

# Identification of regions containing free water in hydrated matrix tablets made of sodium alginate

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# Introduction

Alginates are natural polymers which are extracted from marine algae. They are linear polysaccharides consisting of  $\beta$ -D-mannuronic acid (M) and  $\alpha$ -L-guluronic acid (G) residues. Most of the pharmaceutical applications of alginates are related to hydrogel formation. For this reason, studies that allow to elucidate the mechanisms of interaction between polymer chains and water molecules are of high priority for the understanding of these polymers functionality. The purpose of this work was to identify the regions containing free water in hydrated matrix tablets made of sodium alginate.

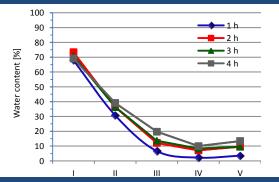
# **Materials**

Sodium alginate (Protanal LF 240 D, Batch No. S13455, FMC BioPolymers, USA);

AQUAMETRIC Composite 5 for volumetric analysis (Panreac AppliChem, Spain);

Methanol according to Karl Fischer (Panreac AppliChem, Spain);

Methanol for UHPLC (Panreac AppliChem, Spain) The distilled water was used to prepare the samples containing sodium alginate for both KF and DSC measurements.



# Figure 1. Water content change acording to the distance from the tablet surface (I – most external layer, V – most internal layer).

## Equipment

EXSTAR DSC 7020 apparatus (SII NanoTechnology Inc.) Karl-Fischer titration equipment (Metrohm, USA) Analytical weight (Sartorius, Germany) EKO laboratory tablet press (Korsch-Erweka, Germany) Home-made device for tablet hydrating and cutting layers

# Methods

#### **Tablets preparation**

12 mm, round flat faced tablets of pure sodium alginate were prepred using EKO laboratory tablet press.

## **Karl-Fischer titration**

The samples for Karl-Fisher titration were prepared in a home-made device which allows unilateral hydration of the tablet. The device contains a micrometric screw which allowes moving up the tablet in the holder to the reguired height and cutting a slice for futher analysis. At specific timepoint (1, 2, 3, 4 h), the device was removed from the medium, the slices were cut and weighted on analytical weight. In such a way 5 slices (of 1 mm each) were obtained. The samples were then put into flasks, filled with methanol, introduced into ultrasounds in order to extract total water amount existing in the sample and measured following Karl-Fischer protocol. The results are shown as the water content (%) in each slice of the hydrated tablet.

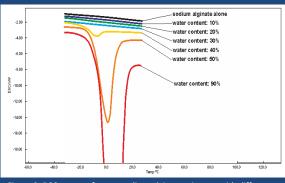


Figure 2. DSC curves of water-sodium alginate mixtures with different concentrations of water in heating run (endothermic process).

## Differential scanning calorimetry (DSC)

The samples were prepared by mixing sodium alginate and distilled water in a mortar in appropriate ratio to obtain final sample weight: 5g. At once 10-15 mg of each sample were put into an aluminum pan, sealed and measured in the temperature program: cooling run at 5°C/min from 20°C to -50°C and heating run at 10°C/min from -50°C to 70°C.

# **Results and discussion**

We observed that above 40% of water content, the DSC peak presenting the free water apeared in both cooling (data not shown) and heating run (Fig.2). Futhermore, unfreezing and freezing bound water was observed up to the concentration of 40%. The water content of 40% corresponded to the second slice of hydrated tablet according to KF experiment (Fig. 1). In this way we identified the regions (the first and the second slice) which contained free water in hydrated matrix of sodium alginate.

## **Conclusion**

In the present work we used KF method to determine the distribution of water content in the hydrated matrix tablets made of sodium alginate with a spatial resolution of 1mm. Additionally, we used DSC to figure out whether the free water exists in the sodium alginate-water mixtures containing rising water concentrations. Combining both methods served to identify the regions containing free water in hydrated matrix tablets made of sodium alginate.

# **References**

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