

Genomics & Microbiome Analysis

to identify the next generation biopesticides

Marcus Meadows-Smith

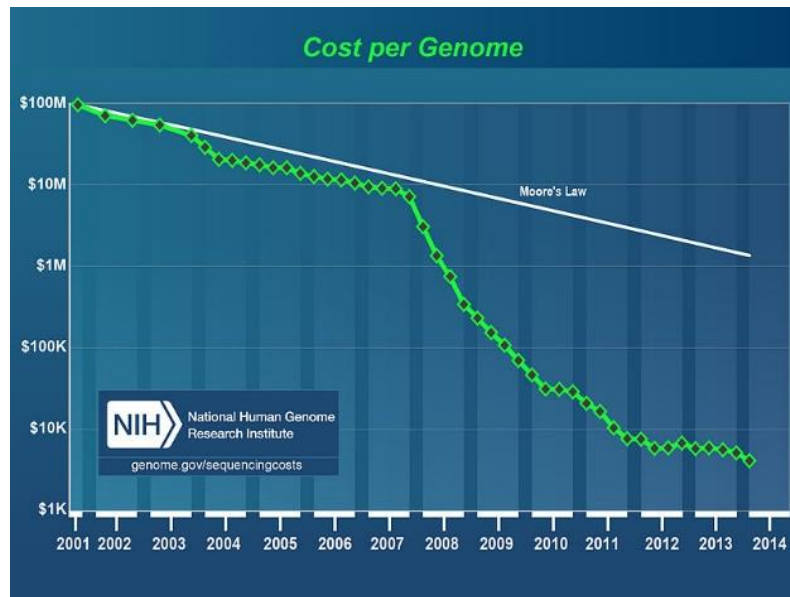
CEO

BioConsortia, Inc.



DNA Driven Discovery

Genome sequencing costs have dropped dramatically in the past decade allowing for consistent use in research programs



First human genome sequenced in 2000₁

\$100 million
9 months

Human Longevity Inc. 2016₁

< \$2,000
15 minutes

Arabidopsis genome sequenced in 2000₂

\$70 million
7 years
500 people

Arabidopsis genome sequencing in 2017₁

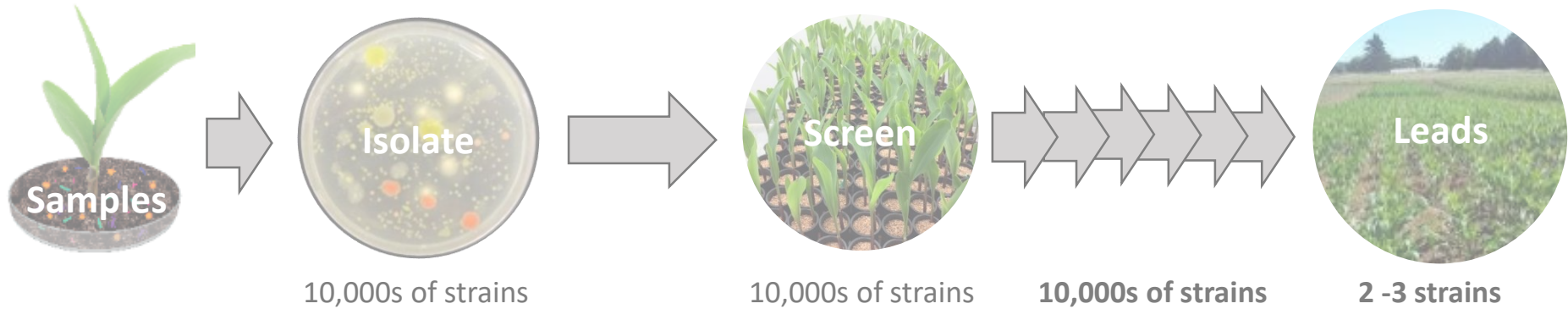
\$99
2 minutes

1. Technology Review: <https://www.technologyreview.com/s/601842/inside-genomics-pioneer-craig-venters-latest-production/>

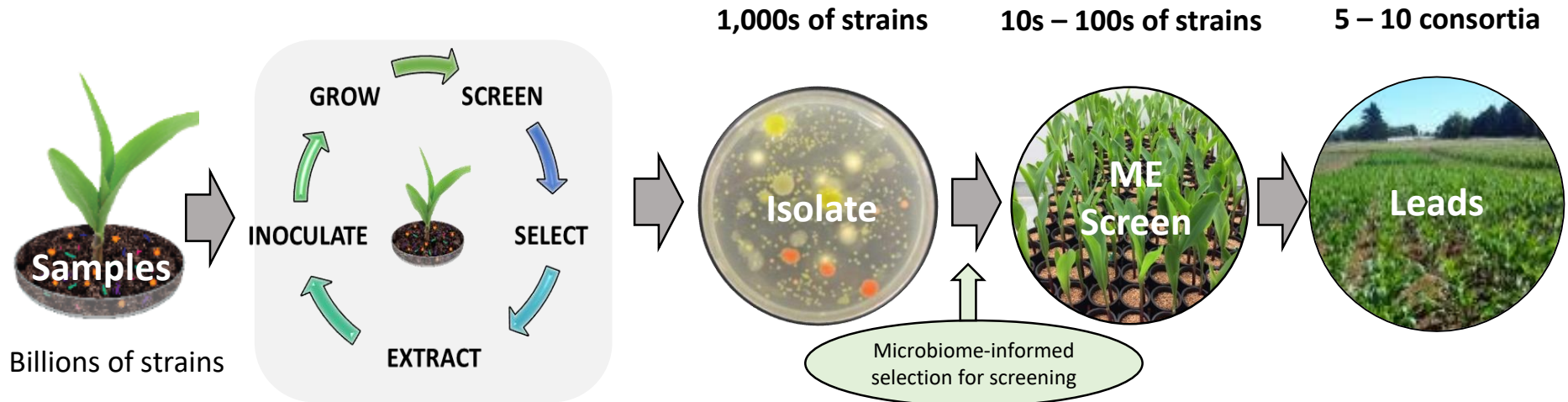
2. Source: Jeff Ross-Ibarra from presentation by Dr. Pamela Roland at University of California Davis
National Human Genome Research Institute

AMS - Superior Discovery Process

Conventional Microbial R&D Process (often years to lead discovery)



BioConsortia's AMS process

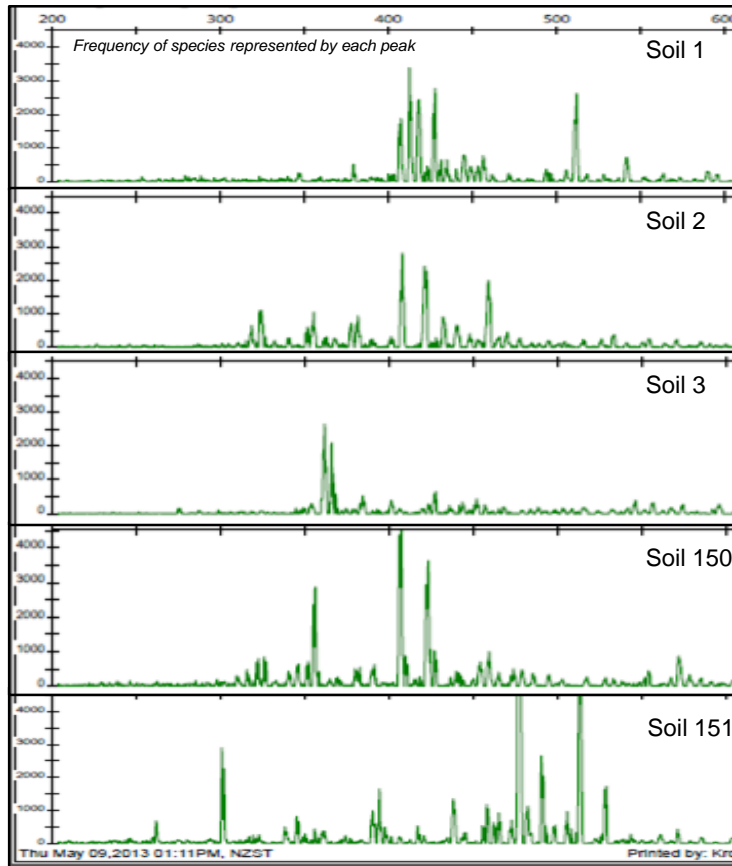


Directed selection of the microbiome, under biotic or abiotic stress, identifying teams of beneficial microbes that improve plant phenotype

Microbiome Analysis

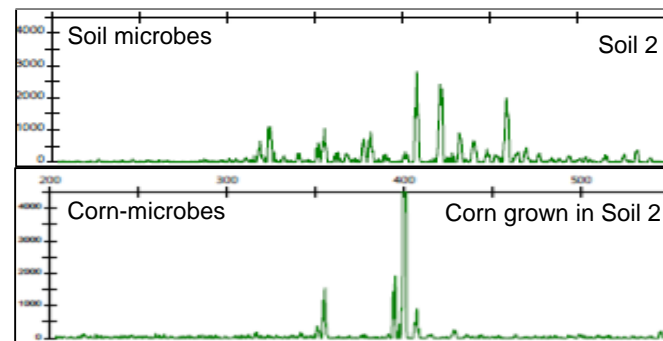
Soil Microbiome Influences Crop Yield

Each soil has a different microbial community



Community Fingerprinting: Automated Ribosomal Intergenic Spacer Analysis (ARISA)

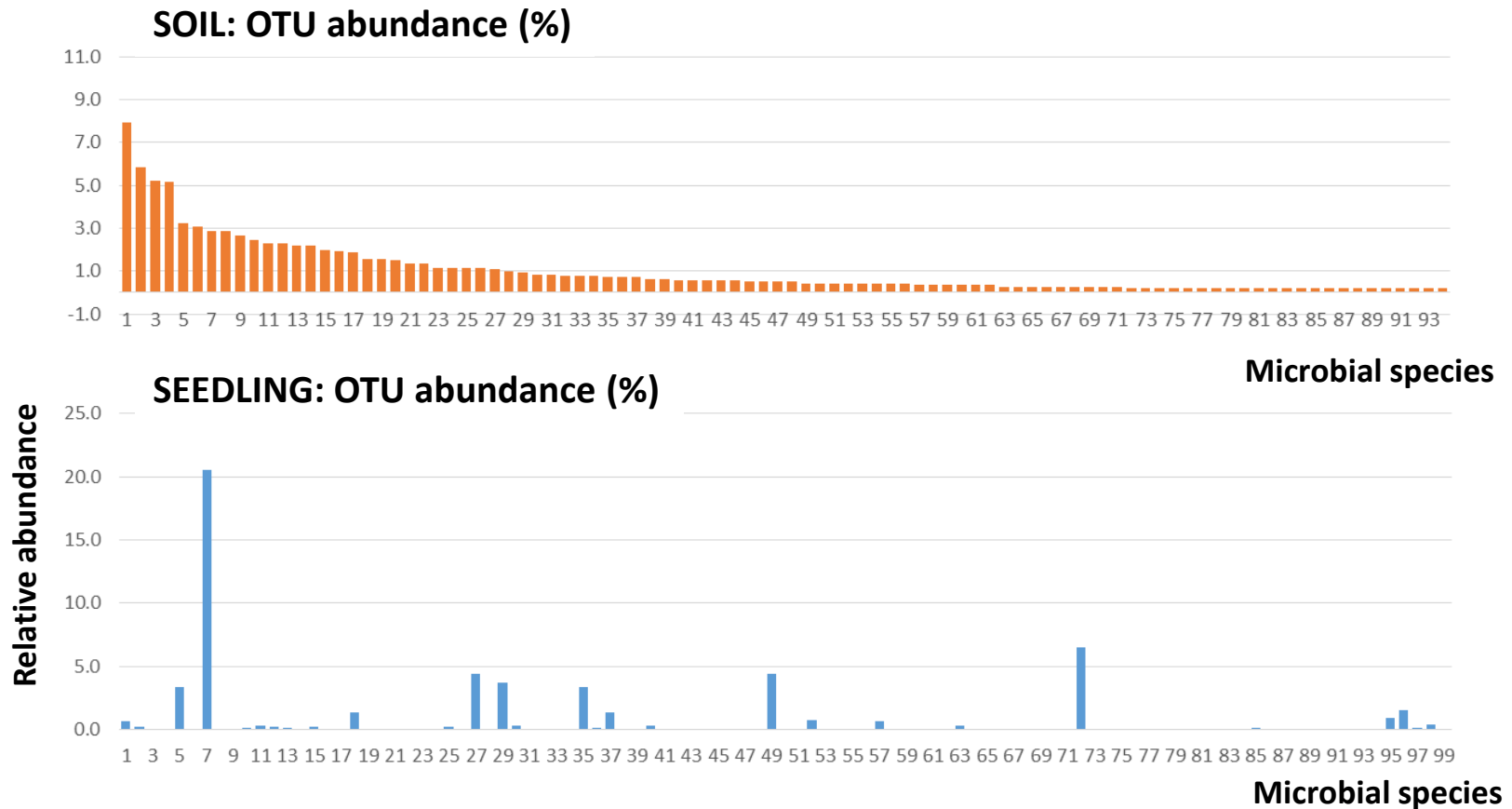
Plants accumulate a different microbial community than the soil in which they are grown



We exploit this natural process identifying the microbial consortia that improve plant traits

Soil and Plant Microbiomes Differ

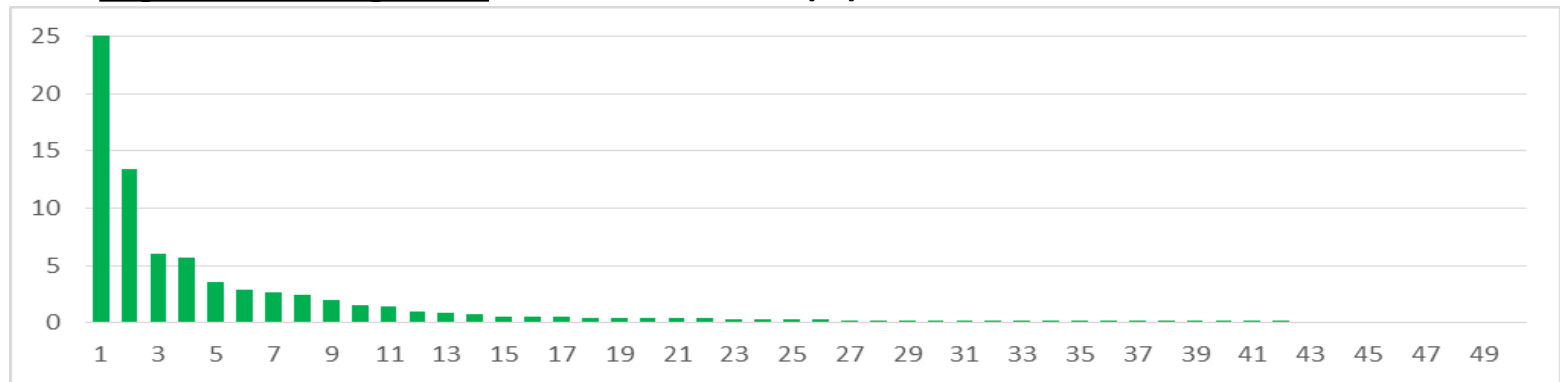
Seedlings accumulate a different microbial community structure than that present in the soil



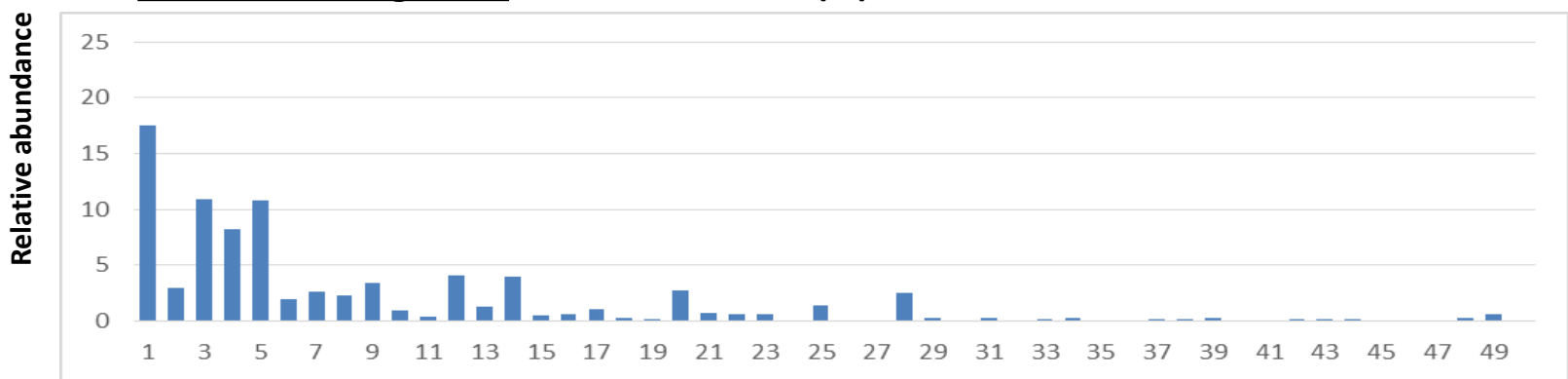
High & Low Performing Plants - Microbiomes Differ


High & low performing plants have a different microbial community
AMS - differences are tracked over successive generations

High Performing Plant: OTU abundance (%)



Low Performing Plant: OTU abundance (%)

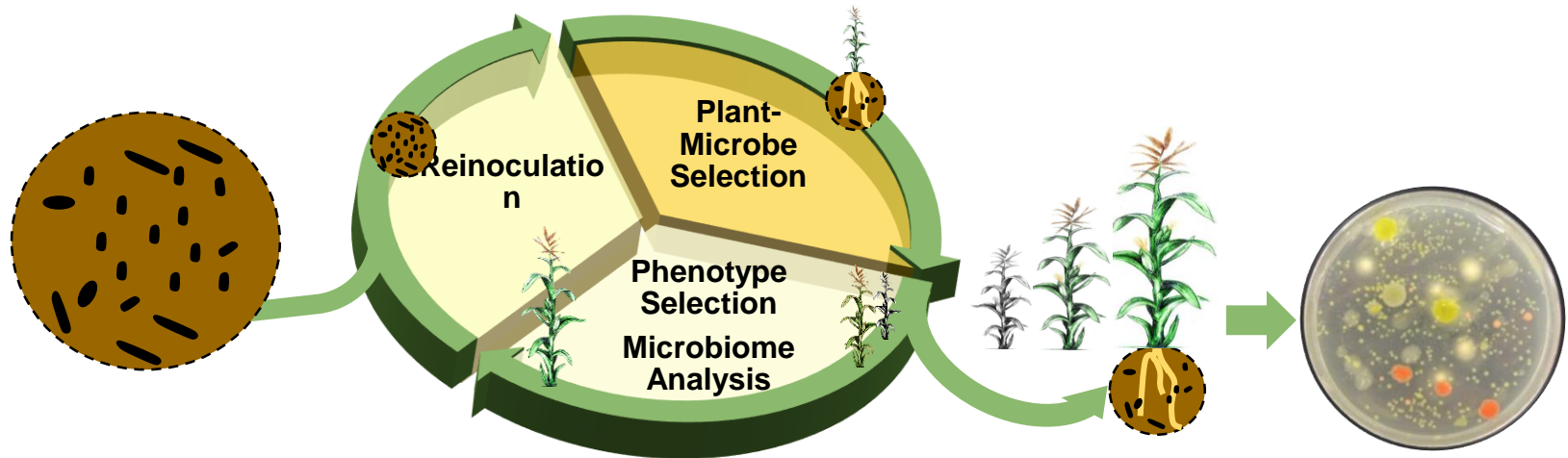




Advanced Microbial Selection (AMS)

Advanced Microbial Selection (AMS)

Directed selection of the microbiome to identify teams of beneficial microbes



Diverse soil microbes

Microbe capture

Directed selection

Final selection

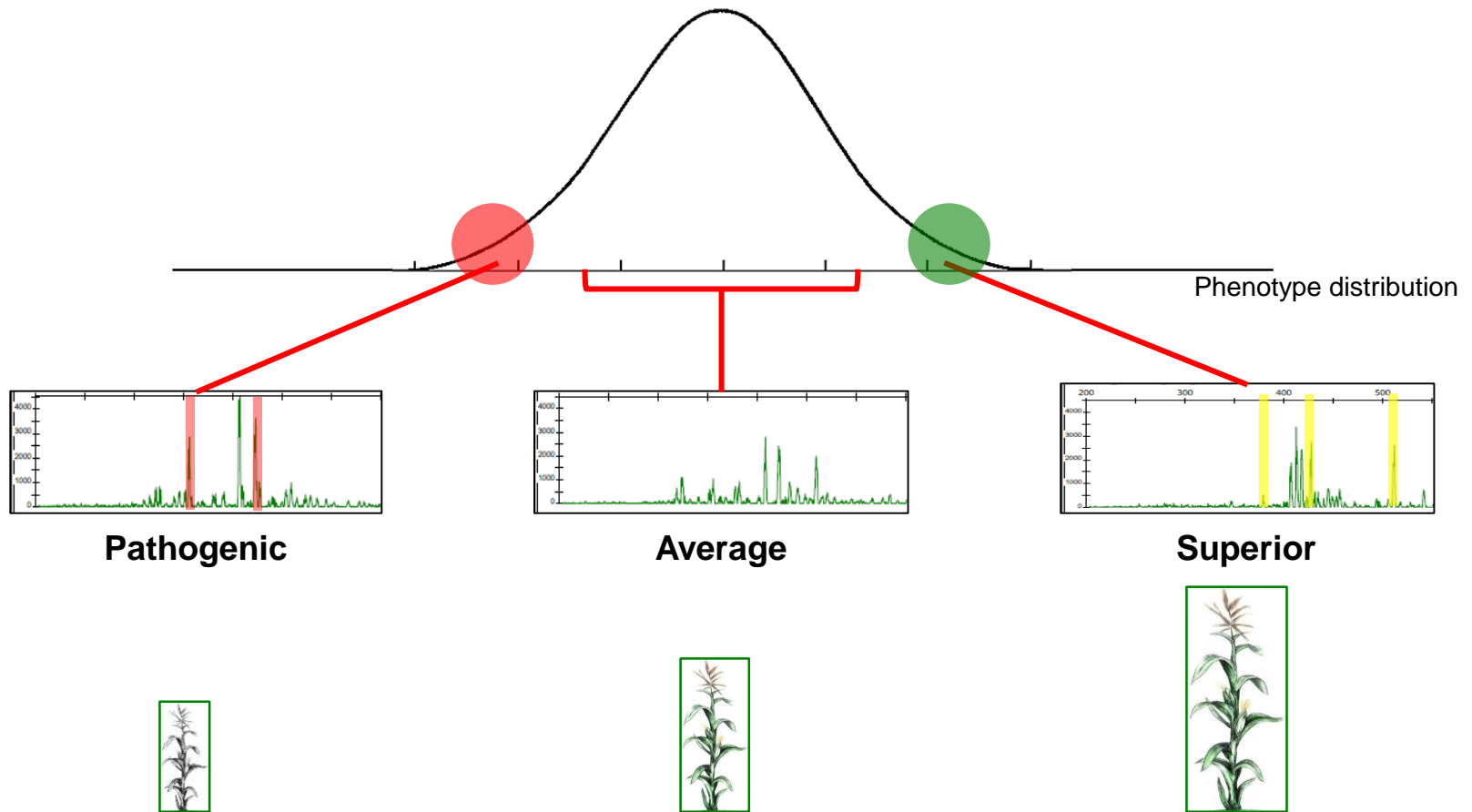
Microbe isolation

Conducted with or without application of stress

Process and selection informed by DNA and microbiome analysis

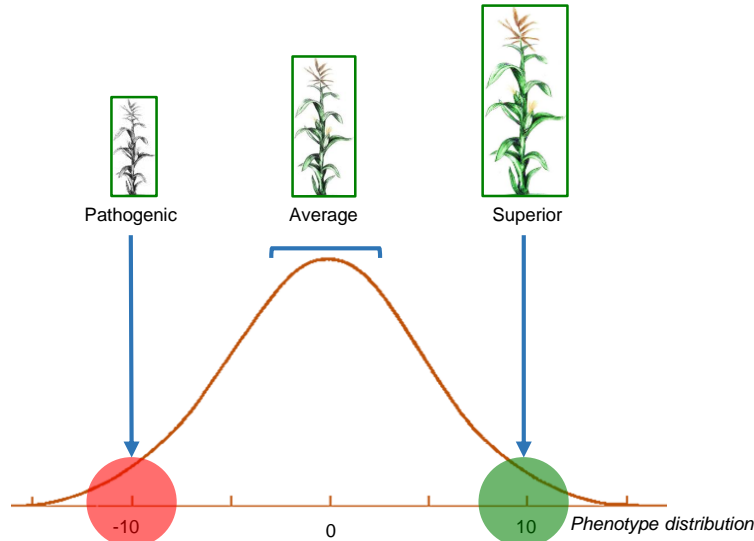
Proprietary, patented process

Selecting Superior Phenotypes



Selection process is completed in both ideal and stressed conditions, such as nutrient deficiency, drought, etc.

Directed Selection

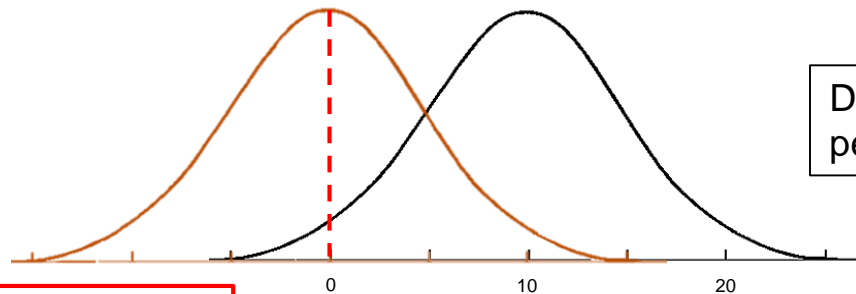
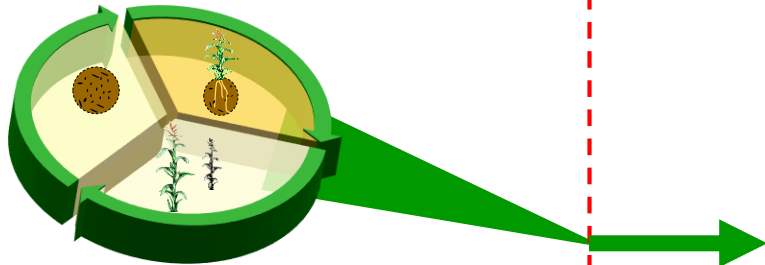


The AMS Process

Selecting the superior phenotype

Changing the microbial community

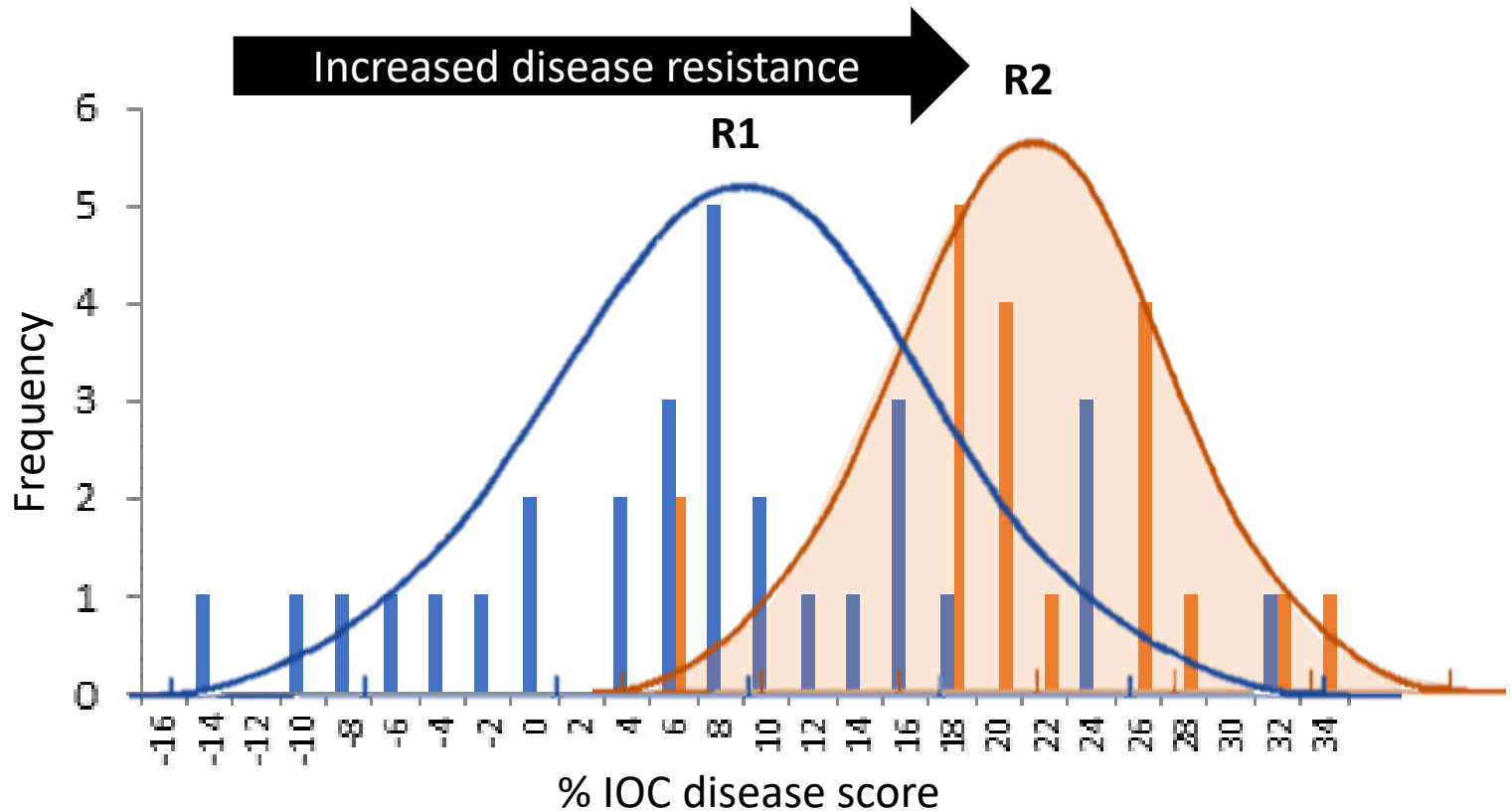
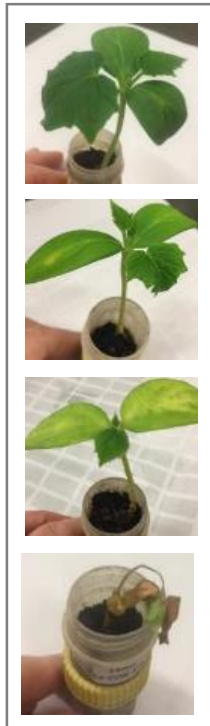
Driving an improvement in trait performance



Proprietary, patent-pending process

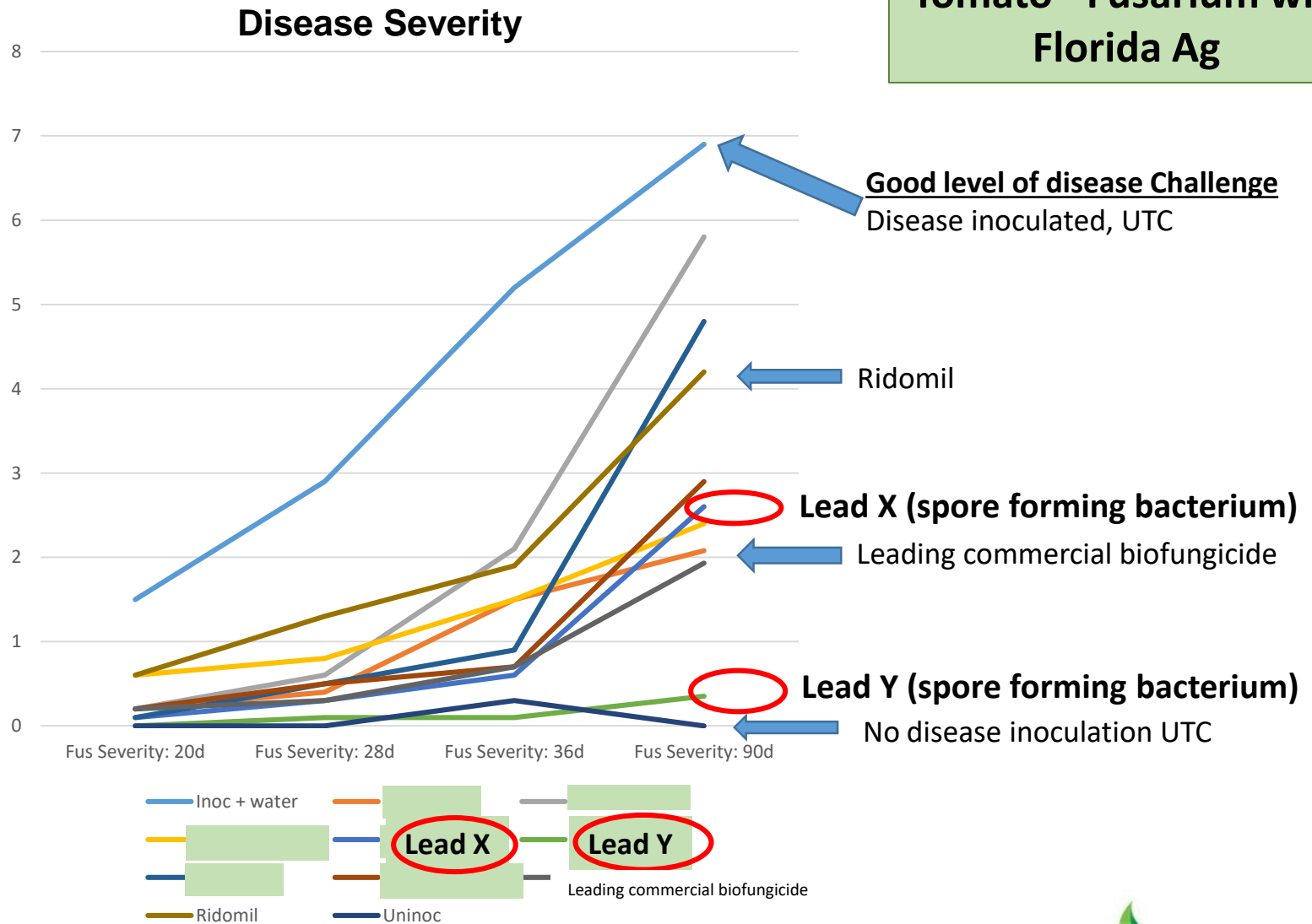
AMS Induces a Shift in Disease Resistance

AMS for Pythium Resistance in Cucumber



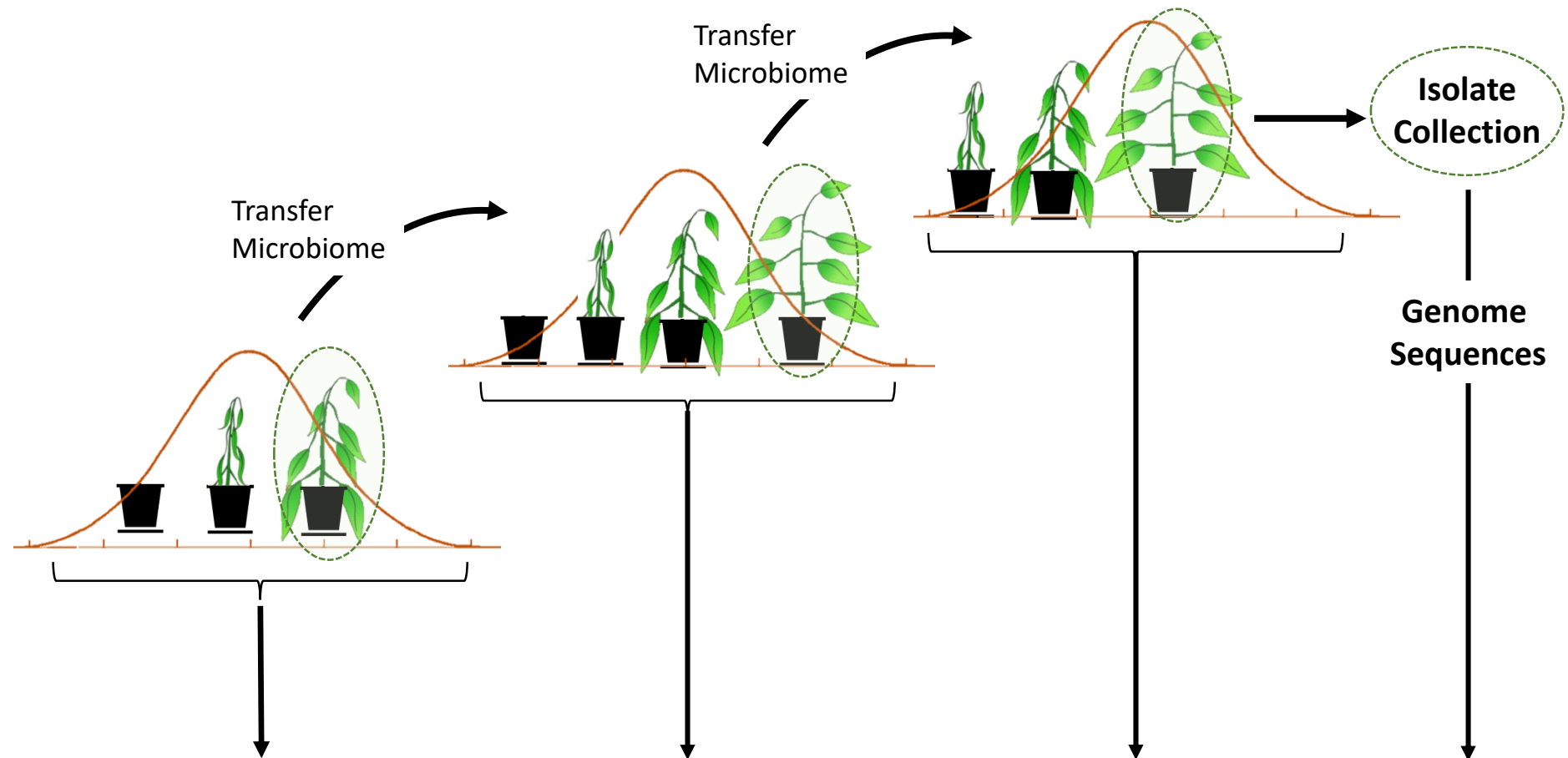
Initial Biofungicide Program

**Tomato - Fusarium wilt
Florida Ag**



AMS + Microbiome Analysis

Microbiome analysis makes AMS more powerful



Microbiome Analysis - Machine Learning Identifies Key Microbes

- Comparative analysis of high and low performing plants across each AMS round
- Key species, strains and consortia (co-occurring) identified

AMS and Microbiome Analysis

AMS helps us narrow in on the members of the microbiome that matter

Filter 1

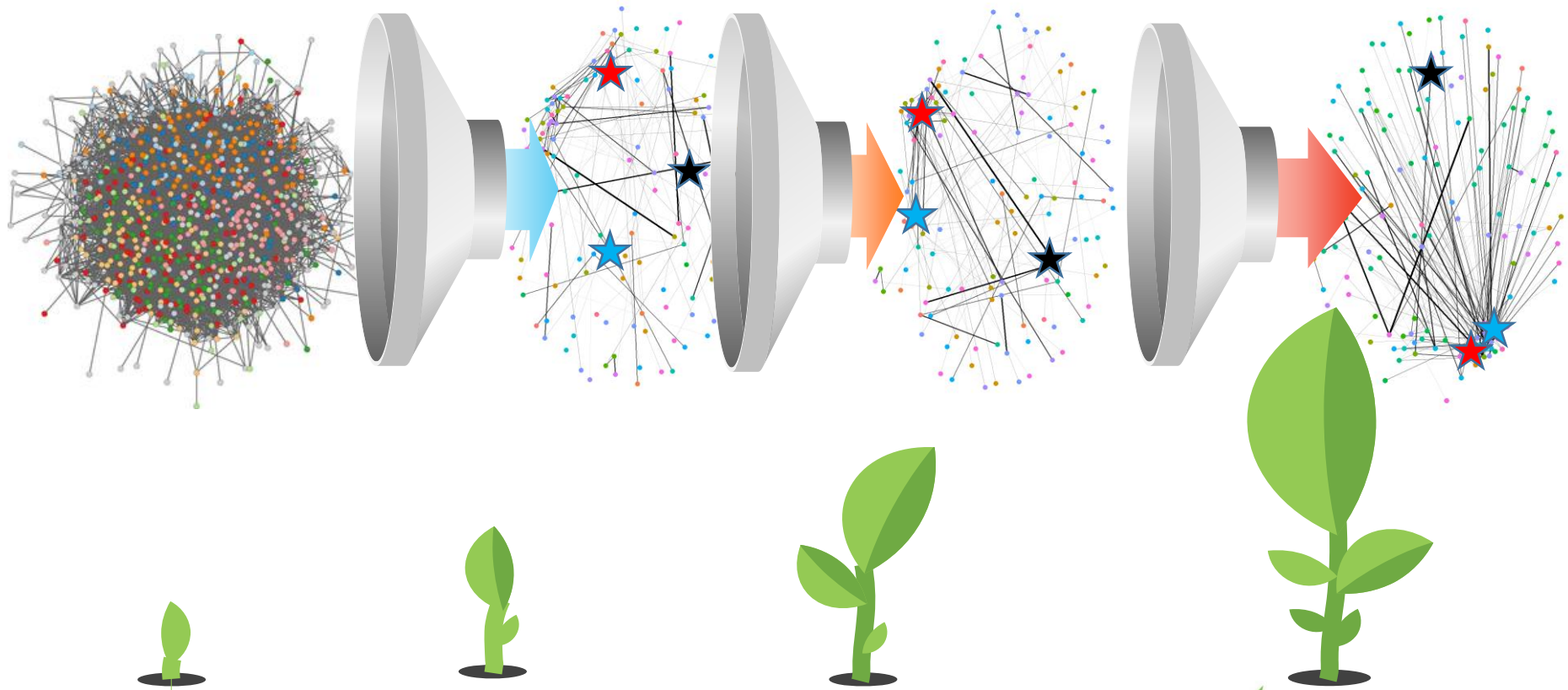
Plant-associated microbes are selected during microbe capture

Filter 2

Microbes that are retained under selective pressure are transferred

Filter 3

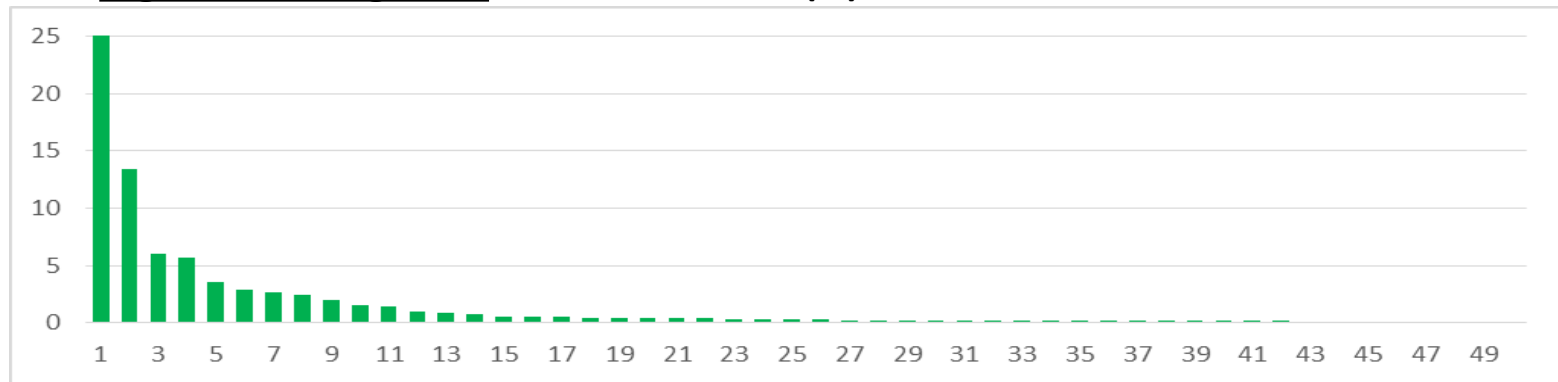
Microbes that consistently confer a positive effect through selection rounds are targeted for isolation



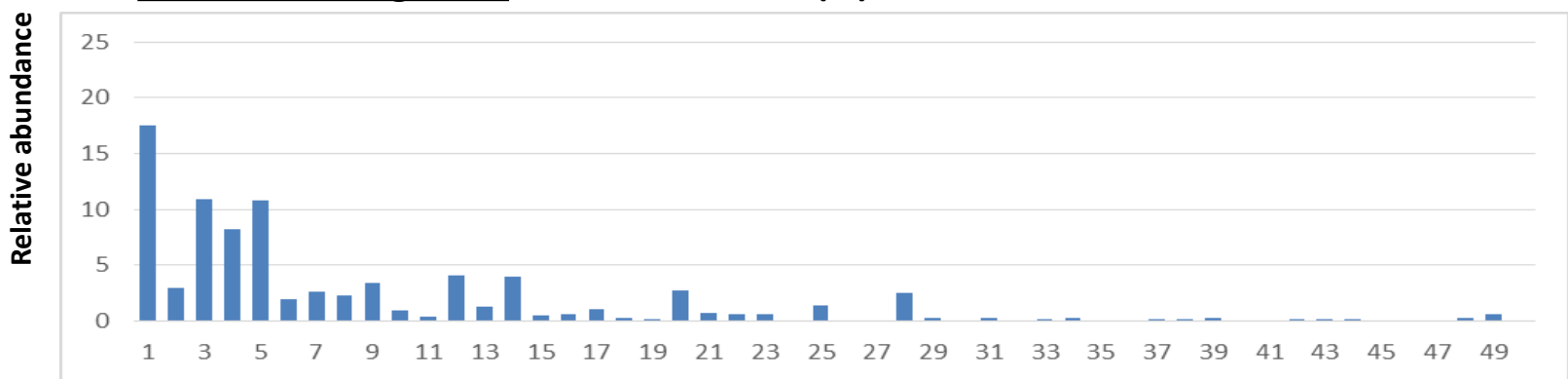
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High Performing Plant: OTU abundance (%)

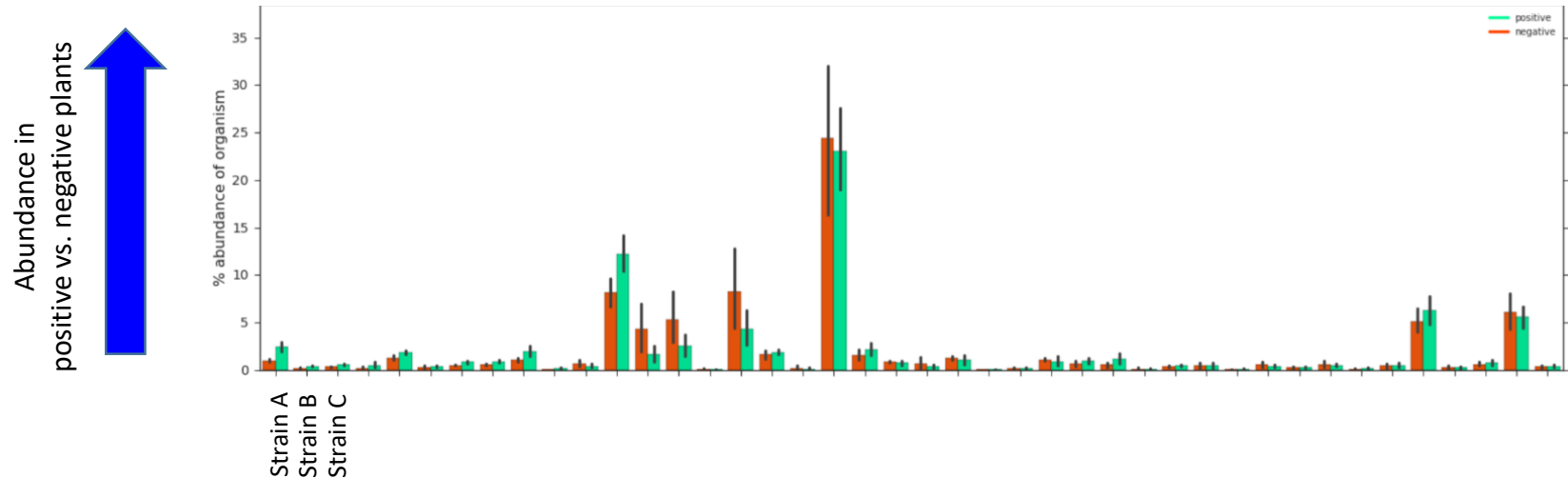
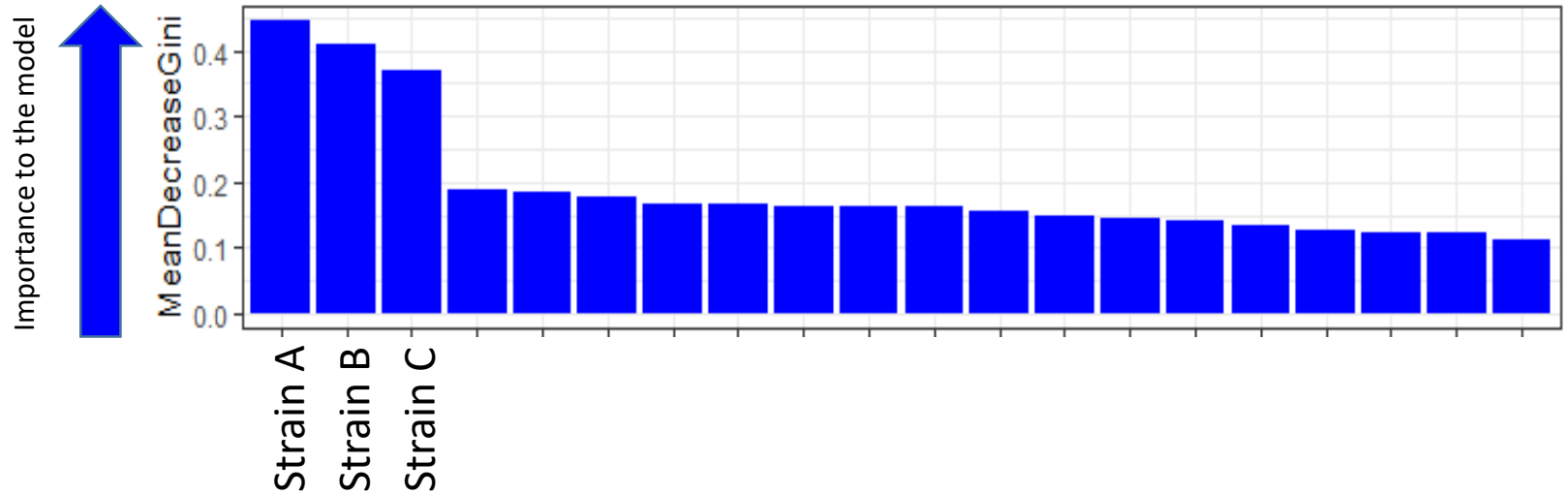


Low Performing Plant: OTU abundance (%)



Machine Learning Identifies Key Microbes

-the most important microbes are not always the most abundant

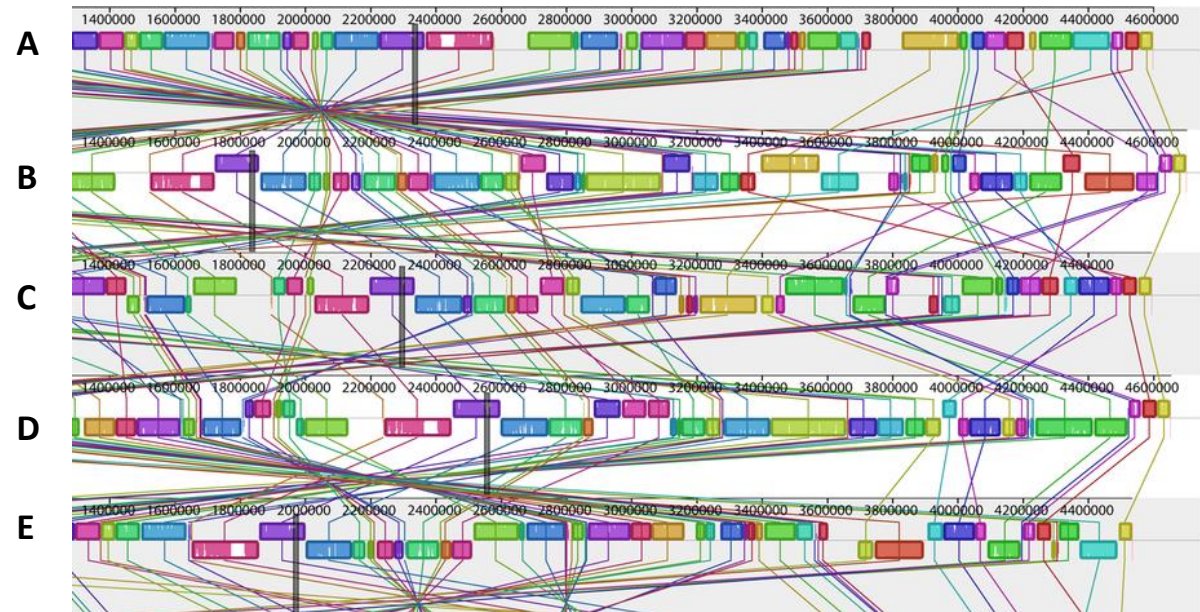
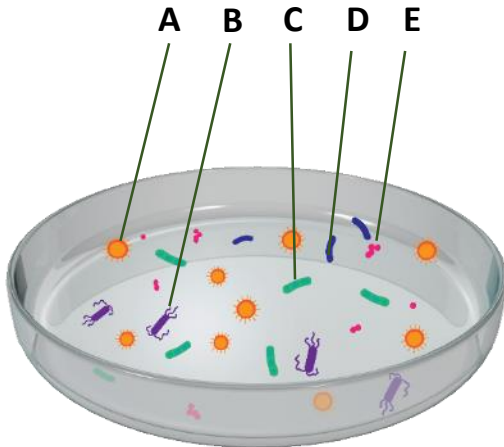


A close-up photograph of a young green plant seedling. The seedling has two large, rounded, light green leaves and a central stem with a smaller, developing leaf at the top. The background is a soft, out-of-focus greyish-green.

Genomics

Genome Sequencing & Comparative Genomics

- Genomic sequencing and comparative analysis of isolates
- Consolidating the link between community analyses, functional prediction and strain selection



Microbial Potential

Genomic analysis can provide insight into microbial potential

01 Nutrient Acquisition

02 Root Colonization

03 Biocontrol

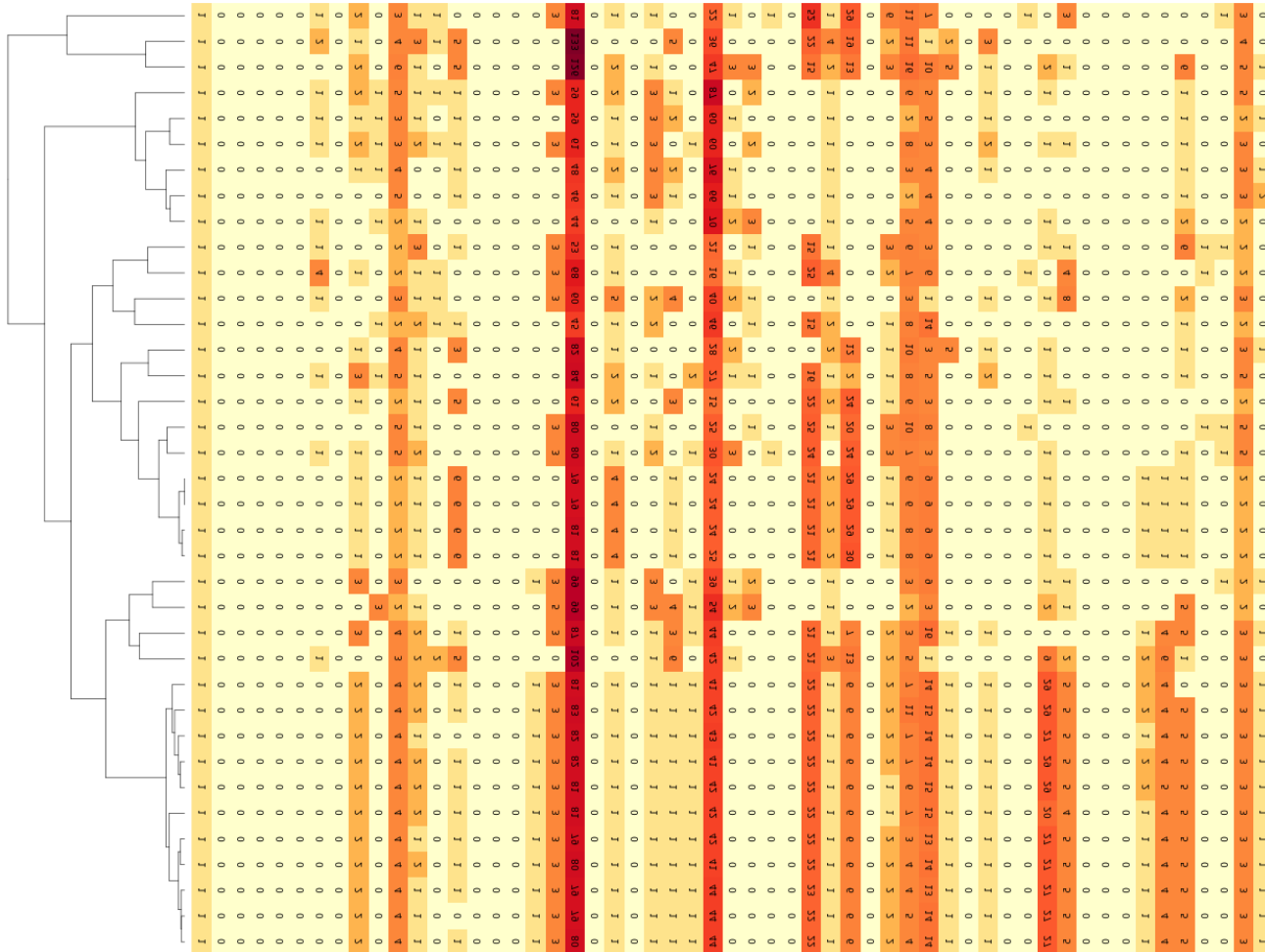
04 Abiotic Stress Resistance

05 Pathogenicity

06 Discovery

Genomics Platform

Combining data from genomics, microbiome, HTP microbial phenotyping, and in planta assays to build a platform for predicting the best consortia

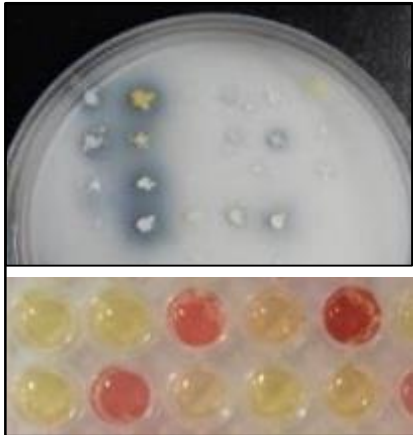


Isolates

Gene Content

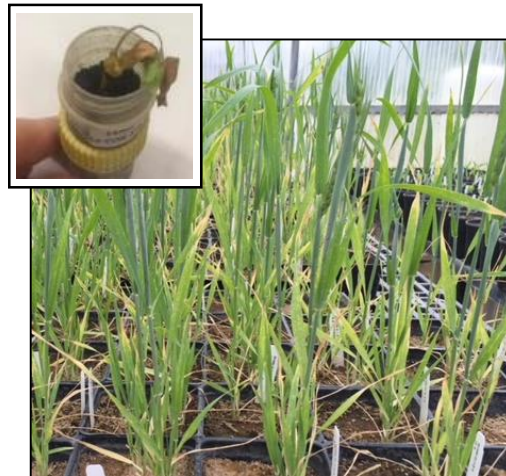
High Throughput Phenotyping (HTP)

HTP microbe phenotyping



- All microbes screened
- Broad spectrum of functions

AMS (focus on easily scored phenotypes)



- Biopesticides
- Abiotic stress

HTP in-planta assays for consortia screening



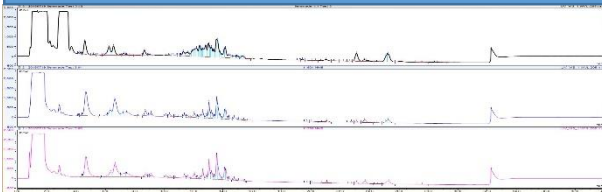
- Growth room & greenhouse assays
- Digital imaging
- Drone-based field imaging

Biofungicide/Biostimulants Leads are Extensively Characterized for Functionality and MoA

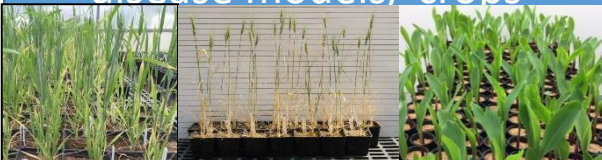
Spectrum of control on plant pathogens



Bioactive metabolite profiling



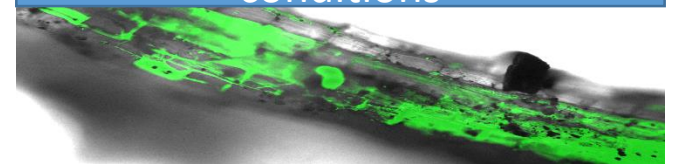
Extensive in-planta evaluations with multiple disease models/ crops



Nutrient solubilization ability
N, P, K, Si, Zn, Fe

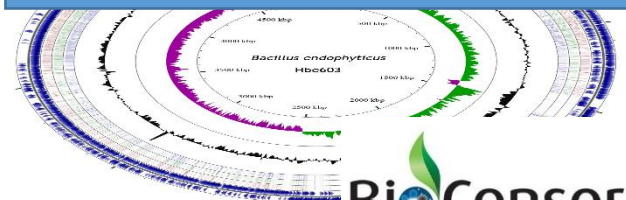


Root colonization robustness
under different soils & conditions



Field
Performance
Consistency
Robustness

Comparative genomics analysis
for beneficial traits /safety



High-Throughput Microbial Screening



One AMS example

150 soil samples

> 10^{13} microbes

> 10^5 different strains

> Multiple seed chemistries

Selection



60 microbial strains

Lab & Growth Chamber Trials

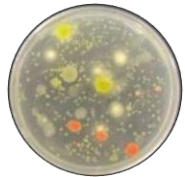
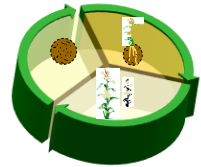


11 microbial consortia

Pot trial

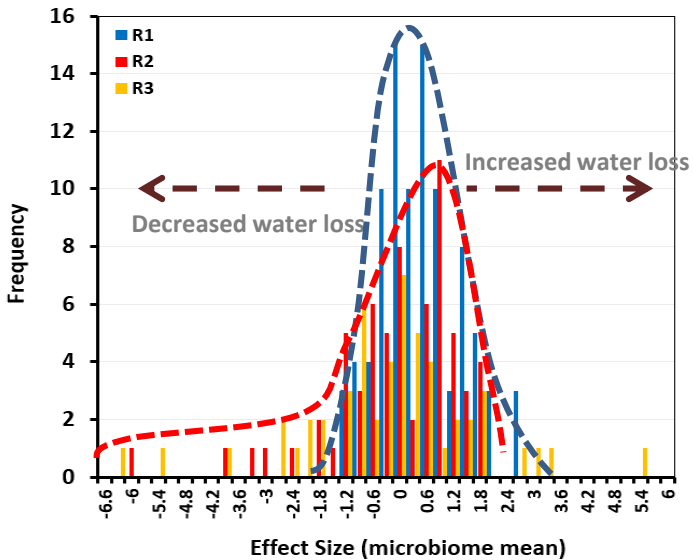


Field trial

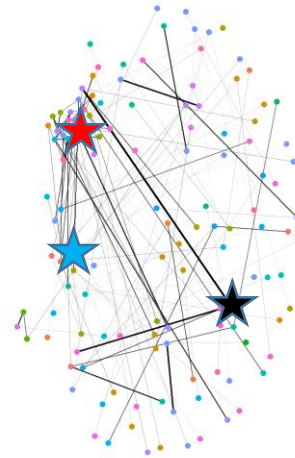


Wheat Water Use Efficiency (WUE)

AMS: Wheat Drought Resilience Multiple positive phenotypes



Consortia Evaluation Water Use Efficiency (WUE)



Machine learning used to identify key microbes in microbiome

Microplot Trials Heat and Water Stress

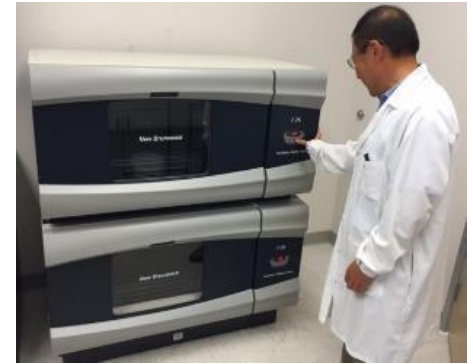
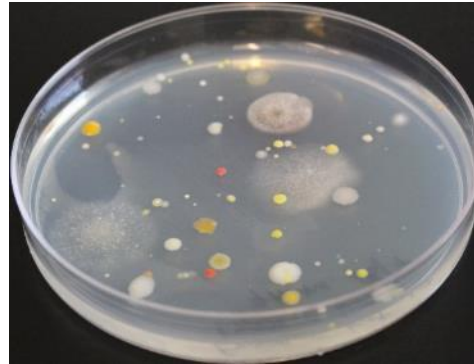
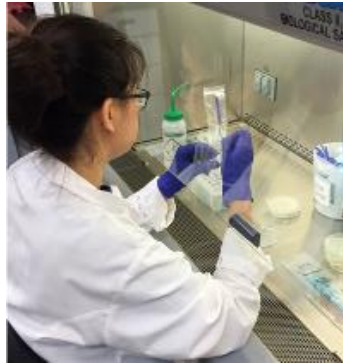
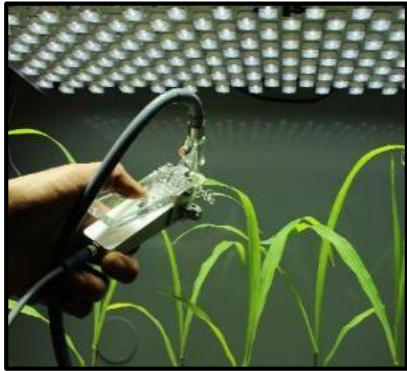


Four consortia (out of 30) showed 15 – 20% yield gain

- Best treatments predicted by microbiome analyses

Initial field trial results

- best lead showed 13.4% yield increase in sub-optimal watering and 8% across all trials



Thank you



Bio**Consortia**