

EXPERIMENT 7 (Organic Chemistry II) Pahlavan / Cherif

Aldol Condensation - Synthesis of Dibenzalacetone



Chemicals

Methanol, CH₃OH
Acetone, CH₃COCH₃
10% NaOH
Benzaldehyde, C₆H₅-CHO
95% Ethanol, C₂H₅OH

Materials

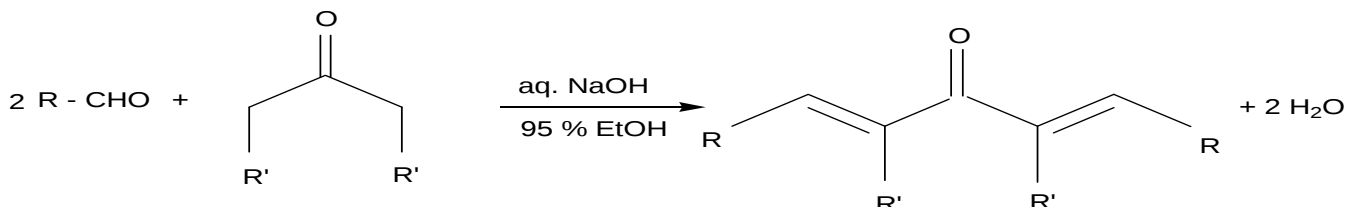
125 mL Erlenmeyer flask
thermometer
test tube rack
mel-temp
ice bath

25 mL graduated cylinder
Büchner funnel
medium size test tube
filter paper

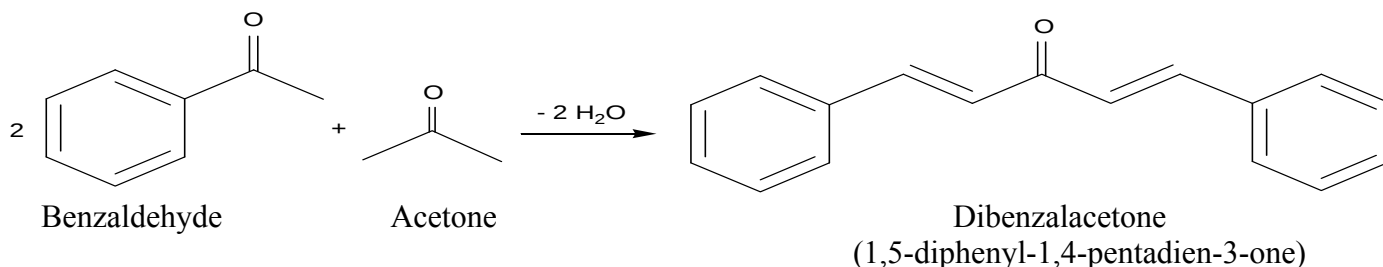
Purpose - The objectives of this experiment are to learn aldol condensation mixture of aldehydes and ketones, which used extensively in organic synthesis to form C-C bonds and make bigger molecules.

Introduction

Like the Grignard reaction, the Aldol Condensation is an extremely useful carbon-carbon bond-forming reaction in organic chemistry. Under the reaction conditions in the experiment, two equivalents of aldehyde will react.



The aldol condensation is a reaction that is named based on the type of product formed when two aldehydes (or ketones), in the presence of dilute base, yields a molecule having both alcohol and aldehyde functional groups. An example of the type of base-catalyzed aldol condensation that you will perform is shown below.



These products are a β-hydroxyaldehyde (or a β-hydroxyketone). This reaction is used extensively in organic synthesis to form C-C bonds and make bigger molecules. In every case, the product results from the addition of one molecule of an aldehyde (or ketone) to a second molecule in such a way that the α-carbon of the first becomes attached to the carbonyl carbon of the second.

MECHANISM OF THE ALDOL CONDENSATION

The acidity of the α -carbon makes β -dehydration of aldols an easy reaction. (This is of course quite different than the chemistry of normal alcohols.) This conjugated enone synthesis is catalyzed by both acids and bases. This shows the mechanism of the experiment performed. The reaction proceeds by an aldol condensation.

Step 1:

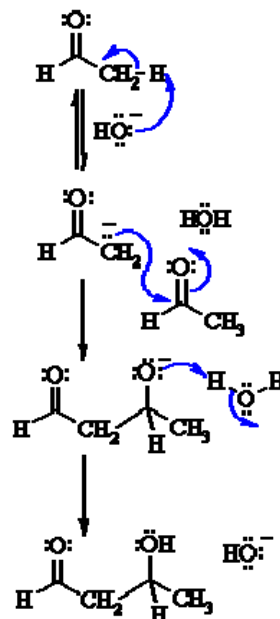
First, an acid-base reaction. Hydroxide functions as a base and removes the acidic α -hydrogen giving the reactive enolate.

Step 2:

The nucleophilic enolate attacks the aldehyde at the electrophilic carbonyl C in a nucleophilic addition type process giving an intermediate alkoxide.

Step 3:

An acid-base reaction. The alkoxide deprotonates a water molecule creating hydroxide and the β -hydroxyaldehydes or aldol product.



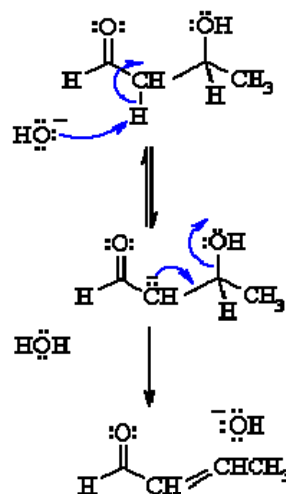
MECHANISM OF THE DEHYDRATION OF AN ALDOL PRODUCT

Step 1:

First, an acid-base reaction. Hydroxide functions as a base and removes an acidic α -hydrogen giving the reactive enolate.

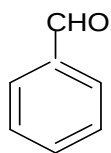
Step 2:

The electrons associated with the negative charge of the enolate are used to form the $\text{C}=\text{C}$ and displace the leaving group, regenerating hydroxide giving the conjugated aldehyde.

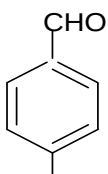


Dehydration generally occurs under slightly more vigorous conditions, such as higher temperature, than the condensation reaction. Thus at higher temperature in base the aldol reaction will go directly to the conjugated enone without any isolation of the aldol intermediate.

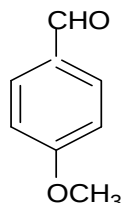
The aldehyde and ketone used for aldol reaction will be selected from the following group:



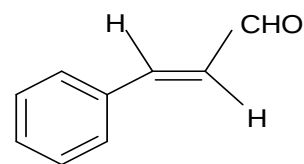
Benzaldehyde



4 - Methylbenzaldehyde
(p - tolualdehyde)



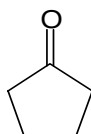
4 - Methoxybenzaldehyde
(p - anisaldehyde)



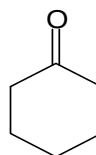
Cinnamaldehyde



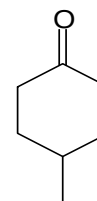
Acetone



Cyclopentanone



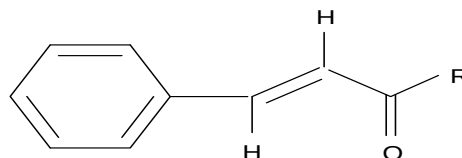
Cyclohexanone



4 - Methylcyclohexanone

Notice that none of the aldehydes shown above contains an enolizable α -hydrogen, so they cannot act as the nucleophilic species in the aldol reaction. Ketones, in general, are less susceptible to nucleophilic attack than aldehydes, so in a reaction mixture containing both an aldehyde and a ketone, the aldehyde will react faster with nucleophiles. Thus, it is possible to perform a "crossed" aldol reaction in which the enolate formed by abstraction of the α -hydrogen on the ketone attacks the carbonyl of the aldehyde. Since we are working with conjugated aldehydes, the resulting β -hydroxyketones readily eliminate water to form enones. Under the conditions used in this experiment (an excess of aldehyde), a "double condensation" occurs by reaction on both sides of the ketone to give the products shown below.

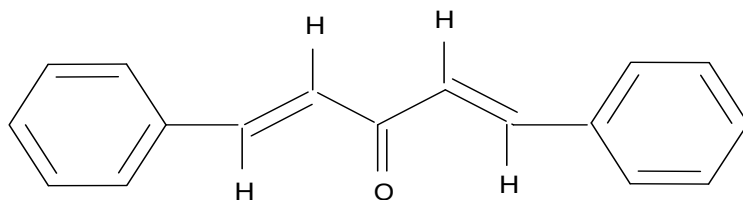
Although a β -hydroxyaldehyde (or a β -hydroxyketone) is produced in an aldol condensation, the ultimate product of these reactions (as shown above) is often the α,β -unsaturated aldehyde and a separate molecule of water. Upon heating, the β -hydroxy aldehyde product of an aldol condensation easily undergoes dehydration to yield an α,β -unsaturated aldehyde (or ketone). Conjugation of the newly formed double bond with the carbonyl group (and of the benzene ring, as shown in the example below) stabilizes the product and provides the thermodynamic driving force for the dehydration process.



In the present case, the reaction—a mixed, or crossed aldol condensation involving an aromatic aldehyde—is referred to as a Claisen-Schmidt condensation. The Claisen-Schmidt condensation always involves dehydration of the product of the mixed addition to yield a product in which the double bond (produced during dehydration) is conjugated to both the aromatic ring and the carbonyl group.

Because this aromatic aldehyde lacks α - hydrogens, only one product is formed, rather than a mixture of four different compounds, as long as the concentration of the second aldehyde is carefully controlled. In this

experiment we will prepare the dibenzalacetone: 1,5-diphenyl-1,4-pentadien-3-one. The equilibrium is shifted toward the product because the compound precipitates from the reaction mixture as it is formed.



1,5-diphenyl-1,4-pentadien-3-one

Dibenzalacetone is a common ingredient in sunscreen. This is because dibenzalacetone absorbs UV light and helps to protect the skin from the sun's damaging rays. The properties that are most valuable in a compound that is used in sunscreen are the compound's abilities to absorb, reflect, or even scatter the harmful UV rays. Another importance is for the compound to not cause an allergic reaction on a person's skin.

This experiment was being performed so that dibenzalacetone could be synthesized from benzaldehyde and acetone. This experiment was performed to show how a ketone and an aldehyde could be added together through the aldol condensation.

The aldol condensation is extremely important because it can form a β -hydroxy aldehyde or ketone from two carbonyl compounds. This type of reaction proceeds through the creation of a resonance-stabilized enolate ion from one of the carbonyl groups. The enolate ion can then act as a strong nucleophile and add to another carbonyl group. It is extremely important that one of the carbonyl groups has an acidic alpha hydrogen (one adjacent to a carbonyl group) so that the enolate ion can be formed. Aldol products can be formed through either acidic or basic conditions and since they are usually exothermic the reaction will be driven to completion.

In this experiment, you will run an aldol condensation between an aldehyde and a ketone and then the product of the reaction precipitates out of solution and can be collected by filtration. The crude product is normally purified by recrystallization. Weigh your product and determine percent yield. What reactant is your percent yield based on? Determine the melting point and compare to the literature value. (Table 1 and 2)

Table 1. Physical properties of compounds

Compound	MW	Mp	Bp	Density
Benzaldehyde	106.13	-56°C	178°C	1.04
Acetone (reagent)	58.08		56°C	0.79
Dibenzalacetone	234.30	113°C		

Table 2. Melting points of aldol condensation products

	Ketones			
Aldehydes	Acetone	Cyclopentanone	Cyclohexanone	4-Methylcyclohexanone
Benzaldehyde	111-113°C	189°C	118°C	98-99°C
4-Methylbenzaldehyde	175°C	235-236°C	170°C	133-135°C
4-Methoxybenzaldehyde	129-130°C	212°C	159°C	141-142°C
Cinnamaldehyde	144°C	225°C	180°C	163-164°C

Safety Note

- a) NaOH in aqueous ethanol is corrosive and particularly dangerous to the eyes. If contacted, remove with plenty of water.
- b) Acetone is highly flammable.
- c) Benzaldehyde is listed as moderately toxic (but contributes to the flavor of almonds)

Experimental Procedure

Transfer 15mL of ethanol into a 125-mL Erlenmeyer flask and add 20mL of 10% NaOH to it. Using a thermometer, cool the solution to 20°C.

In a medium size tube, mix 2mL of benzaldehyde with 15 drops of acetone, and leave it at room temperature for 5 minutes. Then, add the mixture to the ethanol-NaOH solution in small portions and stir with magnetic stirrer (if available) for 30 minutes. Chill the solution in an ice-water bath. Collect the yellow crystals by suction filtration and hand-dry them by pressing them between dry paper towels.

Determine the weight of the dibenzalacetone product, its melting point, and the percent yield. Return the product to your instructor.

NOTE- The amounts of the reagents used in this reaction are very important to ensure the correct product forms. In a given example a student added twice as much acetone as the procedure called for. This error on the student's part would greatly affect the end product. Since there would be such an excess of acetone the benzaldehyde would only see acetone and would not end up adding twice to any acetone molecules. This would give an end product of benzalacetone instead of dibenzalacetone.

EXPERIMENT 7 (Organic Chemistry II)

Aldol Condensation

REPORT FORM

Name _____

Instructor _____

Date _____

Reaction Equation:

-
1. Mass of benzaldehyde used _____ g
(Show calculation)
 2. Moles of benzaldehyde used _____ mole
(Show calculation)
 3. Moles of dibenzalacetone expected (theoretical yield) _____ mole
(Show calculation)
 4. Mass of dibenzalacetone expected _____ g
(Show calculation)
 5. Mass of dibenzalacetone recovered _____ g
 6. Percentage yield of cyclohexene _____ %
(Show calculation)
 7. Structure of dibenzalacetone
 8. Literature melting point _____ Observed melting point _____
Range

Pre-Laboratory Questions–EXP 7 Name: _____

Aldol Condensation

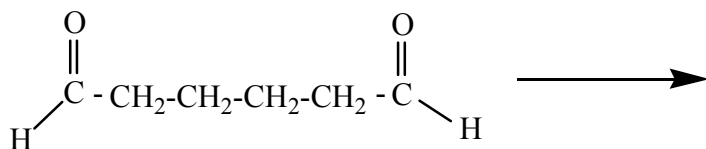
Due before lab begins. Answer in space provided.

1. 0.791 g acetone reacted with 1.044g benzaldehyde to produce experimentally 1.00 g dibenzalacetone. Calculate the theoretical and percentage yield.

2. Using aldol or crossed aldol condensation, suggest a synthesis of the following compounds.



3. What product would you expect to obtain from aldol cyclization of hexanedioal in basic solution.



4. Write an aldol condensation product(s) between the following compounds.

a) acetone + benzophenone (1:1 ratio)

b) cyclohexanone and cyclopentanone (1:1 ratio)

Post-Laboratory Questions–EXP 7 Name: _____

Aldol Condensation

Due after completing the lab.

1. If student added two fold of acetone the acetone will react with itself and the product would be isolated as diacetone alcohol or mesityl oxide. Write a complete reaction to support your result.
2. Draw the structure of the *cis* and *trans* isomers of the compound that you prepared. Why do you imagine that you obtained the *trans* isomer as the major, or even sole, product?
3. Draw a complete electronic mechanism of the aldol product between benzaldehyde and acetophenone.

4. Complete the following reactions.

