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Thematic course: Synthesis and study of the properties of composite materials based on cellulose and chitosan containing various therapeutic agents. Part 3.

## Hydrolytic destruction of dressings based on dialdehydecellulose

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## Abstract

Creating systems for targeted delivery of drugs to the affected organ is one of the most promising areas for the development of systems with controlled release of the active substance. Polysaccharides are widely used as drug carriers. However, most of them are chemically inert and require preliminary functionalization in order to interact with physiologically active compounds (therapeutic agents-TA). A simple and effective method for introducing reactive groups is the periodic oxidation of the polysaccharide by the Malaprade reaction. While cellulose is insoluble in water and resistant to weak solutions of acids and alkalis, dialdehyde cellulose (DAC is the product of the periodic oxidation of cellulose) and its derivatives are destroyed in water and weakly acidic and slightly alkaline solutions. This process is called hydrolytic destruction. The kinetics of hydrolytic destruction is described by semi-logarithmic anamorphosis, which allows us to calculate the rate constants of hydrolytic destruction as the rate constant of first-order reactions. The products of hydrolytic degradation were studied by UV spectroscopy and using 3,5-dinitrosalicylic acid (DNSA). The degradation products of C and DAC were also studied by the phenol-sulfur method. From the data presented and cited earlier, it follows that when our composite material is placed in a liquid medium, the hydrolytic destruction of the drug immediately begins. What can be connected with the breakdown of both the carrier - TA bonds (DAC, C, Ct carriers) and the destruction of the carrier itself. Under the conditions of the organism, biological destruction can also join process. Biodestruction is the process of destruction (both carriers and immobilized TAs) under the action of the body's enzymes. Using IR spectroscopy, cellulose carriers were studied before and after exposure to 1/15M FB medium (pH 6.2 and 37 °C) for 48 hours. As can be seen from the data obtained, primarily for DAC samples, significant changes in the spectrum are visible in the 1800-1600 and 900  $\text{cm}^{-1}$  fields.

The results of experimental toxicological studies of samples of various cellulosic materials allow us to conclude that the samples studied do not have toxic, hemolytic, allergenic effects, as well as mutagenic activity.

## References

- [1] S.N. Marychev, B.A. Kalinin. Polymers in medicine: Textbook. allowance. Vladim. state un-t; Vladimir. **2001**. 68p. (russian)
- [2] A.A. Belov. Development of industrial technologies for obtaining new medical materials based on modified fiber-forming polymers containing biologically active protein substances. Diss. on cois. uch. step. Dr. tech. Science M., MUCTR. 2009. 385p. (russian)
- [3] V.A. Dvatlov, I.S. Kruppa, S.A. Mamaeva et al. Change of polysaccharide molecular-weight distribution and fraction homogeneity after periodate oxidation. Chemistry of Natural Compounds. 2014. Vol.50. No.6. P.973-977.
- [4] Z.A. Rogovin. Chemistry of cellulose. *Moscow: Chimia.* 1972. P.125-244. (russian) (Malaprade L., Bull. Soc. chim. France, 1928. Vol.43. P.683.).
- [5] M.I. Shtilman. Polymers in biologically active systems. Soros educational J. 1998. No.5. P.48-53. (russian)

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- [6] M. Singh, A.R. Ray, P. Vasudevan, K. Verma, S.K. Guha. Potential biosoluble carriers. *Biomaterials*. 1979. Vol.7(4). P.495-512.
- [7] M. Singh, A.R. Ray, P. Vasudevan. Biodegradation studies on periodate oxidized cellulose. *Biomaterials*. 1982. Vol.3. P.16-20.
- [8] L.G. Vlasov, R.B. Virnik. The study of absorbable textile materials containing immobilized enzymes. Adj biochem. and a microbe. 1988. Vol. XXIV. Iss.2. P.264-268. (russian)
- [9] V.V. Ryltsev, R.B. Virnik. Investigation of the kinetics of the release of trypsin immobilized on dialdehyde cellulose during hydrolytic destruction. Antibiotics and chemotherapy. 1989. Vol.34. No.3. P.202-205. (russian)
- [10] Ung-Jin Kim, Masahisa Wada, Shigenori Kuga, Solubilization of dialdehyde cellulose by hot water. Carbohydrate Polymers. 2004. Vol.56. P.7-10.
- [11] Luka's' Mu"nster . Jan Vı'cha . Jir'ı' Klofa et al Stability and aging of solubilized dialdehyde cellulose. Cellulose. 2017. Vol.24. P.2753-2766.
- [12] V.N. Filatov, V.V. Ryltsev. Biologically active textile materials. *Moscow.* **2002**, 248p. (russian)
- [13] A.A. Vaniushenkova, E.E. Dosadina, S.V. Kalenov, N.S. Markvichev, and A.A. Belov. Synthesis and study of the properties of composite materials based on cellulose and chitosan containing various therapeutic agents. Part 2. Effect of chitosan on the destruction of cellulosic carriers and the kinetics of release of the therapeutic agent in the model environment. Butlerov Communications. 2019. Vol.57. No.3. P.105-119. DOI: 10.37952/ROI-jbc-01/19-57-3-105
- [14] E.E. Dosadina, M.A. Kulmetieva, O.E. Dubovikova, A.Yu. Evdokimenko, E.E. Savelyeva, and A.A. Belov. Synthesis and study of the proteinase complex properties immobilized on polysaccharide carriers for medical use. Butlerov Communications. 2016. Vol.46. No.6. P.1-10. DOI: 10.37952/ROI-jbc-01/16-46-6-1
- [15] E.E. Dosadina, M.A. Bikineeva, A.Y. Evdokimenko, E.E. Savelyeva, E.O. Medusheva, and A.A. Belov. Immobilization of proteinases of proteolytic complex of hepatopancreas of crab on some polysaccharides: production, properties, application. Butlerov Communications. 2016. Vol.48. No.12. P.83-93. DOI: 10.37952/ROI-jbc-01/16-48-12-83
- [16] A.A. Belov, A.A. Vanyushenkova, E.E. Dosadina, A.A. Hanafina. New textile dressings based on biodegradable polymers containing proteinases for the treatment of wounds and burns. Wounds and wound infections. Magazine them. prof. B.M. Kostyuchenka. 2018. Vol.5. No.1. P.16-26. (russian)
- [17] E.E. Dosadina, E.E. Savelyeva, A.A. Belov. The effect of immobilization, drying and storage on the activity of proteinases immobilized on modified cellulose and chitosan. Process Biochemistry. 2018. Vol.64. P.213-220.
- [18] V.I. Gumnikova. Synthesis of dialdehydedextran and dialdehyde carboxymethyl cellulose and their chemical transformations. Diss. for a job. student Art. Ph.D., Moscow: RHTU. 2014. P.137.
- [19] G.L. Miller. Use of dinitrosalicylic acid reagent for determination of reducing sugar. *Anal. Chem.* **1959**. Vol.31. No.3. P.426- 428.
- [20] Irina Sulaeva, Karl Michael Klinger, Hassan Amer et al. Determination of molar mass distributions of highly oxidized dialdehyde cellulose by size exclusion chromatography and asymmetric flow field-flow fractionation. Cellulose. 2015. Vol.22. Iss.6. P.3569-3581.
- [21] M. Dubois, K.A. Gilles, J. Hamilton, P.A. Robers, F. Smith. Colorimetric method fordetermination of sugars and related substances. Analyt. Chem. 1956. Vol.28. P.350-356.
- E.E. Savelyeva, A.A. Vanyushenkova, A.A. Belov. The destruction of cellulose carriers in a model [22] environment. Sat scientific Proceedings in chemistry and chem. technology. 2018. Vol.XXXII. No.12. P.27-30. (russian)
- [23] A.A. Vanyushenkova, A.A. Belov, E.E. Savelyeva et al. Bio- and hydrolytic destruction of medical wound healing polysaccharide materials in conditions simulating a purulent wound. Proceedings of the XV annual. int. young conf. IBChF RAS-Universities "Biochemical Physics", M. 2018. P.27-30. (russian)
- [24] M.A. Bychuk. Obtaining and properties of polymer films based on poly-3-hydroxybutyrate and poly-εcaprolactone. Diss. for a job. student Art. Ph.D., M., MGUDT. 2016. 169p. (russian)
- [25] A.P. Popov. System analysis, modeling and control of the batch process of thermo-oxidative degradation of polymers in solution. Diss. for a job. student Art. Ph.D., Voronezh, Voronezh State University. 2015. 242p. (russian)
- V.A. Kaminsky, A.A. Kuznetsov. On the kinetics of polymer destruction according to the law of [26] random chain breaking. Theoretical. basics of chem. technology. 2012. Vol.46. No.4. P.453-457. (russian)

- [27] A.Yu. Budantsev. Fundamentals of histochemistry: Textbook (computer version) section 2. Microtechnology. Pushchino: Pushchino state. University. 2008. (russian)
- [28] E.E. Dosadina, E.E. Savelyeva, A.A. Belov. Interaction Between Chitosan Solutions, Cellulose Carriers and Some of the Multi-enzyme Complexes. International Journal of Bioorganic Chemistry. 2017. Vol.2. Iss.2. P.51-60.
- [29] E.E. Dosadina, M.A. Kulmetyeva, A.A. Belov. The changing of enzymatic activity of hydrolass immobilized on natural polysaccharide matrix for purulent and burn wounds treatment during storing and exploitation. Biointerface Research in Applied Chemistry. 2016. Vol.5. Iss.3. P.1291-1298.
- [30] Q.G. Fan, D.M. Lewis, K.N. Tapley. Characterization of Cellulose Aldehyde Using Fourier Transform Infrared Spectroscopy. Journal of Applied Polymer Science. 2001. Vol.82. P.1195-1202.
- [31] R.G. Zhbankov, R.M. Marunov, N.V. Ivanova, A.M. Shishko. Spectroscopy of cotton. *Moscow:* Science. 1976. P.248. (russian)
- [32] Xihong He, Zhina He, Yan Li et al. Modeling of the bacterial inactivation kinetics of dialdehyde cellulose in aqueous suspension. International Journal of Biological Macromolecules. 2018. No.116. P.920-926.