## **Microbiome Metabolites:**

### **Understanding Microbial Function Within the Host**







6<sup>TH</sup> MICROBIOME R&D & BUSINESS COLLABORATION FORUM

**3<sup>RD</sup> PROBIOTICS** CONGRESS: USA





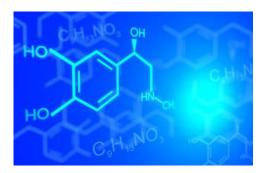
### Microbiome Metabolites: Moving from Correlation to Causality to Innovation



Who is There?



What are they Doing?



How are they Doing it?





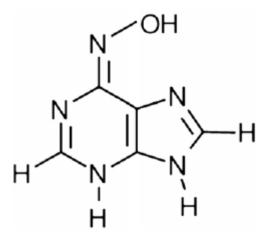


#### SCIENCE ADVANCES | RESEARCH ARTICLE

#### HEALTH AND MEDICINE

# A commensal strain of *Staphylococcus epidermidis* protects against skin neoplasia

Teruaki Nakatsuji,<sup>1</sup> Tiffany H. Chen,<sup>1</sup> Anna M. Butcher,<sup>1</sup> Lynnie L. Trzoss,<sup>2</sup> Sang-Jip Nam,<sup>2</sup>\* Karina T. Shirakawa,<sup>1</sup> Wei Zhou,<sup>3</sup> Julia Oh,<sup>3</sup> Michael Otto,<sup>4</sup> William Fenical,<sup>2</sup> Richard L. Gallo<sup>1†</sup>





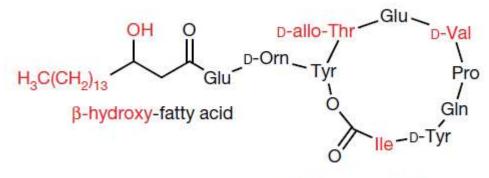


# ARTICLE

https://doi.org/10.1038/s41586-018-0616-y

## Pathogen elimination by probiotic Bacillus via signalling interference

Pipat Piewngam<sup>1,2</sup>, Yue Zheng<sup>1,5</sup>, Thuan H. Nguyen<sup>1,5</sup>, Seth W. Dickey<sup>1</sup>, Hwang-Soo Joo<sup>1,4</sup>, Amer E. Villaruz<sup>1</sup>, Kyle A. Glose<sup>1</sup>, Emilie L. Fisher<sup>1</sup>, Rachelle L. Hunt<sup>1</sup>, Barry Li<sup>1</sup>, Janice Chiou<sup>1</sup>, Sujiraphong Pharkjaksu<sup>2</sup>, Sunisa Khongthong<sup>3</sup>, Gordon Y. C. Cheung<sup>1</sup>, Pattarachai Kiratisin<sup>2</sup> & Michael Otto<sup>1\*</sup>



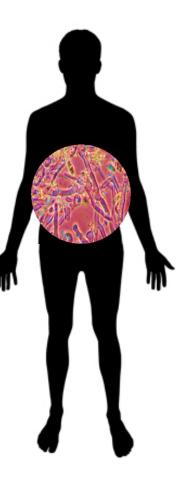
8-Amino-acid lactone ring



### Microbiome Metabolites by the Numbers

# 3,000

The approximated number of biosynthetic gene clusters in the human microbiota identified from the HMP data set predicted to code for small molecules, most of unknown structure and function<sup>2</sup>



2%

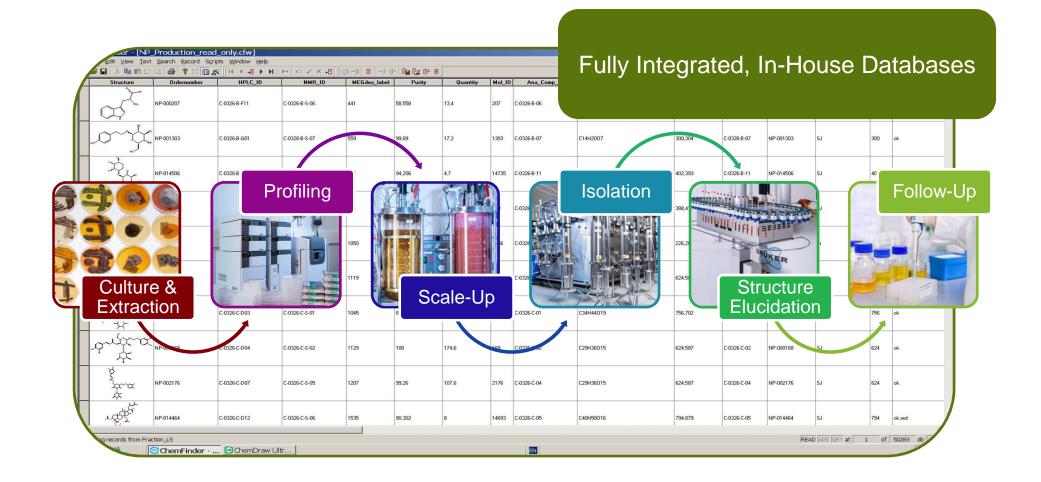
The estimated percentage of the microbiome metabolomics data from mass spectrometry that can currently be annotated<sup>1</sup>

<sup>1</sup> Brown and Hazen. Targeting of Microbe-Derived Metabolites to Improve Human Health: The Next Frontier for Drug Discovery. *J. Biol. Chem.* (2017) <sup>2</sup> Donia, M. S. *et al.* A Systematic Analysis of Biosynthetic Gene Clusters in the Human Microbiome Reveals a Common Family of Antibiotics. *Cell*. (2014).



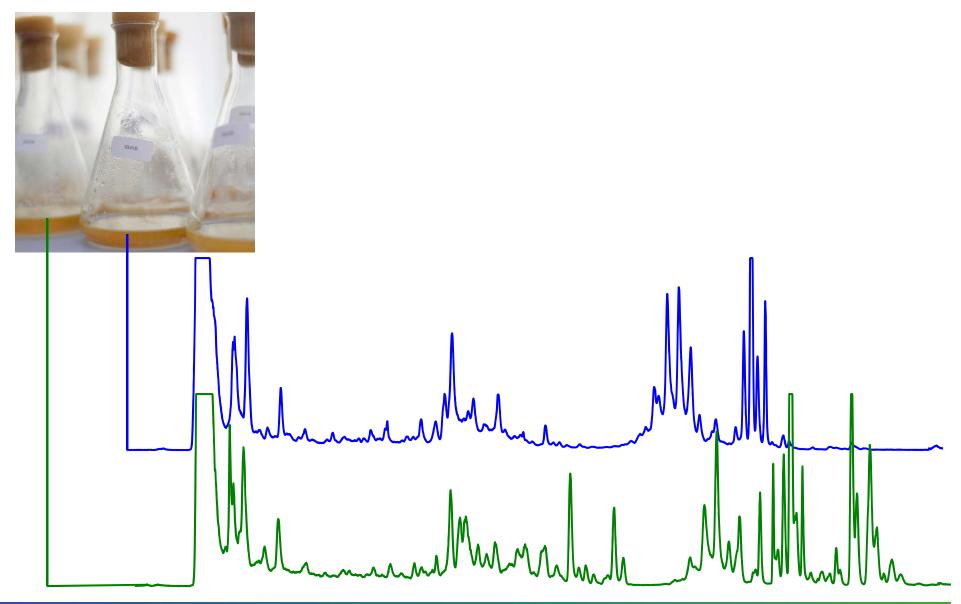


### A Natural Product Approach to Microbiome Metabolites





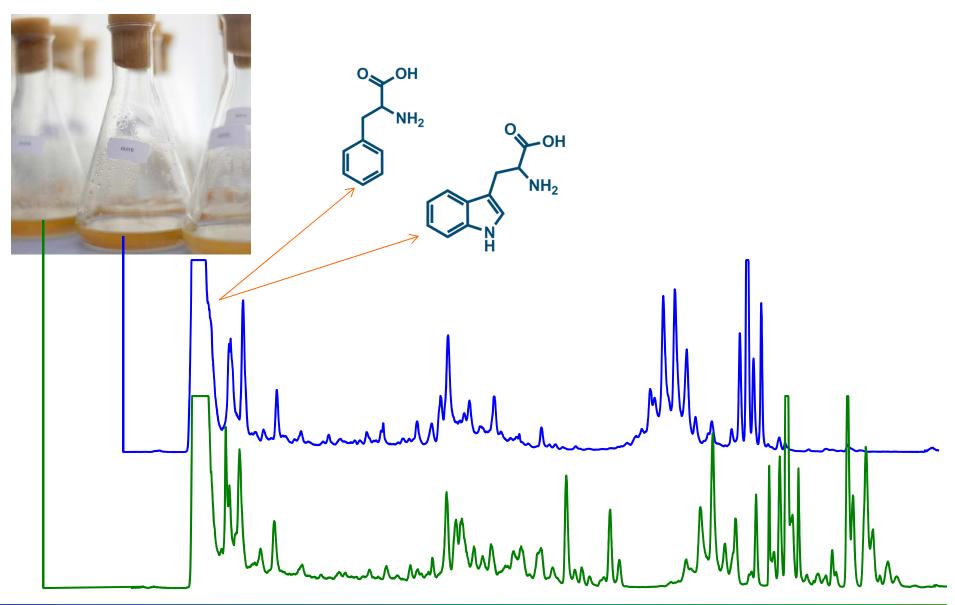
### Case Study: B. subtilis; Culture, Extraction, & Profiling







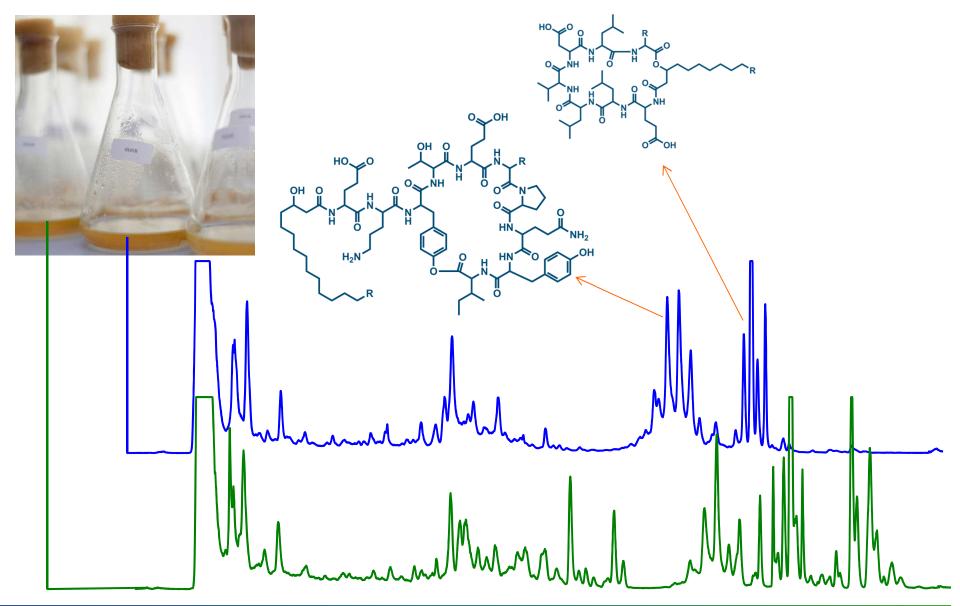
### Case Study: B. subtilis; structure elucidation







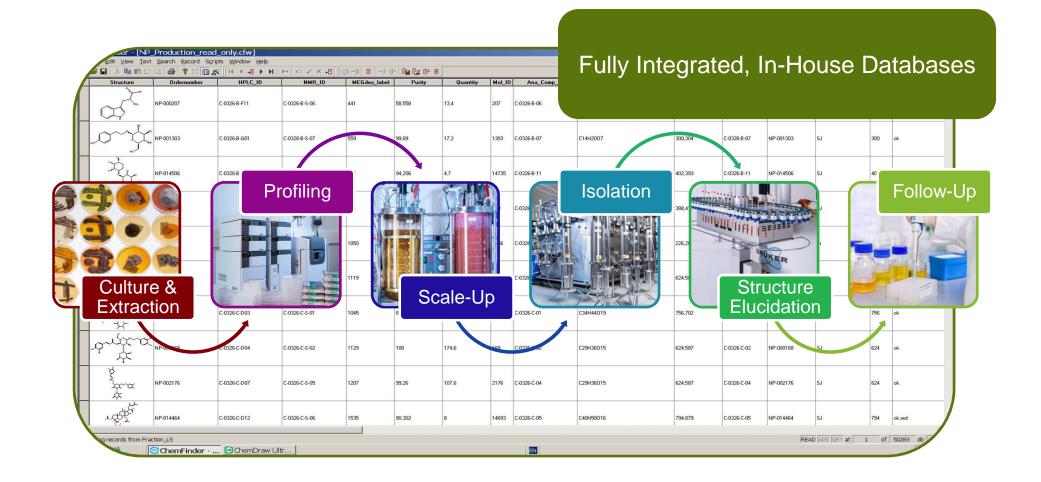
### Case Study: B. subtilis; structure elucidation





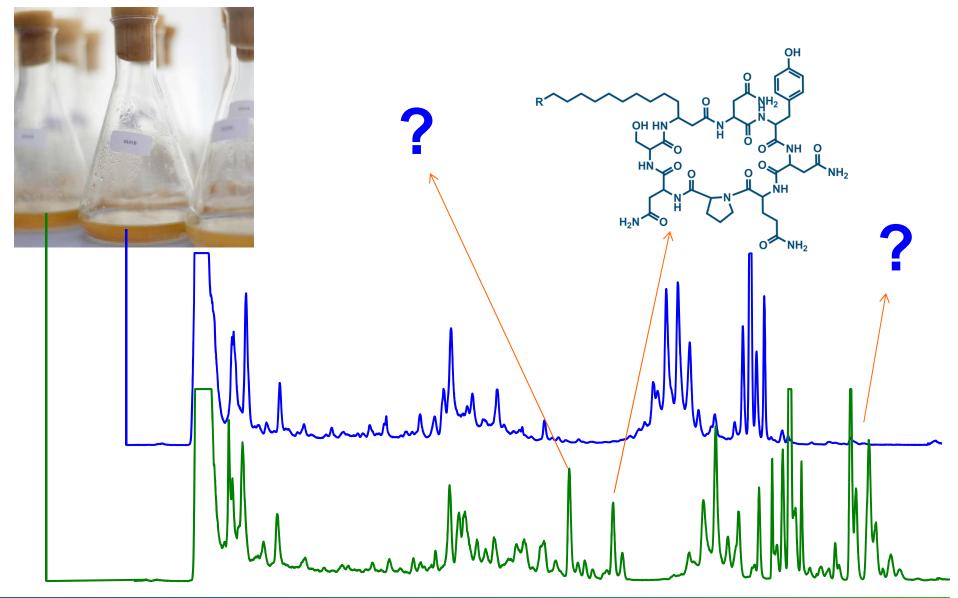


### A Natural Product Approach to Microbiome Metabolites





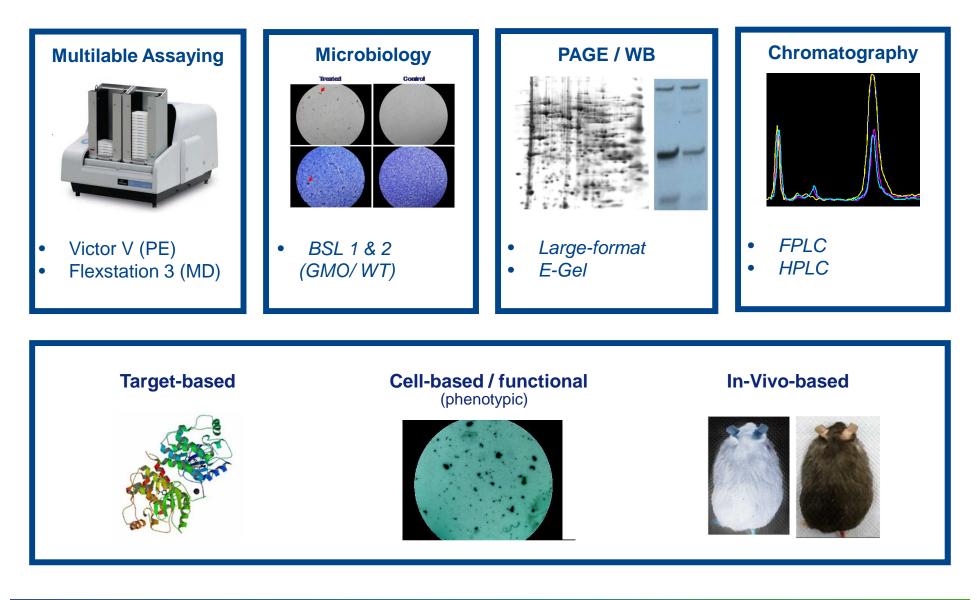
### Case Study: B. subtilis; structure elucidation







### Case Study: B. subtilis; Follow-Up







### Metabolite Solutions for Microbiome Drug Discovery



The Natural Product Company

- Aerobic & Anaerobic Fermentation
- Small- to Large-Scale Fermentation
- Culture & Extraction Optimization
- Extract Profiling & Dereplication
- Preparative Chromatography
- Metabolite Purification & Identification
- Chemical & Biological Assays
- Synthetic & Medicinal Chemistry



9601 Medical Center Drive Rockville, MD 20850, USA +1-240-406-1256 www.ac-discovery.com