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Rehabilitation**

## **Method for Increasing the Production or Activity of Catalase in the Body**

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### **Abstract**

The aim of this study is to develop a new, more sensitive and accurate method for the induction and/or activation of catalase (CAT) in the body. The proposed goal is achieved by using bromo-2- $\{[2-(prop-2-en-1-ylcarbamothioyl)-hydrazinylidene]methyl\}$ phenolacopper – a coordinating compound from the class of thiosemicarbazidates of transition metals, which expands the arsenal of synthetic compounds with high CAT induction and/or activation activity. The synthesis method of this compound and its structural formula are described. It was established that this compound exhibits the highest induction and/or activation of catalase, which exceeded 2.7 the values of the control group and 1.8 the values produced by vitamin D<sub>3</sub> (prototype). This indicates the existence of an excessive synthesis of catalase after exposure to this compound, a particularly important fact established by us for the first time.

This compound can be used in medicine as a therapeutic agent, which, by activating the important production of catalase in the body, can prevent and/or reduce the occurrence of neurodegenerative, renal, cardiovascular pathologies, atherosclerosis and carcinogenesis, inflammatory processes, the development of cellular and tissue damage, associated with excessive accumulation of hydrogen peroxide.



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The obtained data mark a beginning that opens the perspectives of developments, which will diversify the arsenal of effective tools to combat various severe pathological processes.

*Keywords: copper coordinating compounds, thiosemicarbazide derivate, catalase activating, multifactorial diseases*

## References

1. Nandi, A., Yan, L.-J., Jana, C.K., Das, N.: Role of catalase in oxidative stress- and age-associated degenerative Diseases. *Oxidative Med, Cellular Longevity* **2019**, Article ID 9613090, 19 (2019)  
<https://doi.org/10.1155/2019/9613090>
2. <https://doi.org/10.1155/2019/9613090>
3. de Magalhães, C.S., Takarada, J.E., Carvalho, N.C., et al.: The coffee protective effect on catalase system in the preneoplastic induced rat liver. *J. Chem.* **2016**, Article ID 8570321, 9 (2016).  
<https://doi.org/10.1155/2016/8570321>
4. Kim, Y.-H., Chun, Y.-S., Park, J.-W., et al.: Involvement of adrenergic pathways in activation of catalase by myocardial ischemia-reperfusion. *Am. J. Physiol.Regulatory Integrative Comp. Physiol.* **282**(5), R1450-R1458 (2002)
5. Stevens, T.M., Boswell, G.A., Jr., Adler, R., Ackerman, N.R., Kerr, J.S.: Induction of antioxidant enzyme activities by a phenylurea derivative. *EDU. Toxicol Appl. Pharmacol.* **96**(1), 33–42 (1988)
6. Sposoby uvelicheniya proizvodstva ili aktivnosti katalazy [Method to increase the production or activity of catalase] Ru 2012 157 084 a
7. Graur, V.: Designul ,si sinteza compu ,silor biologic activi ai metalelor 3d cu 4- alilcalcogensemicaerbazone ,si deriva ,tii lor [Design and synthesis of biologically active compounds of 3d metals with 4-allylalkogensemicaerbazone and their derivatives]. Autoreferat al tezei de doctor în stiinte chimice [Reference of the doctoral thesis in chemical sciences]. Chişinău, 30 p. (2017)
8. Gulea, A., et al.: Utilizarea compus, ilor coordinativi ai sărurilor de cupru(II) cu 2-(2-hidroxibenziliden)-N-(prop-2-en-1-il)-hidrazincarbothioamida în calitate de inhibitori ai radicalilor superoxizi [The use of coordination compounds of copper (II) salts with 2-(2-hydroxybenzylidene)-N-(prop-2-en-1-yl)-hydrazine carbothioamide as inhibitors of superoxideradicals]. Brevet de invent,ie MD [Patent of invention MD] nr. 4749. Publ. BOPI nr.3/2021. P. 52–53



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9. Gulea, A.E., Zariciuc, E., Graur, V., Tsapkov, V., Rudic, V.: The study of antimicrobial and antitumor activity of biometal coordination compounds of substituted salicylaldehyde 4-allylthiosemicarbazones. In: International Scientific Conference on Microbial Biotechnology, 3 edn, Chișinău, Moldova, 12–13 October, Abstracts, pp. 67–68 (2016)
10. Korolyuk, M.A., et al.: Metod opredeleniya aktivnosti katalazy [Method for determining catalase activity]. In: Labor.Delo, vol. (1), pp. 16–19 (1988)
11. Góth, L.: A simple method for determination of serum catalase activity and revision of reference range. *Clinica Chimica Acta. Inter. J. Clin. Chem.* **196**(2–3), 143–151 (1991).  
[https://doi.org/10.1016/0009-8981\(91\)90067](https://doi.org/10.1016/0009-8981(91)90067) . PMID 2029780
12. Metode de cercetare a metabolismului hepatic. Elaborare metodică [Research methods of liver metabolism. Methodical elaboration]. Gudumac V., Rîvneac V., Tagadiuc O., Sardari V., Rîvneac E., Andronache L., Stîrba O., Pantea V., Popa V. Sub red. Valentin Gudumac; Univ. de Stat de Medicină și Farmacie „Nicolae Testemițanu”. – Ch.: S.n., 2012 (Tipogr., Tehnica-Info”). – 162 p. (2012)
13. Klappenbach, L.: Prokaryotes Vs. Eukaryotes: What Are the Differences? Thought Co (Apr 5 2023).  
thoughtco.com/what-are-prokaryotes-and-eukaryotes-129478
14. Qi, H., Shan, P., Wang, Y., Li, P., Wang, K., Yang, L.: Nanomedicines for the efficient treatment of intracellular bacteria: the “ART” principle. *Front. Chem.* **9**, 7 (2021)  
<https://doi.org/10.3389/fchem.2021.775682>
15. Kobayashi, M., Sugiyama, H., Wang, D.-H., et al.: Catalase deficiency renders remnant kidneys more susceptible to oxidant tissue injury and renal fibrosis in mice. In: *Kidney International*, vol. 68, pp. 1018–1031 (2005)
16. Park, Y.S., You, S.Y., Cho, S., et al.: Eccentric localization of catalase to protect chromosomes from oxidative damages during meiotic maturation in mouse oocytes. *Histochem. Cell Biol.* **146**, 281–288 (2016)
17. Ye, G., Metreveli, N.S., Donthi, R.V., et al.: Catalase protects cardiomyocyte function in models of type 1 and type 2 diabetes. *Diabetes* **53**(5), 1336–1343 (2004)
18. Giordano, C.R., Roberts, R., Krentz, K.A., et al.: Catalase therapy corrects oxidative stress-induced pathophysiology in incipient diabetic retinopathy. *Invest. Ophthalmol. Vis. Sci.* **56**(5), 3095–3102 (2015)
19. Dutta, R.K., Lee, J.N., Maharjan, Y., et al.: Catalase-deficient mice induce aging faster through lysosomal dysfunction. *Cell Commun Signal* **20**, 192 (2022). <https://doi.org/10.1186/s12964-022-00969-2>



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20. Glorieux, C., Calderon, P.B.: Catalase, a remarkable enzyme: targeting the oldest antioxidant enzyme to find a new cancer treatment approach. *Biol. Chem.* **398**(10), 1095–1108 (2017)
21. O’Sullivan, J.D.B., et al.: The biology of human hair greying. *Biol Rev Camb Philos Soc.* **96**(1), 107–128 (2021). <https://doi.org/10.1111/brv.12648>. Epub 2020 Sep 23 PMID:32965076
22. Wood, J.M., Decker, H., Hartmann, H., et al.: Senile hair graying: H<sub>2</sub>O<sub>2</sub>-mediated oxidative stress affects human hair color by blunting methionine sulfoxiderepair. *FASEB J.* **23**(7), 2065–2075 (2009) [arXiv:0706.4406](https://arxiv.org/abs/0706.4406). <https://doi.org/10.1096/fj.08-125435>. PMID 19237503. S2CID 16069417
23. Seiberg, M.: Age-induced hair greying - the multiple effects of oxidative stress. *Int, J, Cosmet, Sci.* **35**(6), 532–538 (2013). <https://doi.org/10.1111/ics.12090>. Epub 2013 Oct 10 PMID:24033376
24. Xue, F., Zhang, Y., Jiang, W., et al.: Regulation of copper transport crossing brain barrier systems by Cu-ATPases: effect of manganese exposure. *Toxicol. Sci.* **139**(2), 432–451 (2014)
25. Nazıroglu, M.: Molecular role of catalase on oxidative stress-induced Ca<sup>2+</sup> signaling and TRP cation channel activation in nervous system. *J. Recept. Signal Transduction* **32**(3), 134–141 (2012)
26. Wingo, T.S, Liu, Y., Gerasimov, E.S., et al.: Shared mechanisms across the major psychiatric and neurodegenerative diseases. *Nat Commun.* **13**(1), 4314 (2022). <https://doi.org/10.1038/s41467-022-31873-5>. PMID: 35882878; PMCID: PMC9325708
27. Li, F., et al.: An antioxidant enzyme therapeutic for sepsis. *Front, Bioeng, Biotechnol.* **23**(9), 800684 (2021). <https://doi.org/10.3389/fbioe.2021.800684>. PMID:34888304; PMCID: PMC8650590
28. Chen, S., Gao, Z., Hu, L., et al.: Association of Septic Shock with Mortality in Hospitalized COVID-19 Patients in Wuhan, China. *Advances in Virology*. Article ID 3178283, 9 pages (2022). <https://doi.org/10.1155/2022/3178283>