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ALGINATE-CHITOSAN MICROCAPSULES AS DELIVERY SYSTEMS FOR TRAMADOL

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Chitosan is a biocompatible, non toxic and easily bioabsorbable polymer [1] with gel-forming ability at low pH. Moreover, chitosan has antiacid and antiulcer activities that prevent or weaken drug irritation in the stomach [2]. Chitosan matrix formulations also appear to float and gradually swell in an acid medium.

Encapsulation with alginates is the most often carried out by dispersing the alginate encapsulant solution into a calcium chloride gelation medium. More recently, microcapsules with an alginate gel core and a polyanion-polycation membrane have been suggested as candidates for the oral delivery of a wide range of compounds including drugs [3,4] passing the stomach to intestinal sites of absorption.

Diffusion and swelling are the most important rate-controlling mechanisms of systemic controlled release products [5]. The power law equation of Peppas can be regarded as a generalization of the observation that the superposition of two apparently independent mechanisms of drug transport, a Fickian diffusion and a case II transport [6], describes in many cases the dynamic swelling of drug release from polymers, regardless of the form of the constitutive equation and the type of coupling of relaxation and diffusion [7].

Tramadol is a centrally acting analgesic with low affinity for opioid receptors where it has selectivity for μ -receptors [8]. Its limiting side-effects in the treatment of acute and chronic pain are reported to be less intense and less frequent than with other opioids. A low drug delivery system is particularly suitable for the formulation of an analgesic agent. Tramadol is a good candidate drug to be formulated in a sustained-release dosage form.

The purpose of the current study was to prepare microcapsules for the controlled release of Tramadol hydrochloride by a simple technique that uses alginate-chitosan as a matrix. We studied two different methods of alginate-chitosan capsule formation and their effect on the release of Tramadol hydrochloride from microcapsules.

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REFERENCES

1. P. Giunchedi, I. Genta, B. Conti, R.A.A. Muzzarelli, *Biomaterials*, 19 (1998)157.
2. W.M. Hou, M. Mayasaki, M. Takada, T. Tomai, *Chem. Pharm. Bull.*, 33 (1985) 3986.
3. P.R. Hari, T. Chandy, C.P. Sharma, *J. Microencapsulation*, 13 (1996) 319.
4. S. Takka, O.H. Ocak, F. Acartürk, *Eu. J. Pharmaceut. Sci.* 6 (3) (1998) 241.
5. R. Langer, N. A. Peppas, *Rev. Macromol. Chem. Phys.* C23 (1983) 61.
6. A. Peppas, in: V.F. Smolen, L. Ball (Eds), *Controlled Drug Bioavailability*, Vol. 1, John Wiley & Sons, New York, 1984, pp. 203-237.
7. H. Frisch, *Polym. Eng. Sci.* 20 (1980) 2.
8. A.A. Obaidat & M. Obaidat, *Eur. Pharm. Biopharm.*, 52 (2001) 231.