

USCCN Workshop, 20 August 2023

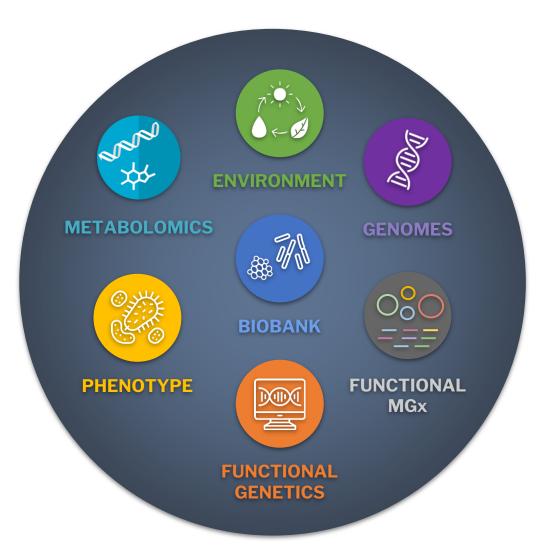
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AGBIOME

- Introduction to the AgBiome Culture Collection
- What we find **Valuable** about the collection
- A few **Examples** of discoveries from our collection
- Public/Private Collaboration opportunities and challenges

AgBiome GENESIS[™] Platform





- Focused on Broad Environmental Samples
- Biobank of 100,000+ Individual Isolates
- Fully Sequenced Genomes for each isolate
- Archived Environmental Samples
- > Metagenomics for unculturable microbes
- **Functional Genomics** on half a billion genes
- > **Phenotypic Screens** with real-world translation
- > Diverse Chemistry via **Biosynthetic Gene Clusters**
- Complete Meta-Data for every isolate

The AgBiome Culture Collection

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Number of strains

per species

<10

10-99

>999

100-999

Phylum

Acidobacteriota Actinobacteriota Bacteroidota Cyanobacteria Deinococcota Firmicutes Firmicutes_A Firmicutes_C Firmicutes_I Proteobacteria -

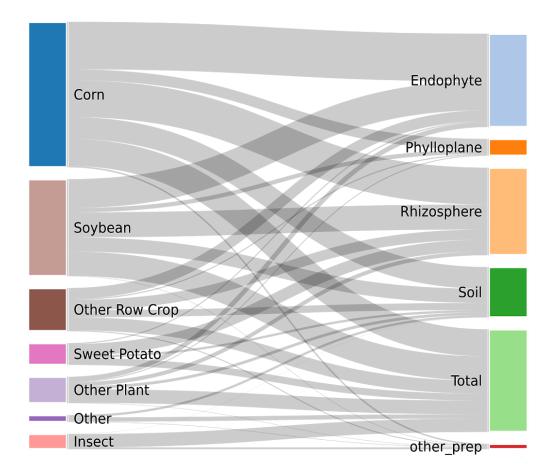
8,000+ unique environmental samples from North America and Africa

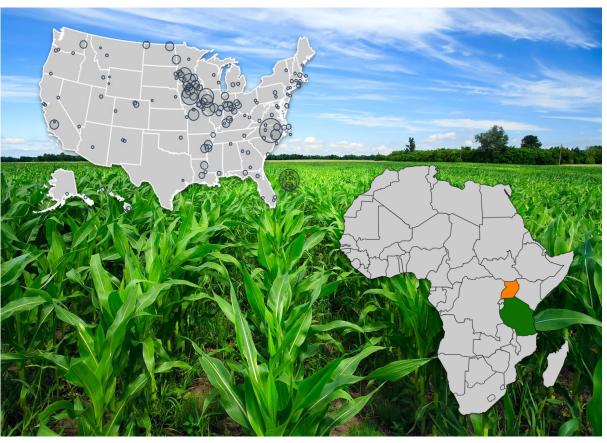
Highly diverse collection of **100,000+** fully sequenced microbes comprising **1,460** species and **431** genera

High quality genomic sequence of each microbe representing **500,000,000+** genes and **1,000,000+** BGCs

Broad access to Functional Diversity Specific Functions per Taxa Ability to find the Best of the Best

To build our collection, we emphasize **Broad Environmental Sampling**

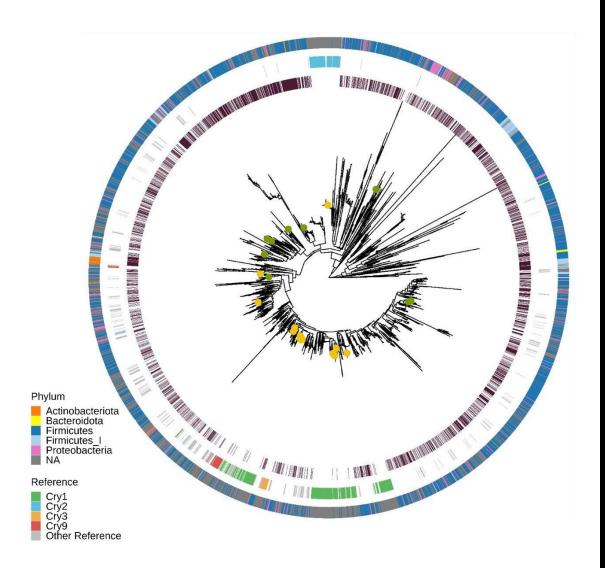




Primarily from Agricultural Fields and High-Diversity Environments Over 8,000 Unique Environmental Samples Archived

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Complete Genomes deliver **Gene Diversity**



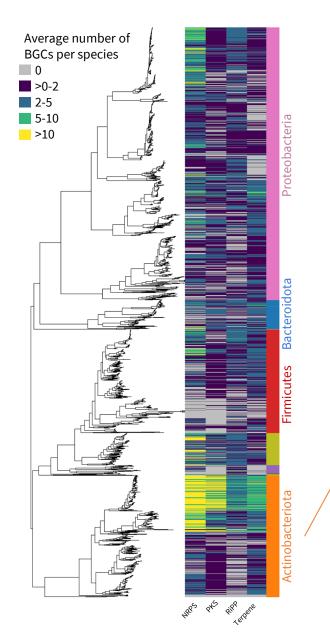
Extensive diversity of putative insecticidal proteins

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→250,000+ Unique Genes

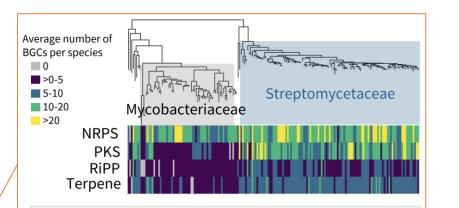
Microbial Universe delivers **Diverse Chemistry**



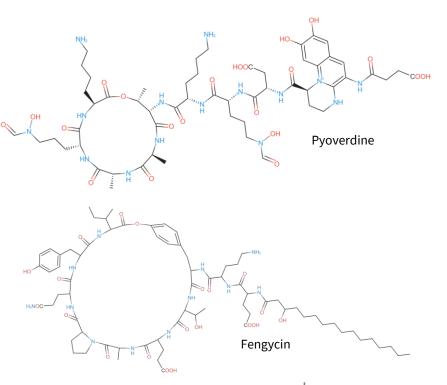


Over 1 million Biosynthetic Gene Clusters

- Extensive Potential for Chemical Diversity
- Novel Natural Product Discovery



Partial representation of Actinobacteriota with two subclades. Streptomycetaceae is gifted with all BGC classes, especially PKS. Mycobacteriaceae are similarly gifted with NRPS, PKS, and Terpenes, but have fewer RiPPs.

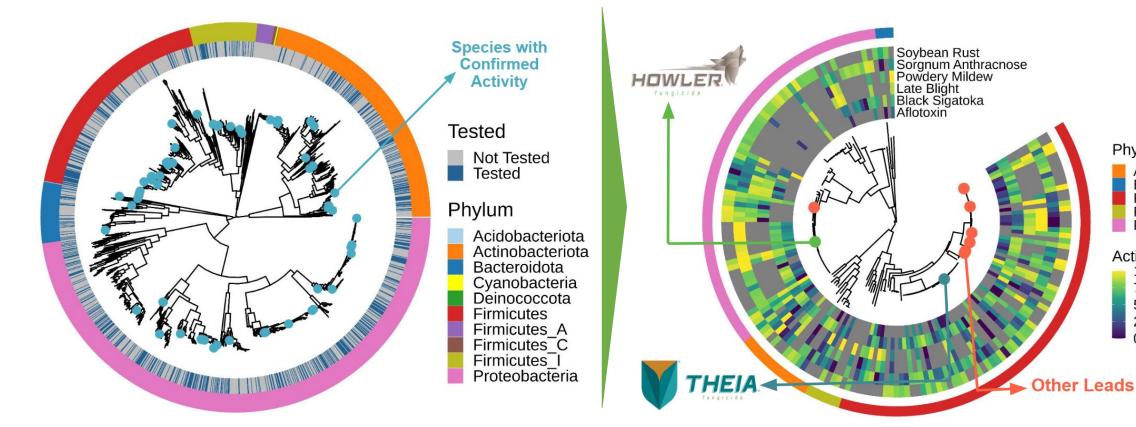






Discovery for Broad Spectrum Disease Control

Diverse Isolates screened across 10 different phyla



Confirmed Activity across 5 Phyla

Diversity of Actives = Diversity of Modes of Action

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Phylum

Activity

100

75

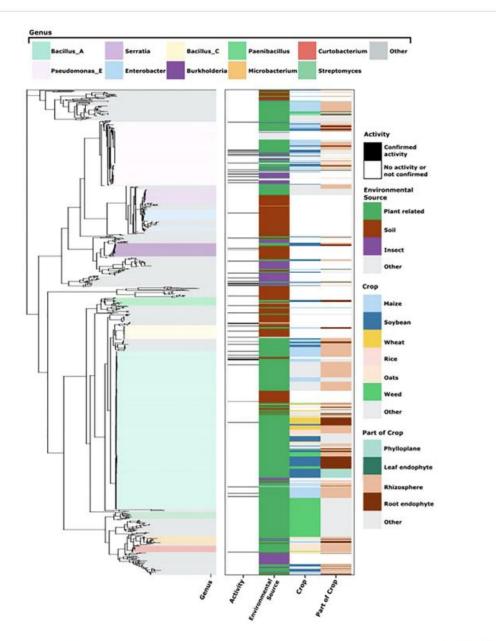
50

25

Actinobacteriota

Bacteroidota Firmicutes

Firmicutes I Proteobacteria



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Diverse environmental bacteria displaying activity against *Phakopsora pachyrhizi*, the cause of soybean rust

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Antifungal activity can be found in any environment and across a broad diversity of taxa

FIGURE 2

Metadata associated with bacterial isolates evaluated in the initial screen. On the left is the SSU phylogenetic tree with the most abundant genera highlighted as classified by the Genome Taxonomy Database (GTDB) (see key at the top). On the right we indicate activity against soybean rust (at least 75% reproducible rust reduction), environmental source of the isolate, and for plant-associated isolates, the crop, and the part of the crop/plant from which bacteria were isolated (see key to the right).



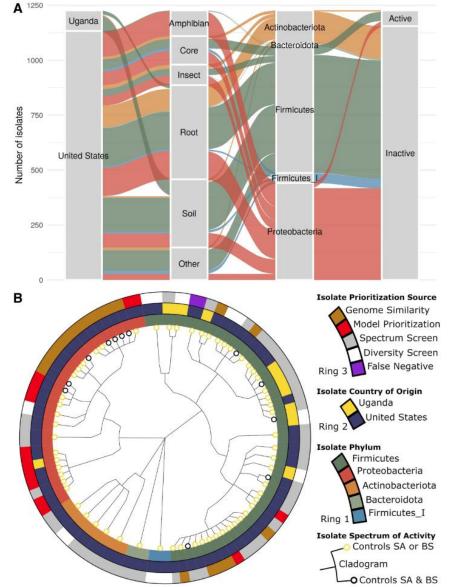


Fig. 2. Scope of screening effort, and diverse isolates with confirmed fungicidal activity. **A**, Alluvial diagram illustrates the source country, sample type (the less abundant types were combined into "Other"), phylum membership, and fungicidal activity of the 1,227 isolates screened in this work. Band colors indicate phylum membership (see legend in panel B). **B**, Cladogram in the center highlights the diversity of the biocontrol isolates discovered in this work (five phyla represented by the center branches, 14 genera, and 22 species). SA = sorghum anthracnose and BS = black sigatoka. The cladogram was constructed using whole-genome-based taxonomic IDs (see Materials and Methods). Multiple isolates displayed activity against both diseases (black leaf nodes). The outer ring shows the isolate prioritization strategies that led to each discovery, including predictive models. The "false-negative" isolate in purple was predicted to be inactive by all modeling approaches and yet was found to be reproducibly active against sorghum anthracnose.

Genomics- and Machine Learning-Accelerated Discovery of Biocontrol Bacteria

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Machine Learning Algorithms developed to accelerate our Discovery:

- ► Identify additional **more effective** isolates
- Counter-select to enrich for additional modes of action

Summary of AgBiome Collection





Breadth and Depth across collection

- 100,000+ Fully Sequenced Isolates
- Complete Metadata for each Isolate
- Archived Environmental Samples
- Documented Ownership
- **Potential value** for many industries

Public/Private Opportunities and Challenges

Maintain Intellectual Property rights Metagenomics "collections" to expand into unculturable space Systematic and consistent connection of all metadata for isolates

