomic number

Atomic mass positive charges (protons) = negative charges (electrons) in the nucleus

• Atomic mass:

positive (protons) and neutral (neutrons) charges in the nucleus

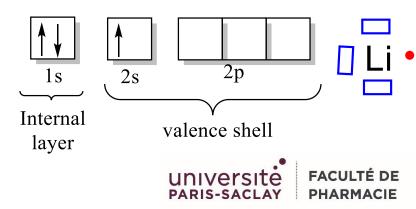
• Electrons occupy various core layers from K layer to M layer



Atoms and chemical bonds

- Hydrogen (H): 1 electron
 1 chemical bond is allowed
 - Helium (He): 2 electrons
 Layer K is full ➤ noble gaz
 No chemical bond allowed

Lithium (Li): 3 electrons
1 chemical bond is allowed
3 electron holes are present



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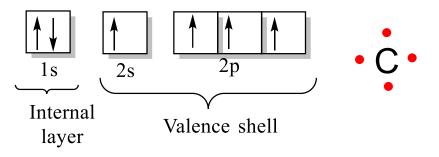
He

1s

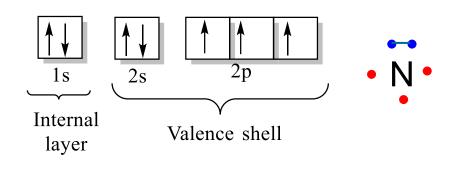
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Atoms and chemical bonds

• Carbone (C): 6 electrons 4 chemical bonds are allowed



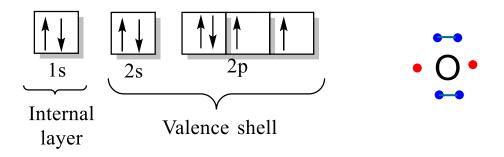
Nitrogen (N): 7 electrons
 3 chemical bonds are allowed
 1 lone pair is present



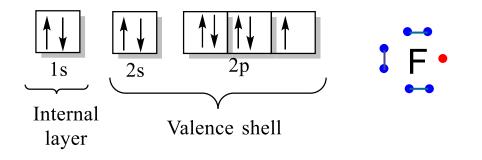


Atoms and chemical bonds

• Oxygen (O): 8 electrons



• Fluorine (F): 9 electrons



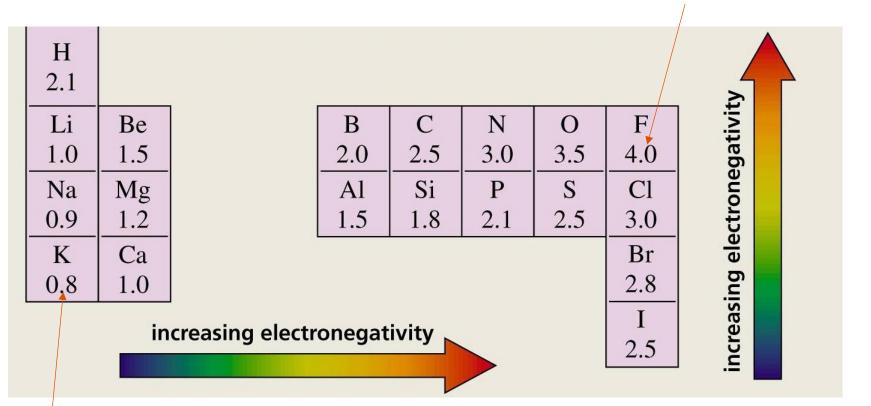


- Chemical bond = lasting attraction between two atoms by means of two electrons
- The type of chemical bonds is dependent of the electronegativities of the two atoms (ability of an atom to attract electrons)



Electronegativity based on the Pauling scale

The most electronegative



The less electronegative

Tricks for the two lines: Happy Hector Likes Beer But Could Not Obtain Food, Never!



 Chemical bond = lasting attraction between two atoms by means of two electrons

lonic bond: chemical bond involving the transfer of one electron from one atom to the other. Concerns atoms possessing very different electronegativities Fluorine electronegativity : 4,0 ⇒ strong attraction of electrons Lithium electronegativity : 1,0

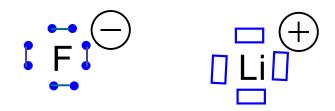




 Chemical bond = lasting attraction between two atoms by means of two electrons

Ionic bond: chemical bond involving the transfer of one electron from one atom to the other. Concerns atoms possessing very different electronegativities Fluorine electronegativity : 4,0 ⇒ strong attraction of electrons

Lithium electronegativity : 1,0





Pure covalent bond: sharing of two electrons from two atoms possessing similar electronegativities

- Carbon electronegativity : 2,5
- Hydrogen electronegativity : 2,1

Polar covalent bond: covalent bond with a partial ionic caracter => Polarized bond Oxygen electronegativity : 3,5 $_{H}^{\delta+}$ $_{H}^{\delta+}$ Hydrogen electronegativity : 2,1 o

> The less electronegative atom $\delta + \delta -$ The most electronegative atom universite FACULTÉ

H'''''''

Type of chemical bonds - Exercise

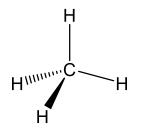
• By considering the atom electronegativity, indicate the bond polarisation. Represent also the lone pairs and electron holes present on the atoms. Is the bond a covalent or a ionic one?

H-F
$$EtO-H$$
 Et_2N-CH_3
H_3C)_3C-CI H_3C-MgI H_2N-Li

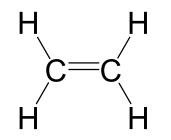


Single and multiple bonds

• When the two atoms share two electrons (one on each atom), you have a single bond.



 When the two atoms share four/six electrons (two/three on each atom), you have a double/triple bond.

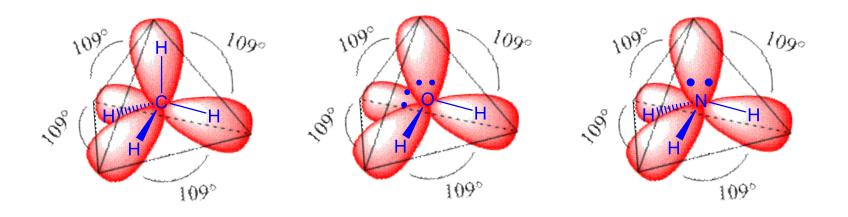






Impact of bonds on chemical structures

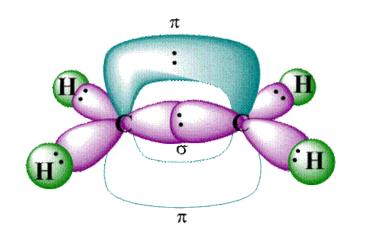
- An atom possesses a tetrahedral structure if he has around it 4 atoms and/or lone pairs
 - Only single bonds and lone pairs
 - Free rotation about single bonds

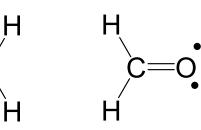




Impact of bonds on chemical structures

- An atom possesses a trigonal (plane) structure if he has around it 3 atoms and/or lone pairs
 - One double bond must be present
 - No free rotation about double bonds (rigid structure)

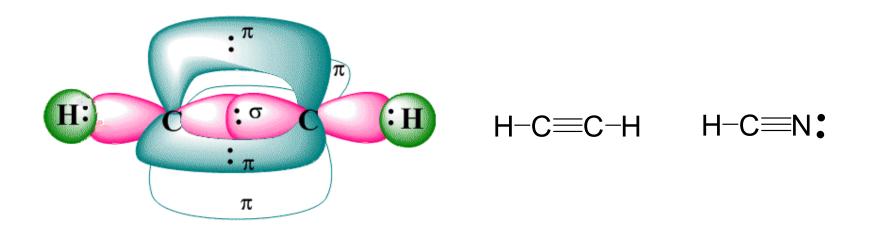






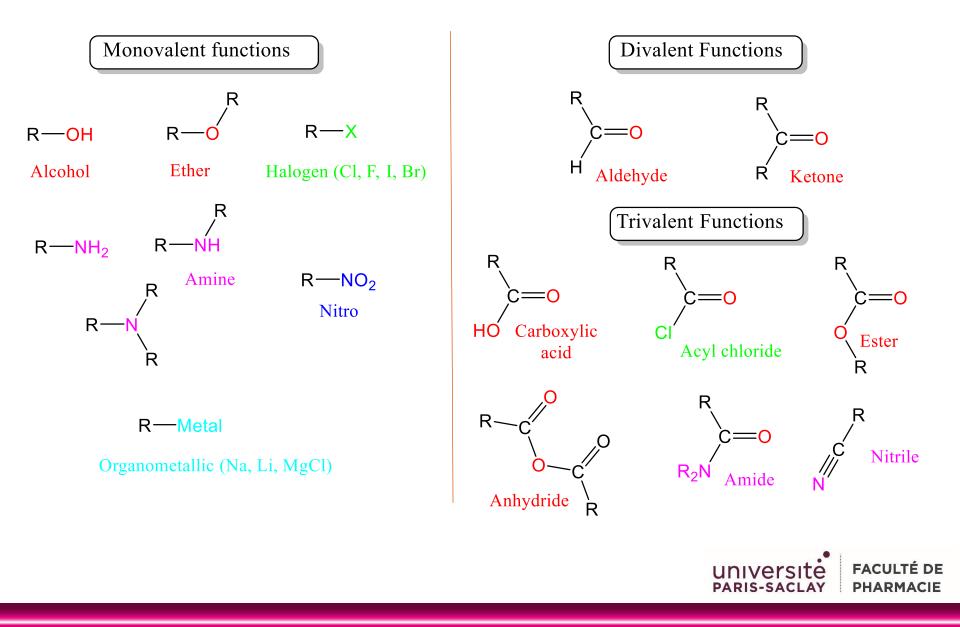
Impact of bonds on chemical structures

- An atom possesses a linear structure if he has around it 2 atoms and/or lone pairs
 - A triple bond must be present
 - No free rotation about double bonds (rigid structure)





Functional groups



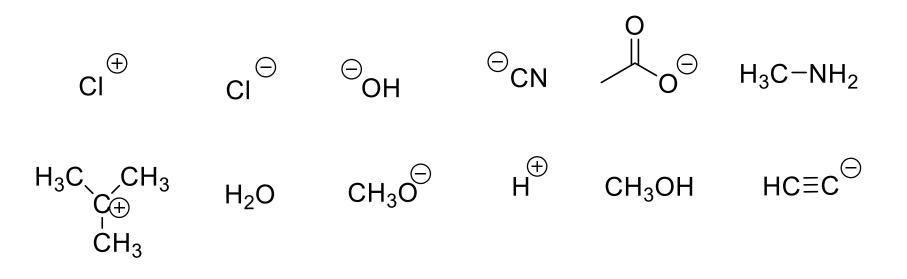




- Nucleophiles and electrophiles
 - A nucleophile is an atom or atoms group possessing a lone pair (eventually with a negative charge). He reacts with electrophiles.
 - An electrophile is an atom or atoms group possessing a hole (eventually with a positive charge or a partial positive charge in the case of carbon). He reacts with nucleophiles.



• Exercice: Represent the lone pairs and electron holes present on the atoms then categorize these compounds as nucleophile or electrophile. Indicate which atom is concerned.

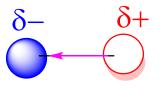




• In complex structures, it depends on the chemical bond polarization and substituants

1A H 1.00007 3 Li	2A Be	 Directions of increasing electronegativity 								SA B	4A C	5A 7 N	6A 0	7A P F	0 He 4000 10 Ne		
6.941	0.0102									10.881	10,0110	14,0007	15.0994	18,5984	20.179		
11 Na 22.6688	12 Mg 24.305	SB	4B	5B	¢B	7B	_	88	_	1B	2B	13 Al 28.2815	14 Si 25.635	15 P	S	17 CI 35.483	16 Ar 35.348
19	20	21	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36
K	Ca	Sc	TI	V	Cr	Mn	Fe	Co	Ni	Cu	Zn	Ga	Ge	As	Se	Br	Kr
43-698	45.05	44,099	47.99	45.842	64,859	94,0552	55.847	65.8532	-88.79	1000	65.55	09.77	72.58	74.0298	28.88	78.004	83.80
37	38	39	40	61	42	43	1	45	46	47	- 45	49	50	51	\$2	53	54
Rb	Sr	Y	Zr	Nb	Mo	TC	Ru	-Rh-	Pd	Ag	Cd	In	Sn	Sb	Te		Xe
85.47	87,62	88.625	41.22	57 554	45.65	(14)	101.07	102.505	198.4	107.868	112,41	114.82	118,69	121.75	127,62	128.5045	121.30
55	- 36	71		73	74	75	76	77	78	79	80	81	82	83	-84	85	85
Cs	Ba	10	Hf	Ta	W	Re	Os	lr l	Pt	AU	Hg	TI	Pb	Bi	Po	At	Rn
102,002	13246	108.91	170.49	180,948	162.65	165.2	193.2	198.2	195.09	196.967	532.59	504.97	207.19	202,592	(0110)	(2190)	(202)
87	88	103	104	105	106	107	108	109	110	111	112	113	114	115	116	117	118
Fr	Ra	Ac	Rf	Db	Sg	Bh	Hs	Mt	Ds	Rg	Cn	Uut	FL	Uup	Lv	Uus	Uuo
(2223)	326.0354	(207)	(837)	689		-											

The most electronegative atom



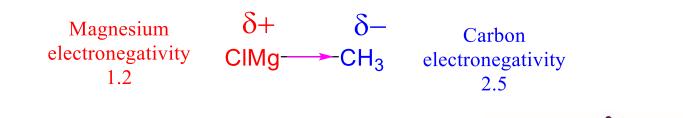
The less electronegative atom



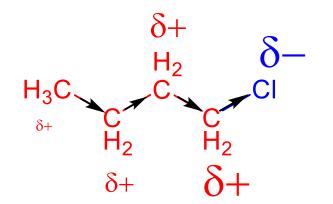
- Electronic effect of substituants Inductive effect
 - Concerns the bond polarization and the impact on the adjacent bonds. The effect is defined relative to a bond formed with a carbon atom.
 - Electro-withdrawing effect –I: When an atom or a group of atoms attracts the electrons of the bond



 Electro-donating effect +I: When an atom or a group of atoms repels the electrons of the bond



- Electronic effect of substituants Inductive effect
 - The inductive effect is transferred through the single bonds but declined rapidly with the distance





• Exercise: Determine if these substituents possess a +l or a –l effect???

-NR ₂ -1	NO_2	-OR
-Halogens	-COR	-Alkyl
-CN	-COOR	-CONR ₂
-Na	-Li	



• Electronic effect of substituants – Inductive effect

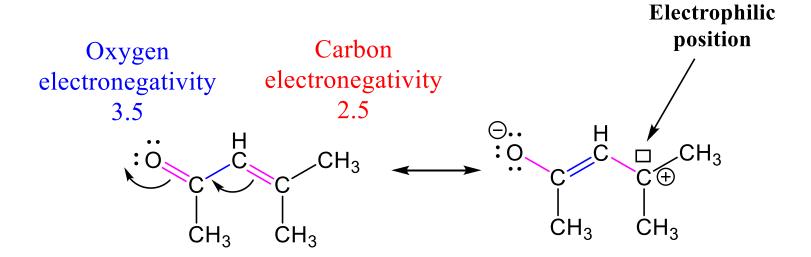
	-1		+				
-NO ₂	-COR		-Alkyl				
-CN	-COOR	-CONR ₂	-Na -Mg(Cl			
-NR ₂	-OR		-Li				
-SR	-Halogènes						



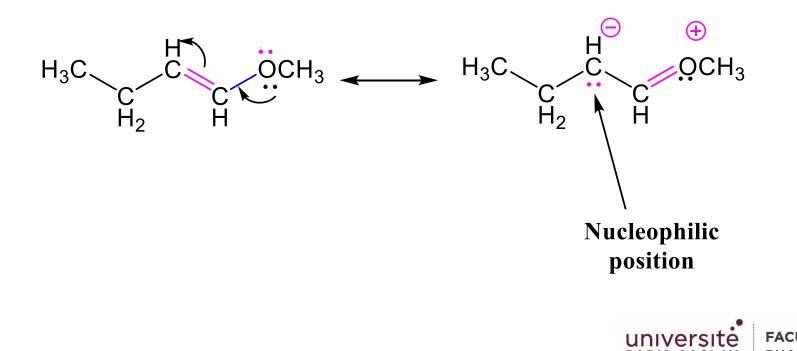
- Electronic effect of substituants mesomeric effect
 - Concerns the delocalization of the π electrons in a molecule ie the lone pairs, electron holes and π bonds (in multiple bonds)
 - The mesomeric effect can only take place when there is one single bond (and only one) between two π electrons:
 - π bond σ bond π bond
 - Ione pair σ bond π bond
 - Electron hole σ bond π bond (or lone pair)
 - Mesomeric effects are superior to inductive effects



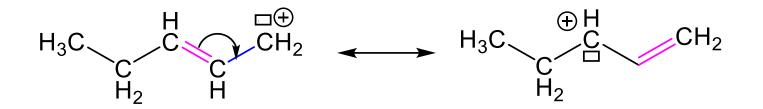
- Electronic effect of substituants mesomeric effect
 - When you have a π **bond** σ **bond** π **bond**, the electrons are displaced from the less electronegative atom to the most negative atom



- Electronic effect of substituants mesomeric effect
 - When you have a **lone pair** σ **bond** π **bond**, the electrons are displaced from the lone pair in direction to to the π bond

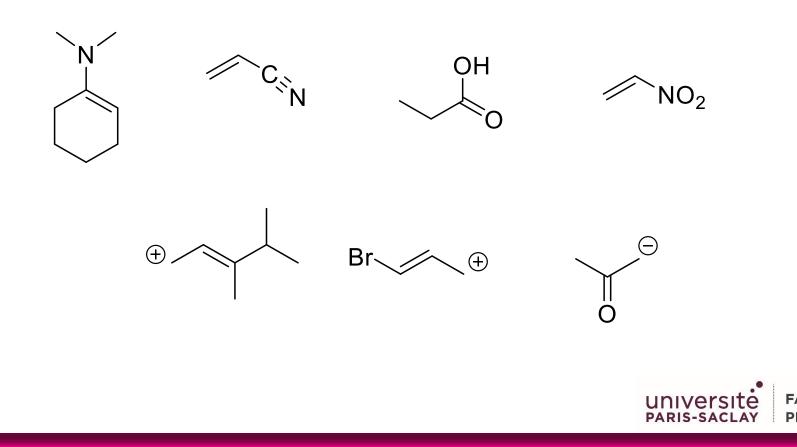


- Electronic effect of substituants mesomeric effect
 - When you have an electron hole- σ bond π bond (or lone pair), the electrons are displaced from the π bond in direction to to the electron hole





• Exercise: For each compound represent the mesomeric forms. Then identify the electrophilic or nucleophilic atoms of the compounds



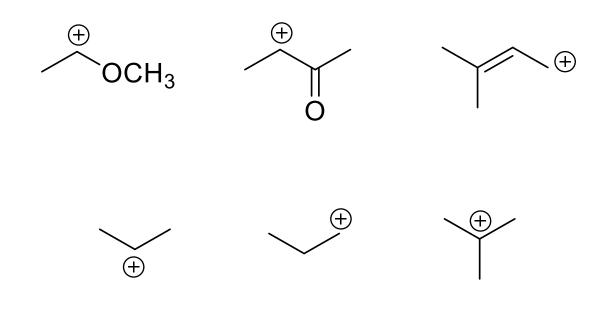
• Electronic effect of substituants – mesomeric effect

	-M		+M				
-NO ₂	-COR		-NF	R_2 -OR			
-CN	-COOR	-CONR ₂	-SR	-Halogènes			

The inductive effects and the mesomeric effects explain the stability and the reactivity of organic molecules

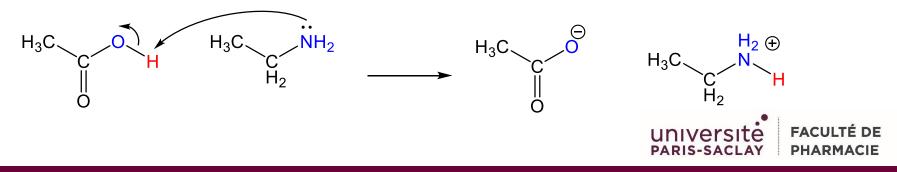


• Exercise: Thanks to the inductive and/or mesomeric effects, classify this carbocation from the less stable to the most stable.

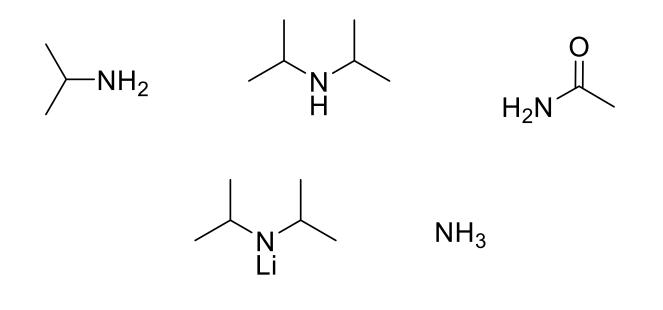




- Acids and bases
 - Acid: molecule able to give a proton (H⁺)
 - Base: molecule able to catch a proton
 - An acid/base pair is defined by its pKa.
 - When pH<pKa the acid form is predominant
 - When pH>pKa the basic form is predominant
 - Acid-base reaction: displacement of the proton from the acid function to the basic one

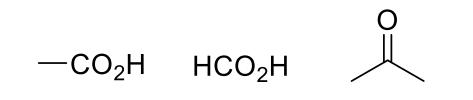


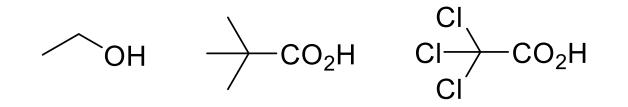
• Exercise: Thanks to the inductive and/or mesomeric effects, classify these nitrogen functions from the less basic to the most basic.





• Exercise: Thanks to the inductive and/or mesomeric effects, classify these molecules from the less acid to the most acid.

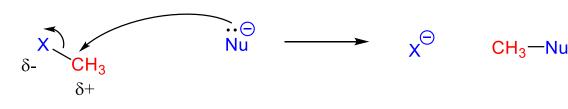






Principal mechanisms

- Nucleophilic substitution
 - When the carbon has only single bonds
 - A nucleophile (Nu) attacks the electrophilic carbon (δ+) And causes the departure of the leaving group X⁻ The nucleophile can be for example:
 - an amine
 - an alcohol
 - The leaving group is in general:
 - water
 - an halide (Cl⁻, Br⁻, l⁻)



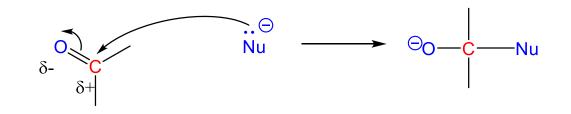


Principal mechanisms

- Nucleophilic Addition
 - When the carbon possesses a double bond.
 - A nucleophile (Nu) attacks the electrophilic carbon (δ +) and the two electrons of the π bond are sent as a supplementary lone pair on the oxygen (δ -) for example (π bond is more fragile than σ bond)

The nucleophile can be for example:

- an amine
- an alcohol (or water)





Principal mechanisms

- Nucleophilic addition/leaving group departure
 - When the carbon possesses a double bond and a leaving group.
 - A nucleophile (Nu) attacks the electrophilic carbon (δ +) and the two electrons of the π bond are sent as a supplementary lone pair on the oxygen (δ -)
 - In a second step the lone pair of the owygen creates a new double bond and causes the departure of the leaving group.

