

## RESIN ACIDS AS RAW MATERIAL FOR THE PREPARATION OF CYCLODEXTRIN COMPLEXES LOADED WITH DEHYDROABIETITIC ACID AND CHROMENOL HYBRID

Marina Zveaghintseva <sup>a</sup>, Eugenia Stingaci <sup>a</sup>, Serghei Pogrebnoi <sup>a,b</sup>, Lucian Lupascu <sup>a</sup>, Alic Barba <sup>a</sup>, Gheorghe Duca <sup>a</sup>, Vladimir Valica <sup>b</sup>, Livia Uncu <sup>b</sup>, Victor Kravtsov <sup>c</sup>, Dumitru Terteac <sup>a,d</sup>, Alexandr Brinzan <sup>e</sup>, Fliur Macaev <sup>a,b\*</sup>

<sup>a</sup>Institute of Chemistry, 3, Academiei str., Chisinau MD-2028, Republic of Moldova

<sup>b</sup>“Nicolae Testemitanu” State University of Medicine and Pharmacy, 165, Stefan cel Mare blvd., Chisinau MD-2004, Republic of Moldova

<sup>c</sup>Institute of Applied Physics, 5, Academiei str., Chisinau MD-2028, Republic of Moldova

<sup>d</sup>Practical Scientific Institute of Horticulture and Food Technology, 59, Vierul str., Chisinau MD-2070, Republic of Moldova

<sup>e</sup>Institute of Biology Bucharest, Romanian Academy of Science, 296, Splaiul Independentei str., Bucuresti 060031, Romania

\* e-mail: [fliur.macaev@ichem.md](mailto:fliur.macaev@ichem.md)

**Abstract.** In this work new methods to obtain complexes from  $\beta$ -cyclodextrin and dehydroabietic acid with chromenol-triazol hybrid with the sizes limits of approximately 0.1-250  $\mu\text{m}$  are reported. Kneading, co-evaporation and co-precipitation for the resolution of racemic 2-tert-butyl-3-(1*H*-1,2,4-triazol-1-yl)-2*H*-chromen-2-ol for obtaining micro- and nanoparticles have been optimized. *In vitro* dissolution studies of the synthesized compounds in phosphate buffer (pH 6.8) showed an improved dissolution rate of chromenol-triazol hybrid in the inclusion complexes compared to the free form. It has been found that  $\beta$ -complexes of  $\beta$ -cyclodextrin loaded with dehydroabietic acid and chromenol hybrid show good antibacterial activity with MIC and MBC values ranging from 0.72 to 44.45  $\mu\text{M}$ . The evaluation results revealed that all compounds showed good antifungal activity with MIC values ranging from 0.02 to 0.4 mM and MFC from 0.07 to 0.52 mM better than the reference drugs ketoconazole (MIC and MFC values at 0.28-1.88 and 0.38 mM to 2.82 mM, respectively), bifonazole (MIC and MFC values at 0.32-0.64 and 0.64-0.81 mM) and nistatin (MIC and MFC values at 0.55-0.65 mM and 0.65-0.79 mM).

**Keywords:**  $\beta$ -cyclodextrin, dehydroabietic acid, 2-tert-butyl-3-(1*H*-1,2,4-triazol-1-yl)-2*H*-chromen-2-ol, chromenol-triazol hybrid, antimicrobial activity.

Received: 22 April 2022/ Revised final: 01 August 2022/ Accepted: 05 August 2022

---