Innovative Solutions to Combat Biofilms and Antimicrobial Resistance: Advancing with Marine Bioactives and Microbiome Research

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● FACULTY OF ● PHARMACEUTICAL SCIENCES





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AGENTSCHAP









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What I will talk about:

- Brief introduction to antimicrobial resistance and clinical biofilms
- Why are biofilm-related infections so difficult to treat?
- Alternative approaches to treatment using marine natural products

Microorganisms have many ways of dealing with antibiotics leading to antimicrobial resistance and therapy failure



Darby et al, 2023, Nature Rev Microbiol

The global burden of antimicrobial resistance - 2019



Murray et al., 2022, Lancet

The burden of antimicrobial resistance in BE and ROK - 2019



Main organisms: Staphylococcus aureus, Escherichia coli, Klebsiella pneumoniae, Streptococcus pneumoniae, Enterocccus faecium, Pseudomonas aeruginosa







2023 Antibacterial agents in clinical and preclinical development

an overview and analysis



Overall, antibacterial agents in the clinical pipeline combined with those approved in the last six years are still insufficient to tackle the ever growing threat of the emergence and spread of drug-resistant infections.

Problem is made worse by microbial biofilm formation What are microbial biofilms?



Sauer et al, 2022, Nature Rev Microbiol

Biofilms occur as <u>attached communities on surfaces</u> and as suspended aggregates







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Biofilm

Microbial diversity and antimicrobial susceptibility in endotracheal tube biofilms recovered from mechanically ventilated COVID-19 patients

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Biofilm-related tolerance also contributes to therapy failure



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- Antimicrobial resistance is a global burden leading to excess mortality worldwide
- The problem I made worse by biofilm-associated antimicrobial tolerance
- There is an urgent need for new antimicrobial agents/antibiotics
- Where do we find these?

The ProBio project (2020-2022)













Overview of workflow

- Sampling
- Focus on approx. 50 species







Overview of workflow

- Sample preparation
- Fractionation (96 tubes/organism)





Dr. Geert Goeminne Dr. Keylla Bicalho

Overview of workflow

- Bioactivity assays
- Testing of growth inhibitory activity against
 - Staphylococcus aureus
 Pseudomonas aeruginosa
 Acinetobacter baumannii
 Candida albicans





• If active against planktonic organisms: anti-biofilm assays

Some fractions show broad antimicrobial activity Example: F2 from *Echiichthys vipera*









Kleine pieterman Lesser weever

Some fractions show strong biofilm <u>inhibition</u> Example: F4 and F5 from *Alcyonium digitatum*



Dead man's fingers

Some fractions show strong biofilm <u>eradication</u> Example: F55, F56 and F83 from *Ophiura ophiura*



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- Various marine organisms from the Belgian part of the North Sea contain compounds with antimicrobial *and* antibiofilm activity against clinically relevant pathogens
- What are these compounds?
- What is the role of the microbiome in the production of these compounds?
- Expanding this approach to marine species found in other parts of the world may lead to discovery of interesting lead antimicrobial compounds

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