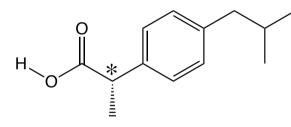
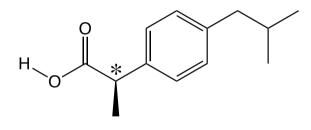


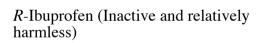
S-Thalidomide (Relieves morning sickness)

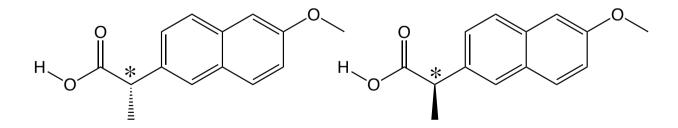


S-Ibuprofen (Advil, Motrin)





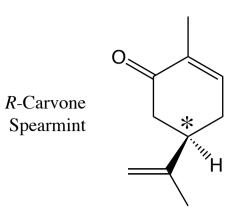




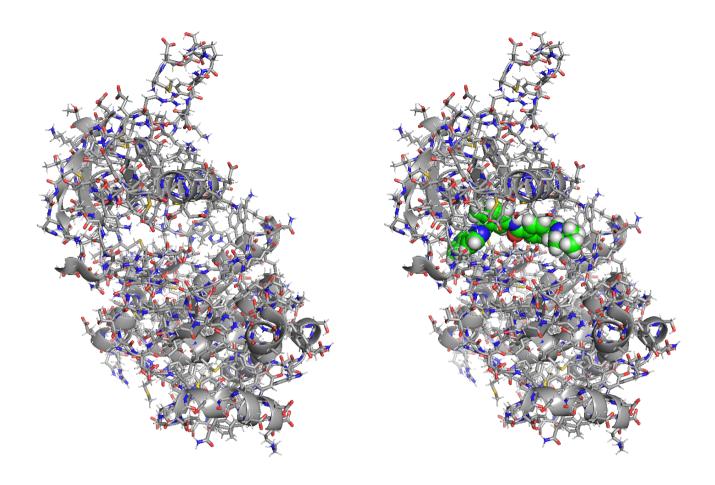
S-Naproxen (Aleve)

R-Naproxen (liver toxin)

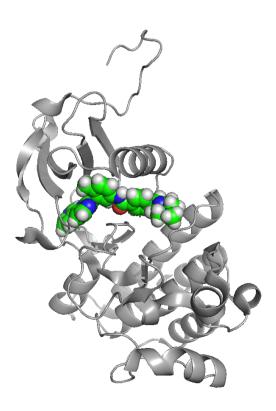


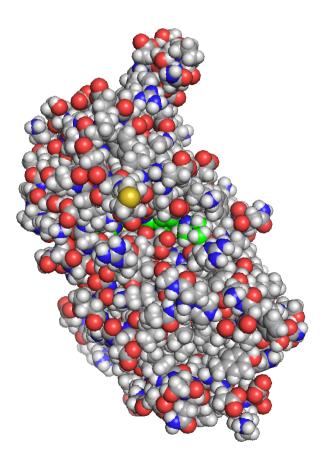






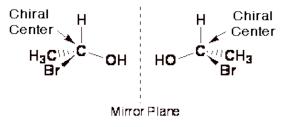
The drug Gleevec (green) bound to its target protein, the ABL kinase.





STEREOCHEMISTRY HOPEFULLY MADE SIMPLER

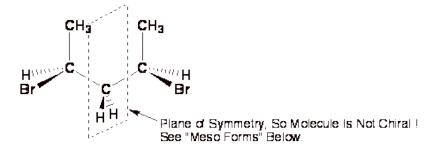
- **Stereoisomers**-molecules that have the same constitution, but different disposition of groups in space. In other words the atoms are connected to each other in the same way, they only differ with respect to relative orientation in three--dimensional space.
- **Chiral-General Definition**-Any object that is not superimposable on its mirror image. Your hands are chiral, that is why you need two different leather gloves, one that only fits your right hand, and one that only fits your left hand. If your hands were superimposable, then you would only need one kind of glove and it would fit both hands.
- **Chiral-Chemistry Definition; Atom**-Any tetrahedral carbon atom that has four different substituents is a **chiral center**. Any tetrahedral carbon atom that has four different substituents is a **chiral center** (it was worth repeating). This is a simple consequence of geometry; there are **two different_ways** to place four different substituents in a tetrahedral arrangement. Looking for four different substituents on a single carbon atom is the easiest way to identify a chiral molecule. It is worth mentioning that a chiral center is a special type of a more general situation called a stereocenter. A stereocenter is any atom in a molecule for which exchanging two groups creates a different stereoisomer. All chiral centers are stereocenters, however, not all stereocenters are chiral centers as we will encounter examples of this in later chapters. Do not sweat this detail at this point. The bottom line is that when you have a tetrahedral carbon atom with four different groups, call it a chiral center and life will be good.



As a further consequence of simple geometry, there are only two different ways to place four substituents around a chiral center, and as shown above the resulting two different molecules are mirror images of each other. They are non-superimposable mirror images of each other. You should make models of the above molecules and prove to yourself they are different and non-superimposable if you haven't done this already. This pair of chiral molecules that are mirror images of each other are called **enantiomers**, a chemistry name given to represent this special mirror image relationship between different molecules.

A carbon atom is **not** a chiral center if even two of the substituents are the same. A carbon atom is **not** a chiral center if even two of the substituents are the same (that was also worth repeating). Historical note: in the past, chiral centers have also been called; stereogenic centers, asymmetric carbon atoms, asymmetric centers, or chiral atoms even though these names can be confusing in certain situations.

Chiral-Chemistry Definition; Molecule- A molecule is not chiral (even if it has chiral centers) if it has a plane or center of symmetry. A plane of symmetry is any plane cutting through the molecule such that one side is a perfect reflection of the other. When looking for a plane of symmetry you have to put the molecule in the most symmetric conformation possible, DO NOT WORRY ABOUT WHICH CONFORMATION IS MOST STABLE, STABILITY OF CONFORMATION IS NOT IMPORTANT FOR STEREOCHEMISTRY. This means you look for symmetry in cyclohexane derivatives that are drawn flat and alkanes that are eclipsed !

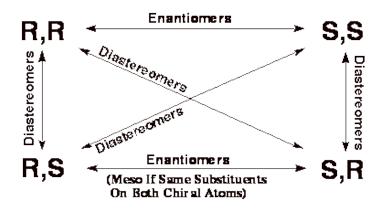


- A **center of symmetry** is any point is space such that any group on the molecule can be reflected back through that point an equal distance but opposite direction and an equivalent group is found. This is relatively rare in organic chemistry.
- Molecules With More Than One Chiral Center If there are two chiral centers in a single molecule, there are four possible stereoisomers. This is because each carbon atom can be in one of two possible forms (R or S) as we saw above, so there are 2x2=4 possible combinations. Now refer to the general figure below as well as the figure on the following page with the four chemical structures. Make sure you understand all

the relationships of the four different molecules shown. It is easiest to remember that **enantiomers must be mirror images of each other**. **Diastereomers** is the chemistry term that describes the relationship beween the pairs of molecules in the figures that are not even mirror images of each other. In other words, each pair of molecules in the figures must be related because they are all stereoisomers of each other, so they are either enantiomers (mirror images) or diastereomers (not mirror images). Thus, **diastereomers are stereoisomers that are not enantiomers**. If a molecule happens to be symmetric so that two of the four possible stereoisomers are identical (the S,R is identical to the R,S; the S,S and R,R will always be enantiomers), this form of the molecule is called the **meso form**. Since this situation requires some special symmetry (usually a plane of symmetry) to be present, it is the unusual exception, not the rule.

To Determine Whether Molecules Are Enantiomers, Diastereomers or Meso Compounds On A Test: *

- 1). Determine the absolute (R or S) stereochemistry at each chiral center. Click here to go to a description of how to determine whether a chiral center is "R" or "S".
- 2). Use the following table to determine relationship.



3). If the molecule happens to be symmetric, check for meso compound, remembering that only the R,S or S,R molecules can be meso (the S,S and R,R forms of even symmetric molecules are not meso compounds, they are enantiomers).

* This method is much easier than simply trying to look for mirror images etc. by rotating molecule in your mind or redrawing structures in various orientations.

Other Important Ideas:

- 1). Different enantiomers can only be distinguished by chiral things (optical activity etc.). Living systems and biological molecules are chiral, so chiral molecules are almost always distinguished by living systems.
- **2).** Different diastereomers can be distinguished by all of their chemical properties.
- **3).** A **racemic mixture** is defined as being a one-to-one mixture of two enantiomers.

