



**C1 Protein Production Platform** 

Addressing Vaccine & Drug Shortfalls Through Better Science
September 2021

## Safe Harbor Regarding Forward-looking Statements

Certain statements contained in this presentation are forward-looking statements within the meaning of Section 27A of the Securities Act of 1933 and Section 21E of the Securities Exchange Act of 1934, including those regarding Dyadic's expectations, intentions, strategies and beliefs pertaining to future events or future financial performance. Actual events or results may differ materially from those in the forward-looking statements as a result of various important factors, including those described in Dyadic's most recent filings with the SEC. Undue reliance should not be placed on the forward-looking statements in this presentation, which are based on information available to us on the date hereof. Dyadic assumes no obligation to update publicly any such forward-looking statements, whether as a result of new information, future events or otherwise. For a more complete description of the risks that could cause our actual results to differ from our current expectations, please see the section entitled "Risk Factors" in Dyadic's annual reports on Form 10-K and quarterly reports on Form 10-Q filed with the SEC, as such factors may be updated from time to time in Dyadic's periodic filings with the SEC, which are accessible on the SEC's website and at <a href="https://www.dyadic.com">www.dyadic.com</a>



## **Our Mission, Transforming Biomanufacturing**

"To improve how we feed1, fuel1, and heal the world by utilizing modern biotechnology to revolutionize science, medicine, agriculture<sup>1</sup>, and engineering. To provide a costeffective solution to increase biomanufacturing outputs and satisfy the growing demand for protein production and unmet needs for affordable biologic drugs, vaccines and other biologic products and processes."





1 Dyadic has achieved certain ports of the above "Mission" through its twenty plus years of experience in industrial biotech. Currently, we are primarily focused on animal and human health applications while opportunistically exploring if and how we may enter/re-enter certain industrial biotechnology applications.

## **Dyadic At-a-Glance**

## Re-engineering GMP Vaccine & Drug Production

- Proprietary & patented biologic and vaccine production platform (the "C1 Platform"), designed to bring biologic vaccines and drugs to market faster, in greater volumes, at lower cost
- The C1 platform is de-risked, with a safe and long track record in industrial protein manufacturing

Value Creation Through Technology Licensing, Co-Development Partnering and Wholly Owned Product Development



Market Capitalization	\$168.5 million (as of 09/07/2021)
Cash & Investment-grade securities, including accrued interest	\$25.8 million (as of 6/30/2021)
Shares Outstanding	~ 28.1 million (as of 8/11/2021)
Debt and Warrants	None
Insider Ownership	~30%
2021 R&D Revenue 6 Months	\$1.4 million (YoY 66% increase)



NASDAQ DYAI



**HEADQUARTERS**Jupiter, Florida



HISTORY Founded In 1979



**RESEARCH LOCATIONS**Finland, Spain, US & Others

## **Investment Highlights**

Solid Financial Position with \$25.8 million in cash and investment securities, no debt<sup>1</sup> Harnessing the power of C1 fungi to transform manufacturing of vaccines and other therapeutics

**Versatile Platform** 

C1 protein production is a platform technology, with the potential to disrupt the manufacturing of sub-unit vaccines, therapeutics, enzymes and other peptide, protein and glycoprotein-based products

**Strong Competitive Advantage & IP Position** 

Robust scientific data demonstrating high productivity, stability, and purity.

DuPont grant back of former Dyadic patents & five provisional/patent applications

**Large Addressable Market Ready for Disruption** 

Vaccine and therapeutic manufacturing for humans and animals remains a broad addressable market with many shots on goal and ready for disruption

**Global Strategic Partnerships** 

Funded partnerships with top-tier global biological R&D organizations, human and animal health pharmaceutical companies, as well as governmental and private agencies

**Experienced Leadership** 

Highly experienced and energized professional management team and world-class Board of Directors & Advisors

(1) As of June 30, 2021

## Management & Directors With Track Record Of Success/Value Creation

Highly Energized Management Team With Deep Industry Expertise & Products In Market Active Board with Decades of Relevant Experience in Biomanufacturing



**DYADIC**® Mark Emalfarb Founder, CEO Serial Entrepreneur, Inventor 25+ U.S. and foreign biotechnology patents, filamentous fugal enzyme product commercialization



Ronen Tchelet CSO 20+ years in Biopharmaceutical **Industry & Recombinant Product** Commercialization





clinical manufacturing

**Ping Rawson CFO** 

Deloitte.

20+ years of finance, accounting & international trade and business development experience



Matthew Jones CCO

20+ years life professional management, business development and leadership of biopharma products





Dr. Arin Bose **Board Member** 34 years bioprocess development and

**Dr. Barry Buckland Board Member** 

**MERCK** 

29 years R&D leadership | US National Academy of Engineering





Seasoned pharma industry finance and operational executive





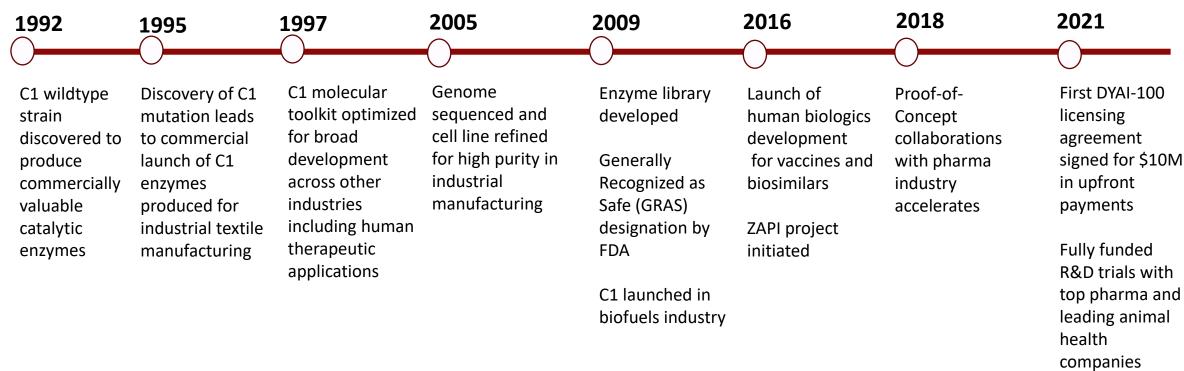
**Board Member** 

20+ years of bioprocess biotech and business development



## **History of the C1 Platform**

"C1" cells are an exceptional strain of genetically engineered fungus — (*Myceliophthora thermophila*) whose usages have expanded through 20 years of commercial engineering



First in human trials are the next step in the C1 platform's commercial evolution



## **Establishing Global Presence with Leading Organizations**

Co-developing C1 enabled COVID-19 (plus variant) vaccines and/or boosters (i.e., tetravalent or quadrivalent COVID-19 vaccines)



Sorrento and Dyadic Announce Binding
Term Sheet to License Dyadic's Lead
COVID-19 Vaccine Candidate "DYAI-100"

Dr. Henry Ji, Sorrento Chairman and CEO, commented, "We look forward to continuing our collaboration with Dyadic, which began last year, initially with a goal of developing and commercializing a protein-based COVID-19 vaccine that can be rapidly manufactured in large quantities in our existing cGMP facilities, and stored and transported at room temperature, in order to increase access and affordability to underserved populations globally."



**Dyadic announces development of COVID-19 Vaccine in India** 



A Biocon company

Mahesh Bhalgat, COO, Syngene International stated, "We look forward to our collaboration with Dyadic to initially explore the development of a COVID-19 vaccine, and to further evaluate the potential of developing a differentiated vaccine platform based on Dyadic's proprietary C1- cell line."



Dyadic Announces Technology Transfer and Licensing Agreement With South Africa's Rubic Consortium

"The need to quickly acquire and commercialize technology and manufacturing capabilities, which addresses the infrastructure necessary to deploy vaccinations for broad populations affordably and timeously has never been a more strategic imperative of African nations than today," said Shabir Madhi, professor of vaccinology, Dean Faculty of Health Sciences at the University of the Witwatersrand, Johannesburg, who is leading COVID-19 vaccine trials in South Africa.



<u>Dyadic and Medytox To Develop Vaccines</u>

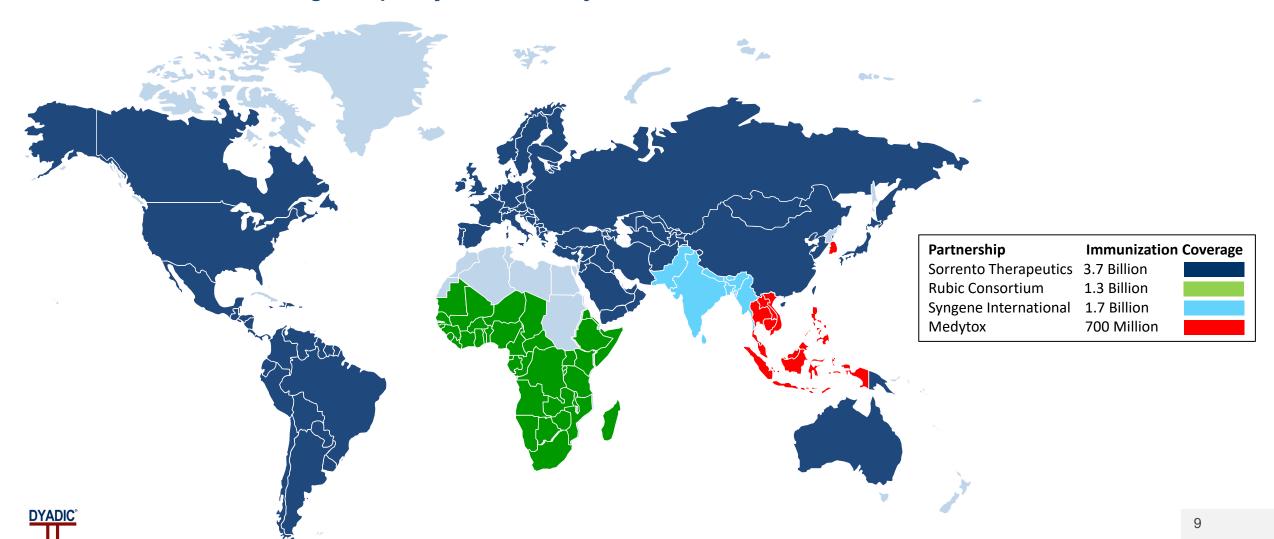
<u>Against COVID-19 Variants</u> (South Korea & SE Asian Countries)

Dr. Gi-Hyeok Yang, Sr, Executive Vice President and Head of Research and Development at Medytox stated, "We have been working closely with Dyadic since July 2020, when we obtained access to their C1 expression platform and experienced the remarkable versatility and high productivity of the C1 platform. We believe that the fungi-derived C1 expression system is the most realistic technology to develop and manufacture multi-valent (i.e., tri-valent, and tetra-valent) vaccines, rapidly and affordably, against COVID-19 mutant viruses without the need for a large-scale bioreactor facility."



## **Dyadic Potential Global COVID Immunization Coverage Nearing 100%**

Signed licensing agreements creates production, commercialization, and distribution pathway for mass-scale immunizations across the globe quickly and efficiently.



## **Development Pipeline**

### Extensive external partnerships are advancing C1 platform for diverse applications

	PROGRAM	STATUS
COVID VACCINE & ANTIBODY		
Sorrento Therapeutics - DYAI-100	SARS-CoV-2-S-RBD antigen and Other Coronavirus applications	DYAI-100 Preclinical animal trial Advancing Towards Clinical Trials
Syngene	COVID-19 variant vaccines	Co-development Program
Medytox	COVID-19 variant vaccines and/or boosters (e.g. multivalent)  Preclinical animal trial	
Rubic South Africa	COVID-19 vaccine candidates Co-development Pro	
ID Biologics (antibody)	COVID-19 mAb vaccine candidate	POC
Epygen <sup>1</sup>	Preclinical & Clinical trials using DYAI-100	Co-development Program
HUMAN (NON-COVID)		
Jiangsu Hengrui	Biologic drugs	POC
Turtle Tree (Growth factor)	Recombinant protein growth factors POC	
Top Pharma	Bispecific antibodies	Co-Development Program
U of Oslo	Influenza vaccine	Internal Program
Other collaborations	Multiple programs for antigens, including bispecific antibodies	10+ POC
ANIMAL PROGRAMS		
ZAPI	SBV and RVFV vaccines	Animal trial
Global Animal Health Co's.	Programs include vaccine for respiratory disease of birds	



## C1-cells Have Broad Advantages Across in Biomanufacturing

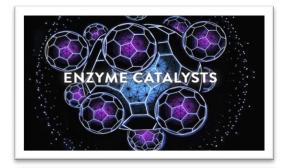
Unique Morphology: Low Cost, High Purity and Yield and Scalable Benefits for Protein Manufacture Across Broad & Growing Applications

















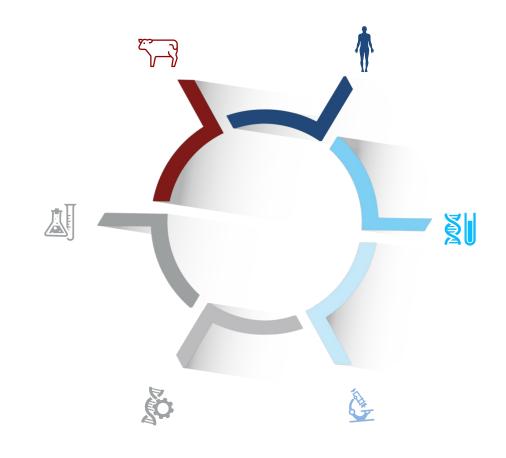




## Broad, Addressable Opportunity to Disrupt Pharma Manufacturing

Human and animal biopharmaceutical manufacturing landscape in need of innovations that improves efficiencies

Market	Global Size
Recombinant Subunit Vaccines and Therapeutic Proteins for Animal health	\$11.3 Billion by 2025 <sup>1</sup>
New Biologics MAbs, Bispecifics, Fc-Fusions	\$319 Billion by 2021 <sup>2</sup>
Recombinant Glycoprotein & Other Antigen Vaccines for Human Health	\$58.4 Billion by 2024 <sup>3</sup>
Biosimilars/Biobetters/Other Biologics	\$69 Billion by 2025 <sup>4</sup>
Vaccines and Therapeutics for environmental (pandemics) and other biological threats, including COVID-19	\$150 Billion by 2022 <sup>5</sup>
Other - Growth Factors, Diagnostics, Metabolites, Reagents, Biocatalysts	

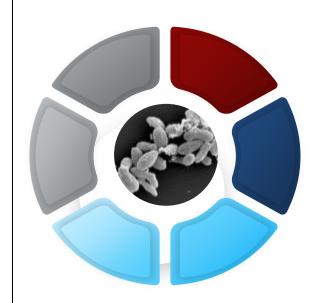




## C1-Thermophilic Filamentous Fungi Key to Platform Technology

The C1 Platform provides safe, highly engineered, thermophilic filamentous fungi ideally engineered to grow under a broad range of conditions.

- 20+-year track record The C1 platform is commercially proven, and has been used to produce large quantities of low-cost enzymes for textiles, biofuels, pulp and paper, food cellulases, etc. at very large industrial scales, up to 500,000 liters
- **Poised to transform healthcare** C1-cells have been carefully re-engineered to enhance scale, purity and yield in therapeutic protein production
- **Genetics toolkit defined -** C1-cell chromosomal genome have been sequenced, full sets of genetic tools for gene engineering and commercial use
- Ready to deploy Multiple genetically engineered C1-cell lines with stable, differentiating properties, ready to deploy across small bespoke discovery, development and commercial cGMP grade production projects
- Generally Recognized As Safe (GRAS) certification from US FDA in 2009
- Toxicology Study Q4 2021
- Advancing Towards First In Human Clinical Trial
- **No commercial retooling needed –** C1 is grown in regulatory friendly, low-cost completely defined synthetic media using standard microbial bioreactors





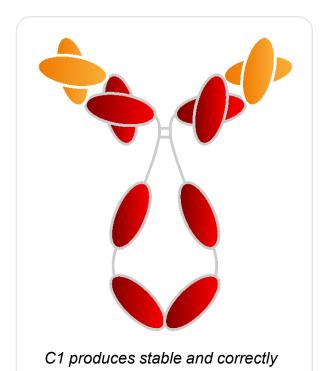
## **C1 Protein Production Platform Competitive Advantages**

Robust Gene Expression Platform Offers A Number Of Competitive Advantages Over Existing Technologies

0-6	Purity	High retention of target secreted protein through downstream processing  No requirement for viral (i.e., CHO) or endotoxin (i.e., E.coli) inactivation
0-3	Productivity	Robust & versatile growth conditions / Low viscosity [unique morphology] High yields of secreted protein
0-33	Robustness	Flexible commercial scales, ranging from laboratory microtiter plates, shake flasks, single use and/or stainless-steel microbial bioreactors.
0-(50)	Speed	Develop stable g/l C1 cell lines in ~7 weeks Production savings of ~30 days over CHO Potential to make ~ 3-4 batches of mAbs in the same time it takes to make 1 batch using CHO cells
DYADIC*	Cost	High yields and fast production times can reduce cost and shrink manufacturing footprint Requires only low-cost cGMP synthetic media  No requirement for viral or endotoxin inactivation, simplifies processing compared to CHO & E.coli saving time, money

## C1 Platform – Potential to Disrupt Conventional mAb Manufacturing

C1 platform produces comparable therapeutic proteins as CHO while overcoming key production limitations



folded mAbs that have binding and

neutralizing properties to those

produced from CHO cells



Flexible production scale; C1 media <1/20 of the cost of CHO media No viral inactivation required



C1 produces product significantly faster (12-14 days) than CHO cells (41-54 days)



C1 produces more product per batch and larger overall quantities

~ Potential to produce three to four batches using C1 in the same timeframe as one batch using CHO cells



## **C1-Cell Recombinant Protein Production: Biologics**

C1-Cells Enable Commercial Manufacture Of Rapid, Cost-Effective, High Value, Safe, Effective Protein Products

High Yields and Purities Demonstrated for Therapeutic Monoclonal Antibodies (mAbs) and Vaccine Antigen GlycoProteins<sup>1</sup>

Fc-Fusion Products	mAb Products	Fab (Certolizumab) Product	Tri-specific Products
15.3 g/l <sup>1</sup>	24.5 g/l <sup>1</sup>	14.5 g/l <sup>1</sup>	6.12 g/l <sup>1</sup>
168 Hours	168 Hours	164 Hours	144 Hours
2.58 g/l/day	3.1 g/l/day	2.1 g/l/day	1.02 g/l/day

### High Productivity for Recombinant Protein Antigen Classes Routinely Used in Vaccines

Influenza HemAgglutinin (HA) Products	Coronavirus Antigen (S-RBD) Products	
413 mg/l <sup>1</sup>	2,000/3,000 mg/l <sup>1</sup>	
137 Hours	120 Hours	
72 mg/l/day	400/600 mg/l/day	





## Regional Collaborations Have Helped Fund Advances in C1

# ZAPI has helped validate the C1 Production Platform which can be used to churn out vaccines and antibodies in unprecedented amounts

• The European Union Zoonosis Anticipation Preparedness (ZAPI) Project, initiated in March 2015 has since worked to develop launch ready methodologies and platform technologies that could be put into production for vaccines and neutralizing monoclonal antibodies in order to efficiently counter emerging or reemerging zoonotic viruses of pandemic proportions.



# Israel Institute for Biological Research (IIBR) Strategic Collaboration Has Helped To Position Dyadic to Enter Human Clinical Trials

- COVID-19 Strategic collaboration with the Israel Institute for Biological Research (IIBR) IIBR has positioned C1 positioned to be able play to important role in combating pandemics.
- Demonstrating that C1-cell produced SARS-CoV-2 receptor binding domain (RBD) can be manufactured in large quantities, at low cost, while neutralizing the COVID-19 Virus.





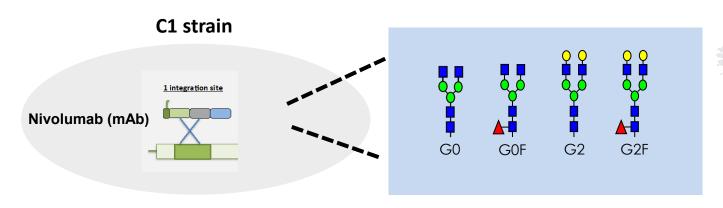
## <u>Human Biopharmaceutical Programs</u> - Nivolumab (Opdivo®) a Biosimilar mAb Candidate

Anti-Cancer mAb, (i.e., Anti-PD-1 IgG) Are Compelling High Value mAb Product Opportunities For Fungal C1-Cell Manufacturing

### **Nivolumab (Opdivo®)**

manufactured by Bristol Myers Squibb, is an immunotherapeutic biologic Mab drug for human metastatic cancers, including melanoma, lung & other cancers

- Opdivo priced at \$12,500 per month or about \$150,000 per year of treatment
- Goal of program is to express Nivolumab (MAb) with a glycoprotein structure similar to Nivolumab produced in CHO cells
- Dyadic has glycoengineered mAb producing C1-cell lines with G0 levels of about 95% and G2 of about 76% as part of its glycoengineering program for glycoprotein Immunoglobulin G (IgG) monoclonal antibodies
- Further C1-cell engineering & manufacturing mAb process development ongoing
- Important Proof Of Concept in successfully manufacturing Optivo mAb biosimilar or biobetter product, C1-cell manufacturing tech applicable to several very high value therapeutic or preventative monoclonal antibodies



Global Sales Example for Anti-PD-1
MAb ImmunoTherapeutic Oncology
Product

\$8.0 B(1)

## **Business Development Strategy**

External development programs help advance C1 recombinant protein platform and fund R&D initiatives

### **Co-Development / Technology Licensing**

Allows Dyadic to develop and advance C1 recombinant protein tech at low cost to Company

### **Big Pharmas**

Funded proof of concept collaborations for specific therapeutic products

Potential for UP-front access fees, milestones and royalty payments

#### Small & Medium Biotech's

Potential for equity, milestones and royalty payments

#### **Grants & Contracts**

Governmental and agency grants and contracts

### **Internal Dyadic Product Development**

Dyadic funds high value product candidate programs where C1 recombinant gene expression overcomes barriers of existing platforms with meaningful technological or commercial impacts

- COVID-19 vaccine candidate, DYAI-100
- Glycoengineering
  - Advance Nivolumab (Opdivo®) Biosimilar MAb
  - As Proof of Concept for Production of mAbs
- Protease deletion and engineering
- SARS-CoV-2 Variants. Influenza other Infectious Diseases
- Metabolites



## Rising Interest in Pharma for Acquisition of Platform Technologies<sup>1</sup>

Recent acquisitions of cell-based gene expression and recombinant protein platforms



### 2006

Merck & Co., Inc. To Acquire GlycoFi, Inc. For \$400 Million <sup>2</sup> **Yeast cell** 



Johnson Johnson

Johnson & Johnson Acquires Vaccine Developer Crucell - \$2.4 billion dollars, moving J&J prominently into the arena of vaccine development <sup>3</sup> *Human cell* 



### 2017

Sanofi completes acquisition of Protein Science for \$750 Million <sup>4</sup>
Baculovirus expression system



### 2020

Ligand Buys Pfenex in \$516 Million Deal to Access Protein Production Platform <sup>5</sup> *Bacterial cell* 

- 1. https://www.forbes.com/sites/mergermarket/2018/10/26/pharma-ma-slumps-as-buyers-seek-platforms-instead-of-single-products/?sh=6fed8c266848/
- 2. https://www.biospace.com/article/releases/merck-and-co-inc-to-acquire-glycofi-inc-for-400-million-/
- 3. https://www.biopharminternational.com/view/johnson-johnson-acquires-vaccine-developer-crucell/
- 4. https://www.pharmaceutical-technology.com/news/newssanofi-completes-acquisition-of-protein-sciences-for-750m-5910136/
- 5. https://www.biospace.com/article/ligand-buys-pfenex-for-516-million/



### **2021 Strategic Objectives**

SARS-CoV-2

Advancing Towards Phase 1 Clinical Trial using a Dyadic owned C1-cell manufactured SARS-CoV-2 Receptor Binding Domain (RBD) recombinant antigen vaccine product candidate in collaboration with CR2O

Demonstrate Safety & Preliminary Efficacy (POC) In Humans

Expanding portfolio of vaccine candidates against SARS-CoV-2 variants through partnerships with Medytox & Syngene Anticipate a Phase 1 Human Clinical Trial in the EU with a C1-cell manufactured SARS-CoV-2 monoclonal antibody

Animal Health Programs Continue to advance commercial recombinant antigen vaccine product candidates in animal trials with global animal health companies, **including for an acute respiratory disease of birds** 

Advancing C1 produced SBV & RVFV recombinant antigen vaccines in additional safety & efficacy studies with additional funding from ZAPI

Human Biopharmaceutical Programs Advancing several recombinant protein growth factors as potential regenerative therapies with TurtleTree Scientific

Advancing mAb with top tier biopharmaceutical company with registrational potential in oncology

Advancing bispecific mAb with top tier biopharmaceutical company with registrational potential in autoimmune disease

Part of the glycoengineering effort we are using Nivolumab (Opdivo®) as a potential biosimilar MAb product candidate

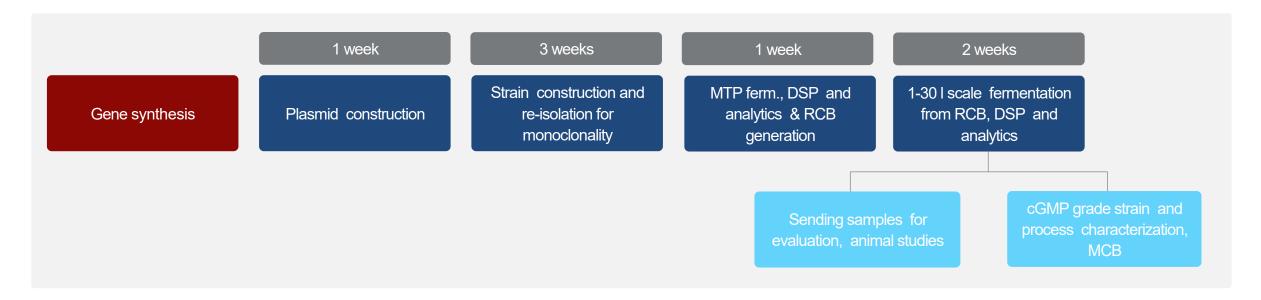


### Infectious Disease Strategy For Animal and Human Health

Potential to Develop Multi-Valent Recombinant Variant Antigens (i.e., SARS-CoV-2 and Influenza) Vaccine Candidates

Already Five (5) Variant of Concern SARS-CoV-2 RBDs have been expressed from C1-cells, including the Delta Variant

Univ. Oslo HA mice trail supports previous Sanofi HA mice trial for potential use of C1 for producing influenza vaccines



Monoclonal antibodies (mAbs) have been expressed from C1-cells for potential use in treating infectious diseases, including SARS-CoV-2, Zika and Rift Valley Fever.



## **Key Takeaways**

### **Next Generation Protein Expression Biotech with Well-established Global Partners**

**Proprietary C1 gene expression technology** 

**Competitive advantages** 

**Validating partnerships** 

**Opportunistic business development** 

**Strong financial position** 

**Experienced management - board of directors** 

Designed to bring biologic vaccines and drugs to market faster, in greater volumes, at lower cost

Compared with other pharmaceutical expression systems supported by robust and growing scientific data on protein expression yield, stability and purity

Well-established, global biological R&D organizations, top-tier animal and human health pharmaceutical companies, as well as governmental agencies

Emphasis on large and growing addressable human and animal health markets; many shots on goal

\$25.8 million in cash and investment securities, no debt and complemented by partner funded on-going R&D collaborations (1)

Driving process and execution excellence

(1) As of March 31, 2021

## Thank-you

Contact: ir@dyadic.com

www.dyadic.com

