



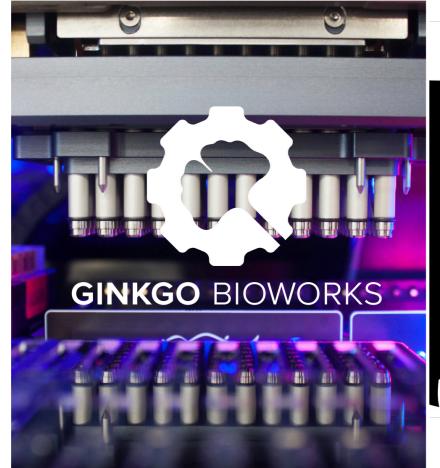


### **Biology by Design**

Will(Yangxiaolu) Cao Design Senior Engineer

### Intro of Ginkgo

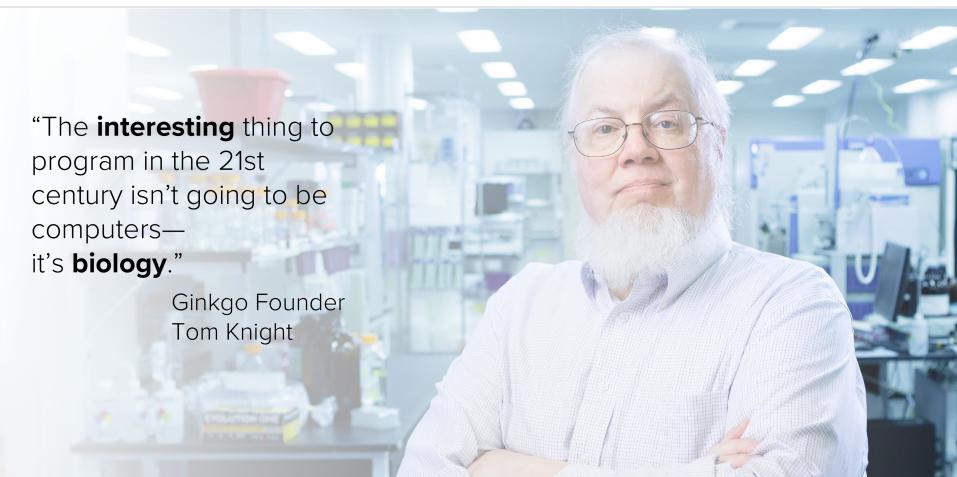






### **How Ginkgo get started?**



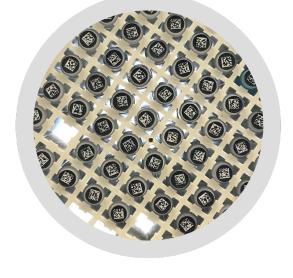




### What do we do?









Chemicals

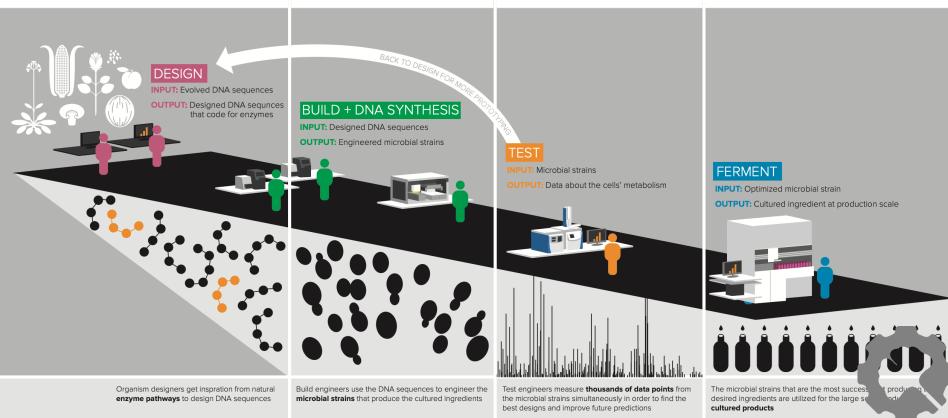
Strain Improvement

Enzymes

### **Foundry**



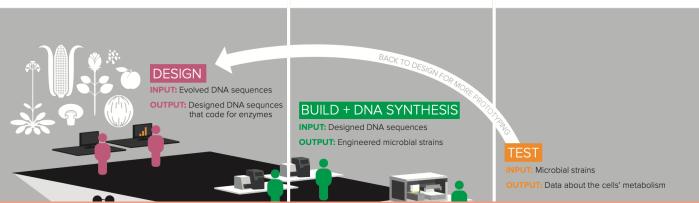
#### LIVING FOUNDRY PROTOTYPING CULTURED INGREDIENTS



### **Foundry**



#### LIVING FOUNDRY PROTOTYPING CULTURED INGREDIENTS



FERMENT

INPUT: Optimized microbial strain

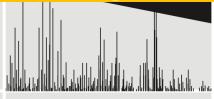
software

#### automation

#### next generation sequencing

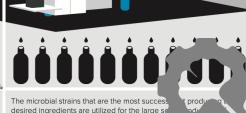






Test engineers measure **thousands of data points** from the microbial strains simultaneously in order to find the best designs and improve future predictions

The microbial strain desired ingredients cultured products

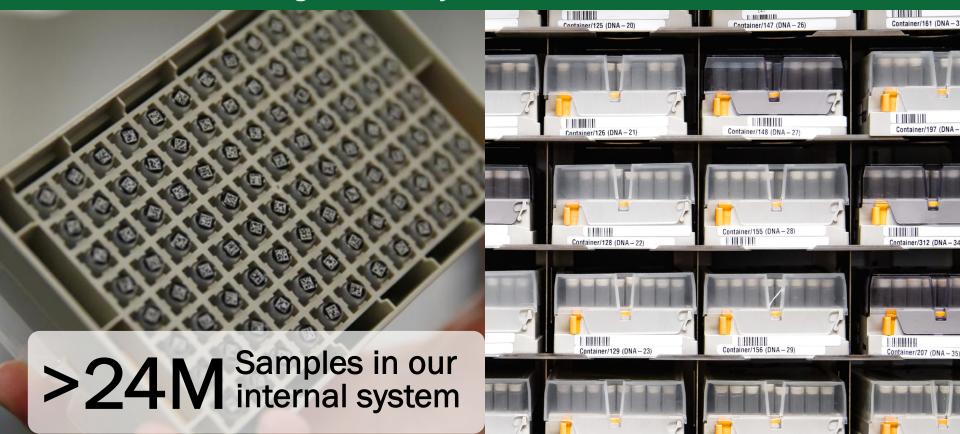


Organism designers get inspration from natural **enzyme pathways** to design DNA sequences

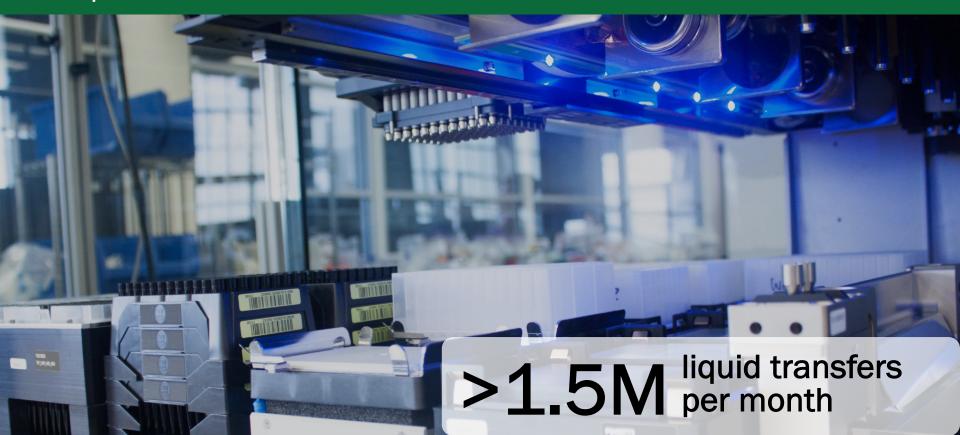
Build engineers use the DNA sequences to engineer the **microbial strains** that produce the cultured ingredients



# Logistics are managed using barcoding and proprietary Information Management System

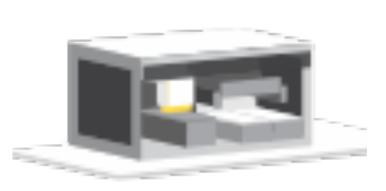


# Automated processing of DNA, proteins, cells, and compounds



### Foundry snapshot in Jan 2020





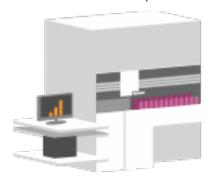
>1 million foundry operations per month



>10K bacterial genomes per week



>80K strain tests per month



>8K new strains

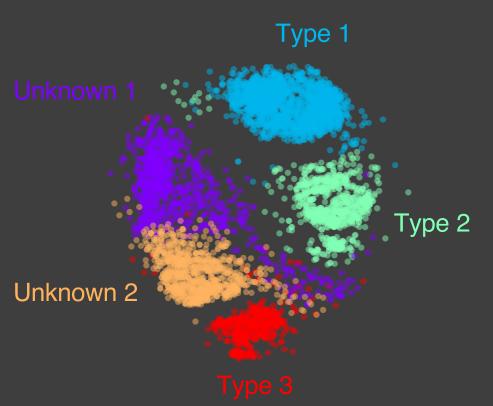
### A case study: enzyme sourcing



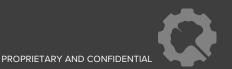




EXPLORING BIODIVERSITY AT FOUNDRY SCALE

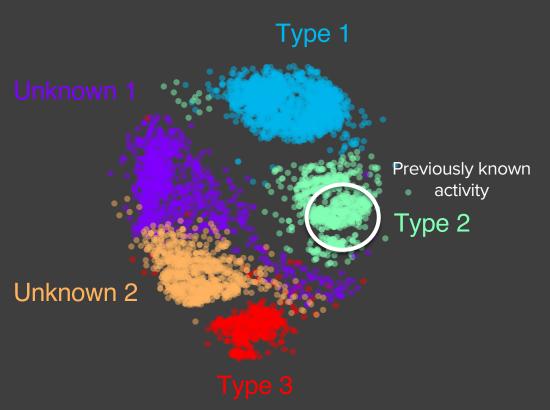


Sequence similarity space of 5449 enzyme homologs

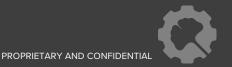




EXPLORING BIODIVERSITY AT FOUNDRY SCALE

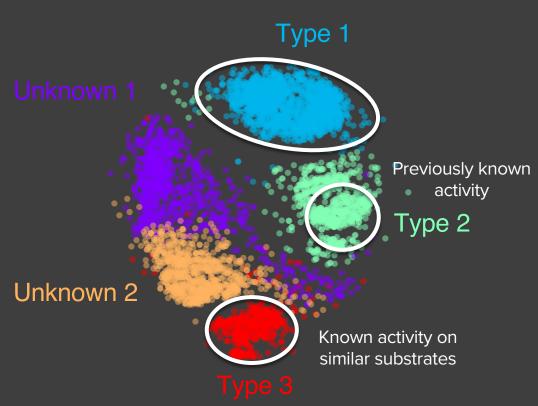


Sequence similarity space of 5449 enzyme homologs

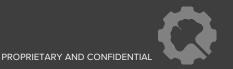




EXPLORING BIODIVERSITY AT FOUNDRY SCALE



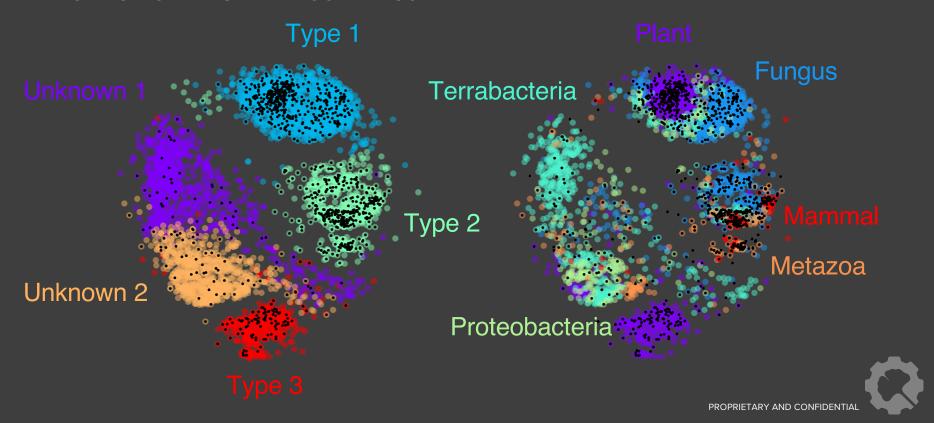
Sequence similarity space of 5449 enzyme homologs



### **ENZYME DISCOVERY**

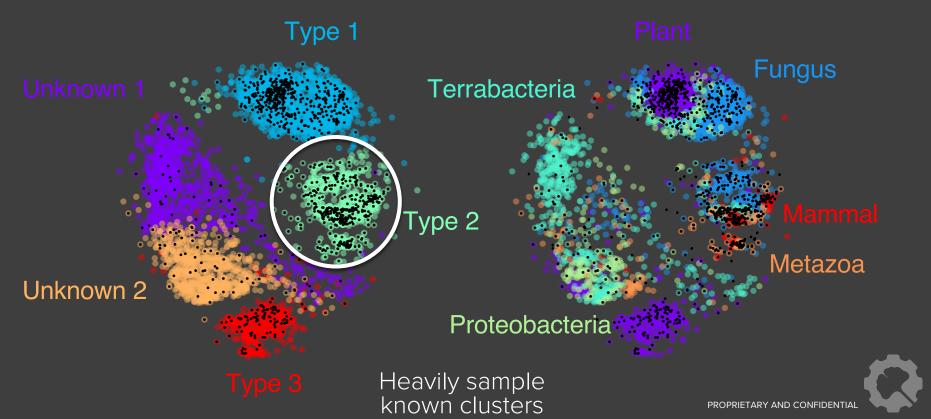


#### EXPLORING BIODIVERSITY AT FOUNDRY SCALE



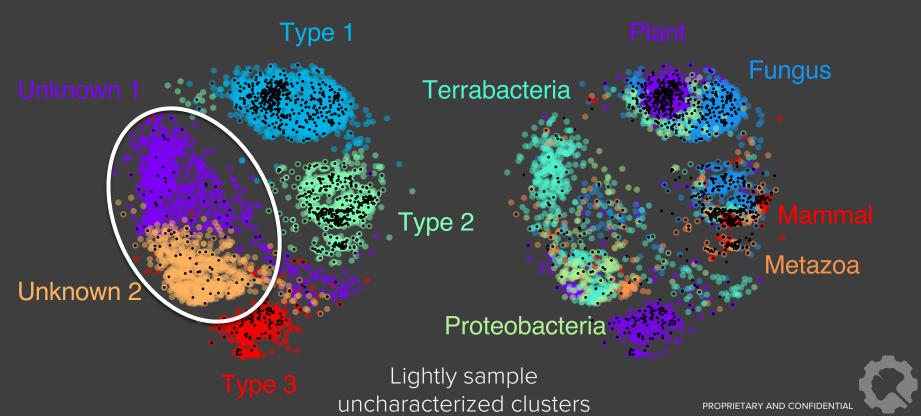






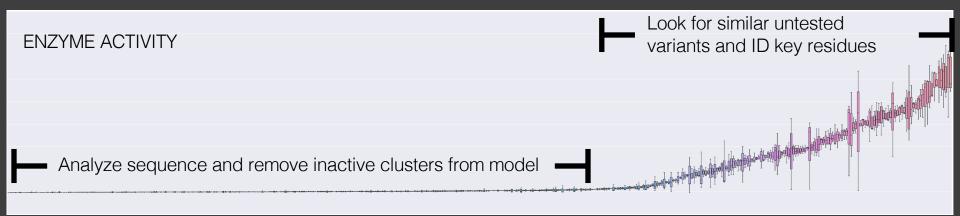


EXPLORING BIODIVERSITY AT FOUNDRY SCALE





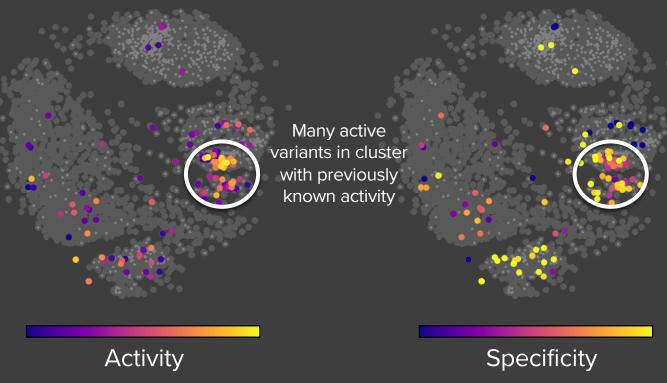
EXPLORING BIODIVERSITY AT FOUNDRY SCALE



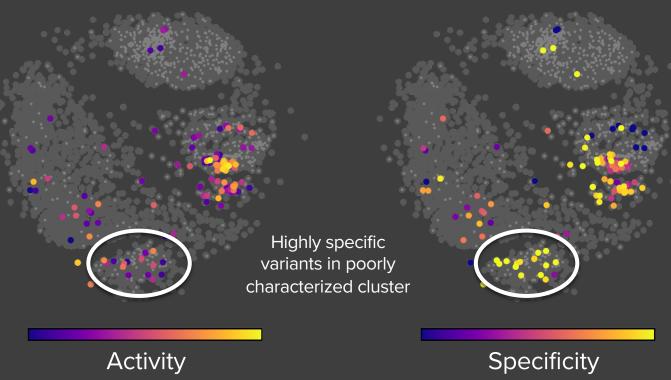






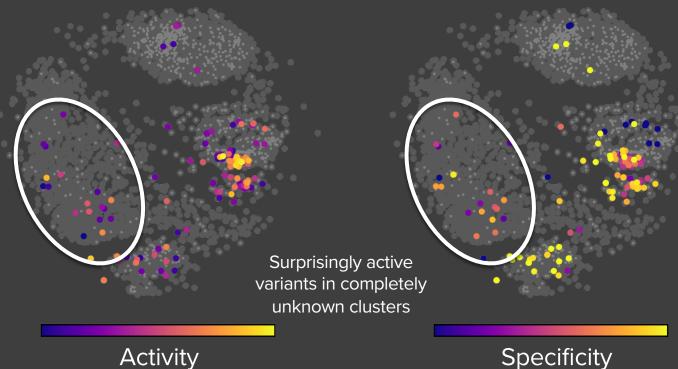






### **ENZYME DISCOVERY**

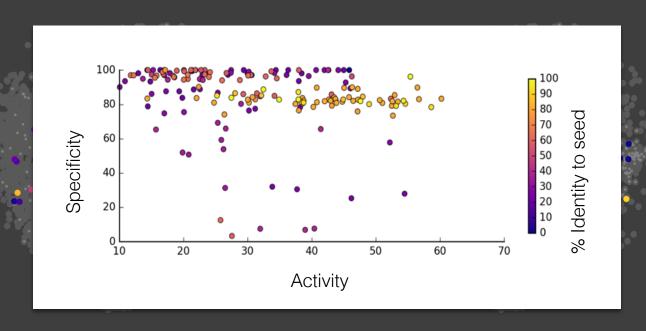








#### METABOLOMICS FOR SPECIFICITY



Activity

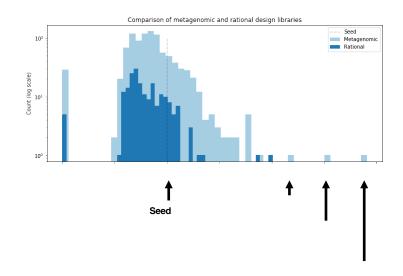
Specificity

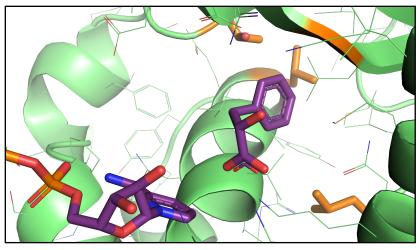


# **Enzyme Discovery+ Engineering**



#### INCORPORATING RATIONAL PROTEIN DESIGN



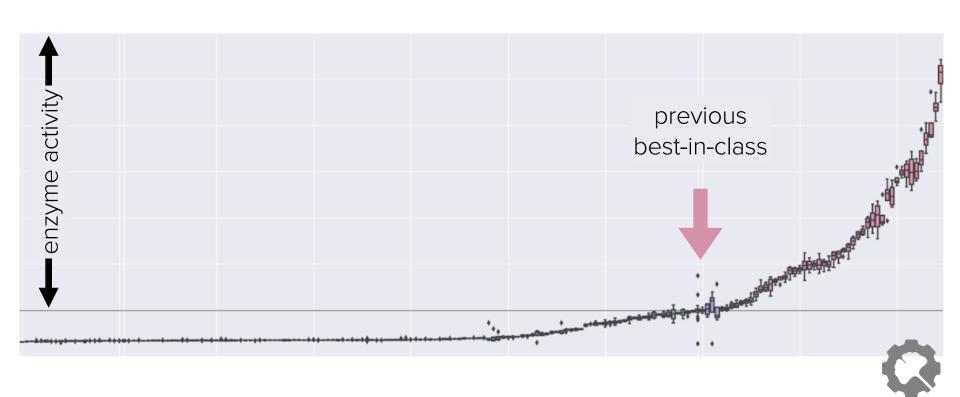


15 positions targeted for site-directed mutagenesis; 4 key positions shown





METAGENOMIC SCREENING



### Pathway balance





### Pathway design





DESIGN

Programmatic design of multiple operon architectures with standard parts

BUILD + SYNTHESIS

500 kb of synthesis Assembly in lowcopy vectors

**TEST** 

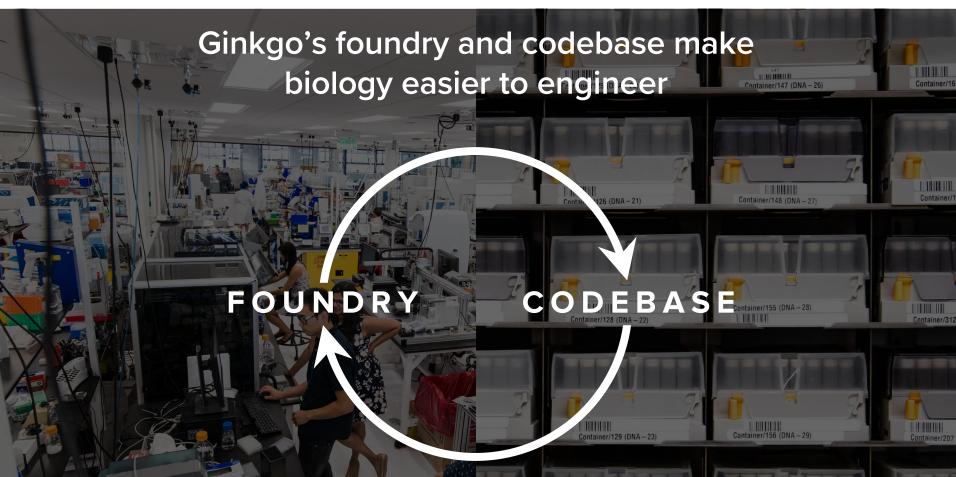
LC-MS metabolomics + 13C fluxomics

**FERMENT** 

Validation of strain performance in scale-down model

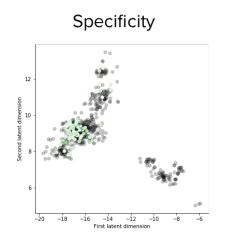
### CodeBase

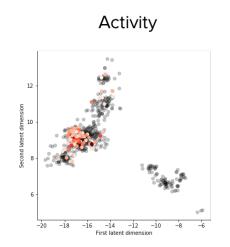


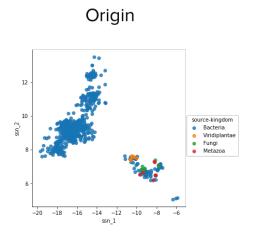


### **Machine and Deep learning**





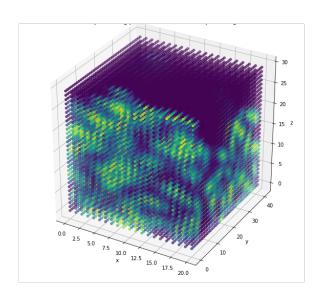




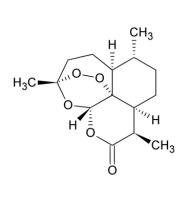


### **Machine and Deep learning**

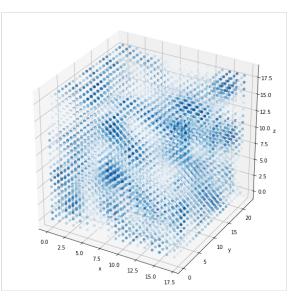




Active site representation of enzyme



Target molecule



Convolutional kernel tensor



### **Codebase benefits**



#### Codebase enables Ginkgo to work across industries more efficiently

FRAGRANCE - - - - - - - BAKING



Enzyme library developed for a fragrance application

Data from fragrance work used to refine and expand the library for an application in electronics

Enzyme library again leveraged to find enzymes for baking applications



FOOD

THERAPEUTIC

AGRICULTUR

M A T E R I A L S

C U L T U R E D I N G R E D I E N T S CONSUMER ELECTRONICS

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INDUSTRIAL BIOTECH

ENERGY & ENVIRONMENT

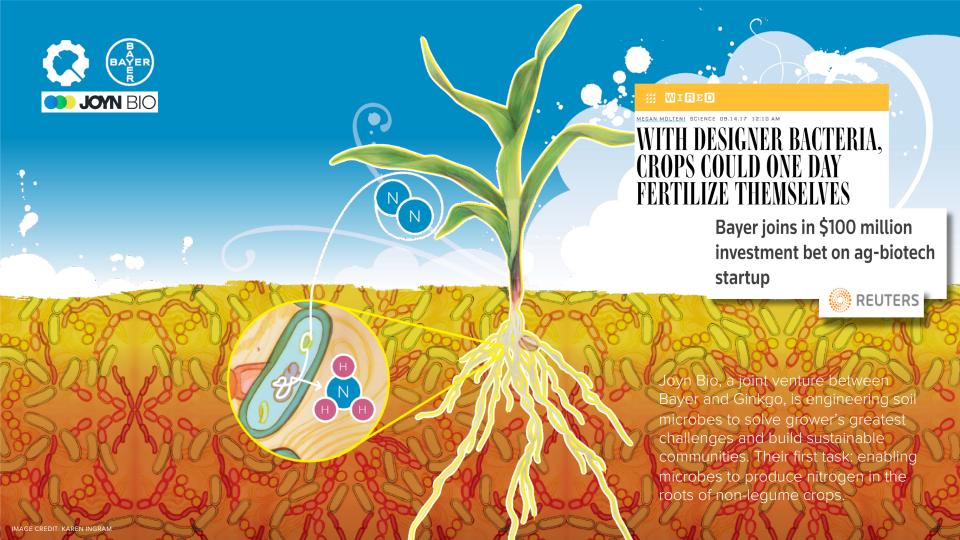
BIODEFENSE



#### **CMN BUSINESS**

Burger King plans to roll out Impossible Whopper across the United States





#### FERMENTED CANNABINOIDS

Enable access to hard-to-source medicinal cannabinoids

Engineered production enables more costeffective and consistent high-quality supply compared to farm-based approaches

Q Search

**Bloomberg** 

**Business** 

**Cronos Partners With Ginkgo to Develop Lab-Grown Cannabis** 

FINANCE POLITICS STRATEGY LIFE INTELLIGENCE ALL

A startup with ties to Bayer has inked a \$122 million deal to make lab-grown marijuana — and it's eyeing the pharma industry





# Thank You

**GINKGO BIOWORKS** 

# BIOLOGY BY DESIGN

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