

P3.6 - OREGANO ESSENTIAL OIL: AN EFFECTIVE AND NON-TOXIC APPROACH FOR PREVENT OR TREAT RESISTANT CANDIDA SPECIES

Liliana Fernandes ^{1*}, **Inês Silva** ¹, **Elena Blázquez** ², **Ainara Tejada** ^{2,3}, **Artur Ribeiro** ^{1,4}, **Sónia Silva** ^{1,4}, **Nuno Mira** ⁵, **Lorena Cussó** ^{3,6}, **Sofia Costa-de-Oliveira** ⁷, **Maria Elisa Rodrigues** ^{1,4}, **Mariana Henriques** ^{1,4}

¹ Centre of Biological Engineering, University of Minho, Braga, Portugal;

² Laboratorio de Imagen Médica. Unidad de Medicina y Cirugía Experimental. Instituto de Investigación Sanitaria Gregorio Marañón, Madrid, Spain;

³ Departamento de Bioingeniería. Universidad Carlos III de Madrid, Madrid, Spain

⁴ LABBELS – Associate Laboratory, Braga/ Guimarães, Portugal;

⁵ iBB, Institute for Bioengineering and Biosciences, Department of Bioengineering, Instituto Superior Técnico, Universidade de Lisboa, Lisbon, Portugal;

⁶ Unidad de Imagen Avanzada. Centro Nacional de Investigaciones Cardiovasculares Carlos III (CNIC), Madrid, Spain;

⁷ Division of Microbiology, Department of Pathology, Faculty of Medicine, University of Porto, Portugal; Center for Health Technology and Services Research – CINTESIS@RISE, Faculty of Medicine, University of Porto, Portugal

(*) e-mail: lilianafernandes@ceb.uminho.pt

Keywords: Vulvovaginal candidiasis, *Lactobacillus* spp; Phytotherapeutic applications; vapor-phase of essential oil; Keratin nanocapsules; Alternative treatment

ABSTRACT

Vulvovaginal candidiasis (VVC) is one of the most prevalent vaginal infectious diseases, and the emergence of drug-resistant *Candida* strains has presented a growing challenge in its treatment. This highlights the urgent need to develop effective and non-toxic alternative treatments. In this context, essential oils (EOs) have emerged as a promising alternative considering low toxicity and high antimicrobial activity.

This work is divided into two parts, the first consists of evaluating the effect of the vapor phase of oregano EO (VP-OEO) on biofilms of antifungal-resistant *Candida* species (*Candida albicans* and *Candida glabrata*) quantified by colony forming units' enumeration and determine their mode of action by flow cytometry. Interestingly, the VP-OEOs has shown to be more effective against *Candida* growth than their liquid form. Indeed, the results revealed high antifungal activity of VP-OEO against these drug-resistant strains, significantly reducing biofilm formation and mature biofilms, with impact on membrane integrity and metabolic activity of the fungal cells. The second part consists of the design and evaluation of nanoencapsulated OEO (KNP-OEO) as another alternative application of OEO for VVC treatment. These nanoparticles provided stability to OEO and controlled release of the EO. The results demonstrated complete inhibition of *C. albicans* growth. Moreover, in *in vivo* assay with BALB/C female mice, a single intravaginal application of KNP-OEO reduced *C. albicans* growth and preserved a healthy vaginal microbiota, including *Lactobacillus* species.

In conclusion, these studies highlight the promising efficacy of OEO as an alternative for VVC treatment. Both approaches, VP-OEO and OEO-KNP, showed effective antifungal activity against drug-resistant strains while preserving vaginal health. These therapeutic options not only combat antifungal resistance, but also potentially propose a safer option for women's health due to their

natural characteristics. However, further research is needed to confirm these promising results and advance the development of these alternative VVC therapies.

Acknowledgements:

This study was supported by the Portuguese Foundation for Science and Technology (FCT) under the scope of the strategic funding of UIDB/04469/2020 unit and grant ref 2020.05720.BD for Liliana Fernandes. Also, this study was supported by LABBELS—Associate Laboratory in Biotechnology, Bioengineering and Microelectromechanical Systems, LA/P/0029/2020 and Maria Elisa Rodrigues thanks FCT for funding through program DL 57/2016—Norma transitória. Sofia Costa de Oliveira acknowledges national funds through FCT, I.P., within the scope of the project "RISE - LA/P/0053/2020. Nuno Pereira Mira acknowledges support from FCT through its funding of research focused on Candida-lactobacillii interactions through LactoCan project (contract number: PTDC/BIA-MIC/31515/2017), iBB (contract: UIDB/04565/2020) and i4HB funding (contract: LA/P/0140/2020). Lorena Cussó acknowledges support by Agencia Estatal de Investigación (PRE2020-095268 MCIN / AEI /10.13039/501100011033 and by "ESF Investing in your future"), by Instituto de Salud Carlos III (PT20/00044) and co-funded by European Union (ERDF, "A way to make Europe").