

# Writing the Future

**EMILY LEPROUST, PH.D., CEO and CO-FOUNDER** 

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# DNA is Changing the World



### Synthetic DNA Is the Future of Everything







**Food** Food Security



Therapeutics
Health



**Diagnostic**Precision Medicine



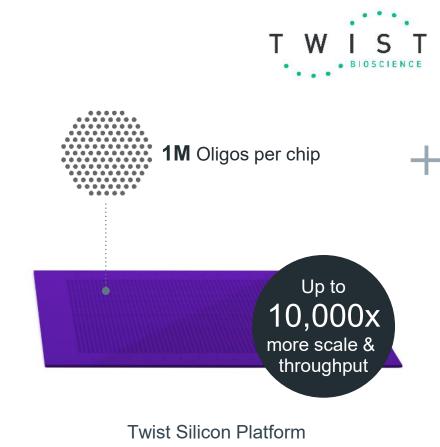
**Data Storage**Preserving Heritage



### **Twist DNA Writing on Silicon Platform**

# **Everyone Else** 1 Oligo per well





can make 9,600 genes

### State of the art commercial infrastructure

Proprietary software

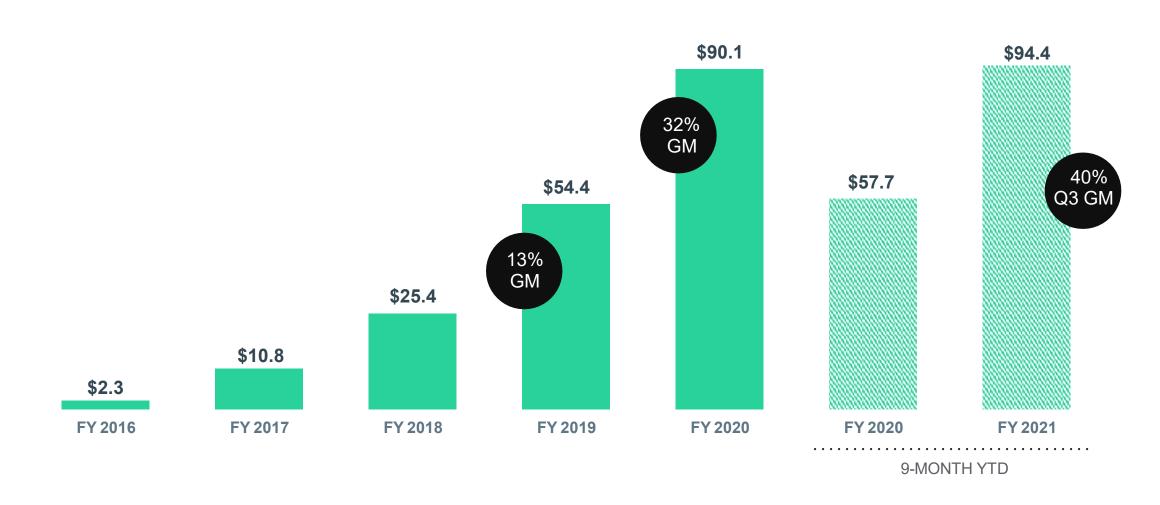
Robotics

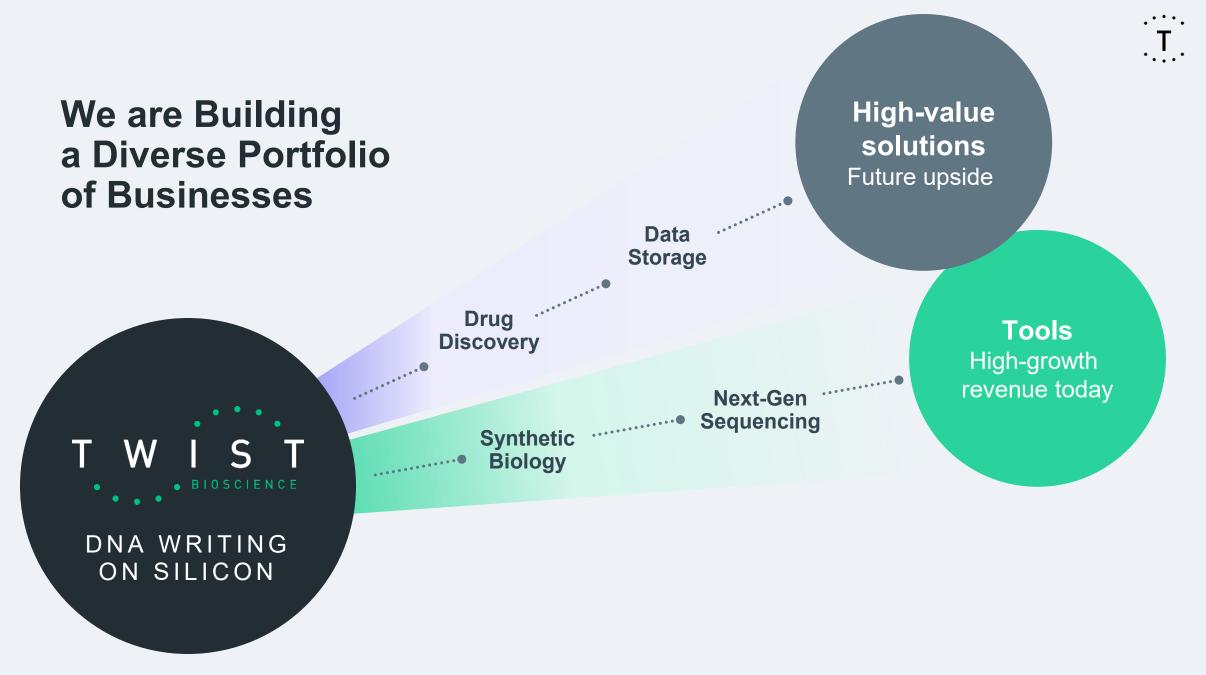
Integrated ecommerce platform

Manufacturing execution system



### **Strong Revenue Growth and Increasing Gross Margin**







### Synthetic Bio: Largest Selection of DNA Offered



#### Genes

Clonal
Non-clonal fragments
Clonal-ready gene
fragments



### **Oligo Pools**

sgRNA



### **Variant Libraries**

Site saturation Combinatorial



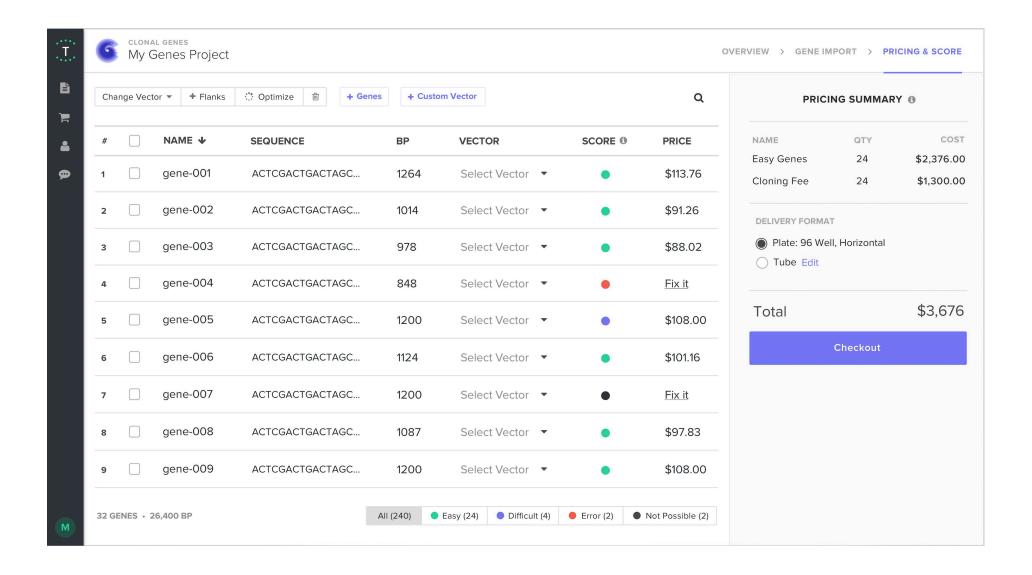
### Pharma / Biotech Solutions

DNA preps IgG



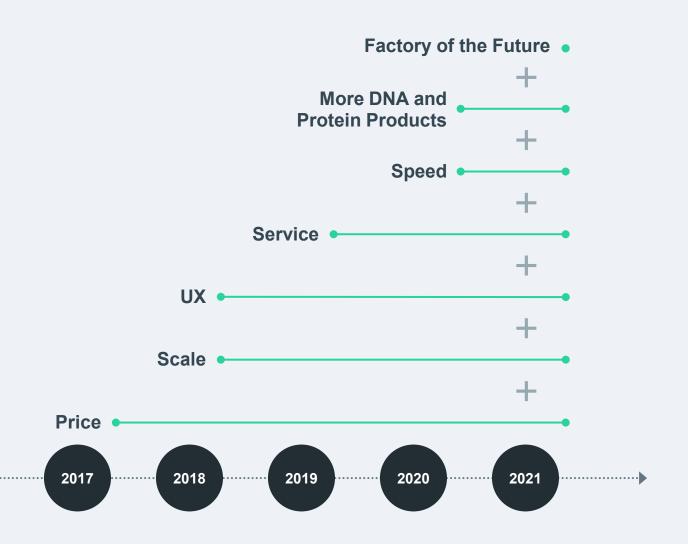


### **Innovative Buying Experience**



### **T**

### **Synthetic Bio: Why We Win**



### We Deliver

- High-quality DNA
- Competitive turnaround times
- Affordability
- High throughput
- Unique customer experience
- Innovative products and solutions



Twist's very high-throughput platform allowed us to quickly and efficiently examine thousands of possible antibodies in order to select the best results faster than ever before.

Robert Carnahan, Associate Director, Vanderbilt Vaccine Center

### **Synthetic Bio: Scratching the Surface**

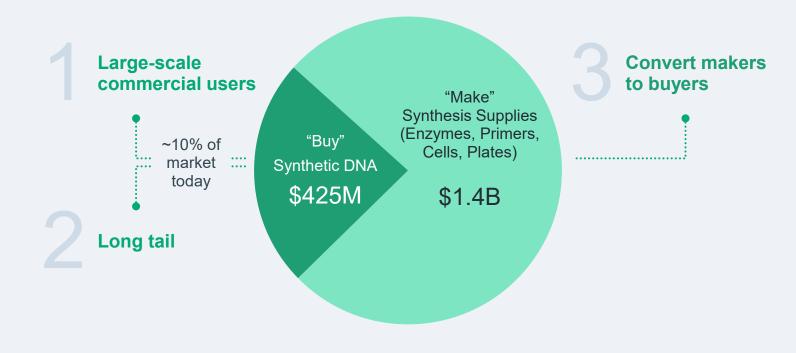
**FY20 Proof Points** 

1,590 **Customers** 

339K **Genes shipped** 

99% Orders via ecommerce **Strategy to Capture Large Market Potential** 

\$1.8B / year





### Synthetic Bio: Roadmap to Expand Our Leadership

**Factory of the Clonal Ready Business to Business** IgG **Future (2022) Gene Fragments** Integration **New Portland facility to:** Serving long tail Software to accelerate Commercial of the market ramp in pharma order processing **Double capacity** Reduce turnaround time **Target new segments** 



### **NGS:** Broad Offering to Meet Expanding Applications

### **Today**



**Human Core Exome** 



**Fixed Panels** 



**Custom Panels** 



**Library Preparation** 



Reagents and Kits



Synthetic Viral Controls



Targeted Methylation



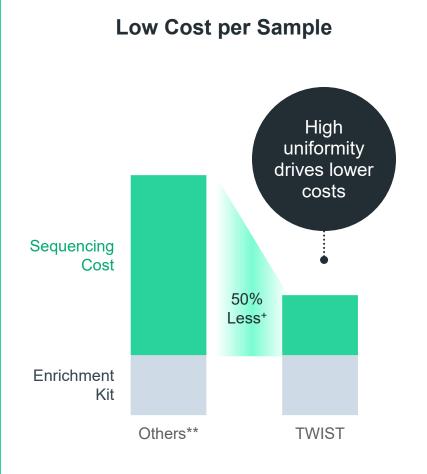
SARS-CoV-2 NGS Assay

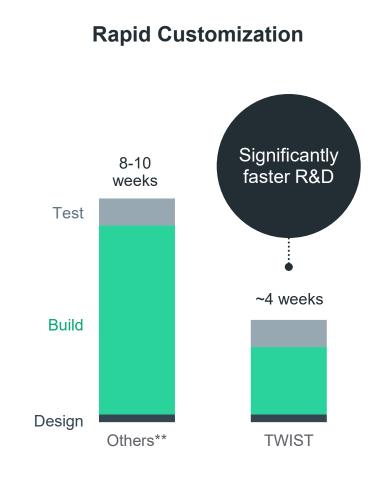
### **Targeted Sequencing**

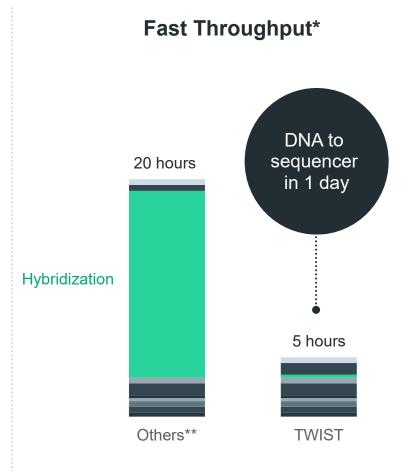
is powering new applications

- Liquid biopsy
- Rare disease
- Oncology
- Population genetics
- Infectious diseases

## NGS: Why We Win







<sup>\*</sup> Based on customer testimonial and Twist's experience

<sup>\*</sup>Includes pooling, pre-hybridization, hybridization, binding, wash steps, amplification, purification, target environment QC, and NGS prep

<sup>\*\*</sup>Illustrative models based on Twist's knowledge of competing technologies.



### **NGS:** Investing in Growth

**FY20 Proof Points** 

>1,000

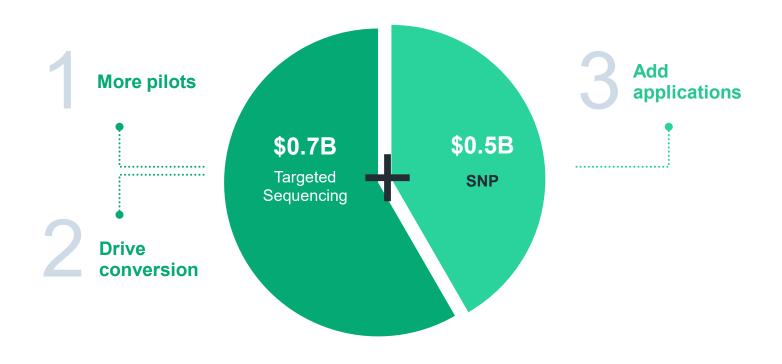
**Customers shipped** 

55 in production

13
OEM partners

**3** SNP microarray conversion

### **Strategy for Expanding Market to >\$1B**



### Т

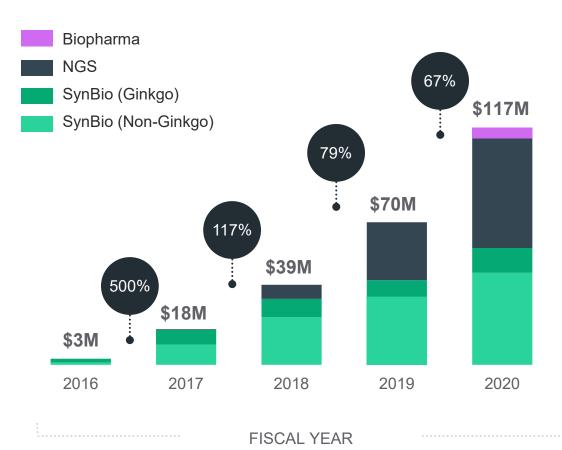
### **Core Business Growth**

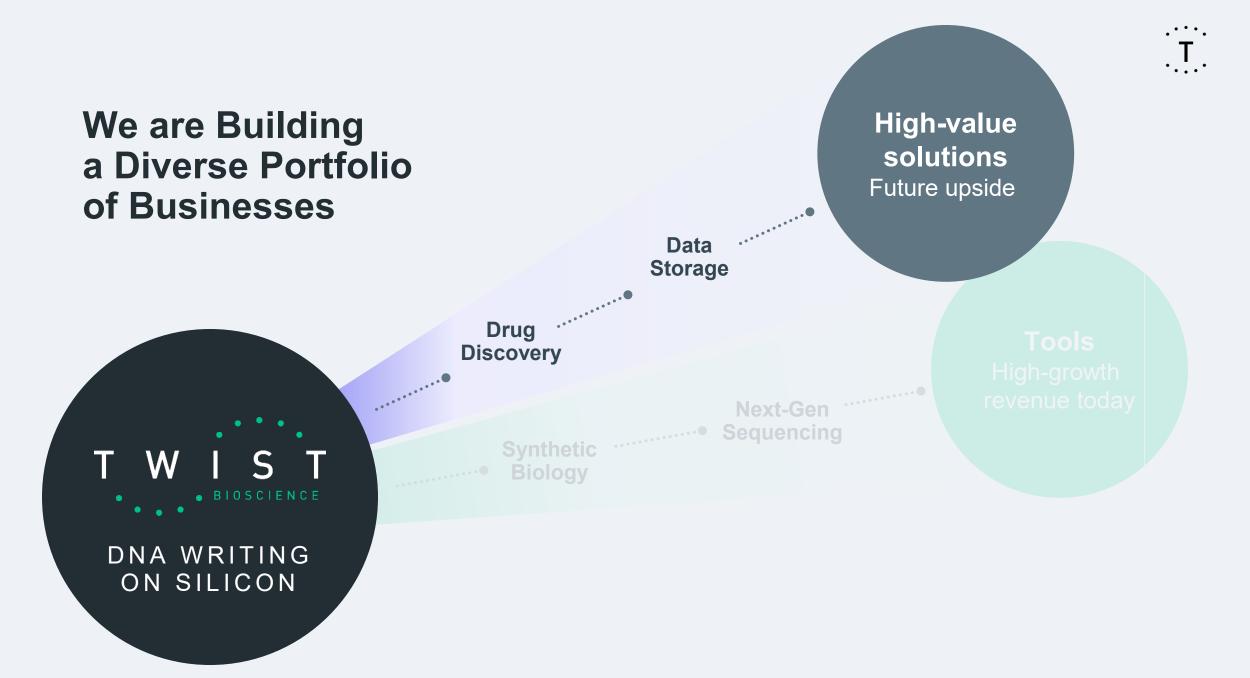
### **Total Customers**

(SynBio & NGS)



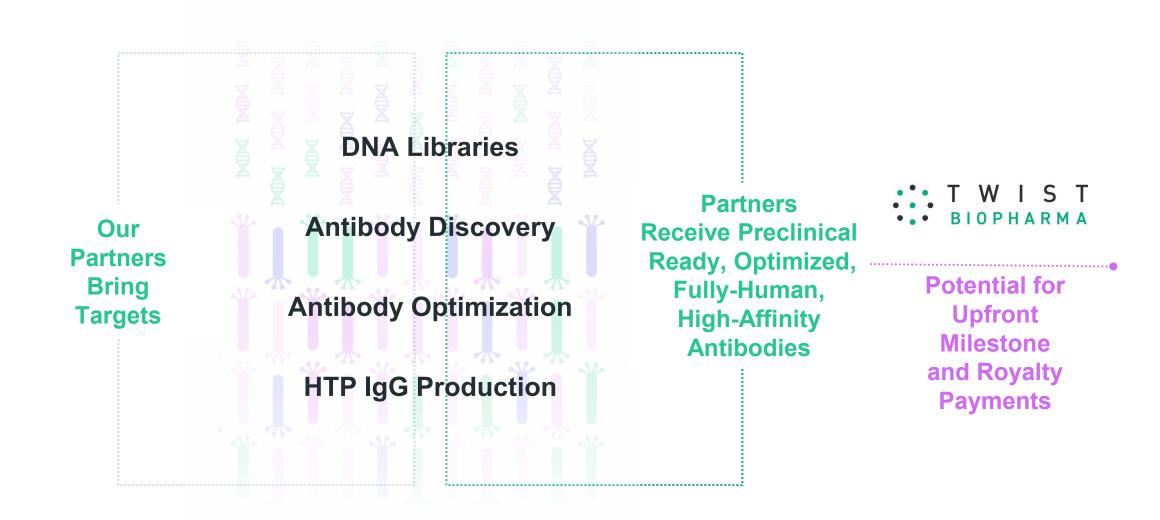
### **Orders**







### **Biopharma: Validated and Expanding Offering**



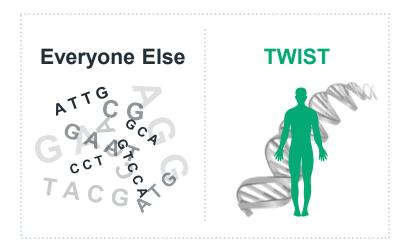


library

### **Biopharma: Why We Win**

fully validated libraries







#### **Broad "DNA" Buffet**

Allowing us more shots on goal with more diversity

# **Every Sequence Always Explicitly Synthesized**

Never random —
All our DNA is human-derived and higher quality, following human repertoire rules

# Automation and Miniaturization

Most of our process, including library production, screening, reformatting, affinity and functional testing increasing efficiency and speed



### **Biopharma Partnerships by the Numbers**

# Broad

**Disease Indications** 

Cancer, Neurology, Immuno-oncology, Infectious Disease, Canine / Feline, Other

Varied

**Modalities** 

mAbs, Bispecific Antibodies, VHH, ADC, Protein Engineering, More

Completed Programs

31 Partners

**43** Active Programs

26

Milestones/ Royalties



### **Biopharma:** Pipeline of Preclinical-Ready Functional Monoclonal Antibodies



TARGET	INDICATION
GLP1R	Diabetes & rare metabolic diseases
PD1	Immuno-oncology
TIGIT	Immuno-oncology
CD3	Immuno-oncology
ADORA2A	Immuno-oncology
CXCR4	Oncology
ACE2	COVID-19
SARS-CoV-2 S1	COVID-19
SARS-CoV-2 N	COVID-19

Leveraging Twist platform to discover functional monoclonal antibodies against high value targets

Pursuing opportunities to maximize value for these assets



### **Biopharma: Immuno-oncology Target ADORA2A**

# **Checkpoint Inhibitor Market** is **Growing Rapidly**

This market is expected to reach \$40 billion by 2025

### Significant Unmet Needs Remain

Only ~20-30% of eligible cancer patients benefit from first generation checkpoint inhibitors

# Large Opportunity for Novel Immunotherapies

Twist can rapidly discover and optimize antibody-based immunotherapy leads across oncology indications



Adenosine pathway is a **master checkpoint** in the tumor microenvironment

Highly expressed in:

Lung
Colorectal
Prostate cancer

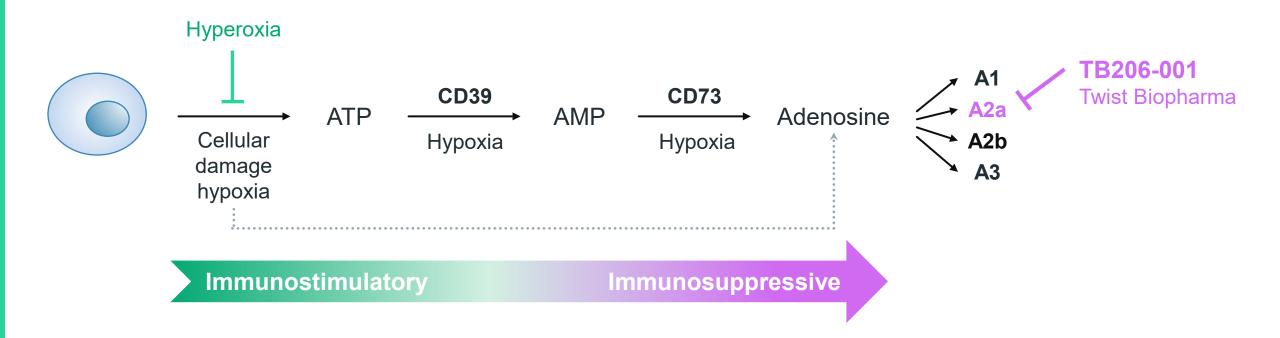
**A2a antibody antagonist** has the potential to have high potency and specificity, improved dosing, and low CNS permeability

Twist's TB206-001 is a Potent A2a Antibody Antagonist Candidate





### **Biopharma:** Adenosine Pathway Targets Master Checkpoint in TME

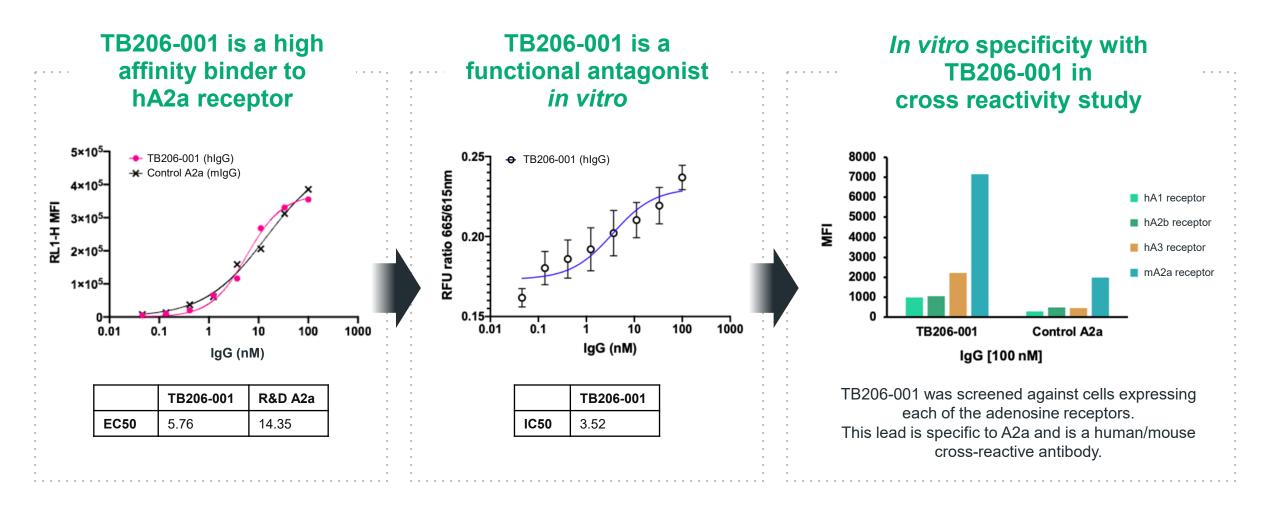


Tumors can evade immune responses from first generation checkpoint inhibitors by usurping the adenosine pathway

Incomplete inhibition of upstream enzymes in this pathway would still produce adenosine and thus drive immunosuppressive signaling TB206-001 directly targets the A2a receptor, the primary adenosine receptor on immune cells, to block immunosuppression



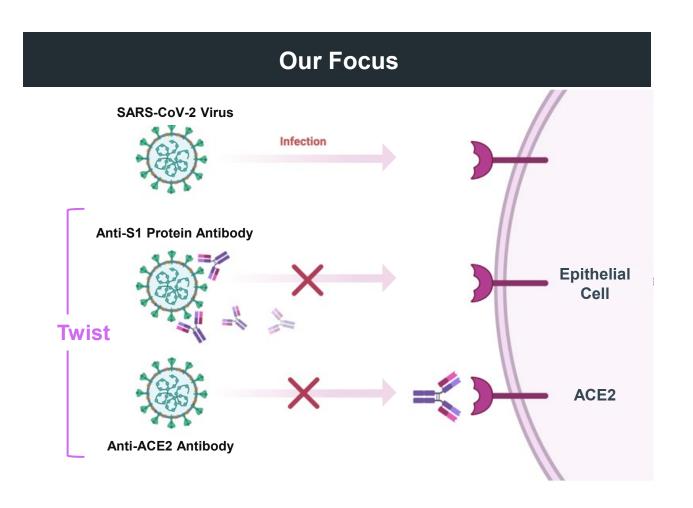
### **Biopharma:** TB206-001 Potent A2a Antibody Antagonist Candidate



We are optimizing and developing this preclinical lead candidate (TB206-001)



### **Biopharma:** Rapid Discovery of SARS CoV-2 Antibodies



# **Novel Therapeutic Antibody Leads**

IgG (TB181-8, 28, 36) VHH (TB201-202)

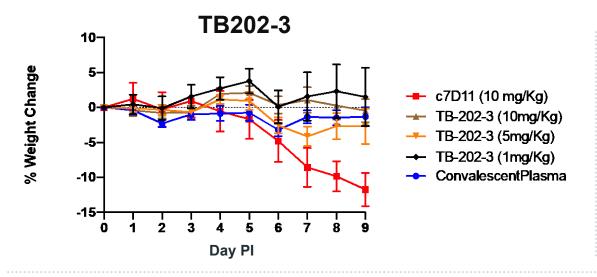
# Well-Characterized and Well-Validated

High-affinity and unbiased, with extensive pseudovirus and live virus, preclinical animal data



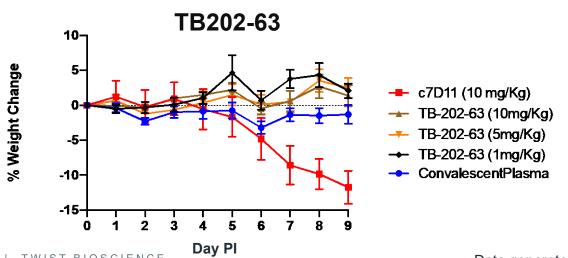
### Biopharma: VHH Single Domain Leads (TB202-3, 63) and IgG Lead (TB181-36) Show Potent In Vivo Activity in Hamster Model

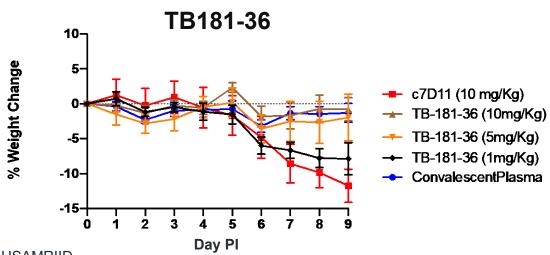
### TB202-3 and TB202-63 Protect Against Weight Loss at the Lowest Dose of 1 mg/kg





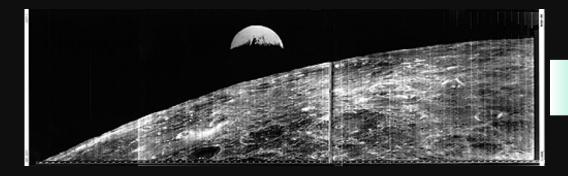
c7D11 = Negative control mAb ConvalescentPlasma = Positive control







Recovered from Lunar Orbiter 1 Tapes







### **Data Storage: How It Works**

Coding

$$\begin{array}{ccc}
00 & \longrightarrow & A \\
01 & \longrightarrow & G \\
10 & \longrightarrow & C \\
11 & \longrightarrow & T
\end{array}$$

Synthesis



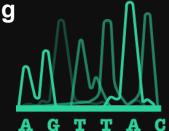
**?** Storage



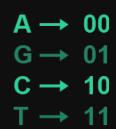
Retrieval



**5** Sequencing

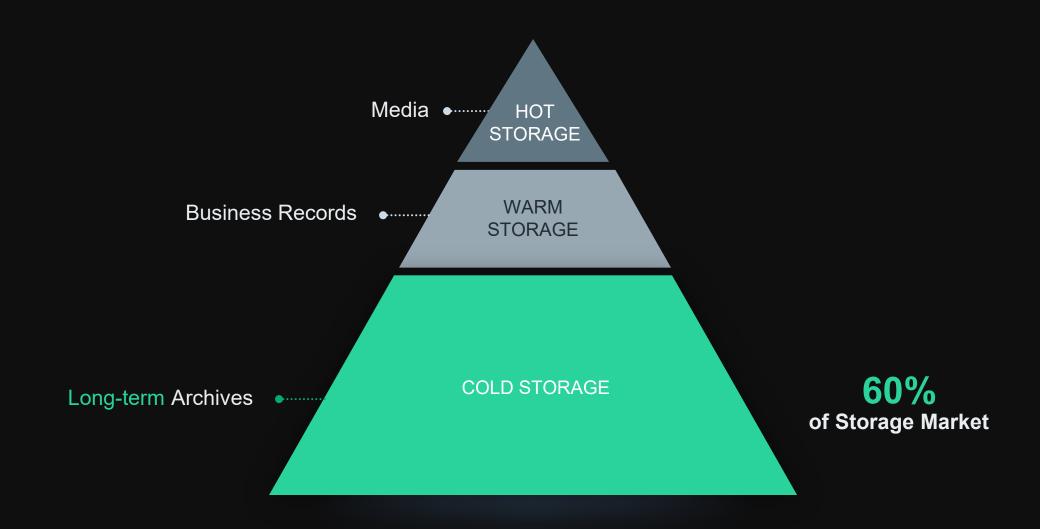


6 Decoding





### **Data Storage:** \$35B Large Market Opportunity

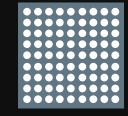




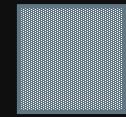
### **Data Storage: Progression**













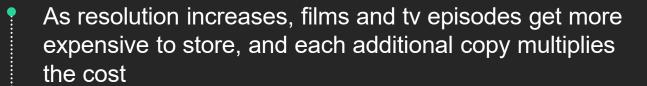
1 Micron Integrated Chip Proof-of-concept

Alpha Integrated Chip Initial Customers

Beta
Integrated Chip
Grow Customer Base

### **DNA Data Storage:**

Cost for Digital Film Preservation



DNA copies are almost free due to the PCR process

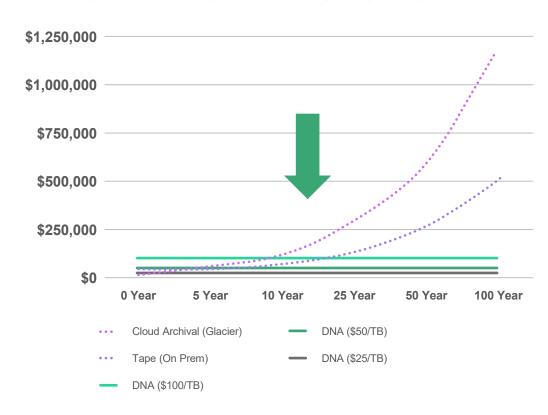
Cost of storage on Tape/Cloud will grow over time due to required data migration while the cost of DNA will remain flat

When DNA reaches \$100/TB it will be more economical to store any data on DNA if it needs to be preserved for 15+ years



#### **TCO COMPARISON**

TCO Analysis for Cloud Archival (Glacier Deep Archive), Tape (On-Prem) and DNA for 1 PB





### **DNA Data Storage Alliance**

# 34 Industry Leaders Joining Forces to Advance DNA Data Storage

Twist Bioscience
Microsoft

Illumina Western Digital

**DNA Script** 

Catalog

**IMEC** 

Molecular Assemblies

PFU, a Fujitsu company

Ansa Biotechnologies

**EPFL** 

Claude Nobs Foundation

University of Washington

ETH Zurich

Iridia

Quantum

Generate industry roadmap

Develop use cases

Educate for broader
awareness and adoption

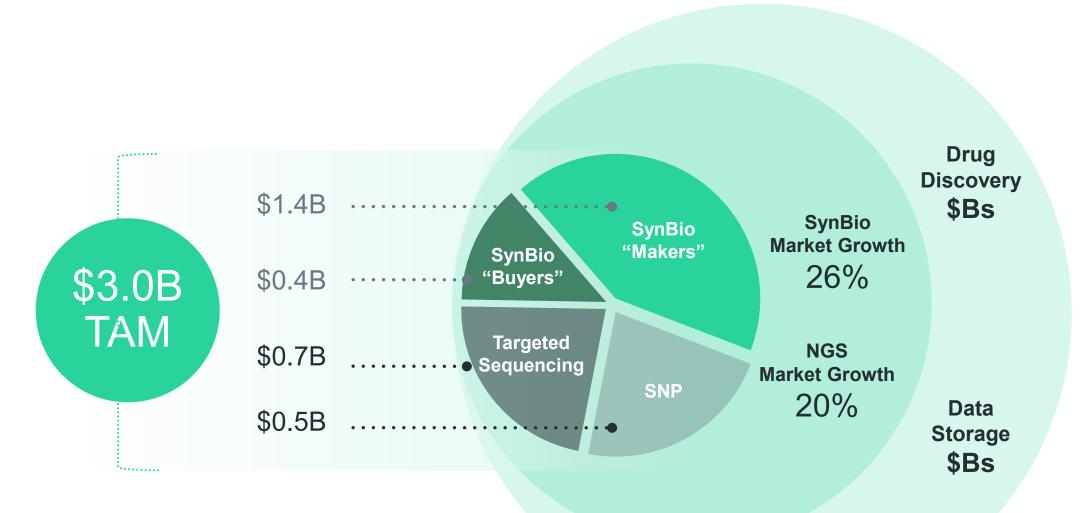
by 2024

30% of digital business will mandate DNA storage trials

- GARTNER, OCT. 2020

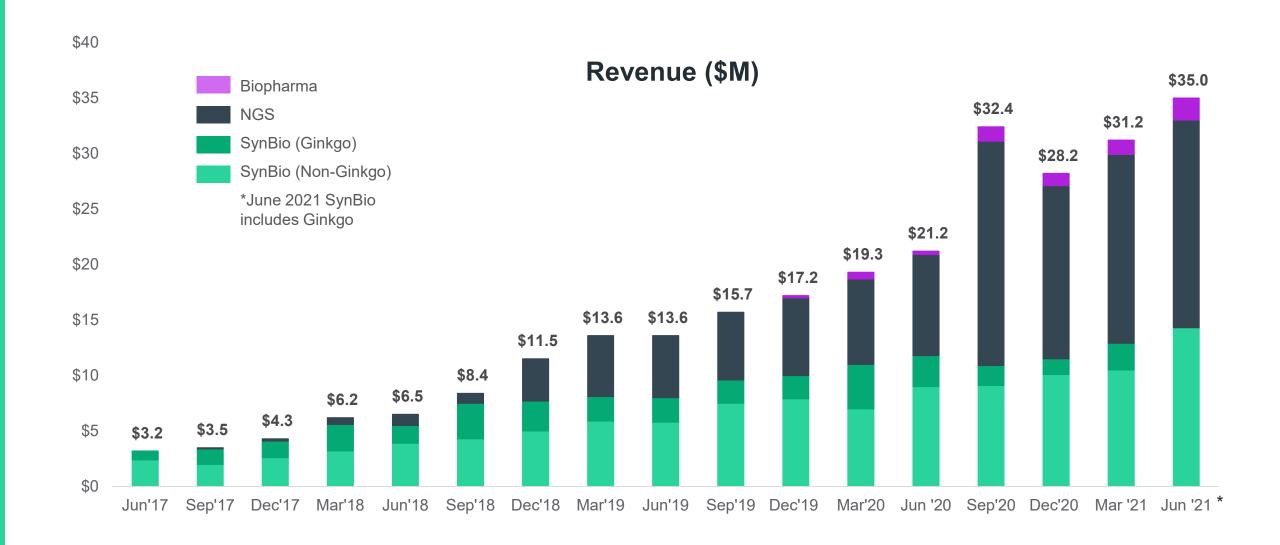


### **Large and Expanding Addressable Market**





### **Strong Revenue Growth**



### Making our Vision a Reality Near Term Objectives

### **Synbio**

- Continued growth and diversification of revenue stream
- Ramping pharma-focused products, including DNA preps and IgG
- ✓ Launch of clonal ready gene fragments
- B2B solutions to allow capture of specific multi-site institutions
- Significant investment in "Factory of the Future"
- Expand OEM strategy

### NGS

- ✓ Continued revenue growth and customer ramping production
- ✓ Full launch of methylation solution
- Relaunch library prep kit
- Continued conversion of SNP Microarrays to NGS + sequencing
- Expand OEM strategy

### Biopharma

- Additional partnerships to expand technology base and generate revenue
- Additional internal pipeline of antibodies, pursuing out licensing opportunities by mid-2022

### **Data Storage**

- Integrated 1-micron pitch chip as proof-of-concept
- Begin design for alpha integrated chip
- Execute on agreement with **IARPA**
- Work with DNA Data Storage Alliance to educate and prepare market

Expand and Accelerate Internal Efforts and Inorganic Investment to Help Ensure Long-term Leadership



# Writing the Future

Platform for writing DNA on silicon

Large, growing markets

**Differentiated** value proposition

Portfolio of high growth businesses

Validated business models

High revenue growth

Track record of execution and innovation



# Writing the Future

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