

co-ordinated with the Director of the Institute / Research Unit

Analytical BioGeoChemistry

PSP-Element:

G-504800-001

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Title of the highlight:

DI-ICR-FT/MS-based high throughput deep metabotyping: A case study of the *Caenorhabditis. elegans* – *Pseudomonas aeruginosa* infection model.

Keywords:

ICR-FT-MS; High-throughput mass spectrometry; deep metabotyping; Infection models; Metabolomics; *Caenorhabditis elegans*

Central statement of the highlight in one sentence:

This study fulfilled two goals. The first was a detailed metabolomics study of *Caenorhabditis. elegans* – *Pseudomonas aeruginosa* infection model to understand interorganismic interactions on a molecular level. The second was a proof of principle that the technology of ICR-FT/MS is throughput compatible and delivered hundreds of organism specific compounds in limited time.

Text of the highlight:

In metabolomics there is an ever-growing need for faster and more comprehensive analysis methods to cope with the increasing size of biological studies. Direct-infusion ion cyclotron-resonance Fourier-transform spectrometry (DIICR-FT-MS) is used in non-targeted metabolomics to obtain high-resolution snapshots of the metabolic state of a system. We applied this technology to a *Caenorhabditis elegans*–*Pseudomonas aeruginosa* infection model and optimized times needed for cultivation and mass-spectrometric analysis. Our results reveal that DI-ICR-FT-MS is a promising tool for high-throughput in-depth non-targeted metabolomics. We performed whole-worm metabolomics and recovered markers of the induced metabolic changes in *C. elegans* brought about by interaction with pathogens. In this investigation, we reveal complex metabolic phenotypes enabling clustering based upon challenge. Specifically, we observed a marked decrease in amino-acid metabolism with infection by *P. aeruginosa* and a marked

increase in sugar metabolism with infection by *Salmonella enterica*. We were also able to discriminate between infection with a virulent wild-type *Pseudomonas* and with an attenuated mutant, making it possible to use this method in larger genetic screens to identify host and pathogen effectors affecting the metabolic phenotype of infection.

Publication:

Witting, M., M. Lucio, D. Tziotis, B. Wägele, K. Suhre, R. Voulhoux, S. Garvis, **Ph. Schmitt-Kopplin**, DI-ICR-FT/MS-based high throughput deep metabotyping: A case study of the *Caenorhabditis. elegans* – *Pseudomonas aeruginosa* infection model. *Analytical Bioanalytical Chemistry*, 2015, forfront paper, 407, 4, 1059-1073

Taking account of the HMGU mission:

The present study shows the state of the art in high resolution non targeted Metabolomics analysis of interorganism interactions with the example of a pathogen of high relevance in infectious diseases.

The internal HMGU co-operation partners with whom the highlight was compiled, if appropriate:

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