



Powering a New Decade of DNA Medicines

May 2021



Forward-Looking Statements

This presentation includes statements that are, or may be deemed, “forward-looking statements,” within the meaning of Section 27A of the Securities Act of 1933, as amended. All statements, other than statements of historical facts, included in this presentation regarding our strategy, future operations, future financial position, future revenue, projected costs, prospects, plans and objectives of management are forward-looking statements. In some cases, you can identify forward-looking statements by terms such as “may,” “might,” “will,” “objective,” “intend,” “should,” “could,” “can,” “would,” “expect,” “believe,” “anticipate,” “project,” “target,” “design,” “estimate,” “predict,” “opportunity,” “proposition,” “strategy,” “potential,” “plan” or the negative of these terms and similar expressions intended to identify forward-looking statements.

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In addition, the forward-looking statements included in this presentation represent INOVIO's views as of the date hereof. INOVIO anticipates that subsequent events and developments may cause its views to change. However, while INOVIO may elect to update these forward-looking statements at some point in the future, the company specifically disclaims any obligation to do so, except as may be required by law. These forward-looking statements should not be relied upon as representing INOVIO's views as of any date subsequent to the date of this presentation.

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Powering DNA Medicines

**15 DNA medicine clinical programs currently in development
(HPV-associated diseases, cancer, and infectious diseases, including COVID-19)**

**Precisely designed plasmids delivered
through proprietary smart device**

**Extensive patent portfolio
protecting technology platform**

**Designed to treat and prevent cancers
& infectious diseases**

**Strong and experienced
management team**



**Well-tolerated and robust immune responses
in more than 3,000 patients**

No anti-vector response

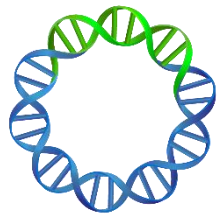
**No frozen storage issues
(room temp storage >1 yr.)**

**Targets multiple antigenic sequences;
combining multiple antigens into single vial**

DNA Medicines Platform Built on INOVIO's Proprietary Technology

OPTIMIZED PLASMID DESIGN AND DELIVERY TECHNOLOGY

**PRECISELY
DESIGNED PLASMIDS**
(SynCon[®])



**PROPRIETARY
SMART DEVICES**
(CELLECTRA[®])

**Intramuscular
Device** for
Pre-Cancers &
Cancers



**Intradermal
Device** for
Vaccines



IN VIVO



INOVIO's Technology Advantages



Clinical Efficacy

- Demonstrated clinical efficacy in Phase 3 study for VGX-3100
- Lead candidate VGX-3100 in Phase 3 for precancerous cervical dysplasia

Tolerability

- Favorable safety profile tested in over 3,000 patients and over 7,000 administrations
- Carries no potential toxicity from viral vectors

Versatility and Boosting

- Targets virtually any antigenic sequence; combining multi-antigens into single vial
- Initiated first-in-human study of optimized dMAb™ plasmid
- **No anti-vector response** – allows for additional boosting

Rapid and Scalable Manufacturing

- “Off-the-shelf” product; **no cold chain required** (room temp storage >1 yr.)
- Rapid development from concept to human in <3 months (COVID-19 vaccine)
- Relatively inexpensive to manufacture; produce large quantities

INOVIO DNA Medicines Pipeline

	PRODUCT	INDICATION	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	PARTNER/COLLABORATOR/FUNDER
INFECTIOUS DISEASES	INO-4800	COVID-19 (Coronavirus)	Internally Funded				
	INO-4700	MERS	Internally Funded				
	INO-4500	Lassa Fever	Internally Funded				
	INO-4201	Ebola	Internally Funded				
	PENNVAX-GP	HIV	Internally Funded				
HPV-TARGETED	VGX-3100	Precancerous Cervical Dysplasia (HSIL)	Internally Funded				 (China; INOVIO maintains global rights)
		Precancerous Vulvar Dysplasia (HSIL)	Internally Funded				
		Precancerous Anal Dysplasia (HSIL)	Internally Funded				
	INO-3107	Recurrent Respiratory Papillomatosis (RRP)	Internally Funded				
	MEDI0457	Head & Neck Cancer	Externally Funded				
Cervical, Anal, Penile, Vulvar Cancers		Externally Funded					
IMMUNO-ONCOLOGY	INO-5401	Glioblastoma Multiforme (GBM)	Internally Funded				
	INO-5151	Prostate Cancer	Externally Funded				
dMab™		COVID-19 (Coronavirus)	Externally Funded				
	INO-A002	Zika	Externally Funded				

INTERNALLY FUNDED



EXTERNALLY FUNDED



Infectious Disease Programs & COVID-19 Vaccine Program



INO-4800 Updates



COVID- 19 Vaccine

INOVIO is developing a two-dose **INO-4800 regimen** for protection against **COVID-19 disease**

Addressing New Variants of Concerns

- Evaluating new strains, with focus on UK, South African and Brazilian, on immune profile of INO-4800
- Using our SynCon® technology, **we are developing a pan-COVID vaccine candidate** to potentially provide better protection against both known and future strains



Clinical Data and Plan

Phase 1 dosing regimen complete

- Showed favorable safety and tolerability profile
- Elicited a broad immune response across multiple assays, preliminary clinical responses
- **Demonstrated binding, neutralizing antibodies & cellular responses at week 6**
- **Boosting with INO-4800 – in process for Phase 1 cohort**

Phase 2 dosing regimen complete

- Fully funded by U.S. DoD
- Showed favorable safety and tolerability profile
- 2mg dose selected - GMFR of binding and neutralizing antibody levels were statistically significantly greater in the 2.0 mg dose group versus the 1.0 mg dose group.

Phase 3 planned

- Global, blinded, case-driven, immunogenicity and efficacy trial

Ongoing Phase 2 clinical trials in China and South Korea



Manufacturing & Scale up

- Scaling up plasmid and device through consortium of CMOs and partnerships globally
- **Excellent stability profile**, room temperature for >1 year, anticipated 5- year shelf life at 2-8°C



INO-4800 Key Differentiators

Favorable Safety & Tolerability:

- INO-4800 has demonstrated favorable safety and tolerability
- Administered intradermally and has caused only very limited side effects (mild injection site reactions)

Immunogenic:

- 100% of Phase 1 participants demonstrated overall immunological response rates
- Balance of neutralizing antibodies and favorable T-cell responses (CD8 and CD4)

Temperature Stable and Transportable:

- Vaccine is projected stable at room temperature for more than a year, at 37°C for more than a month
- Five-year projected shelf life at normal refrigeration temperature and does not need to be frozen during transport or storage

Repeat Administration:

- INO-4800 can be re-administered if immunity wanes
- Potential for seasonal boosting usage with no concerns of generating an anti-vector response, based on observations to date

INO-4800 Ph 1 Trial Data: Regimen was well-tolerated and generated both B and T cell immune response

Data from 40 subject cohort of Phase 1 peer-reviewed and published in *EClinicalMedicine*:

A • Tolerability

- 1.0mg and 2.0mg doses of INO-4800 in a 2-dose regimen is well tolerated in initial cohort of younger (18-50) subjects evaluated in the U.S.

B • Immunogenicity

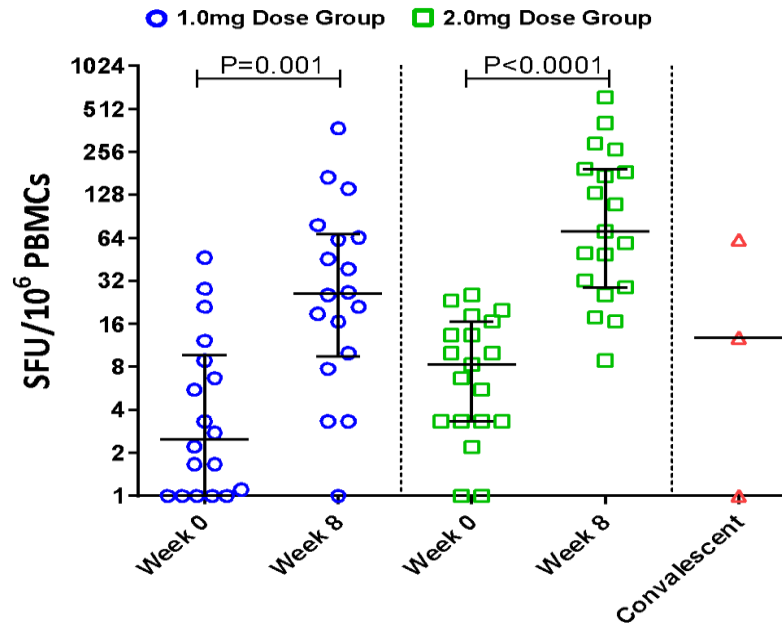
- INO-4800 induced a balanced immune response comprising both B cell (neutralizing and binding antibodies) and T cell (Th1 effector and memory cell) responses

Phase 1 expansion (120 subjects) and booster study is ongoing:

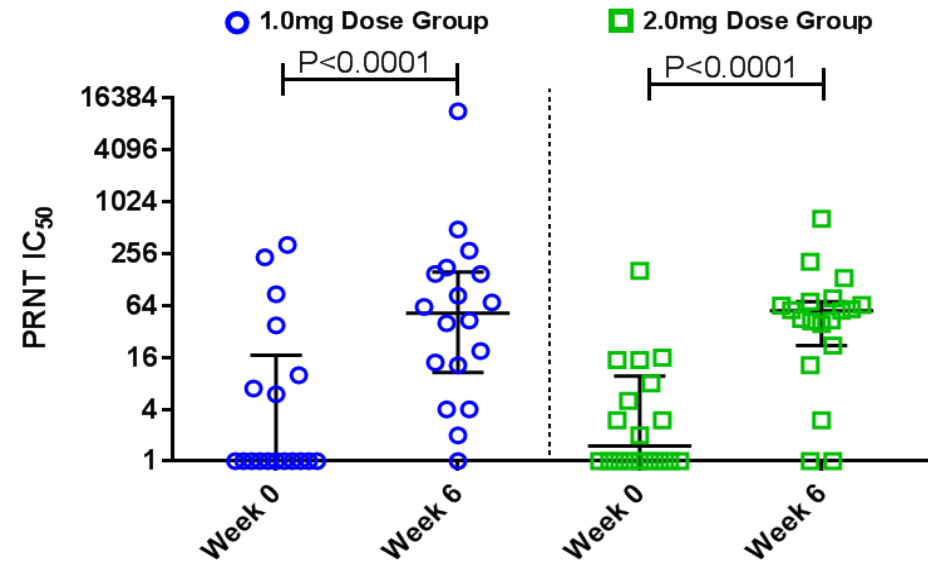
- Tolerability and immunogenicity of 1.0mg and 2.0mg doses in expanded age groups of older subjects (51-64) and elderly (65 years and older)
- Tolerability and immunogenicity of 0.5mg dose in a 2-dose regimen (Days 0, 28) at age groups of 18-50, 51-64 and 65+ years
- 93/120 subjects from Phase 1 were administered a booster at 6-10 months from their second dose

U.S. Phase 1: Week 8 Immunogenicity on 40 Subjects in 18-50 year age group

Induction of Antigen Specific T Cells by ELISpot* 1.0mg vs 2.0mg



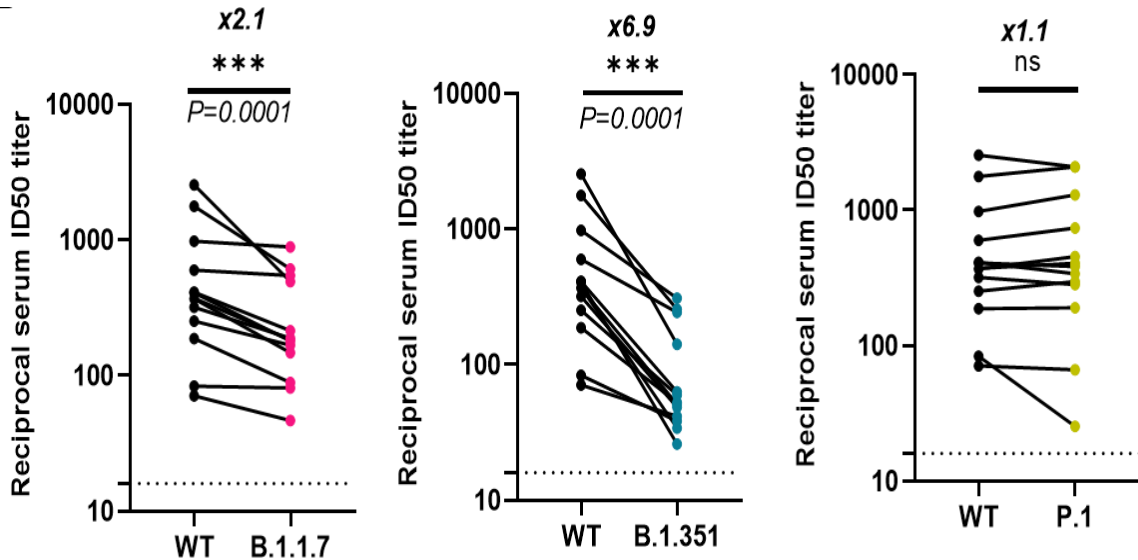
LIVE SARS-CoV-2 Neutralization* 1.0mg vs 2.0mg



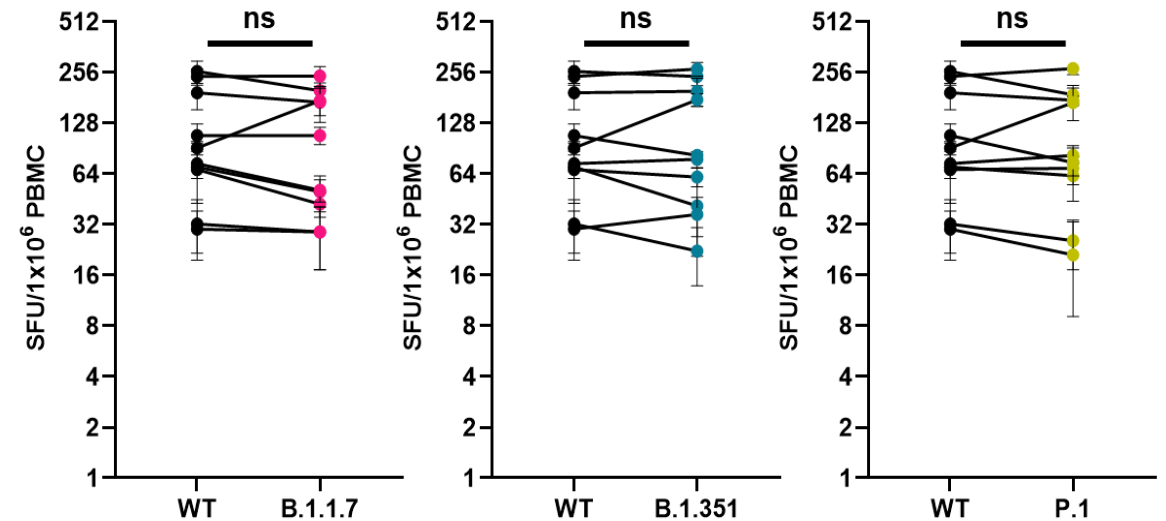
- Strong CD4 and CD8 T cell responses generated to multiple regions of the spike protein
- 74% of the subjects had T cell responses at the 1.0 mg dose group and 100% of the subjects in the 2.0 mg dose group demonstrated cellular responses
- The 1.0 mg and 2.0 mg dose group both demonstrated seroconversion in 95% of the subjects
- 78% demonstrating neutralizing antibodies in the 1.0 mg dose group and 84% demonstrating neutralizing antibodies in the 2.0 mg dose group

Addressing Variant of Concerns: INO-4800 DNA Vaccine Induces Neutralizing Antibodies and T cell Activity Against Global SARS-CoV-2 VoCs

Humoral antibody cross-reactivity responses against SARS-CoV-2 variants. a) Sera from Phase 1 INO-4800 vaccinees were neutralization to WT, B.1.1.7, B.1.351, and P.1 variants.



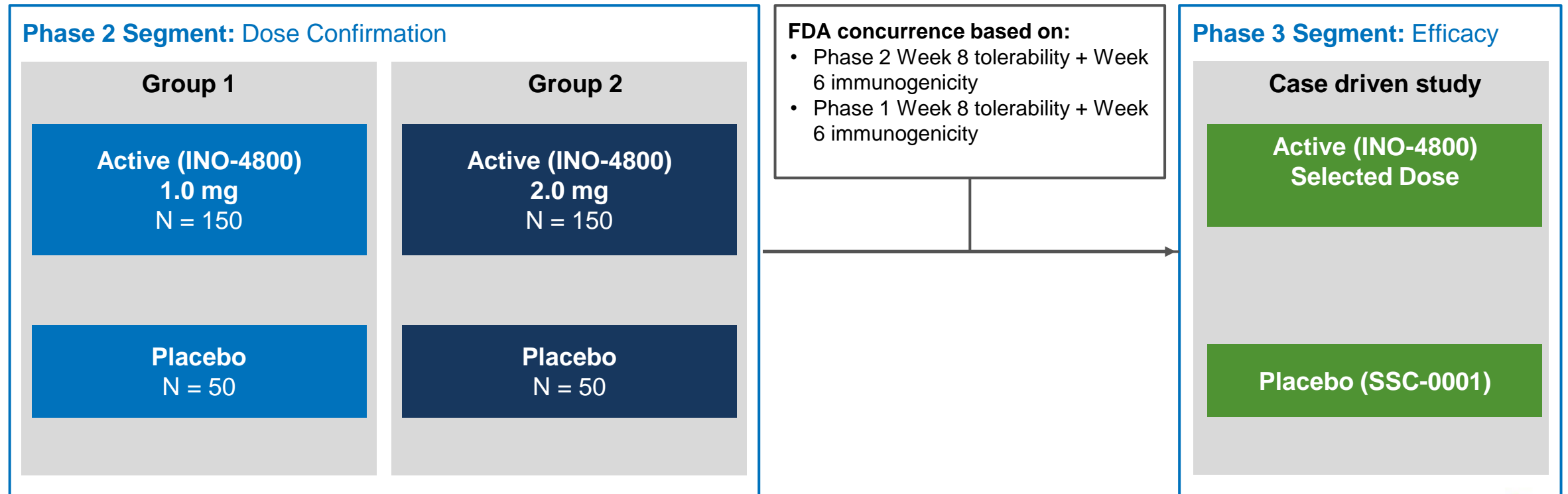
INO-4800 Cellular immune response against SARS-CoV-2 variants. PBMCs from 10 Phase 1 subjects were collected 8 weeks after receiving the second dose of INO-4800.



Phase 2/3 Clinical Trial- INNOVATE (INovio INO-4800 VAccine Trial for Efficacy)

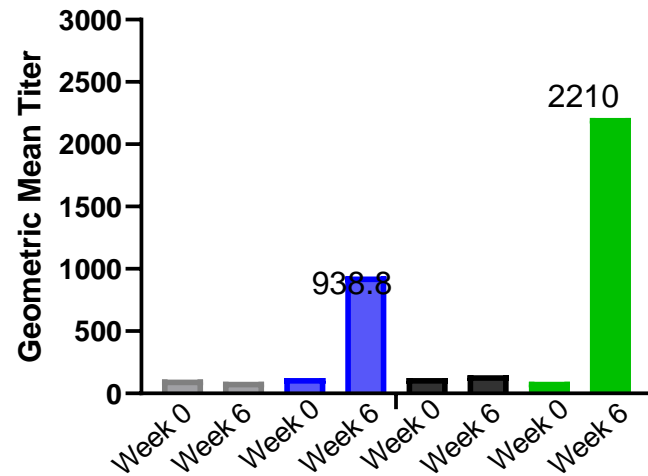
Evaluating efficacy in subjects 18+ years of age with optimal dose for each age group

- Phase 2 segment: to evaluate tolerability and immunogenicity in order to select dose(s) for efficacy evaluation in Phase 3
- Phase 3 segment: to evaluate efficacy using the selected dose(s) from Phase 2 segment in a case-driven fashion

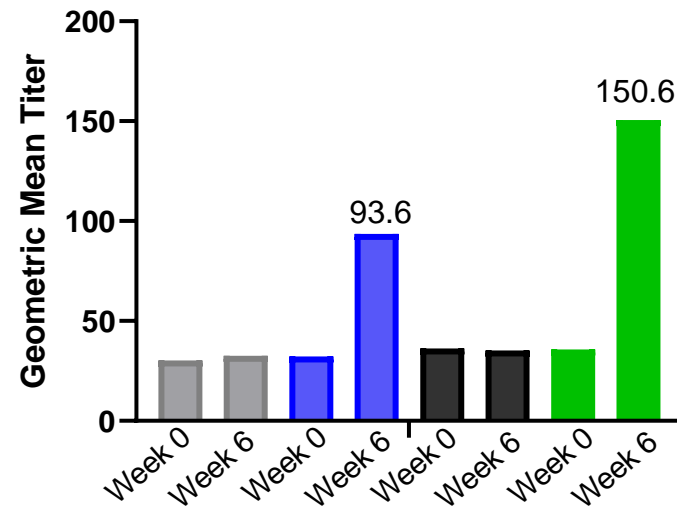


INO-4800 Phase 2 Immune Responses across All Age Groups

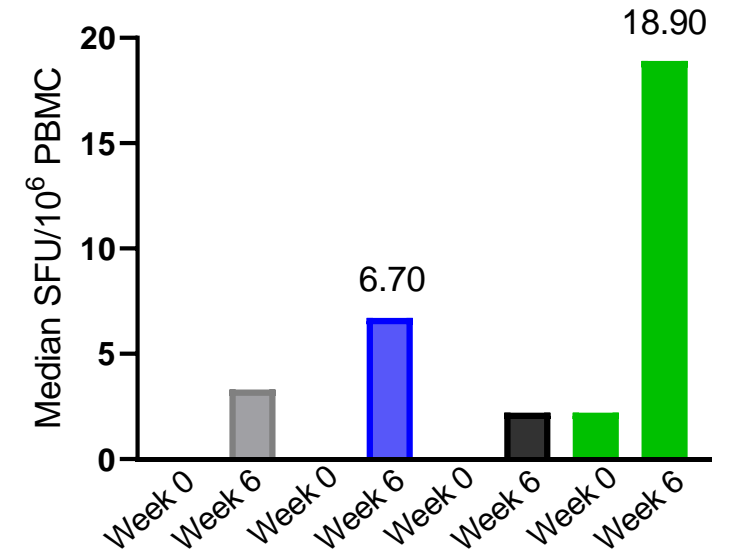
Binding Ab



Neutralizing Ab



T Cell Responses

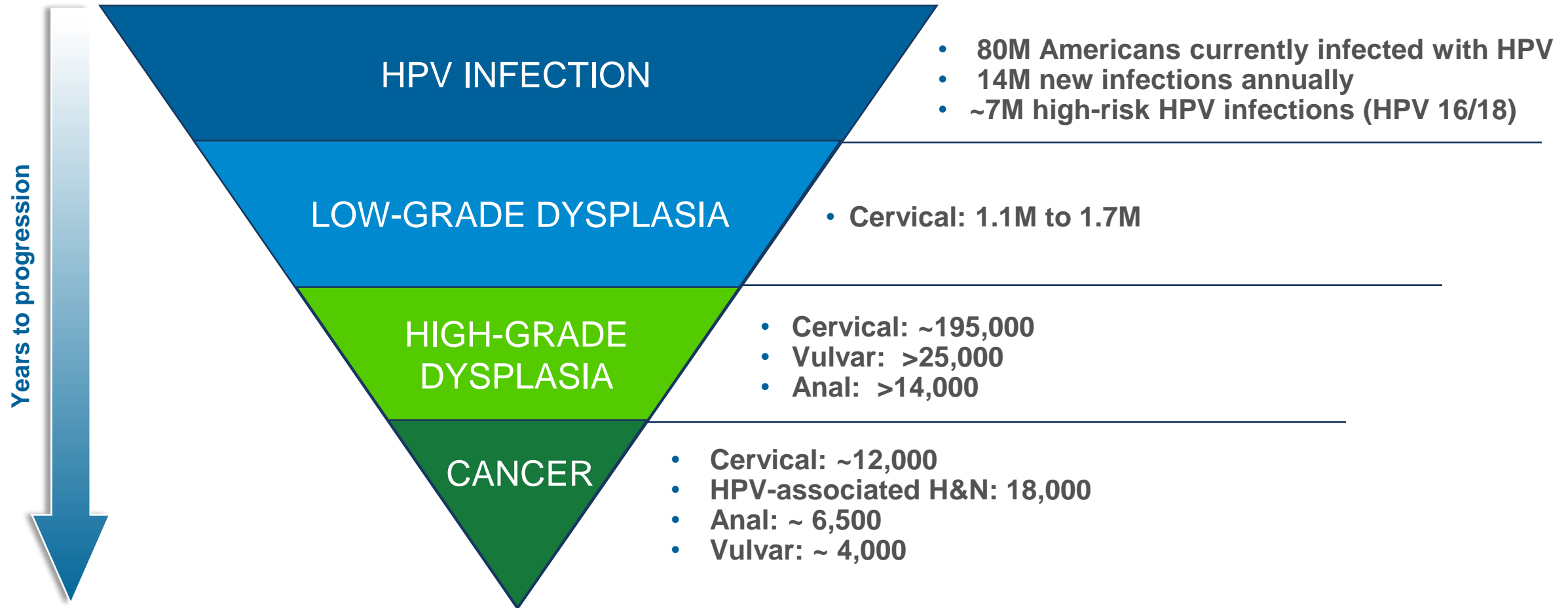


HPV Programs



HPV-Associated Diseases Market Overview

HPV-associated conditions per year in US:



Sources: US CDC (2018) HPV and Cancer, available at: <https://www.cdc.gov/cancer/hpv/statistics/cases.htm> (accessed July 22, 2019); Saraiya M, Unger ER, Thompson TD, Lynch CF, Hernandez BY, Lyu CW, Steinau M, Watson M, Wilkinson EJ, Hopenhayn C, Copeland G, Cozen W, Peters ES, Huang Y, Saber MS, Altekruze S, Goodman MT; HPV Typing of Cancers Workgroup. US assessment of HPV types in cancers: implications for current and 9-valent HPV vaccines. J Natl Cancer Inst. 2015 Apr 29;107(6):dju086; Inovio Pharmaceuticals, internal estimates from published data (2015-16, 2017-18); US CDC, personal communication (2015); NCI SEER Cancer Stat Facts: Cervix Uteri, Vulvar, and Anal Cancers – <https://seer.cancer.gov/statfacts> (accessed 2017-18); *Measured as: Genital Warts – Initial Visits to Physicians' Offices, United States, 1966-2014. Fig. 47; Schiffman M, Solomon D. Findings to date from the ASCUS-LSIL Triage Study (ALTS). Arch Pathol Lab Med. 2003 Aug;127(8):946-9; US CDC. Genital HPV Infection – Fact Sheet.

Complications of Current Standard of Care: Loop Electrosurgical Excision Procedure (LEEP)

Pain^{1,2}

- Local anesthetic injections
- Excision
- Post-procedural cramping



Surgical Complications^{1,3}

- Disfigurement
- Swelling, drainage, bleeding, numbness, redness, burning
- Opening of suture, itching, scarred skin
- Cervical stenosis



Loss of Reproductive Health

Increased risk of:

- Preterm delivery^{6,7}
- Premature rupture of membranes^{6,7}
- 2nd trimester miscarriage⁸
- Terminations⁸



¹Harper DM, et al. *J Family Practice*. 1994;39:249–256.

²Ferenczy A, et al. *Obstet Gynecol*. 1996;87:332–337.

³Mitchell MF, et al. *Obstet Gynecol*. 1998;92:737–744.

⁴Wright TC, et al. *Obstet Gynecol*. 1992;79:173–178.

⁵IARC. *Colposcopy and Treatment of CIN: A Beginner's Manual*. 2003.

⁶Kyrgiou M, et al. *Lancet*. 2006;367:489–498.

⁷Kyrgiou M, et al. *BMJ*. 2016;354:i3633.

⁸Kyrgiou M, et al. *Cochrane Database Syst Rev*. 2015;CD008478.

VGX-3100 Phase 3 Program: HPV-Associated Cervical HSIL/ Precancerous Dysplasia

TRIAL: **VGX-3100**

- Targets HPV 16/18 subtypes; E6/E7 oncogenes
- Designed to treat high-grade squamous intraepithelial lesions (HSIL)



Phase 3 consists of 2 studies in parallel:

REVEAL 1 (primary) n=201 – Enrollment Closed
Study follow-up through week 88 (as in Phase 2b)
Topline efficacy data reported 1Q21

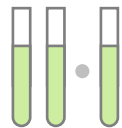
REVEAL 2 (confirmatory) n=198 – Now Enrolling
Study follow-up through week 40

FIRST treatment
for HPV infection of
the cervix

FIRST non-invasive
treatment for cervical
pre-cancer

Primary endpoint:
Regression of HSIL (CIN2/3) AND
clearance of HPV 16/18 in the cervix

2:1 Randomized (2:1), double-blind, placebo-controlled



Dosing: month 0, 1, 3
(as in P2b)

6 mo.

Primary endpoint measured 6 months after completion of dosing (as in Phase 2b)

REVEAL 1: VGX-3100 Phase 3 Pivotal Trial for Cervical HSIL Meets Primary & Secondary Efficacy Objectives for All Evaluable Subjects

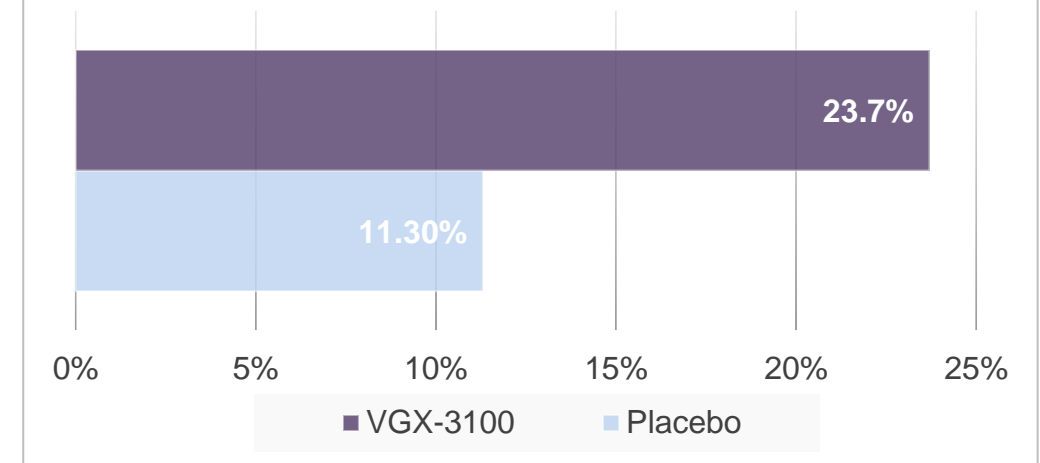
First DNA medicine to achieve efficacy endpoints in a Phase 3 trial

No treatment-related serious adverse events; most adverse events were mild to moderate and self-resolving

Partnership with QIAGEN to develop pre-treatment predictive biomarker to help identify those likely to respond to VGX-3100

- **Achieved statistical significance for primary objective:** regression of cervical HSIL combined with virologic clearance of HPV-16/18, 6 months after administration
 - 23.7% (31/131) in treatment group vs. 11.3% (7/62) in placebo group
 - $p=0.022$; 12.4 difference in percentage, 95%CI: 0.4,22.5
 - mITT includes all subjects w/ endpoint data (N=193) *
- **All secondary efficacy objectives achieved**
- REVEAL 2 is currently ongoing

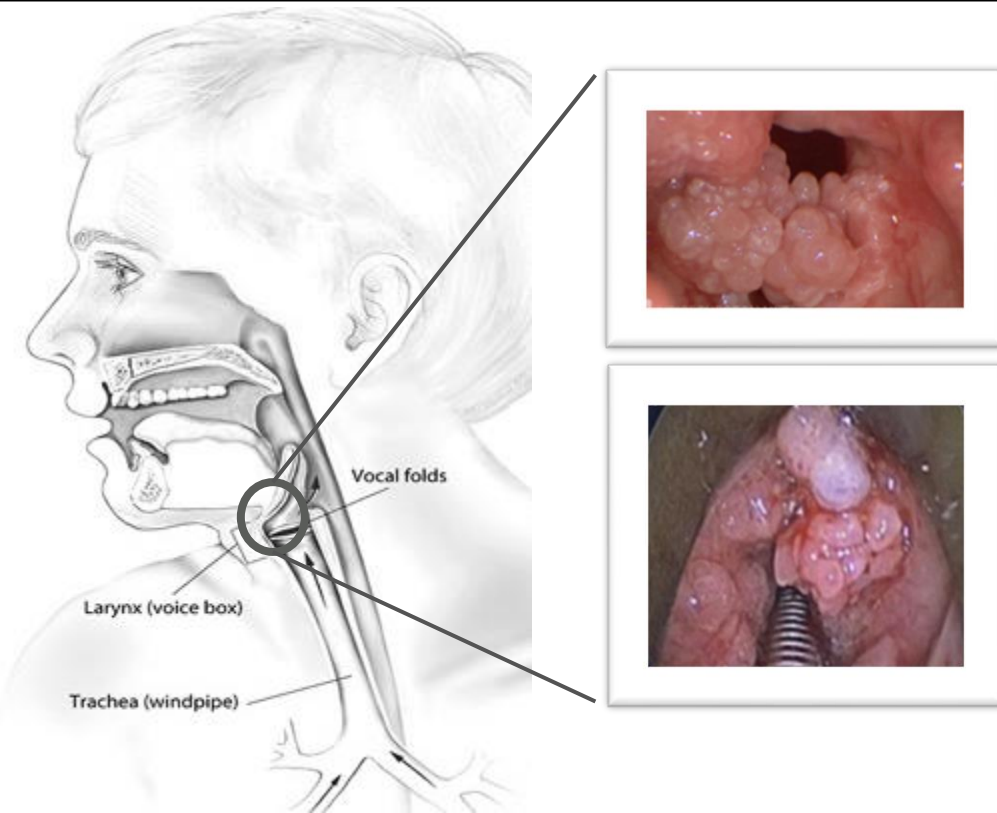
Regression of Cervical HSIL with Virologic Clearance



* mITT, ITT and a third per-protocol (PP) were pre-specified in trial protocol. PP analysis will be performed upon trial completion.

Recurrent Respiratory Papillomatosis (RRP) Caused by HPV 6 and 11

Areas affected by Recurrent Respiratory Papillomatosis (RRP)



- HPV-associated disease; **caused by HPV 6 and 11**
- Rare, orphan disease with **~15,000 total active cases** within the U.S., where **virtually all of those require surgical procedures**
 - **~6,000 new cases per yr. in the U.S.**
- Growths can lead to life-threatening airway obstructions
- **SoC is lifelong surgery (repeated/multiple times per yr)**
 - Currently, disease is incurable and can only be treated by surgery to remove tumors, which temporarily restores the airway
- RRP may occur in adults as well as in children who are thought to have contracted the virus during childbirth

Immuno-Oncology Programs (INO-5401 for Newly Diagnosed GBM)



INO-5401 for Newly Diagnosed GBM in Phase 1/2 Study in Collaboration with Regeneron

TRIAL: **INO-5401** (encoding tumor-associated antigens: hTERT, WT1, PSMA)



Phase 1b/2 open label study for **newly diagnosed glioblastoma (GBM)**



Combination with Regeneron's PD-1 checkpoint inhibitor cemiplimab (Libtayo®)

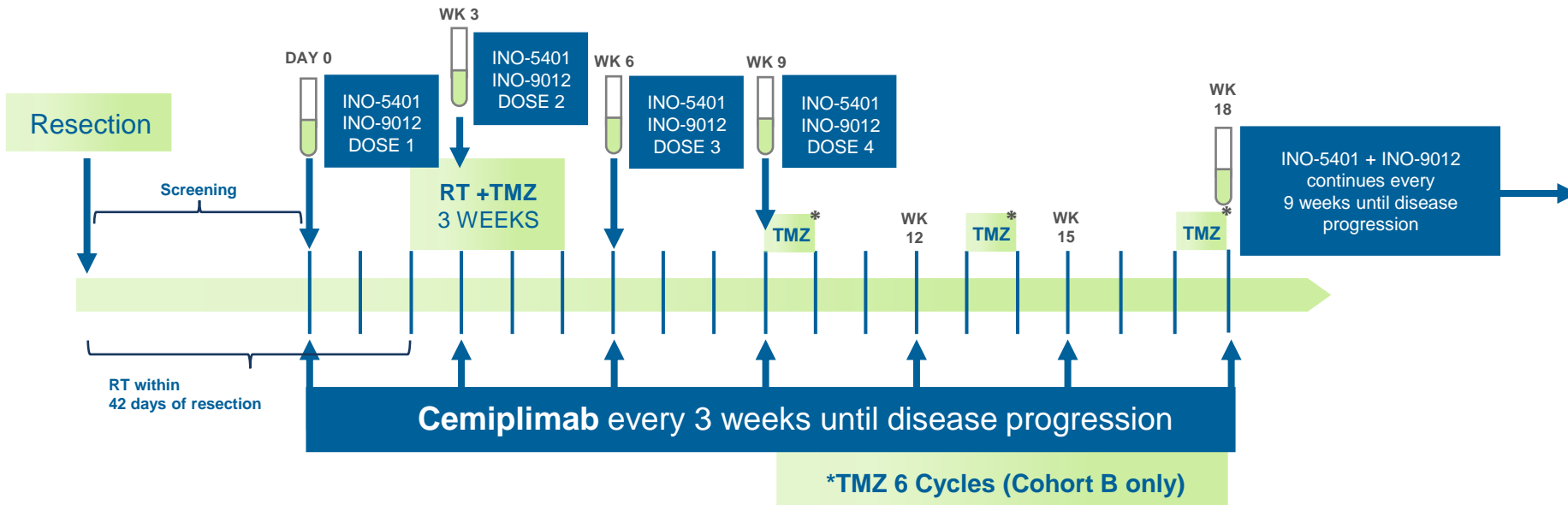
Primary Endpoints:
Safety, tolerability
Secondary Endpoints:
Immunological impact, **PFS and OS**



Cohort A:
MGMT Promoter
Unmethylated:
32 patients



Cohort B:
MGMT Promoter
Methylated:
20 patients

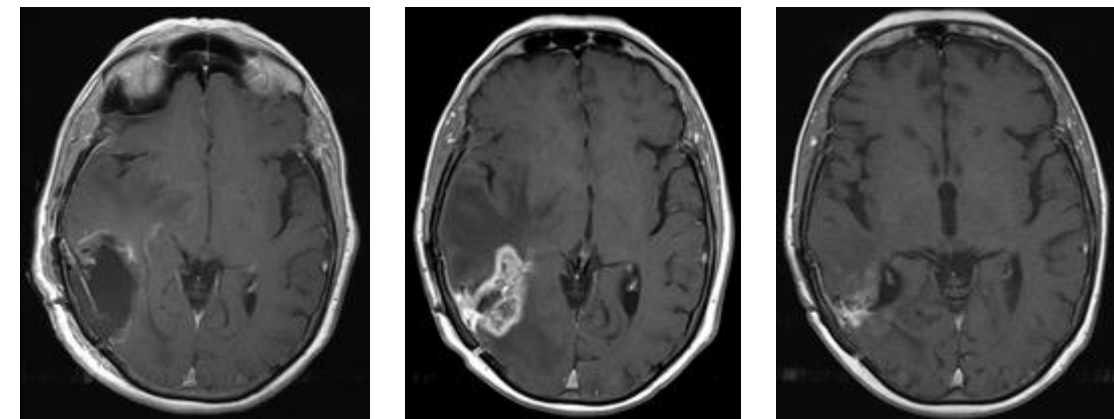
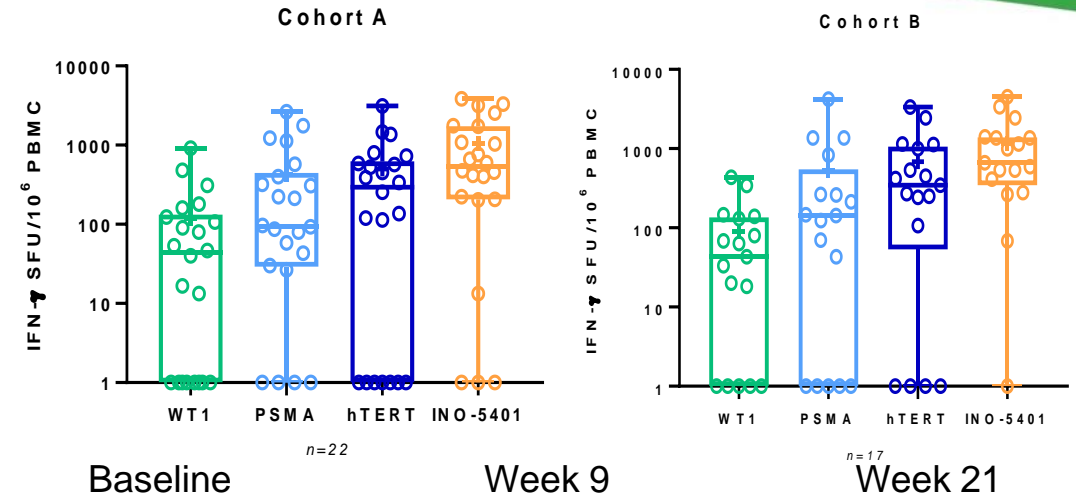


INO-5401 Results: Interim review in newly diagnosed GBM patients

OS18 data, demonstrated immunogenicity and tolerability in a majority of patients

- **Overall survival at 18 months (OS18)** presented at SNO 2020 Annual Meeting:
 - Promoter Methylated OS18 of 70% (14/20)
 - MGMT Promoter Unmethylated OS18 of 50% (16/32)
- **Median overall survival in the unmethylated GBM patients was 17.9 months, which compares favorably to historical controls**
 - Median OS for methylated patients has not yet been reached and the study is ongoing
- This study shows that INO-5401+INO-9012 with cemiplimab and radiation/TMZ have an acceptable tolerability profile, are immunogenic, and may improve survival in newly diagnosed GBM
- *24-month OS data expected later this year, including correlative immunology and tissue data, as well as total study drug exposure and concomitant medication use*

Immunology Output to Date



Several patients have experienced pseudo-progression, with progression by RANO criteria and radiographic evidence of progression on MRI, without evidence of tumor on repeat biopsy

Overall Survival at 18 Months

Median OS; unmethylated (Cohort A)	17.9 mo. (14.5 - NR)	<i>Historical 14.6-16 mo.**</i>
Median OS; methylated (Cohort B)	NR (18.4 – NR)	<i>Historical 23.2-25 mo.**</i>

Overall Survival at 12 Months	n Alive/N Total	OS12% (95% CI)
MGMT Unmethylated (Cohort A)	27/32	84.4 (67.2-94.7)
MGMT Methylated (Cohort B)	17/20	85.0 (62.1-96.8)
Combined	44/52	84.6 (71.9-93.1)

Overall Survival at 18 Months	n Alive/N Total	OS18% (95% CI)
MGMT Unmethylated (Cohort A)	16/32	50 (31.9 - 68.1)
MGMT Methylated (Cohort B)	14/20*	70 (45.7 – 88.1)
Combined	30/52	57.7 (14.5 – 71.3)

NR: not reached

*Two patients in Cohort B withdrew consent for additional follow-up at Week 3 and were considered deceased

**Gilbert et al. J Clin Oncol 2013; Gilbert et al. N Eng J Med 2014. Comparison is limited due to differences in study population

Management & Financials



Experienced Executive Team and Board of Directors



J. Joseph Kim, Ph.D.
President & CEO

- Decades of biotech/ pharma management
- Merck: hepatitis A and B vaccines manufacturing; HIV vaccine (Ad5) R&D



Peter Kies
CFO

- Ernst & Young
- Experience with growth companies



Jacqueline Shea, Ph.D.
COO

- Former CEO/COO of Aeras
- Held management positions at Emergent BioSolutions and Microscience Ltd.



Laurent Humeau, Ph.D.
CSO

- Extensive R&D leadership exp. in vaccine, cell and gene therapy developments at Intrexon and VIRxSYS

Board of Directors

Simon X. Benito

Chairman of the Board, Former SVP, Merck Vaccine Division

Roger Dansey, M.D.

Former Head of Late-Stage Oncology at Merck & Co.

J. Joseph Kim, Ph.D.

President & CEO, INOVIO Pharmaceuticals

Ann. C. Miller, M.D.

Former Head of Sanofi Oncology Global Marketing

Jay Shepard

Former President & CEO, Aravive

David B. Weiner, Ph.D.

Executive VP, Director, Vaccine Center, The Wistar Institute

Wendy L. Yarno, Ph.D.,

Former Executive VP and Chief Marketing Officer, Merck

Lota S. Zoth

Former CFO, MedImmune

Well-Balanced Pipeline of Milestones Supported by Strong Balance Sheet

NASDAQ:INO

\$518.6M

Cash and short-term investments

As of March 31, 2021

209.3M

Common stock shares outstanding

As of March 31, 2021

INO-4800

- ✓ Dec. 2020: Published Phase 1 data from first cohort in The Lancet's EClinicalMedicine
- ✓ January 2021: Fully enrolled 640 patient Phase 2 clinical trial in China
- ✓ 1Q21: Completed enrollment for INNOVATE Phase 2 segment
- ✓ 2Q21: Report data from Phase 2 segment of INNOVATE (INovio INO-4800 VAccine Trial for Efficacy)
- ☐ 2Q21: Initiate Global Phase 3 segment of INNOVATE (*U.S. sites pending clinical hold lift by FDA for device*)

VGX-3100

- ✓ 1H21: REVEAL 1 Phase 3 top-line efficacy & tolerability data
- ☐ 2021: Initiate Phase 3 trials for VIN/AIN; attain orphan drug designation

INO-5401

- ✓ 4Q20: OS18 data from Phase 1/2 GBM clinical trial (INO-5401 plus Libtayo®)
- ☐ 2021: OS24 and additional immunology data

Platform Development

- ✓ 4Q20: Awarded two-year grant from DARPA to advance COVID-19 dMAb candidate
- ✓ 1Q21: Initiate Phase 2 field study for Lassa with INO-4500 funded by CEPI
- ☐ 2021: Initiate Phase 2 MERS study with INO-4700 funded by CEPI



INOVIO
POWERING DNA MEDICINES™