# **Drug-Receptor Bonding**

**Ionic**: the strongest type of non-covalent bond. This results from the attraction of ions with opposite charges.





**Ion-Dipole** : results when there is an attraction between an ion and the partial charge of a dipole of the opposite polarity.



#### ion-dipole bonding

**Dipole-Dipole** : Here a partially positive atom in a dipole is attracted to a partially negative atom in another dipole.



dipole-dipole bonding

Hydrogen Bonding: A dipole-dipole interaction where on of the constituents is a hydrogen attached to a heteroatom.

#### Hydrogen bonds

- Vary in strength
- Weaker than electrostatic interactions but stronger than van der Waals interactions.
- A hydrogen bond takes place between an electron deficient hydrogen and an electron rich heteroatom (N or O).
- The electron deficient hydrogen is usually attached to a heteroatom (O or N).
- The electron deficient hydrogen is called a hydrogen bond donor (HBD).
- The electron rich heteroatom is called a hydrogen bond acceptor (HBA).



#### Hydrogen bonds

- The interaction involves orbitals and is directional.
- Optimum orientation is where the X-H bond points directly to the lone pair on Y such that the angle.
   between X, H and Y is 180°.



#### Hydrogen bonds

- Examples of strong hydrogen bond acceptors
   carboxylate ion, phosphate ion, tertiary amine.
- Examples of moderate hydrogen bond acceptors
   carboxylic acid, amide oxygen, ketone, ester, ether, alcohol.
- Examples of poor hydrogen bond acceptors
   sulfur, fluorine, chlorine, aromatic ring, amide nitrogen, aromatic amine.
- Example of good hydrogen bond donors
  Quaternary ammonium ion.



# Water can act as an H-bond Donor or Acceptor



# Donates H

# Accepts H

# Lone pair electrons

# Examples of H-bonding interactions





The Hydrophobic Effect : when two alkyl chains approach one another, water is extruded from the space in between them, resulting in an increase in entropy, and thus a decrease in energy.



Charge-Transfer Complexes : a lone pair of electrons is "shared" with a neighboring group that has considerable  $\pi$  character.



Van der Waals Forces : one carbon in a chain approaches another carbon on a neighboring chain, causing a perturbation known as an induced dipole. These opposite partial charges then attract one another.



Van der Waals forces

> Drugs may also bind to receptors using covalent bonding. This may be a permanent bond, in which case the receptor or enzyme target is "killed", or it may be transient.



Drug Interaction with Receptor Lock & Key Model - NT binds to receptor NT = key Receptor = lock













OH







## $\beta$ -Adrenoceptor



### ADRENALINE

### $\beta$ -Adrenoceptor



### SALBUTAMOL



### SALBUTAMOL



# SALBUTAMOL