

## Research Week 2023

## Commensal glycerol metabolism determines oral niche development.

June Treerat<sup>1</sup>, David Anderson<sup>1</sup>, Rodrigo A. Giacaman<sup>3</sup>, Justin Merritt<sup>1,2</sup>, and Jens Kreth<sup>1,2</sup>

<sup>1</sup>Department of Restorative Dentistry, School of Dentistry, Oregon Health & Science University (OHSU), Portland, OR 97239, USA. <sup>2</sup>Department of Molecular Microbiology and Immunology, School of Medicine, Oregon Health & Science University (OHSU), Portland, OR 97239, USA. <sup>3</sup>Cariology Unit, Department of Oral Rehabilitation, Faculty of Dentistry, University of Talca, Talca, Chile.

## Keywords

Oral microbiome, glycerol kinase, glycerol dehydrogenase, membrane vesicles, fatty acids, symbiotic interaction

## Abstract

The oral microbiome has coevolved and coexisted in a symbiotic relationship within their own ecology and with the host. Understanding the oral ecosystem and how it promotes a balanced microbiome, is essential to effectively maintain or restore oral health. Of over 700 bacterial taxa, commensals Streptococcus and Corynebacterium are two of the most prevalent members of the oral microbiome. Surprisingly, the inter-species interactions between both commensals and the effect of such interactions on dental diseases have never been investigated. Therefore, in this study, we aim to understand the molecular mechanism(s) between different taxa interactions, in particularly, Corynebacterium durum and Streptococcus sanguinis, and the implications on oral diseases. So far, we have discovered a specifically mutualistic interaction that C. durum was able to substantially influence the chain morphology of S. sanguinis. When Gas Chromatography/Mass Spectrometry (GC/MS) analysis was employed, 3 fatty acids (palmitic, oleic, and stearic acids) were discovered in association with C. durum membrane vesicles, resulting in enhancing S. sanguinis fitness and survival. Global gene expression profiling of dual species cultures using RNAseq was further conducted in order to investigate the regulatory effect of lipid metabolism on S. sanguinis chain morphology. Of approximately 30 genes as being differentially expressed under the influence of *C. durum*, genes involved *S. sanguinis* lipid metabolism, in particular, glycerol dehydrogenase (gldA), its "sister" glycerol kinase (glpK), and the neighboring downstream genes were found to be significantly upregulated. Subsequently, S. sanguinis fatty acid metabolism-related genes were further characterized and revealed a key role of S. sanguinis gldA and a supported role of glpK during the mutualistic-interspecies interaction. Overall, our findings have provided new insights into how glycerol metabolism is involved in oral niche development, potentially shaping symbiotic health-associated biofilm communities.

Acknowledgements: J.K. acknowledges support through NIH-NIDCR grant DE029492 & DE029612, and J.M. is supported by NIH-NIDCR grant DE022083 and DE018893.