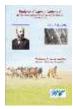


Butlerov Communications C Advances in Biochemistry & Technologies ISSN 2074-0948 (print)

**2021**. Vol.1, No.2, Id.9. Journal Homepage: https://c-journal.butlerov.com/



**Full Paper** 

*Thematic section:* Biochemical Research. *Subsection:* Medical Chemistry.

The Reference Object Identifier – ROI-jbc-C/21-1-2-9 The Digital Object Identifier – DOI: 10.37952/ROI-jbc-C/21-1-2-9 Received 10 June 2021; Accepted 10 June 2021

## Synthesis and study of the properties of chitosan-containing materials including proteases and various therapeutic agents. Part 5. The effect of chitosan on the preservation of the enzymatic activities of polyenzyme preparations during production and operation

## Oleg V. Matiev, and Alexey A. Belov\*+

Department of Biotechnologies. Mendeleev University of Chemical Technology of Russia. Heroes Panfilovtsev St., 20. Moscow, 125480. Russia. Phone: +7 (499) 978-95-15. E-mail: ABelov2004@yandex.ru

\*Supervising author; \*Corresponding author *Keywords:* immobilized proteases, chitosan, hydrolytic degradation, therapeutic agent, wound healing materials.

## Abstract

One of the directions in the development of medical biotechnology is the development and introduction into medical practice of new highly effective drugs for the treatment and prevention of wound healing based on biopolymers. Selective accumulation of the drug in the lesion focus allows solving several problems at the same time: increasing the effectiveness of the drug, reducing its consumption, eliminating the undesirable effect of the drug on healthy organs and tissues. One of the problems in the modification of a therapeutic agent with a polymer is the possible loss of biological activity immediately after modification, or during storage, or during operation (liquid medium, pH and temperature 37 °C).

An important aspect in the use of polymers as drug delivery systems is their place in human metabolism or biodegradability. Polysaccharides are widely used as drug carriers. At present, all over the world, there is an increase in the interest of specialists in preparations based on chitin.

The most important derivative of chitin is chitosan, which, unlike chitin, is soluble in dilute acids, which expands the possibilities of its practical application for practical use. Due to its chemical nature, chitosan is capable of various types of interactions with the formation of 4 main types of bonds. Therefore when chitosan in a cationic form is added to aqueous solutions (dispersions) of mineral, organic, or living objects, depending on the concentration, either flocculation or stabilization of particles occurs in an aqueous medium.

The work studied the effect of cysteine, glucose, glucosamine, N-acetyl glucosamine and chitosan on the preservation of the enzymatic activity of proteases in the process of obtaining composites. The multidirectional effects of the studied factors on the enzymatic activities of the investigated proteases were established.

**For citation:** Oleg V. Matiev, Alexey A. Belov. Synthesis and study of the properties of chitosan-containing materials including proteases and various therapeutic agents. Part 5. The effect of chitosan on the preservation of the enzymatic activities of polyenzyme preparations during production and operation. *Butlerov Communications C.* **2021**. Vol.1. No.2. Id.9. DOI: 10.37952/ROI-jbc-C/21-1-2-9

## References

- A.A. Vanyushenkova, E.E. Dosadina, A.A. Hanafina, S.N. Ivanova, 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 1. The effect of drying and shelf life on the properties of chitosan composites. *Butlerov Communications*. 2019. Vol.57. No.2. P.130-143. DOI: 10.37952/ROI-jbc-01/19-57-2-130 (Russian)
- [2] V.N. Gypsy, Zhogolev K.N., Nikitin V.Yu. Chitosan as a Pharmacist. Nutritional supplements market. *St. Petersburg: "Faros Plus"*. 2002. No .2(4). P.8. (Russian)
- [3] K.G. Skryabin, G.A. Vikhoreva, V.P. Varlamov. Chitin and Chitosan. Obtaining, properties and application. *Moscow: Science*. **2002**. P.368. (Russian)
- [4] D.A. Gryadskikh. Synthesis of composite affinity sorbents with magnetic properties and their technological use in the manufacture of plague immunobiological preparations. *PhD Thesis (Candidate Level on Biological Sciences). Stavropol.* 2004. P.153. (Russian)
- [5] E.E. Dosadina, L.L. Brkich, N.V. Pyatigorskaya, M.A. Bikineeva, A.Y. Evdokimenko, E.E. Savelyeva, E.O. Medusheva, A.S. Kulagina, L.A. Pavlova, and A.A. Belov. Use of chitosan as a carrier for proteinases and Miramistin for obtaining of enzyme-containing gel. *Butlerov Communications.* 2016. Vol.48. No.10. P.49-59. DOI: 10.37952/ROI-jbc-01/16-48-10-49 (Russian)
- [6] T.M. Safronov, T.M. Boytsova. Chitosan as a flocculant of native fish protein. *Moscow.* 1999. P.251-252. (Russian)
- [7] N.V. Melnik, N.D. Skichko, E.A. Shubina. The use of chitosan and its derivatives for the concentration of bulk suspensions. *Moscow.* 1999. P.170-171. (Russian)
- [8] A. Domard. Some physicochemical and structure basis for applicability of chitin and chitosan. *Thesis 2-th Asia Pacific symposium Chitin and Chitosan. Bangkok.* **1996**. P.1-12.
- [9] A.I. Slivkin, V.L. Lapenko, A.P. Arzamastsev, A.A. Bolgov. Aminoglucans as biologically active components of drugs (review for the period 2000-2004). VSU Bulletin. Ser. Chemistry. Biology. Pharmacy. 2005. No.2. P.73-87. (Russian)
- [10] A.I. Slivkin, V.L. Lapenko, A.A. Bolgov. Synthesis of medicinal analogues of chitosan. VSU Bulletin. Ser. Chemistry. Biology. Pharmacy. 2005. No.2. P.205-208. (Russian)
- [11] A.A. Belov. Development of industrial technologies for obtaining new medical materials based on modified fiber-forming polymers containing biologically active protein substances. *PhD Thesis (Candidate Level on Techlogical Sciences). Moscow: RCTU.* 2009. 385p. (Russian)
- [12] Barbara Krajewska Application of chitin- and chitosan-based materials for enzyme immobilizations: a review. *Enzyme and Microbial Technology*. **2004**. Vol.35. P.126-139.
- [13] Maximiliano L. Cacicedo, Ricardo M. Manzo, Sofi'a Municoy et al Immobilized Enzymes and Their Applications. *Advances in Enzyme Technology*. 2019. P.169-200. doi.org/10.1016/B978-0-444-64114-4.00007-8

- [14] E.E. Dosadina, A.A. Belov. Interaction between chitosan solutions, cellulose carriers and some of the multi-enzyme complexes. *International Journal of Bioorganic Chemistry*. 2017. Vol.2. No.2. P.51-60. doi: 10.11648/j.ijbc.20170202.12
- [15] E.E. Dosadina, M.A. Kulmetyeva, A.A. Belov. The changing of enzymatic activity of hydrolases immobilized on natural polysaccharide matrix for purulent and burn wounds treatment during storing and exploitation. *Biointerface Research in Applied Chemistry*. 2016. Vol.6. Iss.3. P.1291-1298.
- [16] E.E. Dosadina, E.E. Savelyeva, L.L. Brkich, A.A. Hanafina, A.A. Vaniushenkova, A.Yu. Evdokimenko, Al Okbi Hidayer Mahmoud Ali, and A.A. Belov. The effect of drying and shelf life on the biological properties of chitosan composites containing enzymes and various therapeutic agents. *Butlerov Communications*. 2018. Vol.55. No.7. P.64-73. DOI: 10.37952/ROI-jbc-01/18-55-7-64 (Russian)
- [17] E.E. Dosadina, A.A. Belov. Possible mechanisms of interaction of chitosan, cellulose carriers and proteins of the proteolytic complex from the crab hepatopancreas. *Advances in Chemistry and Chemical Technology*. **2015**. No.8. P.82-84. (Russian)
- [18] E.E. Dosadina, M.A. Bikineeva, A.Yu. Evdokimenko, E.E. Savelyeva, E.O. Medusheva, A.A. Belov. Study of the interaction of chitosan and some hydrolases. *Proceedings of Universities. Applied Chemistry and Biotechnology.* 2017. Vol.7. No.1. P.62-71. DOI: 10.21285/2227-2925-2017-7-1-62-71
- [19] 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 (Russian)
- [20] V.V. Mosolov. Proteolytic enzymes. Moscow: Science. 1971. 414p. (Russian)
- [21] G.K. Ziyatdinova, L.V. Grigorieva and G.K. Budnikov. Reactions of papain with electrogenerated halogens and their analytical applications. *Butlerov Communications*. 2007. Vol.12. No.7. P.16-19. ROI-jbc-01/07-12-7-16 (Russian)
- [22] A.M. Pendzhiev, A. Abdullaev. Efficiency of using proteolytic enzymes of papaya in medical practice. *Scientific Review. Medical Sciences.* 2017. No.1. P.57-72. (Russian)
- [23] N.A. Efimenko, M.V. Lysenko, Yu.I. Sternin et al. Proteolytic enzymes in surgery: historical aspects and modern concepts of application. *Russian Medical Journal. Surgery.* 2011. Vol.19. No.5. P.368-372. (Russian)
- [24] G.N. Rudenskaya, V.A. Isaev, A.M. Shmoylov et al. Preparation of proteolytic enzymes from kamchatka crab *Paralithodes Camchatica* hepatopancreas and their application. *App. Biochem. Biotech.* **2000**. Vol.22. P.175-184.
- [25] Methods of optical spectroscopy. Ed. Kulakova I.I., Fedorova O.A., Khoroshutina A.V. Moscow State University. 2015. P.117. (Russian)
- [26] Ira B. Klein and Jack F. Kirsch. The mechanism of the activation of papain. *Bioch. and Bioph. Res. Com.* 1969. Vol.34. No.5. P.575-581.
- [27] Mireille St-Vincent and Michael Dickman. Chemical modification of papain and subtilisin: an active site comparison. *Journal of Chemical Education*. 2004. Vol.81. No.7 July, P.1048-1050.
- [28] T. Vernet, P.J. Berti. Chantal de Montigny et.al Proctssing of the papain precursor. J. of Biol. Chem. 1995. Vol.270. Iss.18, 5 May. P.10638-10646.
- [29] M. David. Greenberg and Theodore Winnick Plant proteases I. Activation-inhibition reactions. *J. of Biol. Chem.* **1940**. Vol.135. Iss.2, 1 September. P.761-773.
- [30] Ira B. Klein and Jack F. Kirsch. The Activation of Papain and the Inhibition of the Active Enzyme by Carbonyl Reagents. J. of Biol. Chem. 1969. Vol.244. No.21. Iss.10. P.5928-5935.
- [31] S.D. Varfolomeev. Chemical enzymology. *Moscow: Academy.* 2005. 472p. (Russian)
- [32] Ali KILINC, Se\_cil "ONAL and Azmi TELEFONCU Stabilization of Papain by Modification with Chitosan. *Turk J Chem.* **2002**. Vol.26. P.311- 316.

- [33] V.V. Chernova. Destruction of chitosan under the influence of certain enzyme preparations for medical purposes. *PhD Thesis (Candidate Level on Chemical Sciences). Ufa.* **2011**. 129p. (Russian)
- [34] Vincenzo Guarino, Tania Caputo, Rosaria Altobelli, Luigi Ambrosio. Degradation properties and metabolic activity of alginate and chitosan polyelectrolyte's for drug delivery and tissue engineering applications. *AIMS Materials Science*. 2015. Vol.2(4). P.497-502.
- [35] E.I. Kulish, V.V. Chernova, R.F. Vildanova et al. About the reason for the enzymatic hydrolysis of chitosan under the action of some nonspecific enzymes. *Bulletin of the Bashkir University*. 2011. Vol.16. No.3. P.681-683. (Russian)
- [36] A.V. Bakulin, A.A. Lvov, A.I. Albulov, Tran Din Tbay. Depolymerization of high molecular weight chitosan by the enzyme phytopaine. *New perspectives in the study of chitin and chitosan: Mat. Ninth Int. Conf. Moscow: VNIRO.* **2008**. P.241-243. (Russian)
- [37] Kumar A.B. Vishu, Tharanathan R.N. A comparative study on depolymerization of chitosan by proteolytic enzymes. *Carbohydrate Polymers*. **2004.** Vol.58. P.275-283.
- [38] I.S. Boyko, O.A. Podkolodnaya, S.G. Lysachok, S.L. Shmakov. Viscous degradation of acidic solutions of chitosan and its study by the ion probe method. Proseeding of *Saratov University. New Ser. Chemistry. Biology. Ecology.* 2015. Vol.15. Iss.4. P.21-30. (Russian)
- [39] Emilia Szymańska and Katarzyna Winnicka Stability of Chitosan A Challenge for Pharmaceutical and Biomedical Applications. *Mar. Drugs.* 2015. Vol.13. P.1819-1846. doi:10.3390/md13041819
- [40] Nazma N. Inamdar and Vishnukant Mourya. 6 Chitosan and Low Molecular Weight Chitosan: Biological and Biomedical Applications. Ashutosh Tiwari and Anis N. Nordin (eds.) Advanced Biomaterials and Biodevices. 2014. P.183-242.
- [41] Oleg V. Matiev, Alexey A. Belov. Synthesis and study of the properties of chitosancontaining materials including proteases and various therapeutic agents. Part 5. The effect of chitosan on the preservation of the enzymatic activities of polyenzyme preparations during production and operation. *Butlerov Communications*. 2021. Vol.66. No.6. P.76-84. DOI: 10.37952/ROI-jbc-01/21-66-6-76 (Russian)