



Genomatica

- Sustainable chemicals
- Better economics
- Smaller footprint

Pacific Rim

Summit on Industrial

Biotechnology & Bioenergy

October 10-12, 2012

Nelson Barton, VP R&D

A Process for the Production of Bio-based 1,4-Butanediol (BDO)

Genomatica at a Glance

The Company

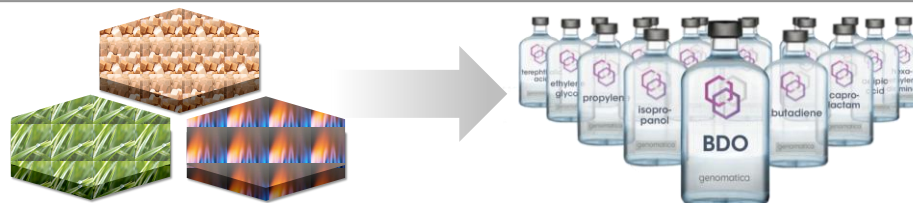
Private company



*San Diego (USA)
105 people*

Bio-Based Chemicals

Basic/intermediate chemicals
Technology provider



Top Financial Investors

Raised \$125 MM to date



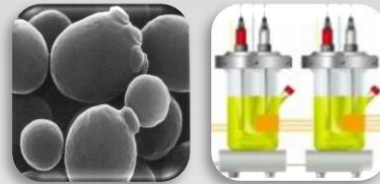
Partnering with Industry

Global commercialization
feedstocks, scale-up, applications



Genomatica: Focused on Bio-based Chemicals

Low-Cost
Renewable Feedstock



High Performance Organisms

Basic & Intermediate
Chemicals



Existing products
sold globally

High titer,
Tolerance

High
yield

High
productivity

- Exact same chemicals
- Better economics
- Equivalent performance
- Reduced energy
- Lower carbon footprint
- Feedstock diversification

Direct production of chemicals likely to afford lowest cost process

Bio-based 1,4-Butanediol

Crude Oil
& Nat. Gas



Acetylene

Propylene

Butane, Butadiene



Sugars



Direct Production

3 B lb/yr existing market



spandex

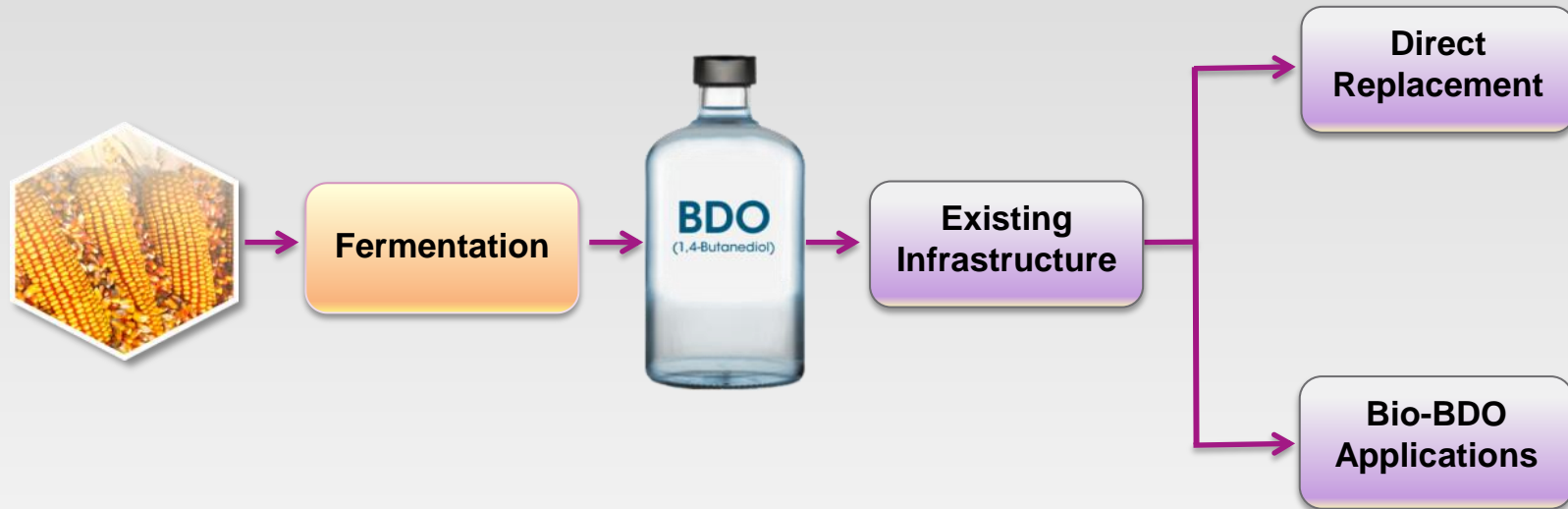


polyurethans



PBT

Advantages of Identical Drop-in Bio-BDO

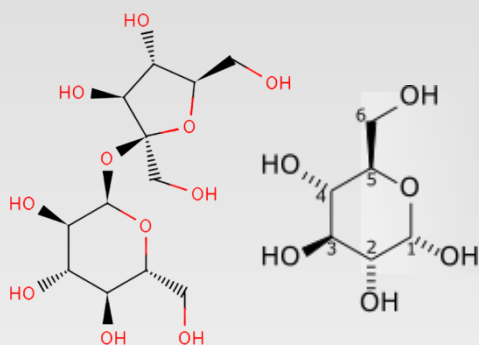


PBT Pellets
from Bio-BDO

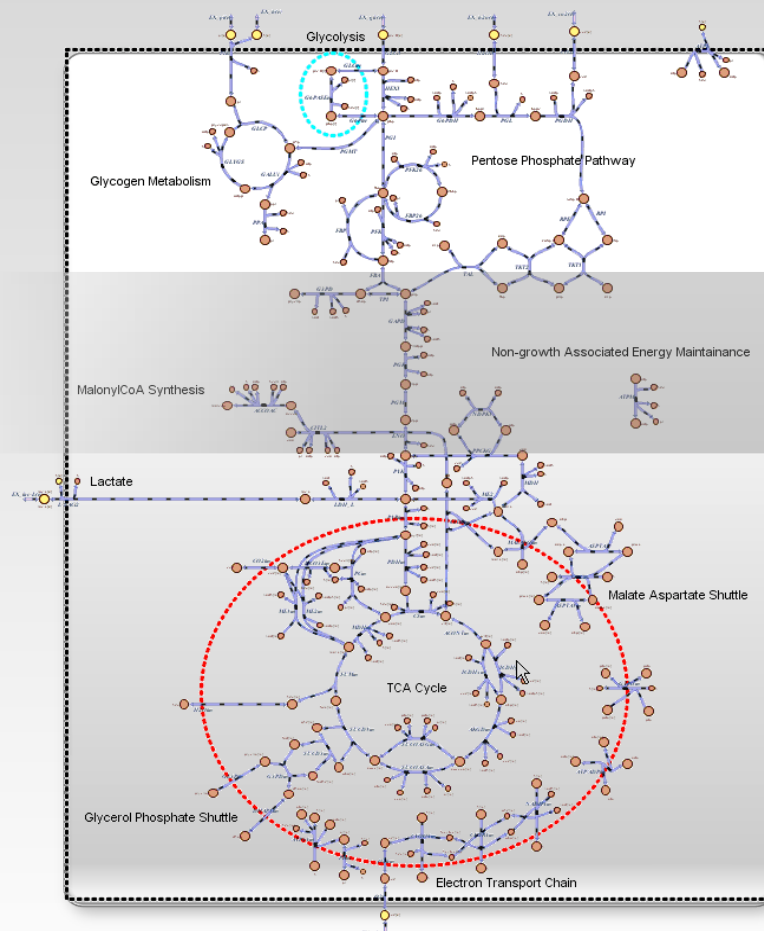
- Downstream infrastructure in place
- Can blend with existing BDO
- Faster adoption
- Meet customer needs for renewable products

Genomatica's BDO Process in Engineered *E.coli*

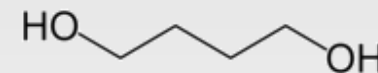
Sugars



BDO-producing *E. coli*



**1,4-Butanediol
(Bio-BDO)**



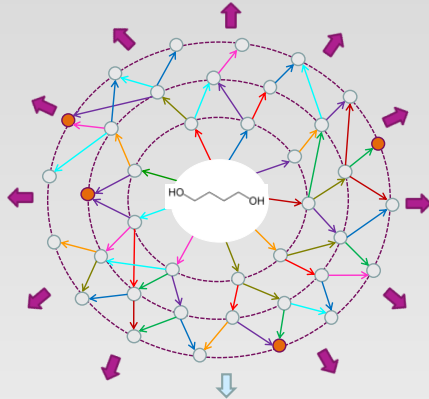
100% bio-based content



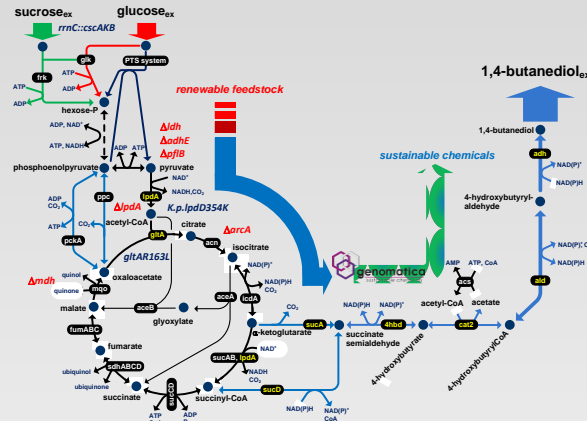
Tate & Lyle facility

Strain, fermentation, process engineering → deliver BEP

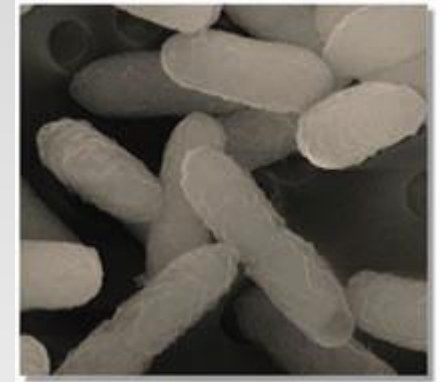
Journey to a BDO Production Strain



Pathway Identification and Engineering



Strain Design and Metabolic Engineering



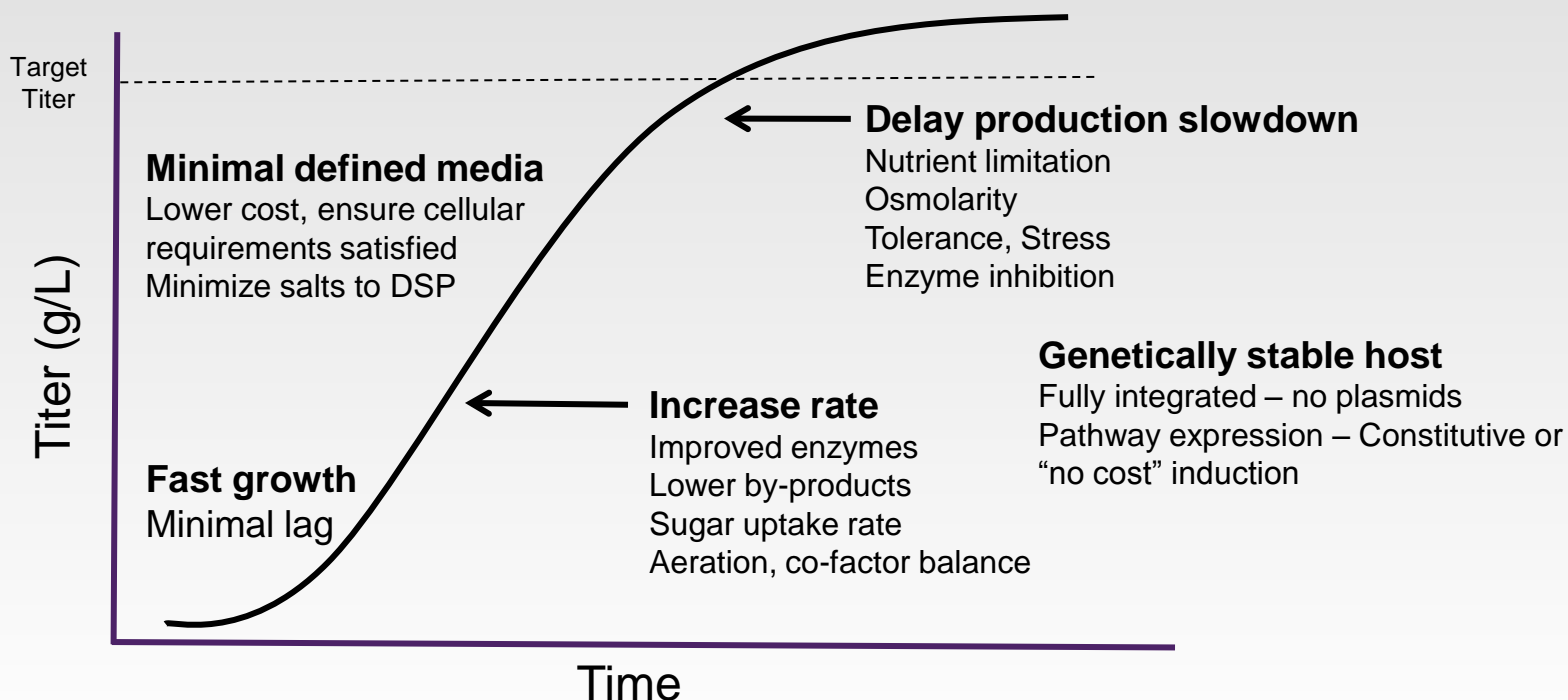
Commercial Strain for BDO Production

Fermentation Metrics → Higher TRY = Lower COGS

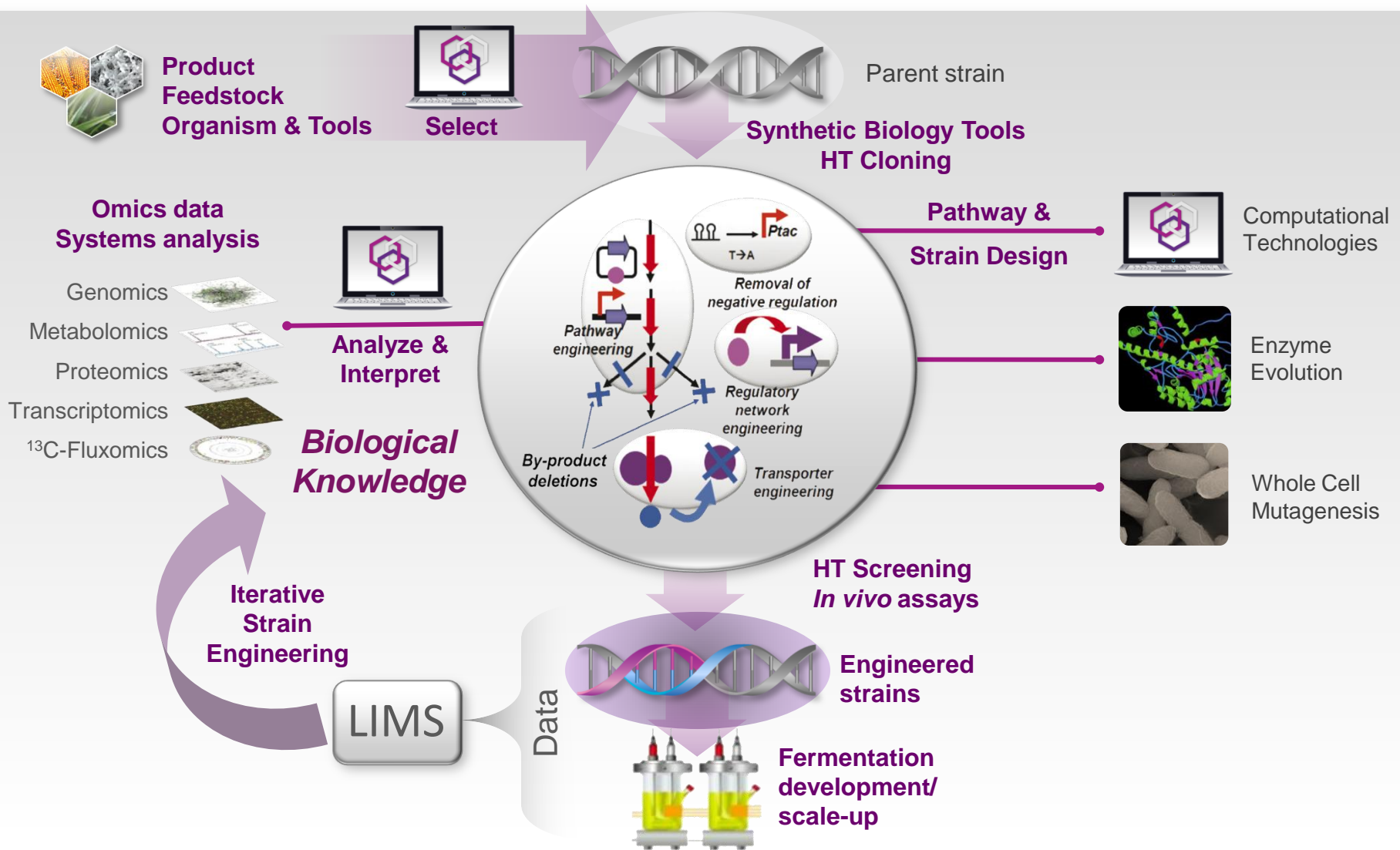
- **Titer (g/L)** - Impacts equipment sizing and energy needs
- **Rate (g/L/h)** - Impacts # of fermentors, plant capacity
- **Yield (g/g)** - feedstock cost, by-product cost in DSP

Commercial Strain Engineering Challenges

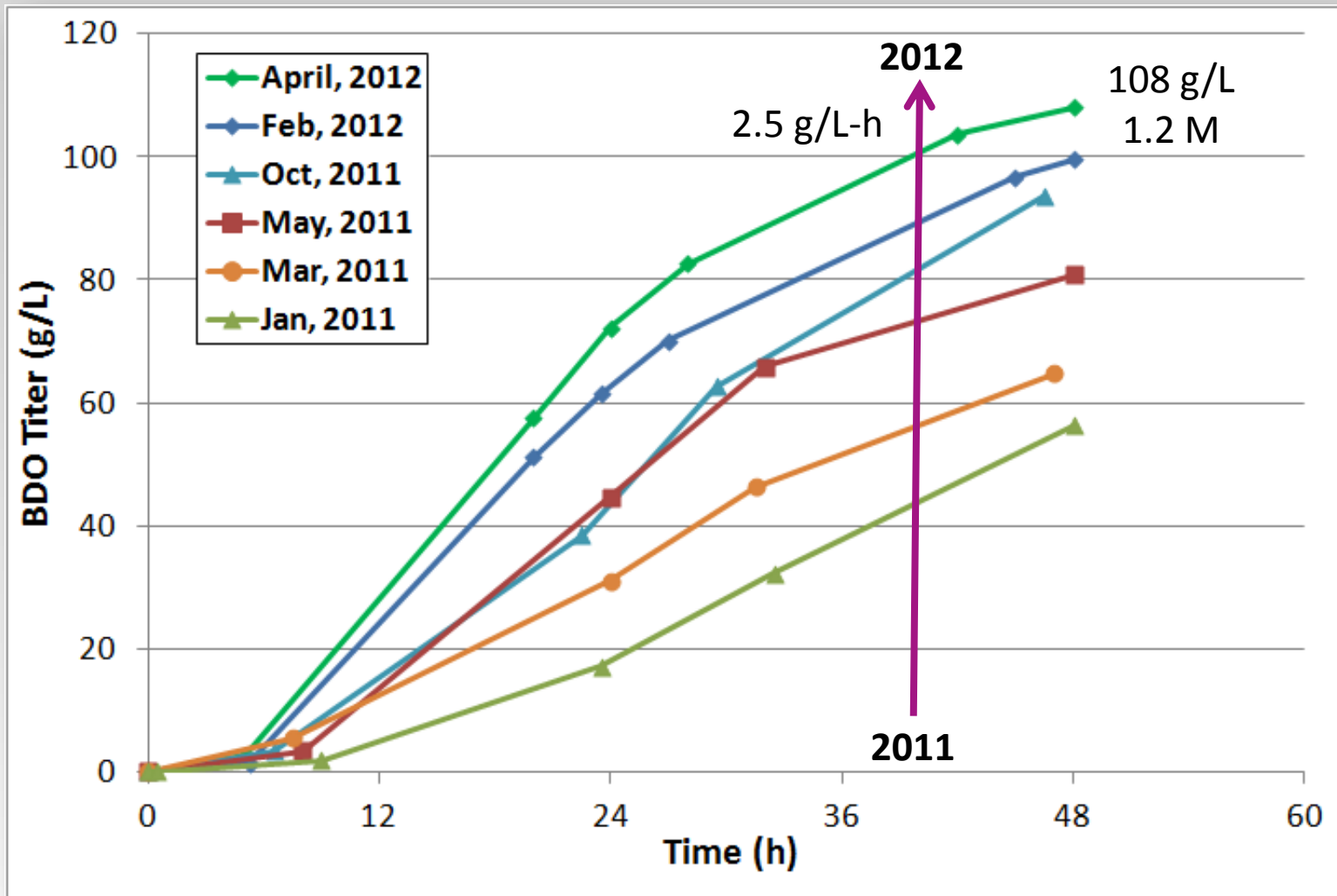
- Major Challenge – High Titer, Rate, Yield – “It is all about the Bug”
- Define commercial TRY targets needed to achieve desired economics



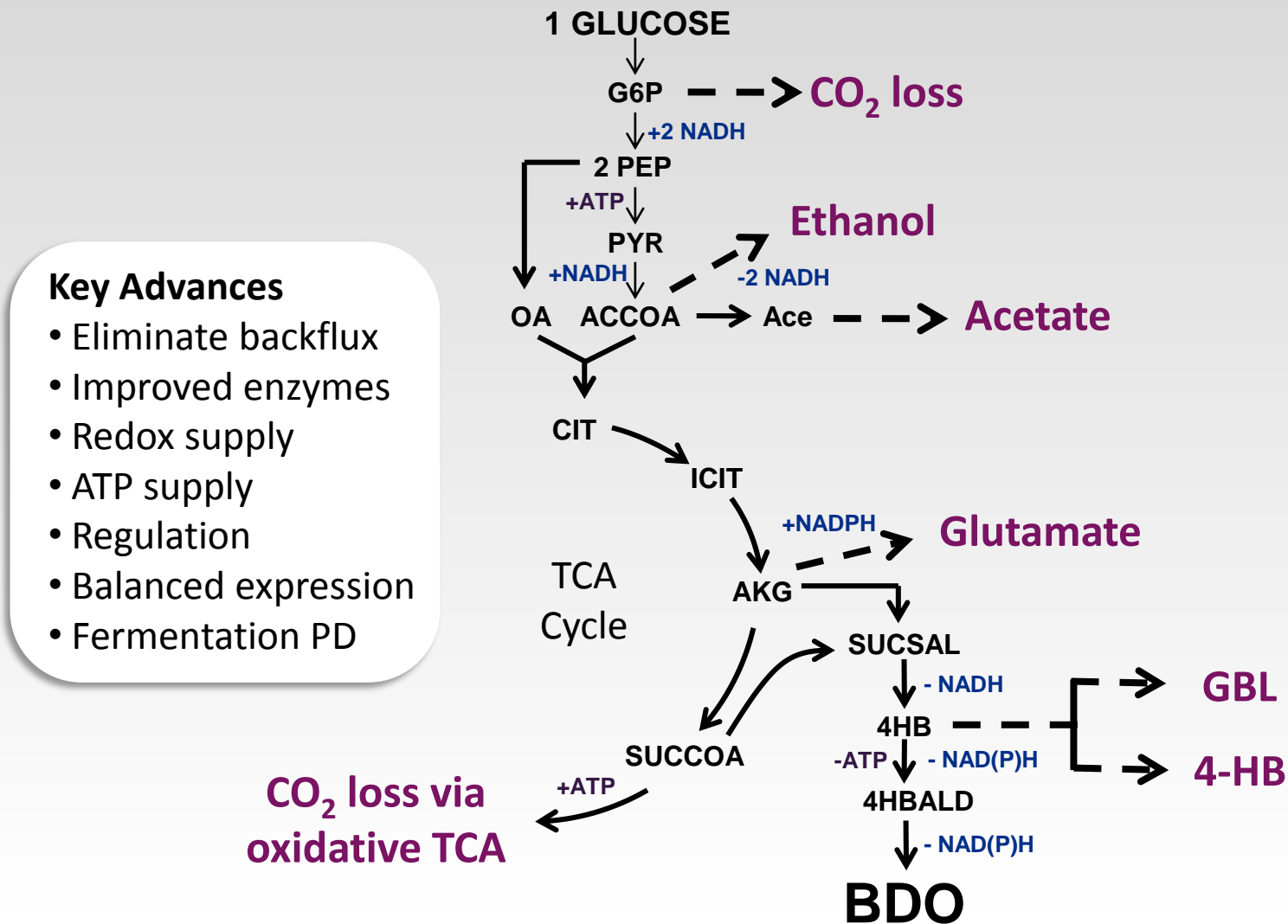
Genomatica's Systems-Based Strain Engineering



BDO Strain Engineering Progress

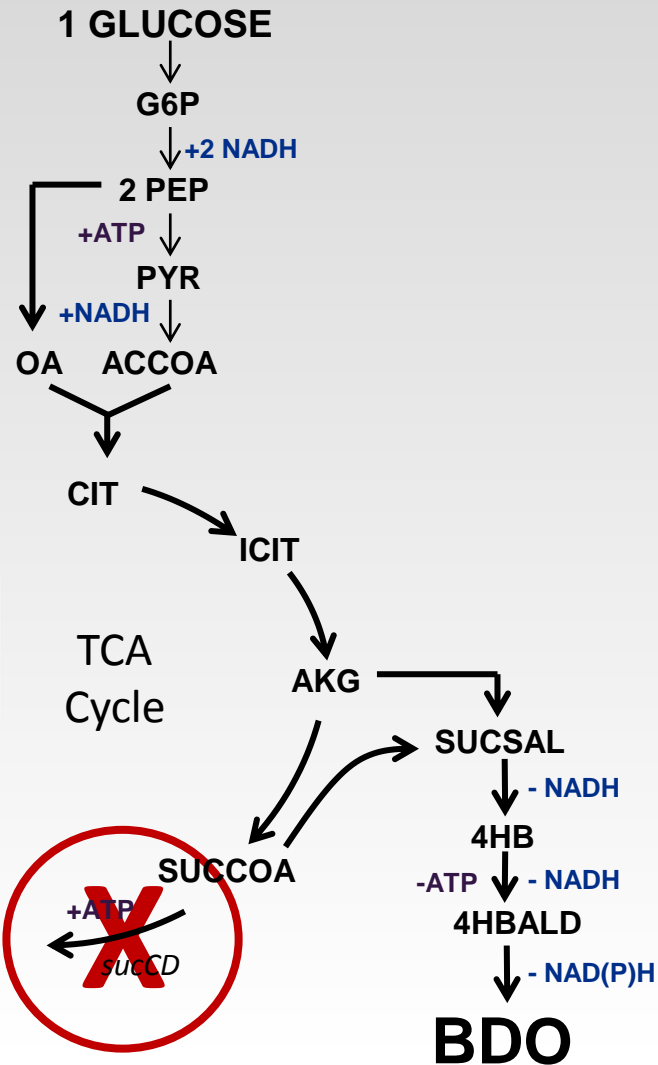
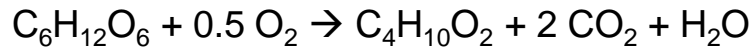


Increasing Rate and Lowering By-products



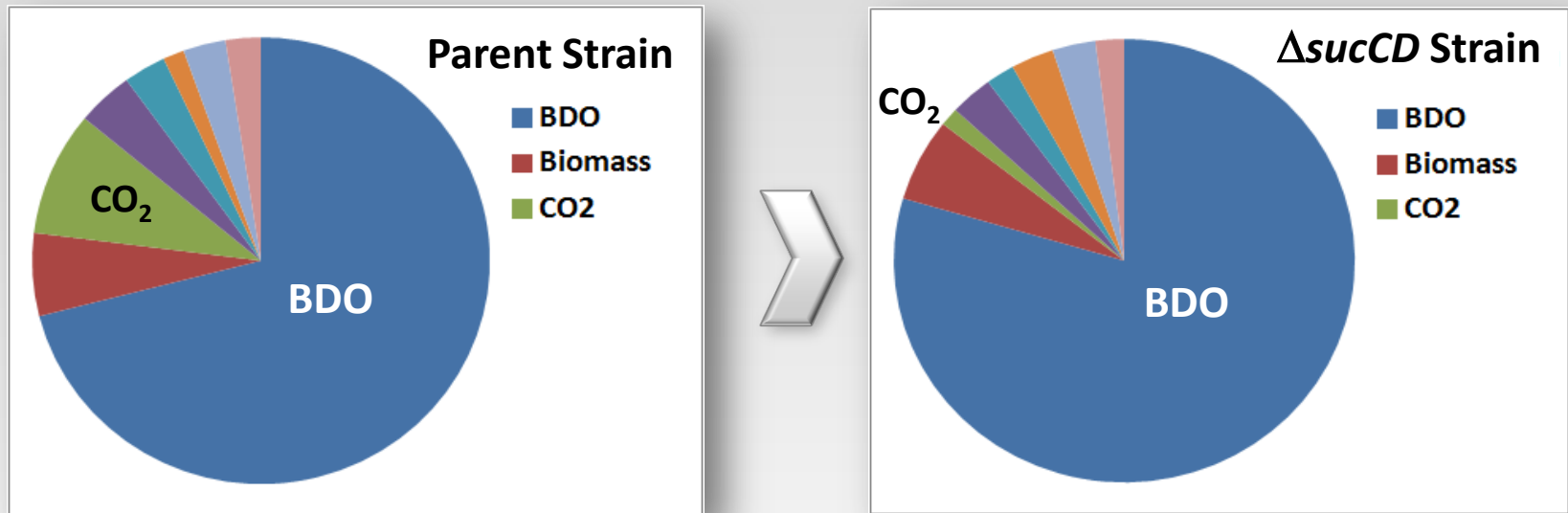
More metabolic steps in a pathway increases avenues for by-products

Lowering By-Products: Excess CO₂



- Excess CO₂ from complete TCA
- Delete *sucCD* gene – lose 1 ATP
- Eliminated “futile” energy drains

Lowering Excess CO₂ via *sucCD* Deletion



ΔsucCD Strain: higher BDO, much lower CO₂

BDO Scale-up and Commercialization

Joint Development Partnership with Tate & Lyle

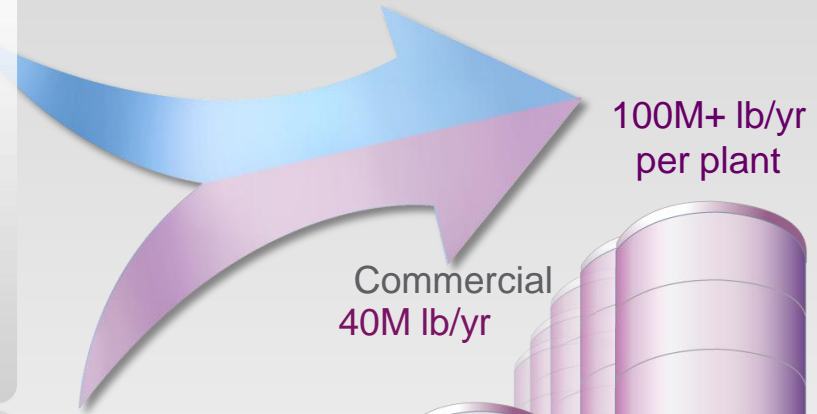


TATE & LYLE

- \$4.3B per year
- Operates four corn wet mills
- JV with DuPont: PDO, 100-140M lb/yr

Genomatica taking proven path:

- ✓ Same base organism
- ✓ Same scale-up factor
- ✓ Similar chemical
- ✓ Similar cost model

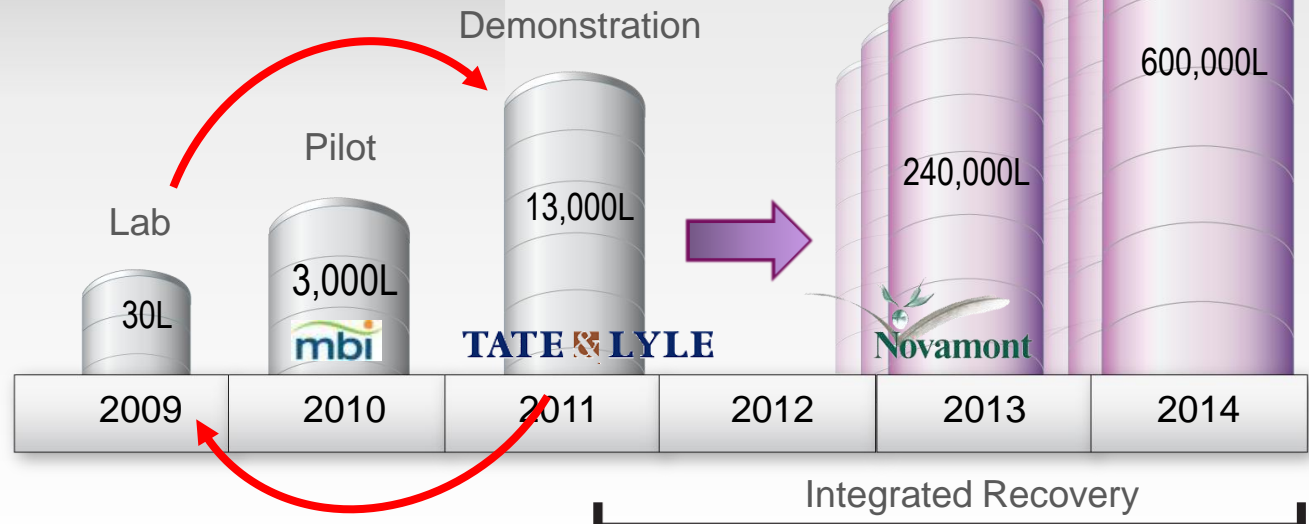


100M+ lb/yr
per plant

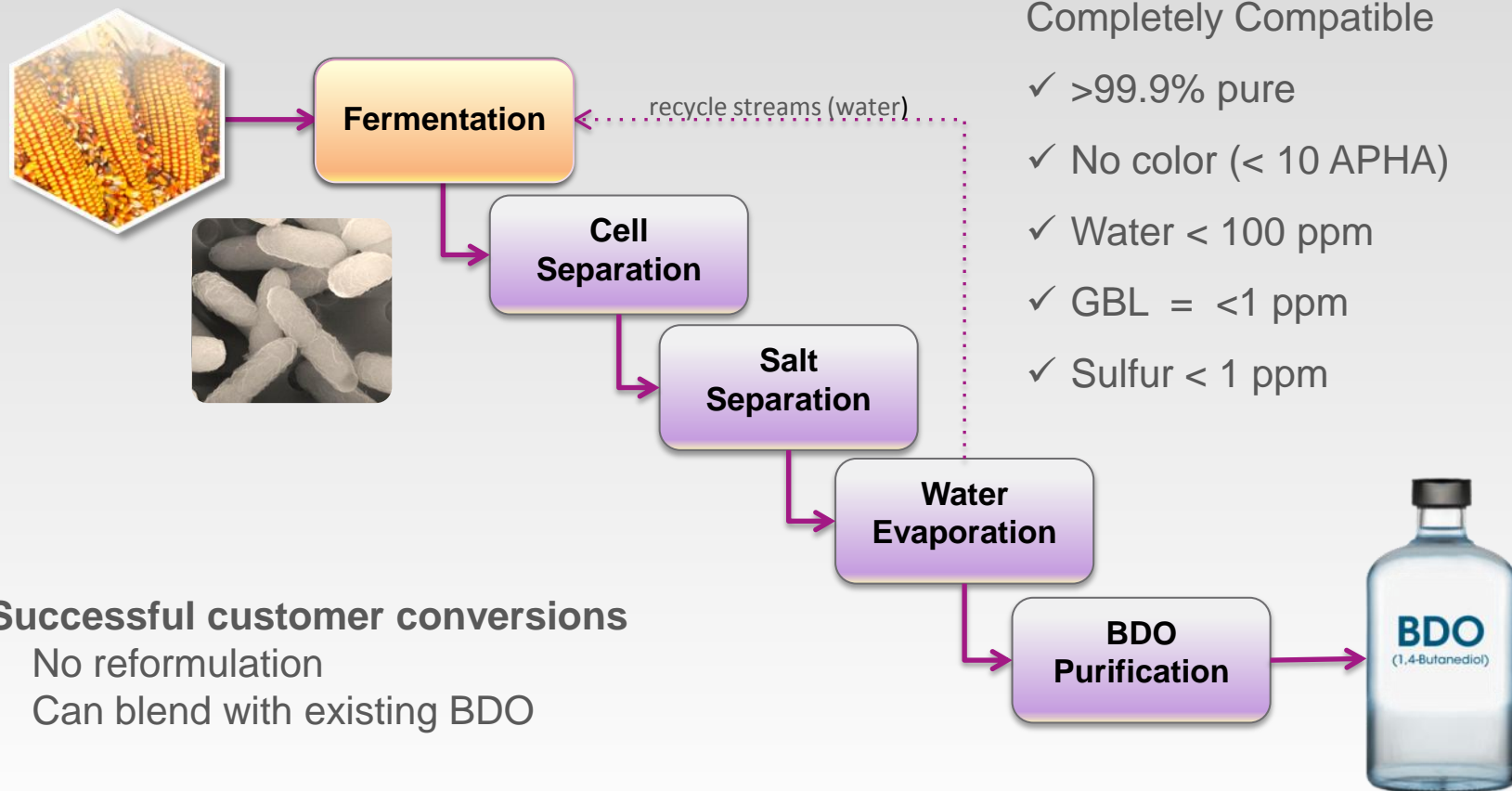
Commercial
40M lb/yr



13,000 L fermentor
in Decatur demo plant



Complete Process Technology for Bio-BDO



High Purity Bio-BDO

Completely Compatible

✓ >99.9% pure

✓ No color (< 10 APHA)

✓ Water < 100 ppm

✓ GBL = <1 ppm

✓ Sulfur < 1 ppm

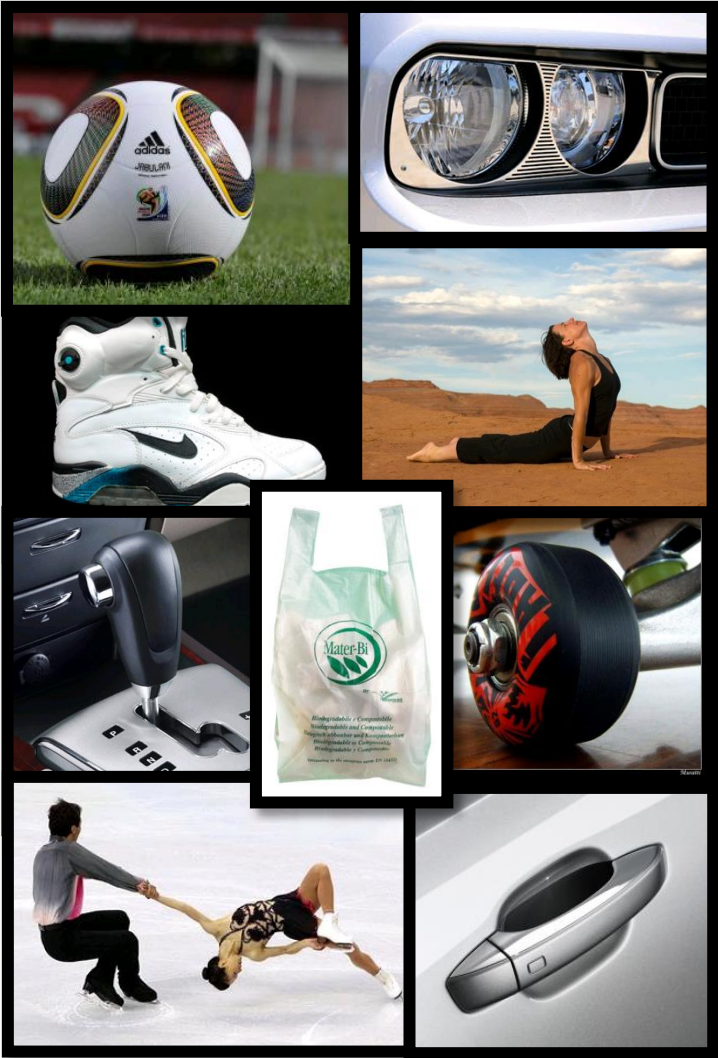
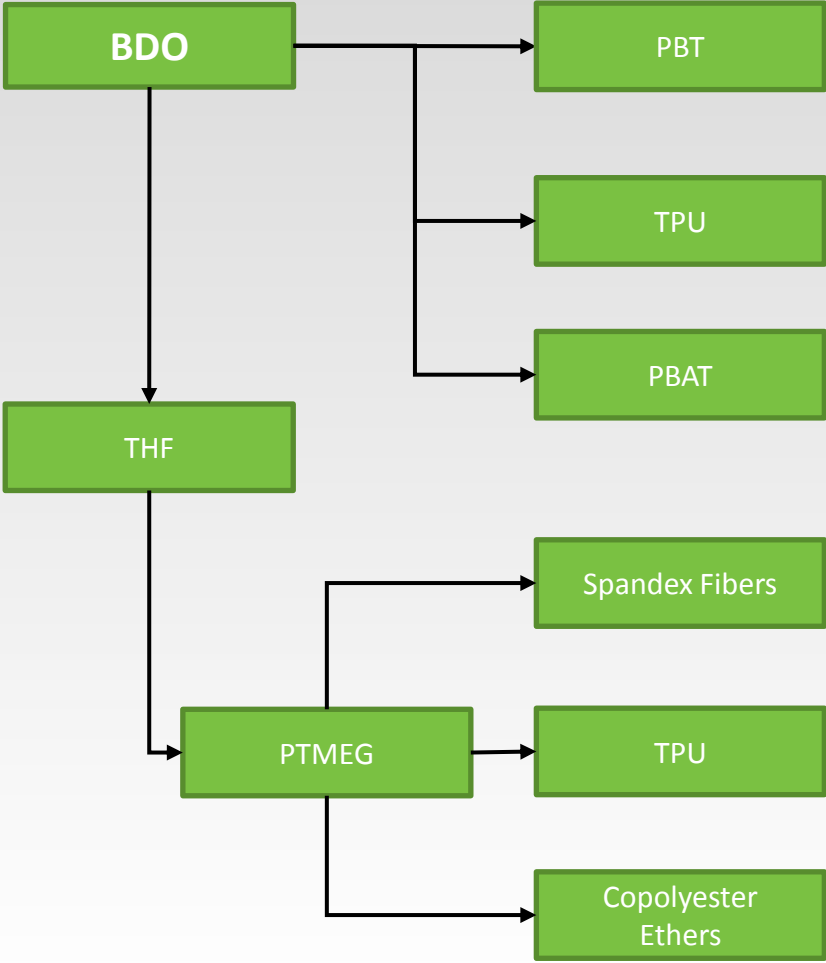
Successful customer conversions

- No reformulation
- Can blend with existing BDO

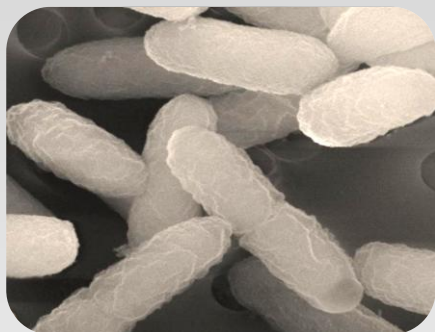
Shipping tons at a time...



BDO Applications



Bio-BDO[®] Becoming a Commercial Reality

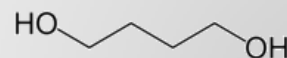


PBT from
Bio-BDO

High quality Bio-BDO converted into all downstream derivatives by partners

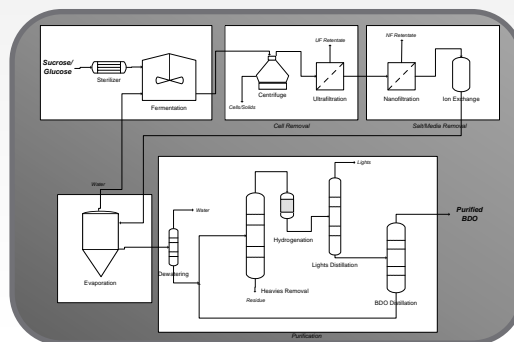
2008

first production of 1,4-BDO
from carbohydrates



2013

commercial scale
production (40M lbs/yr)



Novamont: BDO in Europe

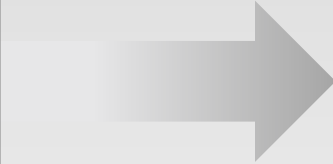


- Joint venture for commercial-scale production of BDO
- Novamont financing 100% of construction costs (~\$50M)
- Begin production 2013
- 40 million pounds per year
- Committed to purchase 100% of production for captive use
- Option for second plant

Biomass-to-Chemicals



Dedicated feedstock



Bio-based Chemicals



Biomass sugars offer:
Lower price volatility
Lower cost potential
No food vs chemicals

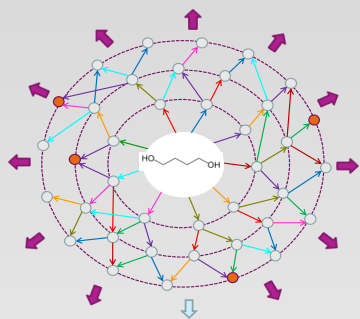
Biomass-to-BDO

\$5 M DOE Grant

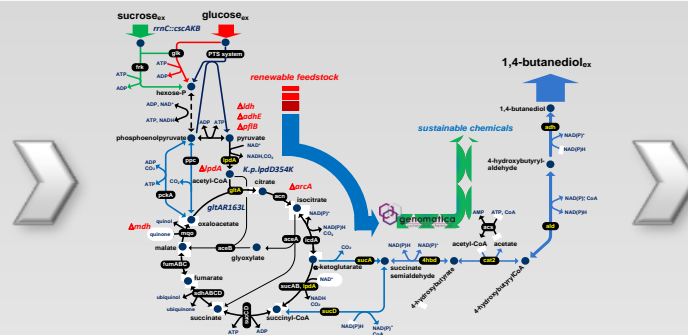


Currently producing BDO in *E. coli* from PROESA biomass sugars at ~85% of performance vs dextrose

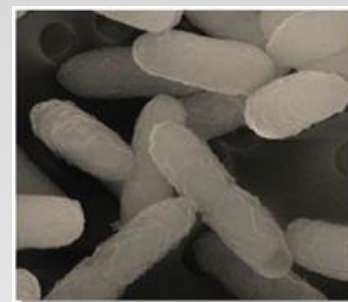
Development of a Robust BDO Production Strain



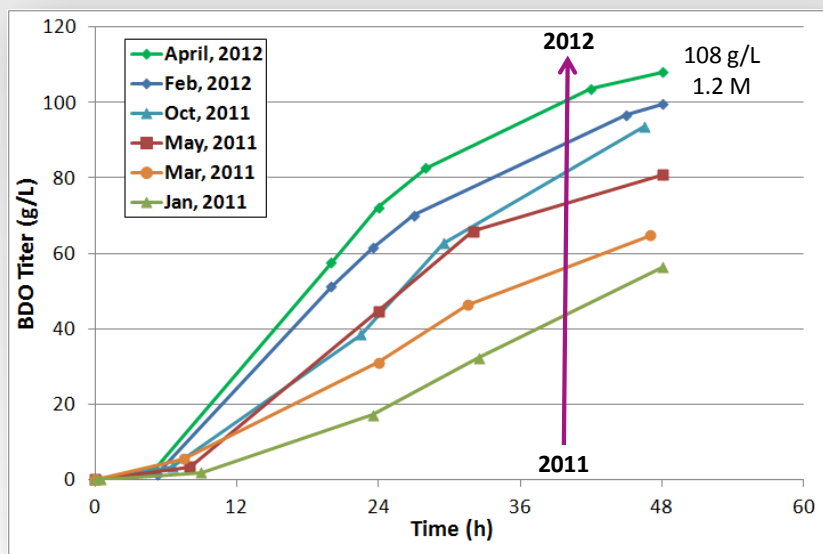
Pathway Identification and Engineering



Strain Design and Metabolic Engineering



Commercial Strain for BDO Production



- ✓ Systems-based approach
- ✓ Over 35 genetic manipulations
- ✓ BDO from dextrose, sucrose, biomass
- ✓ Achieved commercial TRY targets
- ✓ Process scale-up to 13,000 L (1 ton/wk)

Genomatica BDO Team - Past and Present

Molecular Biology

Harry Yim
Steve Van Dien
Bob Haselbeck
John Trawick
Wei Niu
Jeff Boldt
Laura Peiffer
Eric Van Name
Chris Wilson
Stephanie Culler
Brandon Chen
Kevin Hoff
Ewa Lis
Fannie Chau
Hongmei He
Shawn Bachan
Jingyi Li
Luis Reyes

Microbiology

Catherine Pujol-
Baxley
Jazell Estadilla
Jesse Wooton
Jabus Tyerman
Jonathan Moore
Lars Knutstad
Paul Handke
Jonathan Joaquin

Analytical Sciences

Julia Khandurina
Rosary Stephen
Lucy Zhao
Ahmed Alanjary
Blanca Ruvalcaba
Rainer Wagester
Korki Miller

Enzymology

Brian Steer
Stefan Andrae
Cara Tracewell
Mike Kuchinskas
Wayne Liu
Brian Kinley
Amit Shah
Jacqueline Fritz

Process Engineering

Joe Kuterbach
Michael Japs
Janardhan Garikipati
Fasil Tadesse
Ben Adelstein
Rachel Pacheco
Daric Simonis
Arvind Kaul
Ishmael Sonico

Computational

Tony Burgard
Priti Pharkya
Robin Osterhout
Jun Sun
Tae Hoon Yang
Wyming "Lee" Pang

Fermentation

Dan Beacom
Sy Teisan
Brett Schreyer
Laurie Romag
Joseph Woodcock
Don Miller
Gian Oddone
Amruta Bedekar
Rebecca Bratcher
Jason Crater
Akhila Raya
Alex Navarro

SAB

Bernhard Palsson
Sang Yup Lee
Jens Nielsen
George Church
Lee Hood
Harvey Blanch
Bernhard Hauer

Christophe Schilling, CEO Bill Baum, CBO
Mark Burk, CTO Jeff Lievens, EVP, Process Development

Questions?



Thank you