

## Poster 4

# Targeting quorum sensing: ferulic and sinapic acids compromise *Pseudomonas aeruginosa* biofilm architecture and virulence

**Miguel M. Leitão**<sup>1,2,3</sup>, **Fernanda Borges**<sup>3</sup>, **Manuel Simões**<sup>1,2,4</sup> and **Anabela Borges**<sup>1,2,4,\*</sup>

<sup>1</sup> LEPABE—Laboratory for Process Engineering, Environment, Biotechnology and Energy, Faculty of Engineering, University of Porto, Rua Dr. Roberto Frias, s/n, 4200-465 Porto, Portugal

<sup>2</sup> ALICE—Associate Laboratory for Innovation in Chemical Engineering, Faculty of Engineering, University of Porto, Rua Dr. Roberto Frias, s/n, 4200-465 Porto, Portugal

<sup>3</sup> CIQUP-IMS—Department of Chemistry and Biochemistry, Faculty of Sciences, University of Porto, Rua do Campo Alegre, 4169-007 Porto, Portugal

<sup>4</sup> DEQB—Department of Chemical and Biological Engineering, Faculty of Engineering, University of Porto, Rua Dr. Roberto Frias, s/n, 4200-465 Porto, Portugal

\* Correspondence: apborges@fe.up.pt

## Abstract

**Background:** *Pseudomonas aeruginosa* is a Gram-negative pathogenic bacterium that is frequently associated with chronic infections in immunocompromised individuals, such as those with cystic fibrosis or burns [1]. The pathogenicity and virulence of *P. aeruginosa* are primarily regulated by quorum sensing, with the *las* system playing a key role in this process. This system is essential for the formation and maintenance of the biofilm and for controlling various virulence factors [2]. Phenolic acids, such as ferulic and sinapic acids, are plant secondary metabolites, well known for their biological properties and have shown promise in modulating bacterial communication [3]. **Objective:** The aim of this study was to evaluate the potential of ferulic and sinapic acids to inhibit the *P. aeruginosa las* QS system and its underlying effects on biofilm structure and virulence factor production. **Methods:** The inhibitory effect on the *las* system was evaluated using bioreporter strains and bioluminescence-based assays. Biofilm architecture was analyzed using optical coherence tomography, while virulence factors (pyoverdine, pyocyanin, total proteases, lipases, gelatinases and siderophores) production and motility were investigated by absorbance measurement and plate agar method. **Results:** Ferulic and sinapic acids inhibited *las* QS activity by 90 % at a concentration of 1000 µg mL<sup>-1</sup>. The N-3-oxododecanoyl-homoserine lactone production was reduced by 70 % at just 6.25 µg mL<sup>-1</sup> of the phenolic acids. These compounds significantly changed biofilm architecture, reducing biofilm thickness from 25 µm to 9 µm. They also reduced the production of key virulence factors and impaired swarming motility. **Conclusion:** Ferulic and sinapic acids demonstrated strong inhibitory effects on the *las* QS system, leading to altered biofilm structure and reduced virulence. These findings support their potential as antipathogenic and antivirulence agents for prevention/treatment of *P. aeruginosa* biofilm-associated infections.

**Keywords:** biofilm architecture; quorum-sensing; *Pseudomonas aeruginosa*

## Acknowledgments/Funding

This work was supported by: Project InnovAntiBiofilm (ref. 101157363) financed by European Commission (Horizon-Widera 2023- Access-02/Horizon-CSA); Project MultAntiBiofilm (ref. COMPETE2030-FEDER-00852000; N° 17121), and LEPABE, UIDB/00511/2020 (DOI: 10.54499/UIDB/00511/2020)) and UIDP/00511/2020 (DOI: 10.54499/UIDP/00511/2020); ALICE, LA/P/0045/2020 (DOI: 10.54499/LA/P/0045/2020); CIQUP, UIDB/00081/2020 (DOI:10.54499/UIDB/00081/2020); IMS, LA/P/0056/2020 (DOI:10.54499/LA/P/0056/2020); funded by national funds through the FCT/MCTES (PIDDAC; Lisbon, Portugal). Miguel M. Leitão acknowledges individual PhD fellowships from FCT (2021.07145.BD).

## References

1. Gonçalves, A.S. et al. The action of phytochemicals in biofilm control. *Natural Product Reports* **2023**, *40*, 595-627, doi: 10.1039/D2NP00053A
2. Leitão, M.M. et al. Dual action of benzaldehydes: Inhibiting quorum sensing and enhancing antibiotic efficacy for controlling *Pseudomonas aeruginosa* biofilms. *Microbial Pathogenesis* **2024**, *191*, 106663, doi: 10.1016/j.micpath.2024.106663

3. Borges, A. et al. Antibacterial activity and mode of action of ferulic and gallic acids against pathogenic bacteria. *Microbial Drug Resistance* **2013**, *19*(4), 256-265, doi: 10.1089/mdr.2012.0244.



In *Scientific Letters*, works are published under a CC-BY license (Creative Commons Attribution 4.0 International License at <https://creativecommons.org/licenses/by/4.0/>), the most open license available. The users can share (copy and redistribute the material in any medium or format) and adapt (remix, transform, and build upon the material for any purpose, even commercially), as long as they give appropriate credit, provide a link to the license, and indicate if changes were made (read the full text of the license terms and conditions of use at <https://creativecommons.org/licenses/by/4.0/legalcode>).