

CHARACTERIZATION OF B-CYCLODEXTRIN COMPLEXES WITH NATURAL DYE

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ABSTRACT

Many recent reports on curcumin, a polyphenol from *Curcuma Longa*, provide mounting evidence on the pharmacological activity of this natural product. However, the pharmaceutical use of this molecule is hampered due to its poor solubility in the aqueous media, inclusion complex formation with cyclodextrins has been reported as a means to enhance its aqueous solubility. Most of the studies provide Infrared (IR) spectroscopic data as an evidence to support inclusion complex formation. In this study, fully water soluble complexes of curcumin with two β -cyclodextrins were isolated and characterized.

Keywords: Natural dye, β -cyclodextrin, Hydroxyl propyl β -cyclodextrin, FTIR.

1. INTRODUCTION

Turmeric, derived from the rhizome of *Curcuma Longa* has been used by people of Indian subcontinent for centuries with no known side effects, not only as a component of food but also for a wide variety of ailments¹. Curcumin is the photochemical that gives yellow colour to turmeric. Extensive research within the last half century has proven that most of these activities, once associated with turmeric are due to curcumin². Curcumin is reported to have a number of pharmacological activities including antioxidant, HIV anti proteases activity, anti-inflammatory, analgesic, anticancer, etc.,³. Of late, its potent anti amyloidogenic effects in treating Alzheimer's disease have ignited wide spread research interest on this drug⁴. Pre-clinical and chemical trials have revealed that curcumin is safe even up to a dose level of 8.0g and this makes it all the more important⁵. But the pharmaceutical use of this pharmacologically potential molecule is restricted due to its poor aqueous solubility resulting in reduced bio availability⁶.

One way to increase its aqueous solubility is to form inclusion complex i.e., to encapsulate curcumin as a guest within the internal cavity of a water soluble host⁷. Cyclodextrin have been used extensively in pharmaceutical research and development, and there are products numerous cyclodextrin containing pharmaceutical

marketed worldwide⁸. The most common pharmaceutical application of CDs is to enhance drug solubility in aqueous solutions. CDs are also used for increasing stability and bioavailability of drugs, and other additional applications⁹. Many studies on enhancement of solubility of curcumin with cyclodextrin have been reported¹⁰⁻¹³. Recent reports on curcumin cyclodextrin complexes show the vast amount of research going on in this field worldwide¹⁴⁻¹⁷. In recent papers¹⁵⁻¹⁶ also FTIR spectroscopy was used for studying the complex formed between curcumin and hydroxyl propyl β -cyclodextrin.

2. EXPERIMENTAL

Curcumin (1, 7-bis (4-hydroxy-3-methoxyphenyl)-1, 6-heptadiene-3, 5-Dione), β -Cyclodextrin and Hydroxyl Propyl β -Cyclodextrin were purchased from Sigma Aldrich Company Bangalore. The chemicals were used without further purification. FTIR spectra of curcumin, cyclodextrins and the physical mixtures complexes were recorded using **Thermonicolet iS5 FTIR spectrometer**. The scanning range used was 400-4000 cm^{-1} .

3. RESULT AND DISCUSSION

Although many studies have been carried out on curcumin-cyclodextrin complexes, characterization of fully water soluble

complexes have not been reported yet. Characterization techniques used in earlier studies especially FTIR, could not provide adequate evidence for inclusion complex formation [15, 16]. The method used for preparing inclusion complexes is another important factor to be considered. In the present study, the soluble curcumin-cyclodextrin complexes were filtered through 0.45 μm filter in order to remove any insoluble curcumin present. The filtrate thus obtained was freeze dried to obtain solid complexes. In all earlier reported studies the investigations had not filtered out their complexes with curcumin and as a result the characterized products were mixtures of uncomplexed curcumin, complexed curcumin and uncomplexed cyclodextrin. Due to this reason FTIR spectral study used in the earlier reported works is not likely to provide adequate information on the inclusion complex formation.

3.1. FTIR spectrum of curcumin

A detailed study on the vibration spectra of curcumin has been reported earlier by Kolev et al. [18]. The FTIR spectrum of curcumin is shown in Fig. 1. The sharp peak at 3435 cm^{-1} indicates the presence of OH. The strong peak at 3001 cm^{-1} attributed to CH stretching. The 1492 cm^{-1} peak is assigned to the γ (C-H) bending, while, OH bend was obtained at 1423 cm^{-1} . C-O stretching at 1317 cm^{-1} , C-O-H stretching was observed at 1234 cm^{-1} and C-H bending at 690 cm^{-1} .

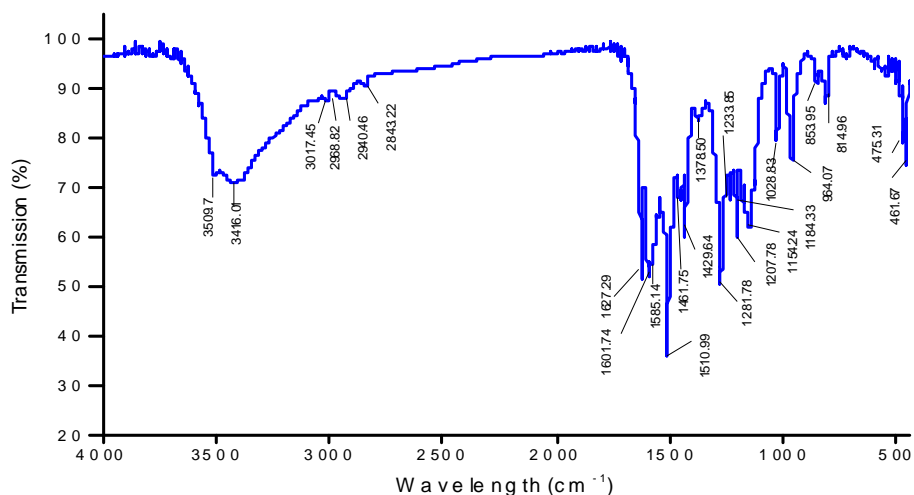


Fig. 1: FTIR spectrum of curcumin

3.2 FTIR spectra of cyclodextrins

The overlaid FTIR spectra of β CD and H β CD are given in Fig. 2. The FTIR peaks of the two β -cyclodextrins are almost identical with slight variations in their intensities. The OH group stretching, the H-bonded OH stretching observed at 2455 cm^{-1} and C=O asymmetry stretching at 1797 cm^{-1} . Since there are no significant variations in the spectra of different cyclodextrin complexes of curcumin with cyclodextrins were selected for detailed analysis.

3.3 FTIR spectra of (Cur+ β CD) and (Cur+H β CD) complexes

FTIR spectra of curcumin, mixture of curcumin and β CD and the mixture of curcumin and H β CD are shown in Fig. 3. The peak assignments of all the spectra are given in Table 1. A little significant changes in the IR spectra of complexes have been observed, show the complex formation of β -cyclodextrins and curcumin. All the major peaks of curcumin are hidden by the cyclodextrin peaks in the similar region.

4. CONCLUSION

FTIR spectroscopy provided clear and distinct evidence for inclusion complex formation of curcumin with β -cyclodextrin and hydroxyl propyl β -cyclodextrin. Complexes were found to be formed when β CDs and curcumin were taken in the ratio 1:1.

Table 1: FTIR peak assignment of curcumin, β CD, H β CD and complexes

Curcumin	β CD (γcm^{-1})	H β CD (γcm^{-1})	Cur+ β CD (γcm^{-1})	Cur+H β CD (γcm^{-1})	Peak assignment
3509.7	3425.58	3435.22	3429.43	3433.29	OH stretching of phenol group OH stretching
2968.82	2921.37	2921.37	2935.30	2937.30	Asymmetric CH stretching OF CH ₃
2940.46	1656.85	1658.78	1656.85	1656.85	CH stretching of OCH ₃ Asymmetric CH stretching OF CH ₂
1627.29	1423.47	1433.11	1423.47	1423.47	HOH of water of crystallization, C=C stretching
1601.74	1186.22	1026.13	1317.38	1317.38	
1510.99	1029.99	952.84	1188.15		
1429.64	948.98	801.75			C=O, Cring-C=C stretching Aromatic C=C stretching
1281.78	813.96	763.81	1033.85		
1154.24	705.95	704.02	948.98	1037.70	C=O stretching, CCC, CC=O in plane bending In plane bending of aromatic (CCC, CCH), enolic (COH), CH in plane
1028.83				948.98	In plane bending of CH, enolic COH, skeletal CCC CH in plane bending of C=CH, aromatic C-O stretching
853.95					CH overtone stretching C- O-C stretching In plane bending of aromatic CCH, skeletal CCC C-O, C-C, CCO, C-O-C stretching of glucose units C-O-C stretching out of plane of CH ₃ , in plane bending of aromatic CCH C=O stretching, in plane bending of CCH CH out of plane of aromatic CCH C-O-C stretching Skeletal C-C stretching CH out of plane bending In plane bending of skeletal CCH and aromatic CCH, C=C stretching

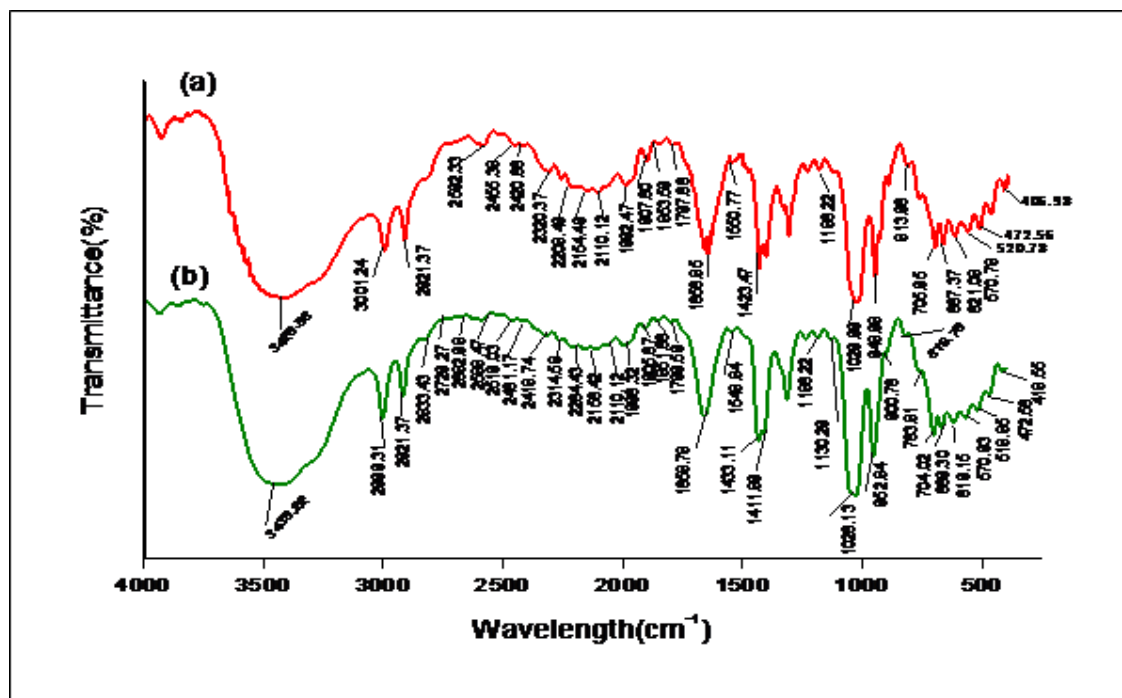


Fig. 2: FTIR spectrum of cyclodextrins (a) β CD (b) H β CD

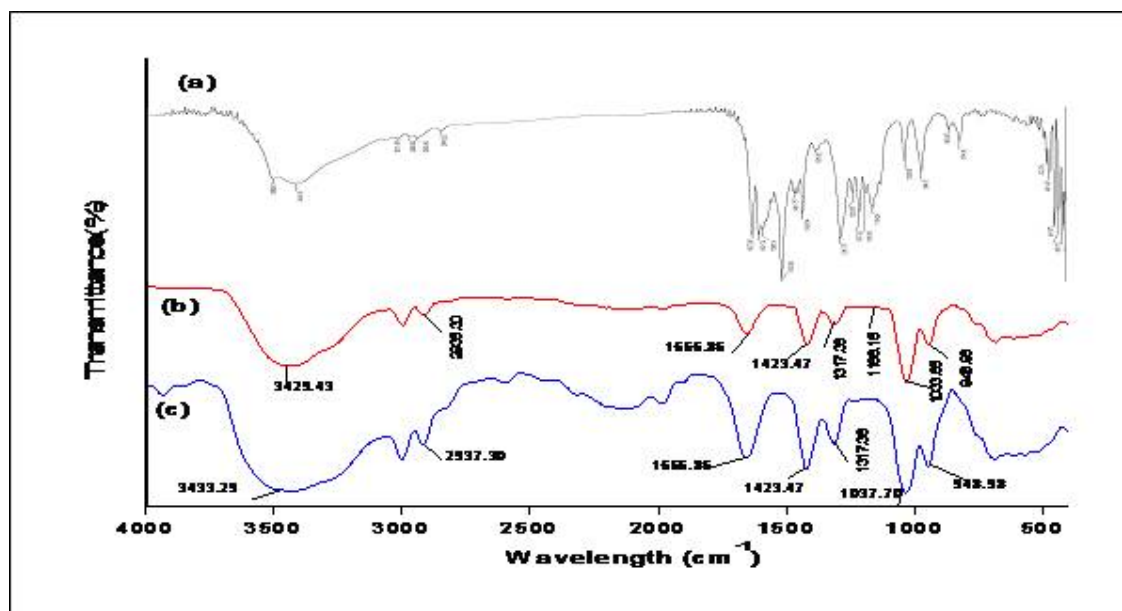


Fig. 3: FTIR spectrum of curcumin complexes (a) Curcumin, (b) Curcumin+ β CD (c) Curcumin+H β CD

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