



IMUNON

***Harnessing the Power of
the Immune System***

Nasdaq: IMNN

**R&D Day
September
14, 2023**

Corinne Le Goff, Pharm.D, M.B.A.
President & CEO IMUNON



Safe Harbor Statement

This presentation and any statements made during any presentation or meeting contain forward-looking statements related to Imunon, Inc. (“Imunon”) under the safe harbor provisions of Section 21E of the Private Securities Litigation Reform Act of 1995 and are subject to risks and uncertainties that could cause actual results to differ materially from those projected. These statements may be identified by the use of forward-looking words such as "anticipate," "planned," "believe," "forecast," "expected," and "intend," among others. There are many factors that could cause actual events to differ materially from those indicated by such forward-looking statements. Such factors include, among other things, unforeseen changes in the course of research and development activities and in clinical trials; possible changes in cost, timing and progress of development, preclinical studies, regulatory submissions; Imunon’s ability to obtain and maintain regulatory approval of any of its product candidates; possible changes in capital structure, future working capital needs and other financial items; changes in approaches to medical treatment; introduction of new products by others; success or failure of our current or future collaboration arrangements, possible acquisitions of other technologies, assets, or businesses; the ability to obtain additional funds for operations; the ability to obtain and maintain intellectual property protection for technologies and product candidates and the ability to operate the business without infringing the intellectual property rights of others; the reliance on third parties to conduct preclinical studies or clinical trials; the rate and degree of market acceptance of any approved product candidates; possible actions by customers, suppliers, potential strategic partners, competitors, and regulatory authorities; compliance with listing standards of The Nasdaq Capital Market; and those risks listed under “Risk Factors” as set forth in Imunon’s most recent periodic reports filed with the Securities and Exchange Commission, including Imunon’s Form 10-K for the year ended December 31, 2022.

While the list of factors presented here is considered representative, no such list should be considered to be a complete statement of all potential risks and uncertainties. Unlisted factors may present significant additional obstacles to the realization of forward-looking statements. Forward-looking statements included herein are made as of the date hereof, and Imunon does not undertake any obligation to update publicly such statements to reflect subsequent events or circumstances except as required by law.

Today's Agenda

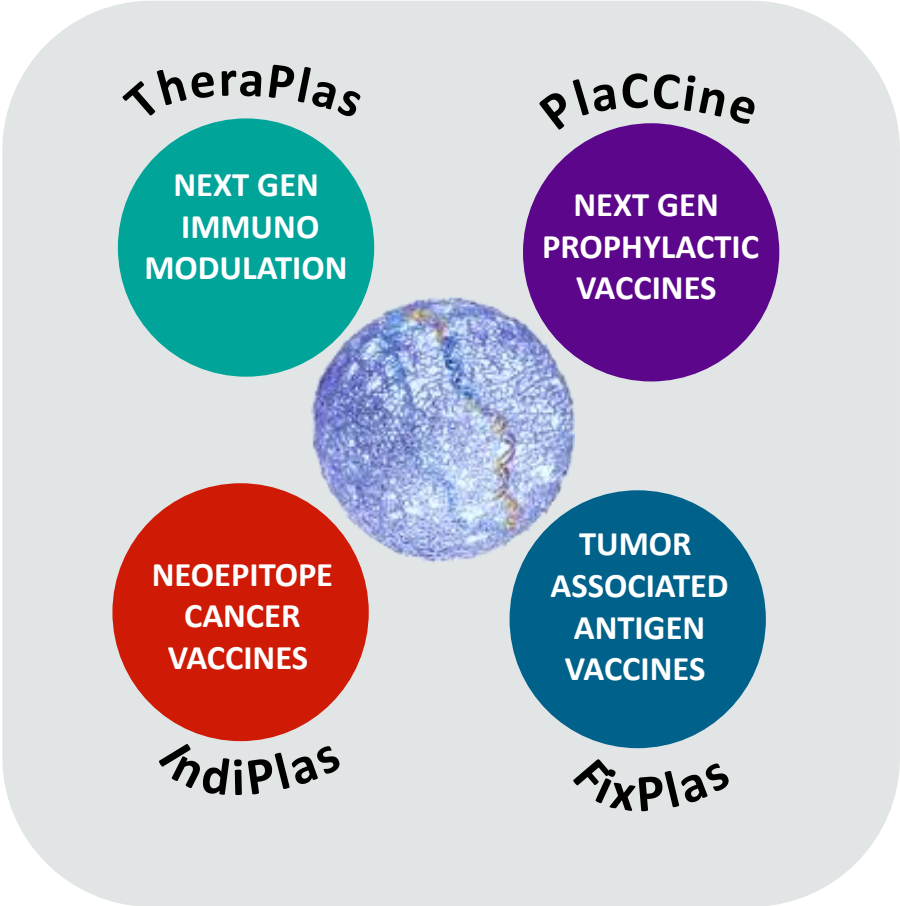
Introduction	Corinne Le Goff, Pharm.D, PhD <i>President & CEO IMUNON</i>	<i>4:00 – 4:15pm</i>
The Vaccines of the Future	Sallie Permar, MD, PhD <i>Nancy C. Paduano Professor and Chair Weill Cornell Medicine</i>	<i>4:15 – 4:35pm</i>
IMNN-101 Development Plan	Khursheed Anwer, PhD <i>CSO</i>	<i>4:35 – 4:45pm</i>
Immuno-Oncology: The remaining unmet need	Patrick Ott, MD, PhD <i>Associate Professor, Medicine, Harvard Medical School Clinical Director, Melanoma Center, Dana-Farber Cancer Institute</i>	<i>4:45 – 5:05pm</i>
Intra-tumoral immuno-oncology and cancer vaccine programs	Khursheed Anwer, PhD <i>CSO</i>	<i>5:05 - 5:15pm</i>
Q&A and Closing Remarks		<i>5:15 – 5:30pm</i>

Transforming Medicine with Our Disruptive Non-Viral DNA Technology

Proprietary Synthetic Delivery and Facilitating System, that promotes DNA Protection, Uptake, Bioavailability and Enhanced Antigen Expression

Gene Therapy

Personalized Cancer Vaccine



Prophylactic Vaccine

Off- the-shelf Cancer Vaccine

IMUNON Strategy to Build a Fully Integrated Biotech Company



FOCUS ON IMMUNO-ONCOLOGY
Cytokines Coding & Cancer Vaccines



GMP MANUFACTURING
In-house Early Development Scale



FOCUS ON INFECTIOUS DISEASES
Pathogen Antigens Coding Vaccines



WORLD CLASS PARTNERS
To Expand our R&D Capabilities



M&A POTENTIAL
Synergistic with our Capabilities



STRONG BALANCE SHEET
To Support our Strategy into 2025

IMUNON's Technology Advantages in Prophylactic Vaccines



Durability of protection

Durable antigen expression

Induces robust immunological response



Speed

Non-viral DNA is a platform

Ability to go from sequence to the clinic to approved products in record time

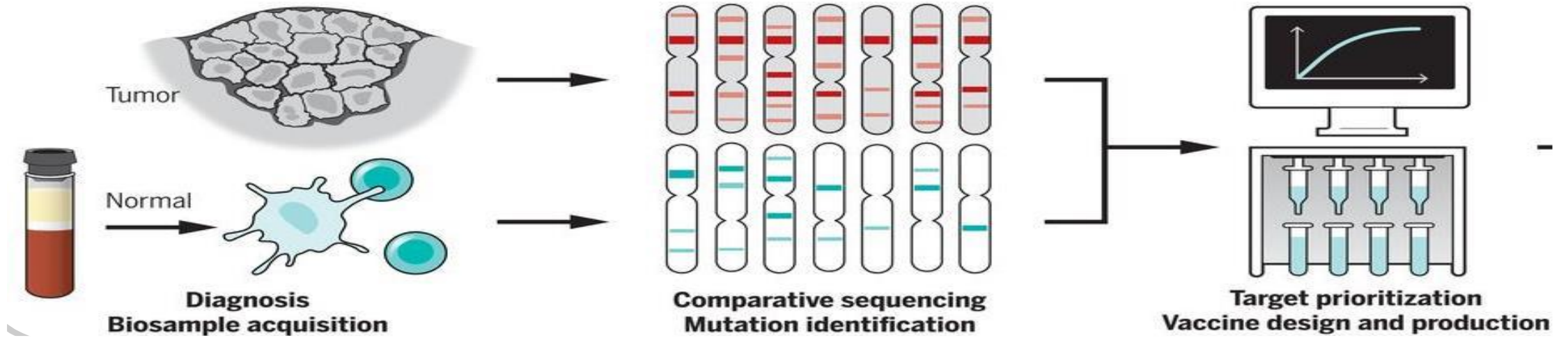


Flexible manufacturing

Simple handling & distribution

Stability and long shelf-life at workable temperatures - Greater Capital Efficiency

IMUNON's Technology Advantages in Cancer Vaccines






Non-Viral DNA Inducing Durable Antigen Expression

Potent T cell response
Repeat Administration

DNA encoded Cancer Vaccines

Shared Antigens
Individualized Antigens

IMUNON's Pipeline of DNA-based Transformative Medicines

Modality	Program	Indication(s)	Discovery	IND enabling	Phase 1	Phase 2	Partnerships	
TheraPlas	IL-12 (OVATION) Intraperitoneal (IP)	Advanced Ovarian, Fallopian Tube or Primary Peritoneal Cancer	IMNN-001 (formerly GEN-1)					
	IL-12 IP in combination with bevacizumab	Advanced Ovarian, Fallopian Tube or Primary Peritoneal Cancer	IMNN-001 + bevacizumab					 #RadicalCollaboration
PlaCCine	Multicistronic SARS-CoV-2. Clinical Proof-of-Concept	COVID-19 Seasonal Vaccine	IMNN-101					
	Prophylactic Vaccine	Infectious Disease target	PL-X					 
FixPlas	Cancer Therapeutic Vaccine	Trp2 /NYESO-1 Tumor Associated Antigen in Melanoma	IMNN-201					
IndiPlas	Individualized Neoantigen Cancer Vaccines		IP-Y					

Financial Summary & Upcoming Key Milestones:

Robust Flow of Value Creating Activities



Cash , Cash Equivalents & Investments

\$24.1M

As of June 30, 2023



Shares Outstanding

10.4M



Estimated Operating Expenses per quarter

\$4.5M

As of March 31, 2023

IMNN-001 OVATION 2
Interim Data

IMNN-001 OVATION 2
Topline Results

Interim Results
IMNN-001 + bevacizumab

IMNN-102
Pre-IND

IMNN-101
SARS-CoV-2 Booster IND
And start of Phase 1/2

IMNN-201
POC Data

2H
2023

1H
2024

Dr. Sallie Permar, MD, Ph.D.
Nancy C. Paduano Professor and Chair
Weill Cornell Medicine



Dr. Khursheed Anwer, Ph.D.
CSO IMUNON





Dr. Patrick Ott, MD, Ph.D.
Associate Professor, Medicine
Harvard Medical School
Clinical Director, Melanoma Center
Dana-Farber Cancer Institute

Dr. Khursheed Anwer, Ph.D.
CSO IMUNON





Vaccines of the Future

Sallie Permar, MD, PhD

**Nancy C. Paduano Professor and Chair, Weill Cornell Medicine
Pediatrician-in-Chief, NewYork-Presbyterian/Weill Cornell Medical Center**

[@salliepermar](#)



**Weill Cornell
Medicine**

NewYork-Presbyterian
Komansky Children's Hospital

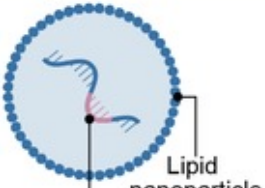
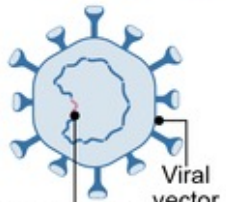

Dr. Permar is a consultant for Pfizer, Merck, Moderna, GSK, Hoopika, and Dynavax CMV vaccine programs



**Weill Cornell
Medicine**

NewYork-Presbyterian
Komansky Children's Hospital

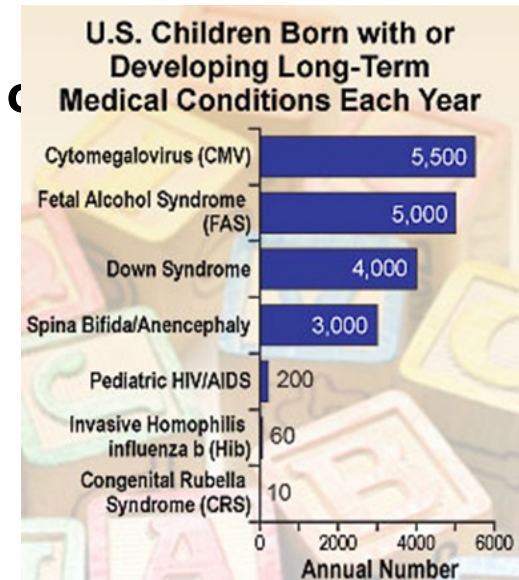
Highly successful SARS-CoV-2 mRNA/LNP vaccine developed in “warp speed”

	mRNA platform	Replication-defective live-vector platform	Recombinant-subunit-adjuvanted protein platform
	 <p>Lipid nanoparticle SARS-CoV-2 spike RNA</p>	 <p>Viral vector SARS-CoV-2 spike gene</p>	 <p>SARS-CoV-2 spike protein</p>
Description	Encapsulated genetic instructions that allow vaccinated individuals to produce the spike protein of SARS-CoV-2 to stimulate immune system but cannot cause COVID-19.	Non-replicating virus that delivers genetic instructions to allow vaccinated individuals to produce the spike protein of SARS-CoV-2 to stimulate immune system but cannot cause COVID-19.	Fully-formed spike protein of SARS-CoV-2 delivered with adjuvant, which helps to stimulate immune system of vaccinated individuals but cannot cause COVID-19.
Operation Warp Speed candidates (most advanced clinical trial phase)	Moderna (phase 3) Pfizer/BioNTech (phase 3)	Janssen (phase 3) AstraZeneca (phase 3)	Sanofi/GSK (phase 2) Novavax (phase 3)


Source: GAO (analysis). Adaptation of images depicting vaccine technologies with permission from Springer Nature: *Nature* ("The Race for Coronavirus Vaccines: A Graphical Guide," Ewen Callaway) © 2020. | GAO-21-319

Congenital Cytomegalovirus (CMV)

- **Most common congenital infection and cause of birth defects worldwide**
 - 1/200 (0.5%) live-born infants globally
 - 40,000 cases annually in US, ~5,000 have permanent sequelae
 - 4 billion cost annually in US
- **Leading cause of pediatric neurologic deficits**
 - Up to 25% of infant hearing loss due to CMV
- **Tier 1 priority vaccine by the National Academy of Medicine for >20 years**



50+ years of CMV vaccine development, yet we still await an approved vaccine

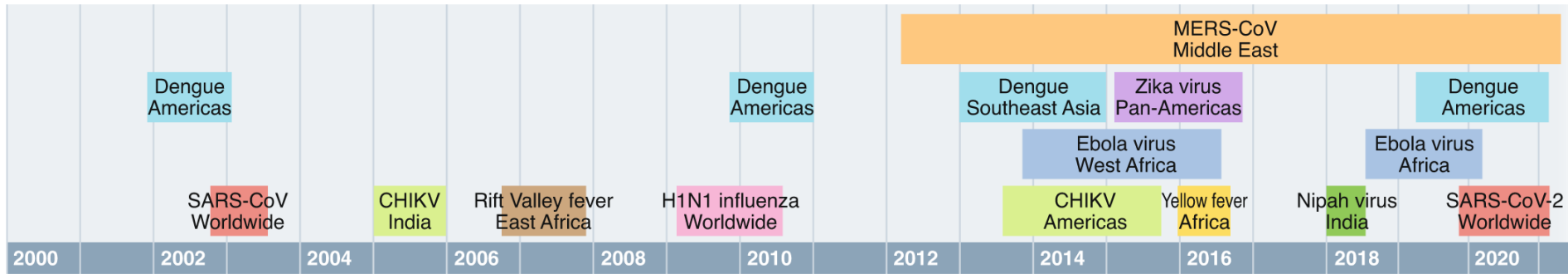


Vaccine Platform	Phase	Efficacy
Live attenuated	Phase I,II	Reduction of disease in renal tx
Live viral vectors	Phase I	-
gB subunit	Phase I,II	50%
eVLP	Phase I	-
Single round DISC vectors	Phase I,II	42%
PC subunit/MVA	Phase I	-
DNA	Phase I,II,III	ongoing
mRNA (gB + PC)	Phase I,II,III	ongoing

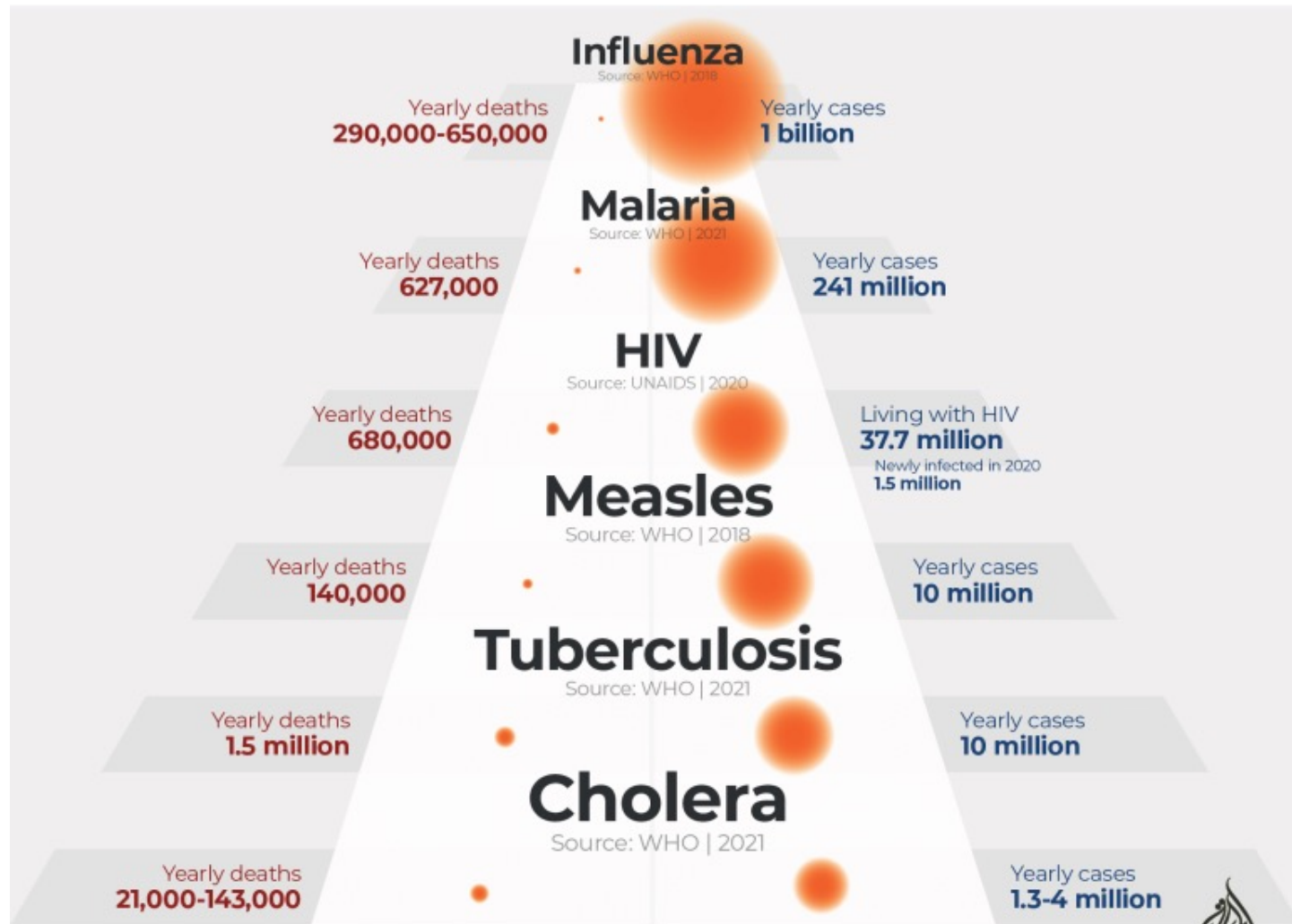
Modified from Schleiss et al, 2017

The rate of pandemics is increasing

a 21st century viral disease outbreaks



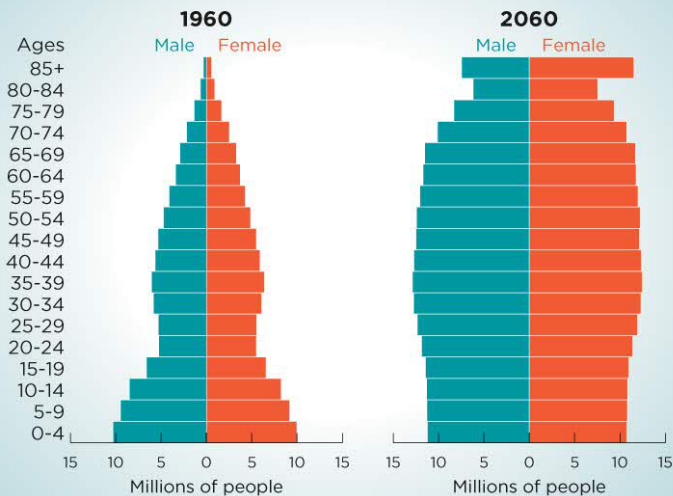
Yet, deadly endemic infections still require novel vaccine efforts



Populations vulnerable to infectious diseases are increasing: elderly and immunocompromised

From Pyramid to Pillar: A Century of Change

Population of the United States

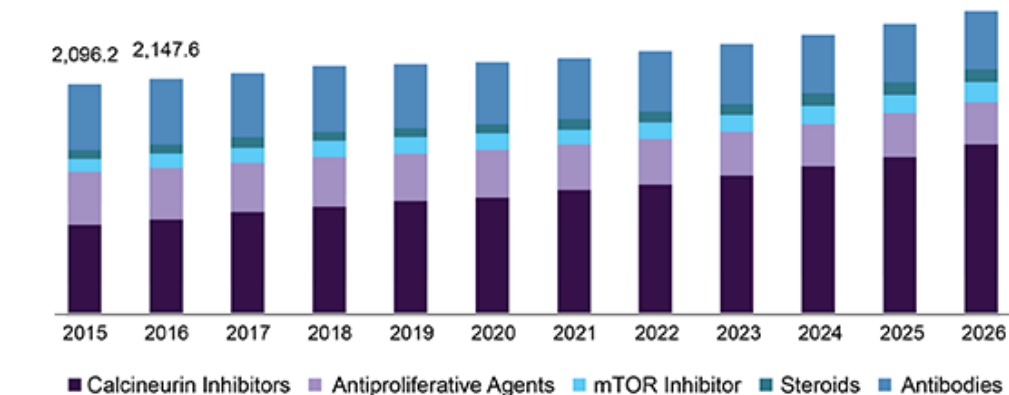


United States[®]
Census
Bureau

U.S. Department of Commerce
U.S. CENSUS BUREAU
census.gov

Source: National Population
Projections, 2017
www.census.gov/programs-surveys
/popproj.html

North America organ transplant immunosuppressant drugs market size, by drug class, 2015 - 2026 (USD Million)



Source: www.grandviewresearch.com

Pregnant women and children frequently more vulnerable to pandemic variants

- **Pregnant/lactating women and children left out of early phase vaccine development**
 - **Prevents innovations from reaching these groups**
- **Children require specific safety and dose ranges, often done late**
 - **A health disparity that children had delayed access to the SARS-CoV-2 vaccine**




<https://www.aamc.org/news-insights/are-covid-19-vaccines-safe-during-pregnancy-experts-weigh>



Ideal Immunity: Vaccination in infancy for life-long prevention

Why do we vaccinate newborns against Hepatitis B?



Hep B is the most common liver disease in the world

Babies can be infected during delivery from an infected mother, breastfeeding, toothbrushes, nail clippers or child to child through open sores or wounds


The virus can live in a single drop of blood, outside of the body for over a week

In developed countries, approximately half of people with Hep B don't know they have it

Infants infected with Hep B have a 90% chance of the disease becoming chronic

The Hep B vaccine is included on the childhood vaccination schedule as a long term prevention strategy to reduce the illness and death from complications due to the disease and to eventually eliminate HepB altogether

Prior to the development of the vaccine, approximately half of babies with Hep B, were infected through casual contact with people other than their mother

 www.facebook.com/PCCVGN www.hepatitisaustralia.com.au

Hepatitis B vaccine

- 1st dose: Birth
- 2nd dose: 1-2 months
- 3rd dose: 6-18 months

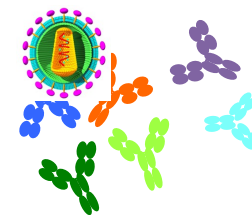
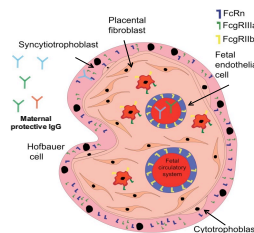
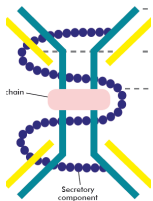
High risk newborn:

- HIBlg + HepB vaccine (passive + active)

Leverage rapidly designed/produced DNA vaccine platforms to design vaccines to protect the next gen

Two Novel Approaches:

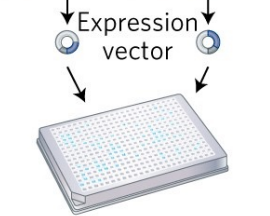
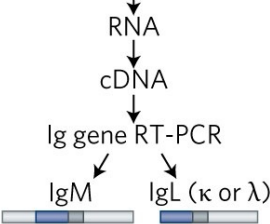
1. **Reverse vaccinology**: Isolation of potent mAbs from pre-screened, infected patients, use to design antigens that will elicit potent, protective responses
2. **Protective Transfer** : Passive immunization with designer mAbs delivered by DNA/RNA technology that are durable and doesn't require IV infusion



New Era of Vaccines: Pandemic Readiness via Reverse Vaccinology

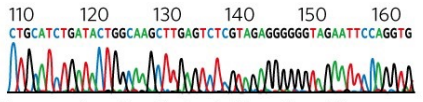
Antigen discovery

Isolation of human monoclonal antibodies from single B cells



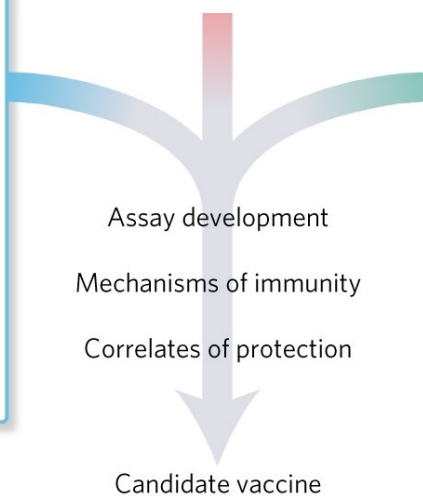
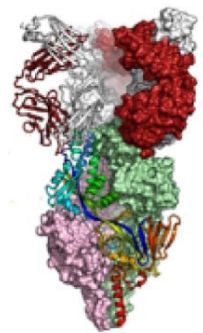
Co-transfection or development of stable cell line for antibody production

Sequencing for viral diversity and escape mutations



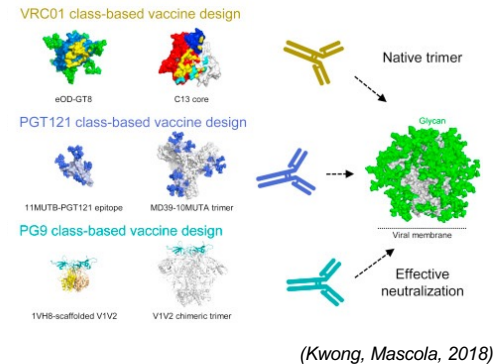
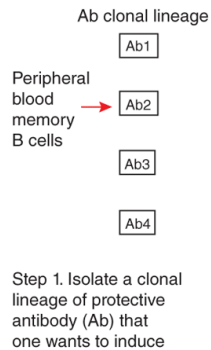
Sequencing B cells to define clonal lineages; TCR-and BCR-specific transcriptome

Structural analysis of antigenic sites on viral surface glycoproteins



Challenging Vaccines: Where natural immunity is not protective

Reverse Engineering 2.0 Strategy: Identify immunogens that can select for B cells with key mutations required for neutralizing IgG development

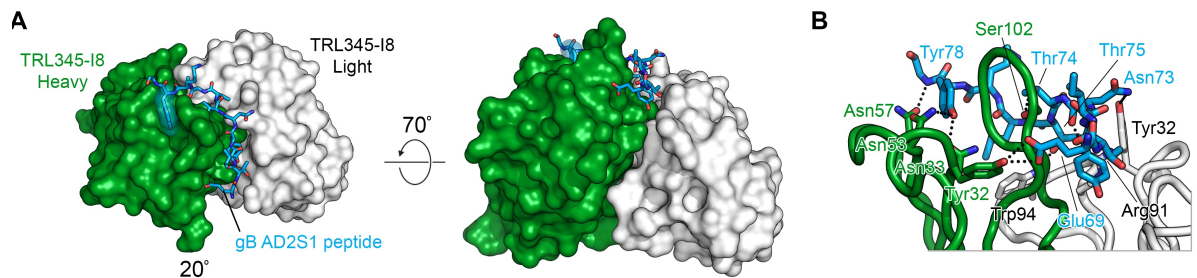
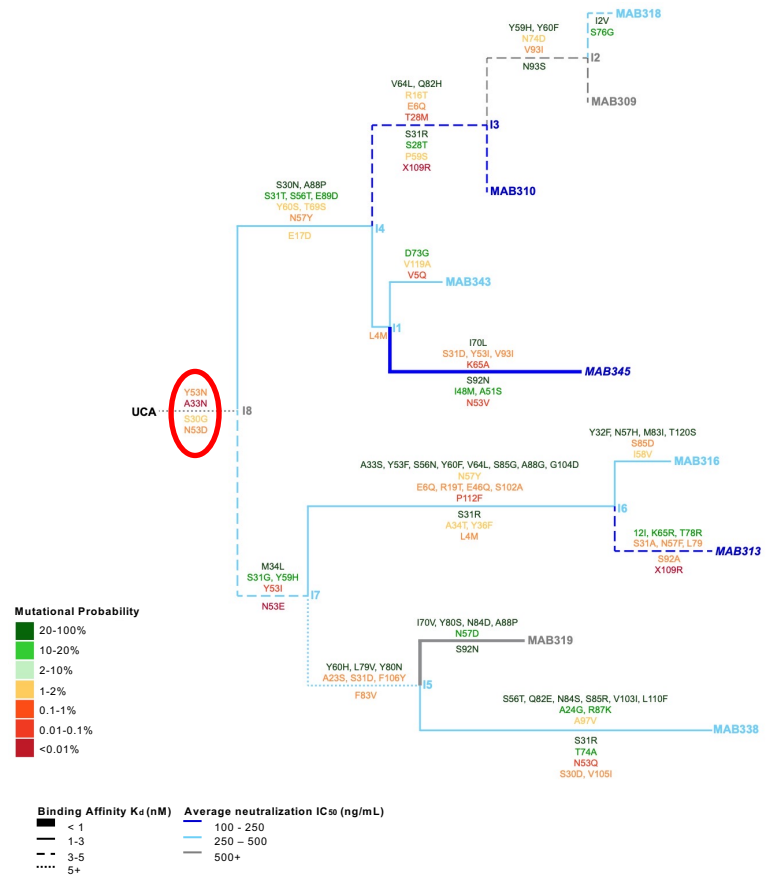


(Haynes, et al. 2012)

Improbable B cell mutations prevent development of potent antibodies after vaccination

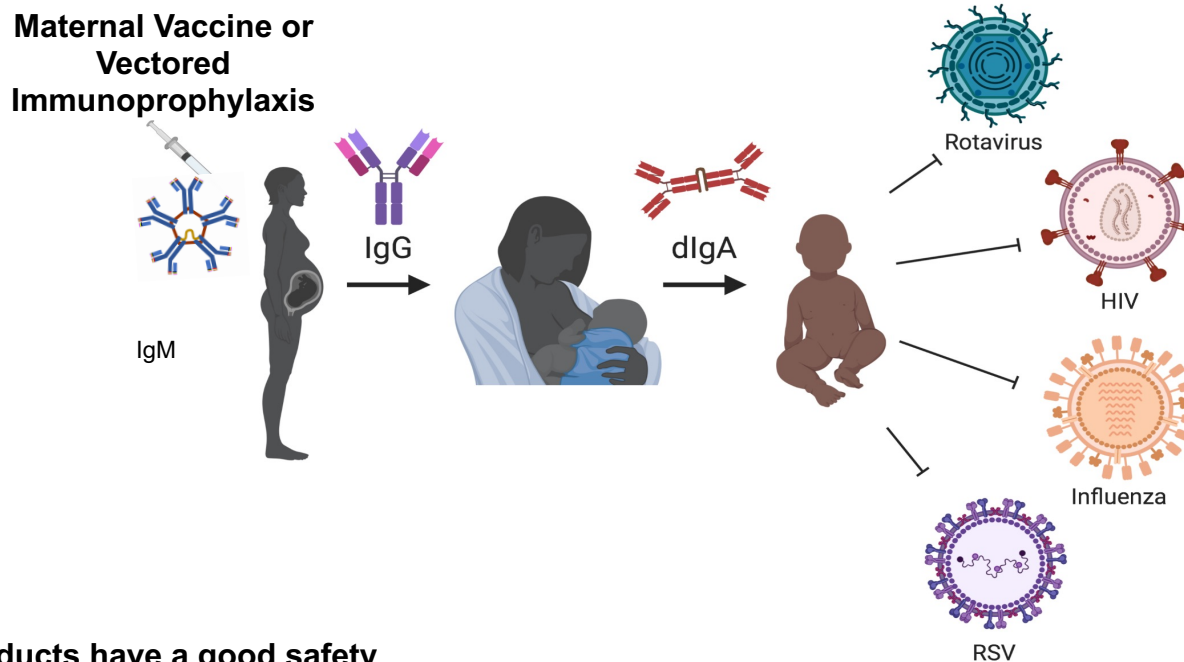
Problem: Improbable B cell mutations may impose a bottleneck on B cell lineage maturation

Solution: Rapidly iterative vaccine design for engaging early/rare B cell lineages



Protective Transfer: Design protective antibodies for delivery to specific sites of virus acquisition

Abs at the maternal/fetal interface



Antibody products have a good safety profile and precedent in pregnancy

- Rh factor incompatibility
- Varicella (VariZIG)

Vertical transmission of Zika Virus

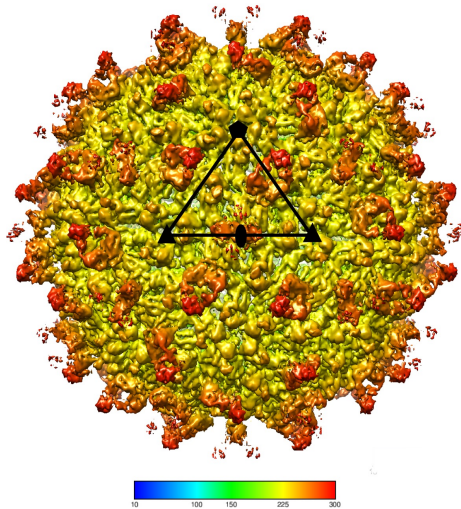
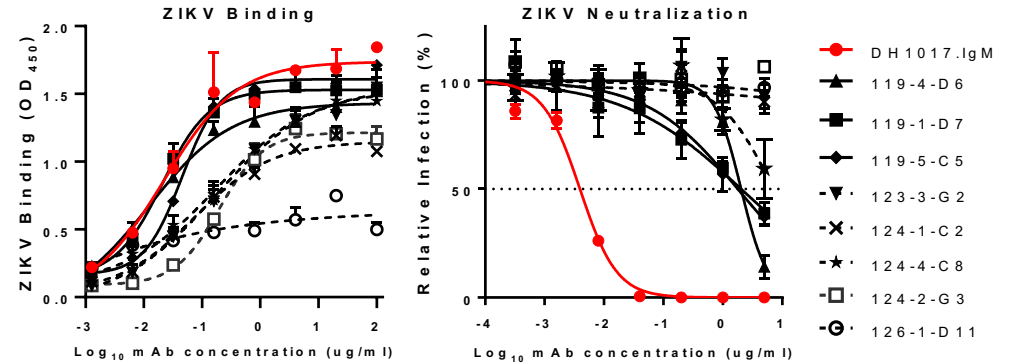
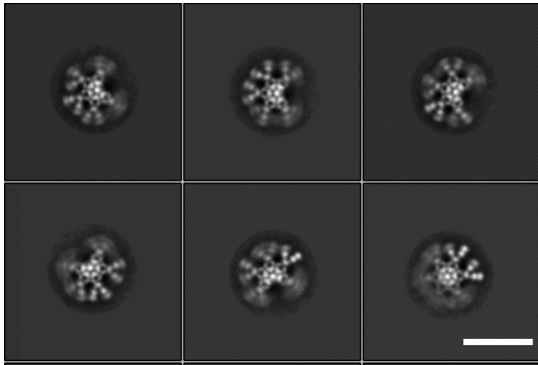
- **1 in 10 pregnant women** with a confirmed Zika infection had a baby with congenital Zika syndrome (US population, CDC)
 - **>11,000 cases** of microcephaly in 2015-16 epidemic
- **Congenital ZIKV symptoms:**
 - Microcephaly
 - Visual and hearing impairment
 - Neurodevelopmental defects
- **No licensed vaccine**
 - Phase 1 trials only performed in non-pregnant subjects



<http://www.bbc.com/news/world-latin-america-37112639>

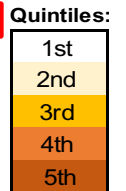
**Need interventions to block
ZIKV infections during
pregnancy**

Ultrapotent ZIKV-neutralizing IgM mAb DH1017.IgM



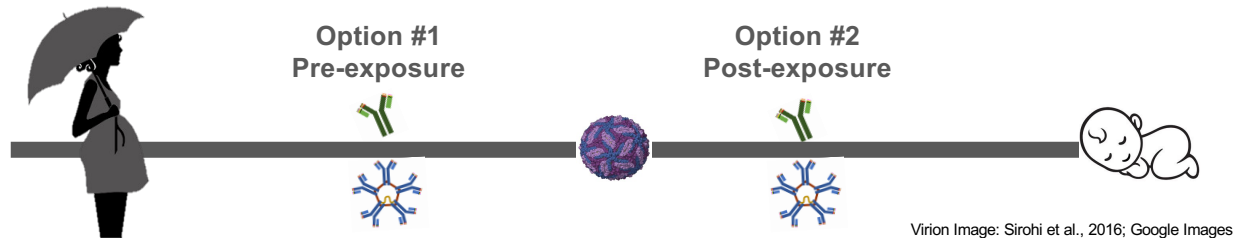
Virus Binding (AUC)

Subject (DPS)	mAb ID	ZIKV	DENV1	DENV2	DENV3	DENV4
B cell derived mAbs						
P73 (71)	DH1017.IgM	187	8	9	8	10
P34 (162)	119-4-D6 (IgG)	144	23	23	71	24
P34 (162)	119-1-D7 (IgG)	153	15	15	26	11
P34 (162)	119-5-C5 (IgG)	163	35	36	63	94
Memory B cell derived mAbs						
P73 (28)	123-3-G2 (IgG)	141	156	226	146	35
P56 (19)	124-4-C8 (IgG)	143	179	218	185	78
P56 (19)	124-1-C2 (IgG)	113	49	180	56	13
P56 (19)	124-2-G3 (IgG)	119	139	197	73	17
P54 (77)	126-1-D11 (IgG)	63	29	25	40	13
Control	Synagis (-)	0.6	0.6	0.6	0.7	0.5

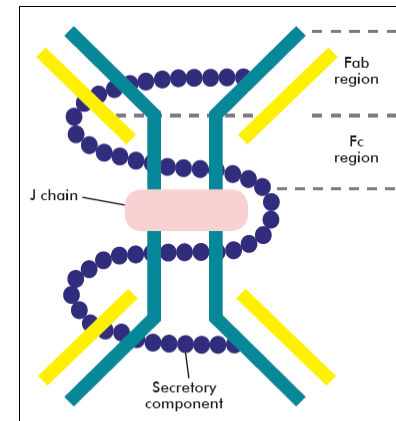
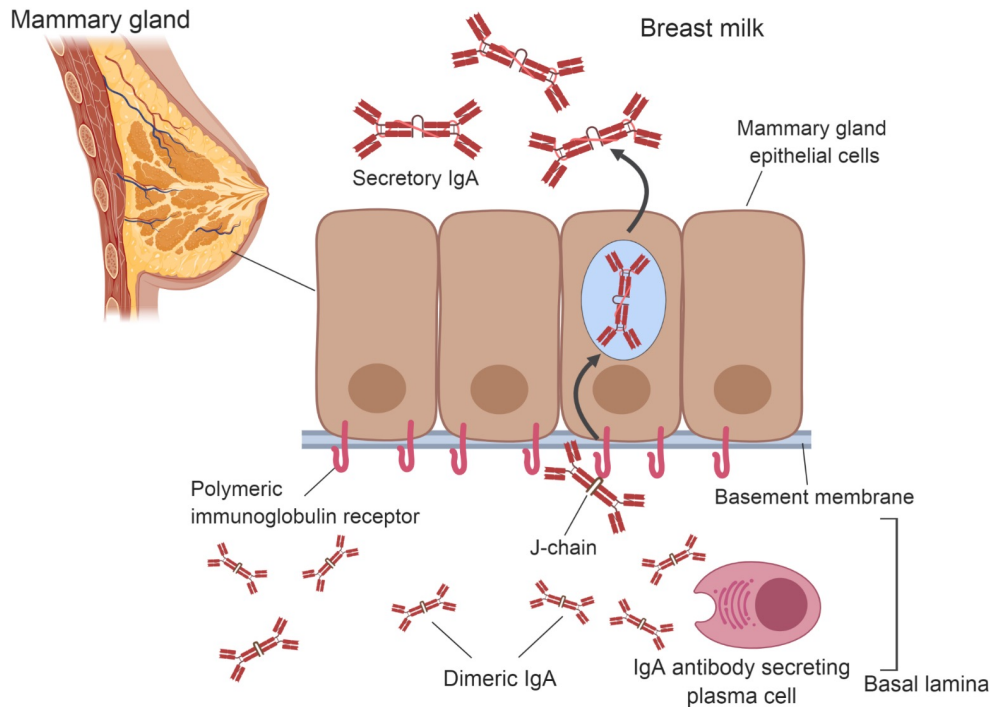


Maternal potent IgM as Zika post or pre-exposure prophylaxis that does not cross the placenta

Potential uses of a ZIKV immunoprophylaxis:



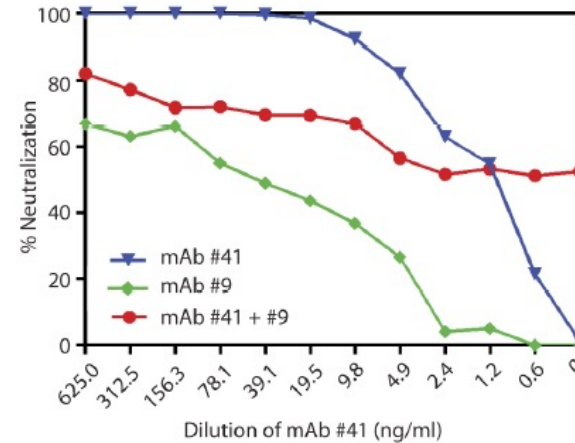
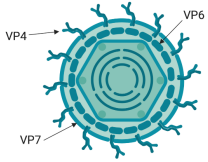
Protective Transfer: maternal dIgA for transfer into breast milk to protect against neonatal enteric pathogens



Design of a maternally administered dIgA to traffic to breast milk and block rotavirus transmission to the infant

mAb#41

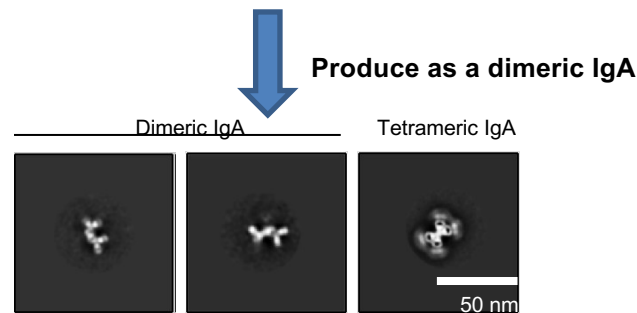
- Isolated from IgA⁺ antibody secreting cells in the intestine
- VP4-specific
- Neutralizing *in vitro* and *in vivo*



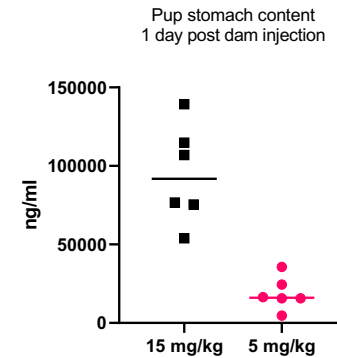
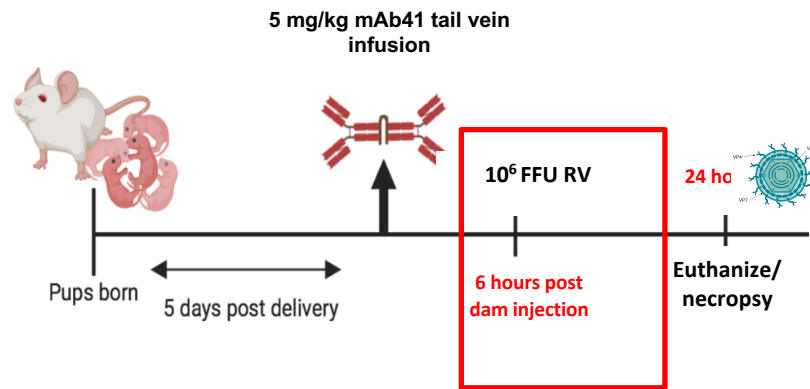
Nair et al., 2017



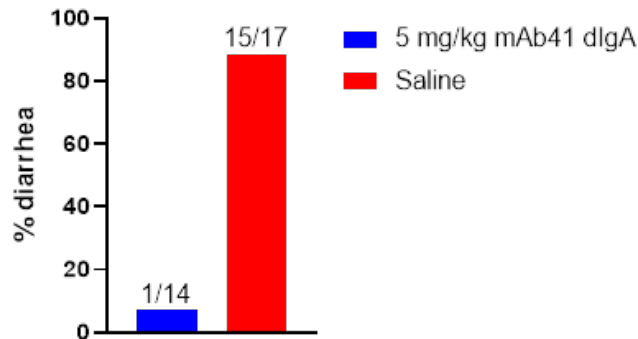
Stephanie Langel, PhD
Maria Blasi, PhD



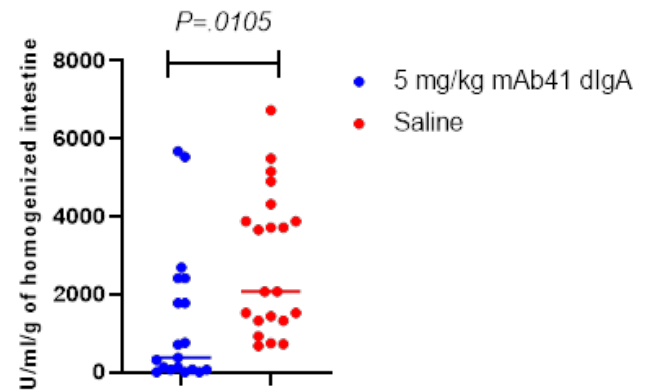
Systemically administered maternal dIgA protect against rotavirus challenge



Protection against diarrhea



Intestinal Viral Load



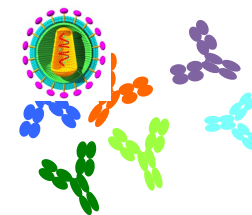
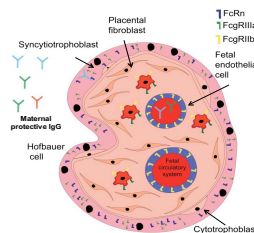
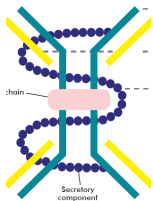
Leveraging rapidly designed/produced DNA vaccine platform for vaccines of the future

Reverse vaccinology: Isolation of potent mAbs from pre-screened, infected patients, use to design antigens that will elicit potent, protective responses

- DNA vaccines could be rapidly and iteratively developed for this approach

Protective Transfer : Passive immunization with designer mAbs

- Delivered by DNA technology that is durable and doesn't require IV infusion



IMNN-101 Development Plan

Dr. Khursheed Anwer, PhD, MBA

CSO IMUNON

IMNN-101– The Lead PlaCCine Product

Next-Generation Covid-19 DNA Vaccine

Addressing the limitations of current vaccines

Key Distinguishing Attributes

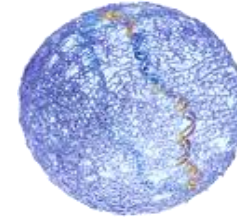
Durable Ag expression

Stable at working temp

Plug & Play design for rapid response

Antigen DNA Plasmid

Omicron XBB1.5 Spike Antigen



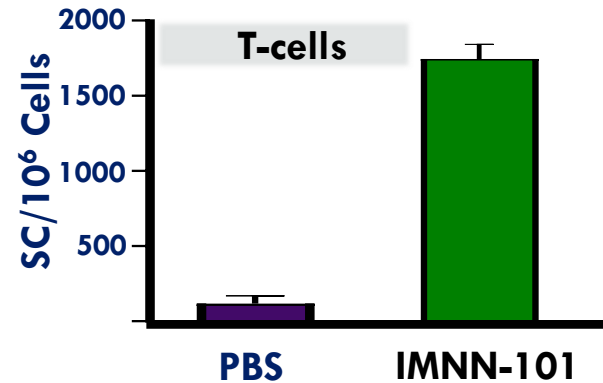
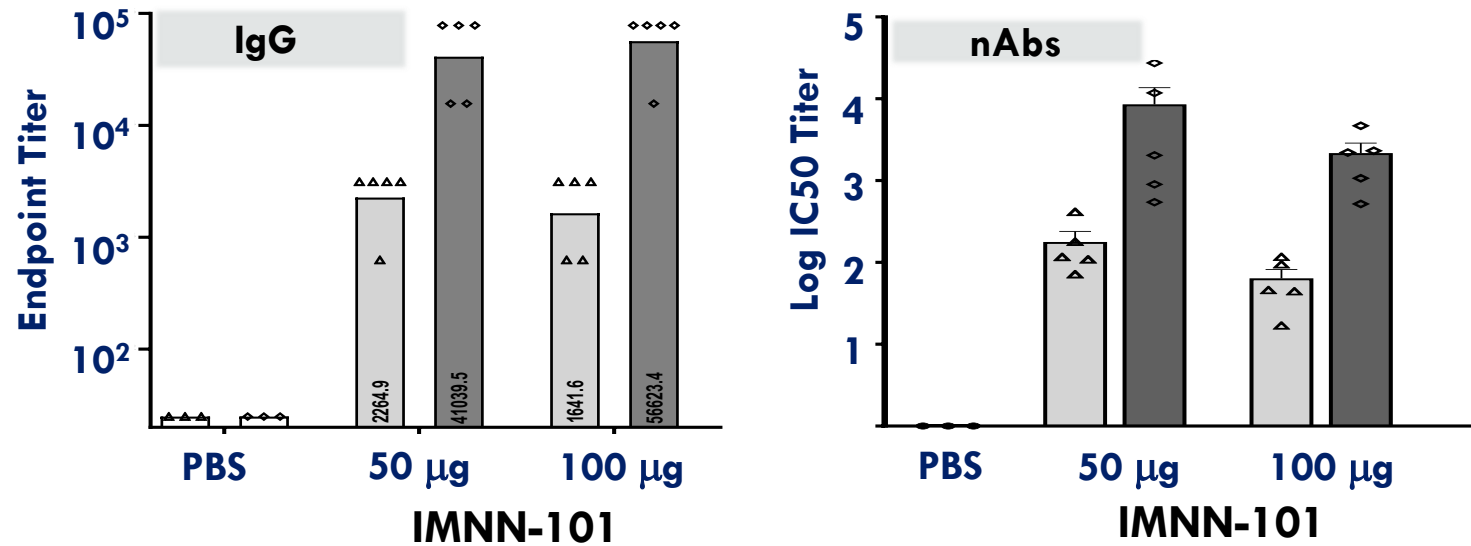
Synthetic DNA Delivery System

Safe & efficient delivery
- non-viral, non-device

IMNN-101– Potentially the First Vaccine Capitalizing on DNA Advantages

IMNN-101– Evidence of Robust Immunogenicity in a Mouse Model

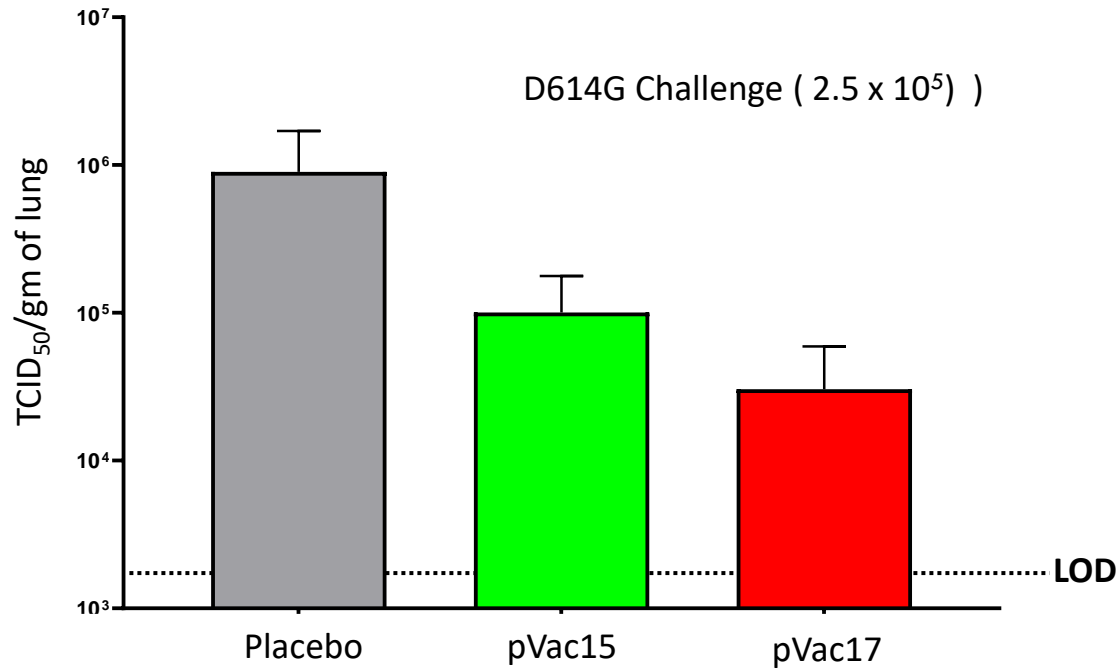
Prime & Boost



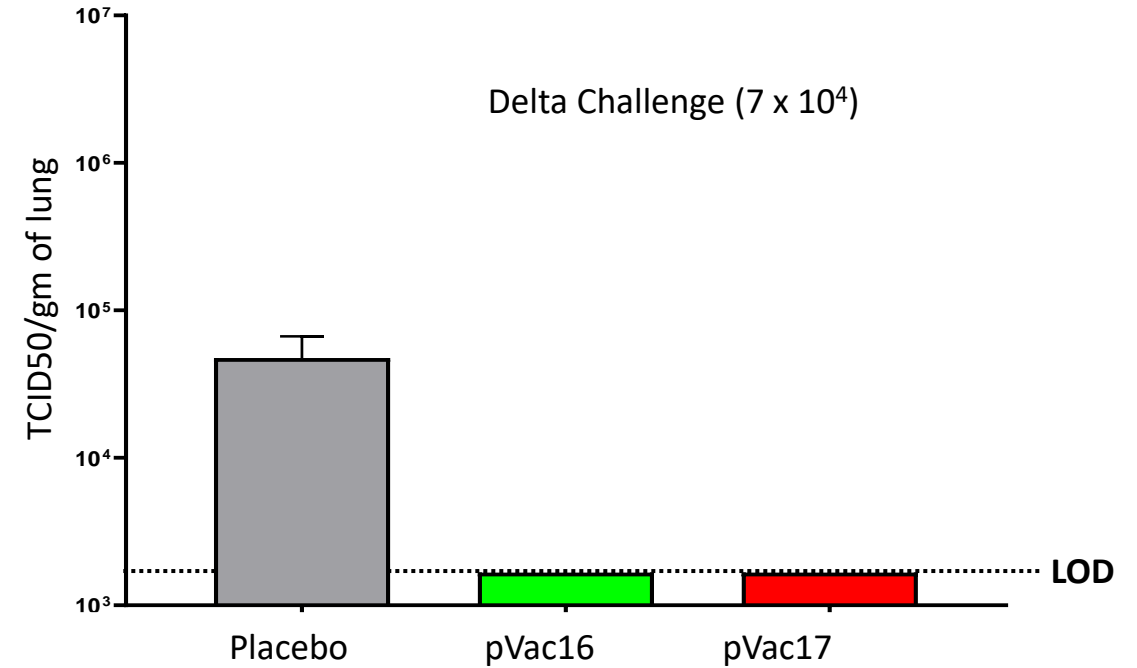
Over 90% Protection From Live Viral Challenge in Mice

IMNN-101 Prototypes – Early Spike Variants

TCID50 Tissue Culture Infection Dose



IMNN-101 Prototype Vaccines

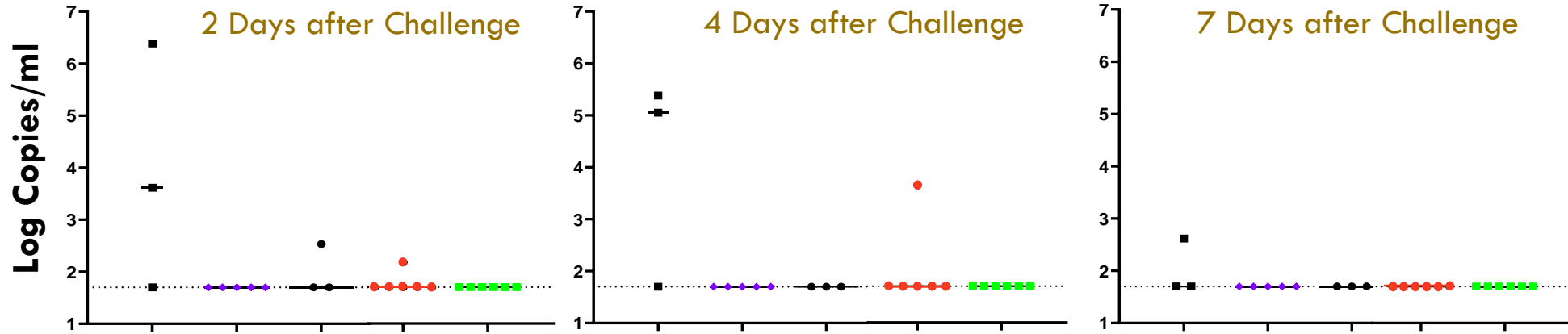


IMNN-101 Prototype Vaccines

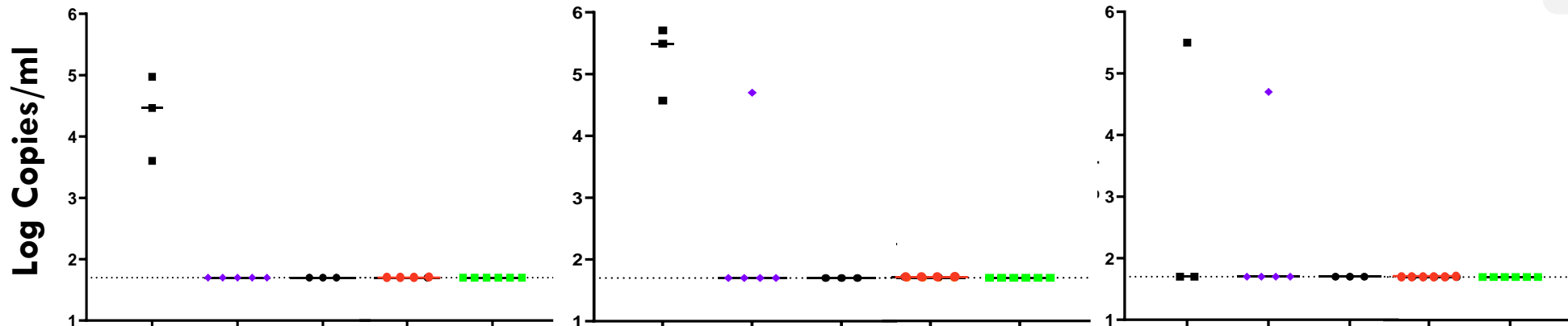
Complete & Comparable Viral Clearance to mRNA Vaccine in Monkeys

IMNN-101 Prototypes- Early Spike Variants

Bronchoalveolar Lavage



Nasal Swab



IMNN-101 Prototype Vaccines

- Placebo
- ◆ pVac-15 (1, 1 mg)
- pVac-16 (2, 2 mg)
- pVac-15 (5, 1 mg)
- mRNA (100, 100 µg)

Prime + boost vaccination

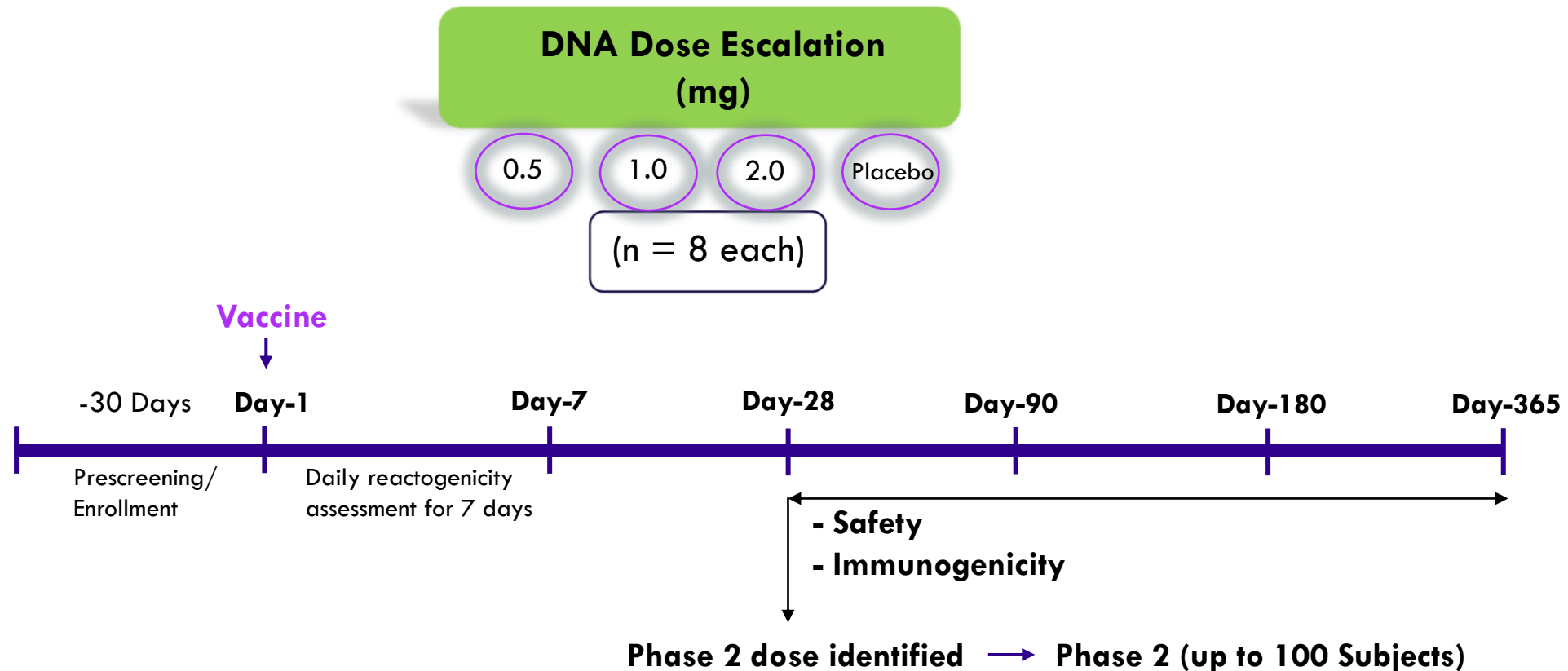
IMNN-101 – Development Status

Preclinical Proof of Concept	
IND-Enabling Studies	
Safety Tox	4Q 2023
Biodistribution	4Q 2023
Clinical Lot	1Q 2024
IND Filing	1Q 2024
Phase 1	April 2024
Phase 2	June 2024

IMNN-101 – FDA-Reviewed Clinical Development Plan

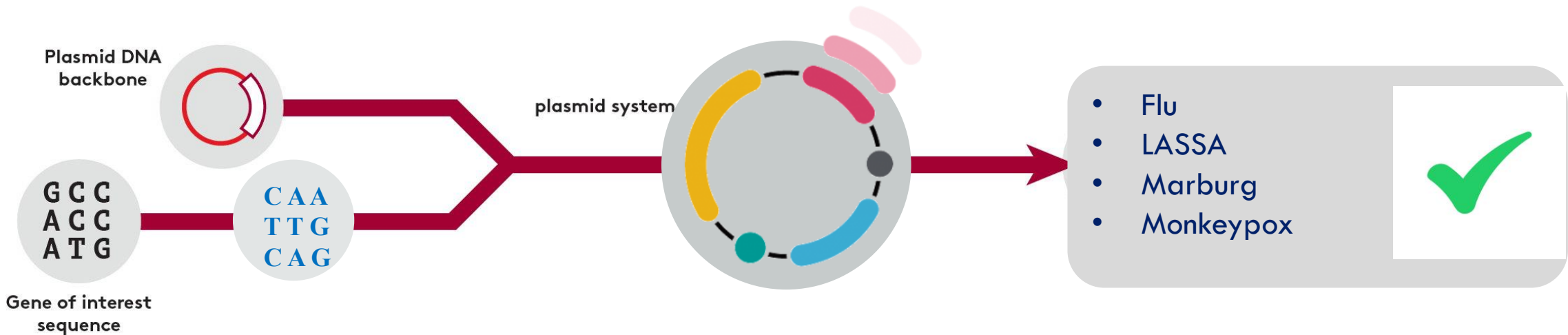
Phase 1/2 Trial in Healthy Subjects: Single Dose, Placebo Controlled

Rapid Dose Escalation Followed by Expanded Phase 2 Approach for Speedy Completion



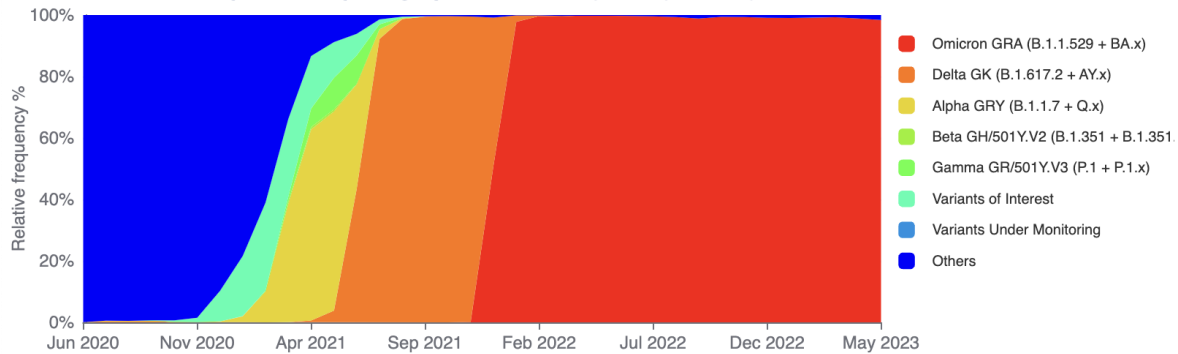
Plug & Play Allows for Rapid Response to New & Urgent Vaccines

Proof of Concept Against Multiple Pathogens



Antigen Coding Cassettes

SARS-CoV2 Variants



Active Vaccine Partnerships

- NIAID/NIH – LASSA
- Wistar Institute – Flu, Marburg

Personal(ized) Immunotherapy

Patrick A Ott, MD, PhD

Clinical Director, Melanoma Center
Dana Farber Cancer Institute
Associate Professor of Medicine
Harvard Medical School



BRIGHAM AND
WOMEN'S HOSPITAL



Dana-Farber
Cancer Institute



HARVARD
MEDICAL SCHOOL

Disclosures

Advisory Role: Alexion, Array, Bristol-Myers Squibb, Celldex, CytomX, Genentech, Merck, Neon Therapeutics, Novartis, Pfizer, TRM Oncology, Evaxion, Immunetune, Imunon, Servier

Grants to institution: Armo Biosciences, AstraZeneca/MedImmune, Bristol-Myers Squibb, Celldex, CytomX, Genentech, Merck, Neon Therapeutics, Novartis, Pfizer

Speaking Engagement: Medscape

What is personalized Immunotherapy, really?

Personalized Immunotherapy in the neoadjuvant setting: Use of biomarkers to adapt therapy

Baseline **biomarker**

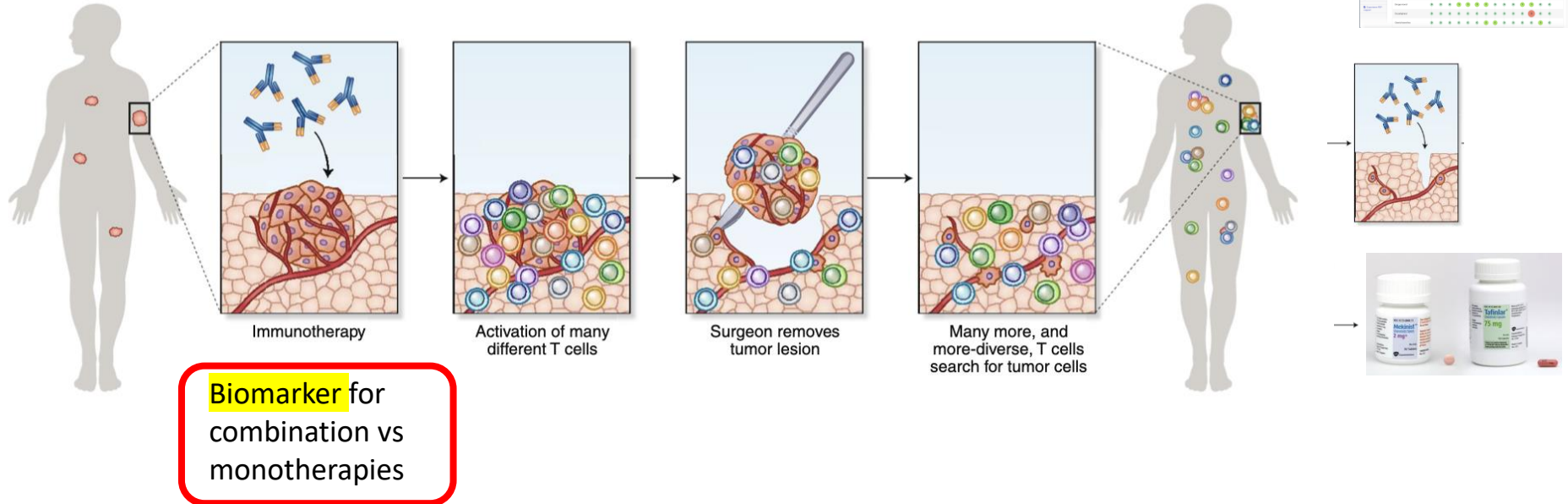
- response
- toxicity

On treatment **biomarkers** for

early Tx adjustment

Pathologic response

- **prognosticator**
- +/- adj therapy



Personalized Immunotherapy in the neoadjuvant setting: Use of biomarkers to adapt therapy

Baseline **biomarker**

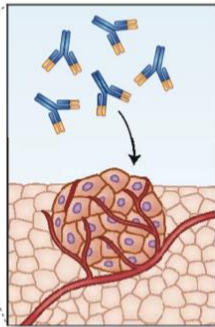
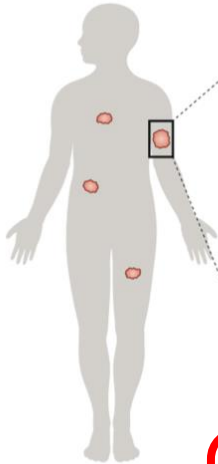
- response
- toxicity

On treatment **biomarkers** for

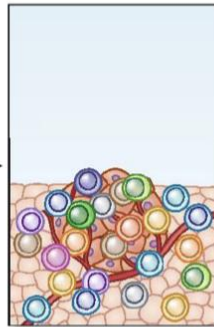
early Tx adjustment

Pathologic response

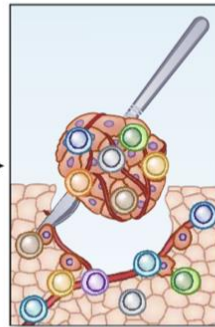
- **prognosticator**
- +/- adj therapy



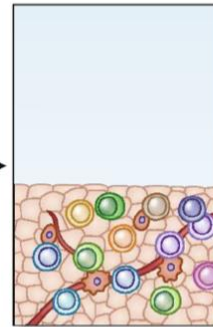
Immunotherapy



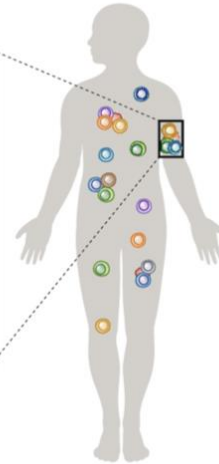
Activation of many different T cells



Surgeon removes tumor lesion

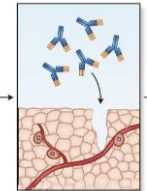
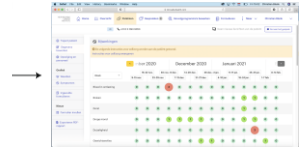


Many more, and more-diverse, T cells search for tumor cells



Biomarker for combination vs monotherapies

Extent of surgery



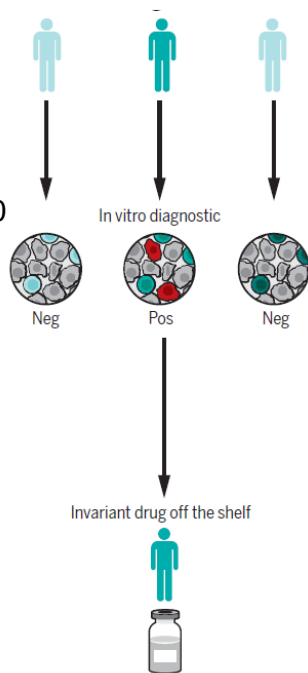
Adjuvant therapy

Monitoring

“Stratified” vs. “Personalized Therapy”

Stratified

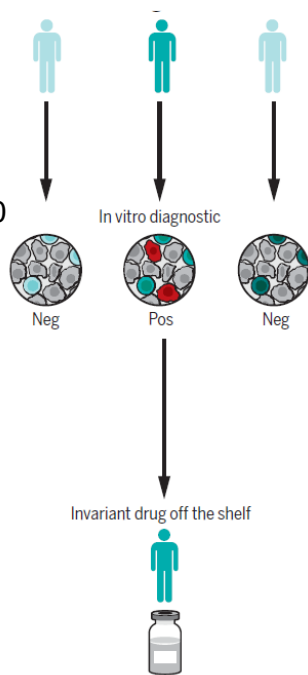
- Tebentafusp – HLA-A2
- BRAF/MEKi – BRAF^{V600}
- EGFR/ALK/METi



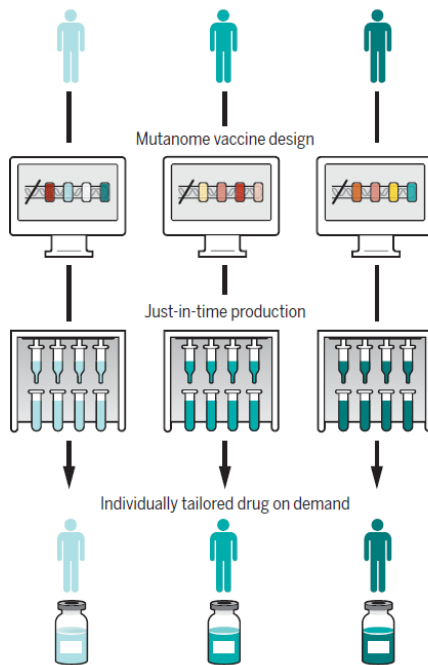
“Stratified” vs. “Personalized Therapy”

- Tebentafusp – HLA-A2
- BRAF/MEKi – BRAF^{V600}

Stratified

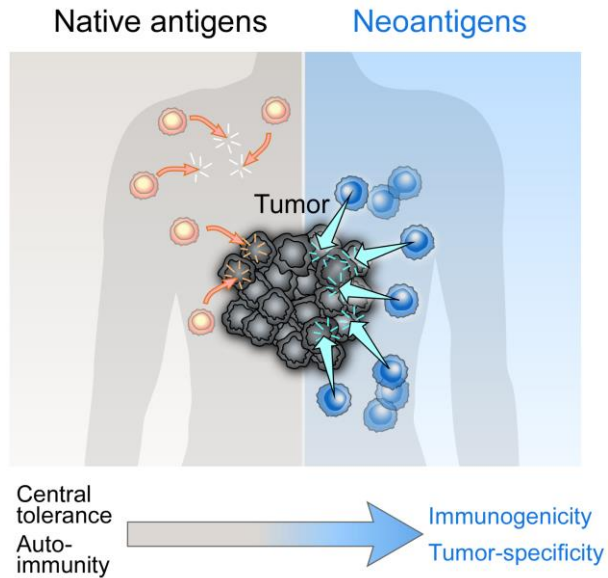


Personalized – the drug is custom-made for each patient

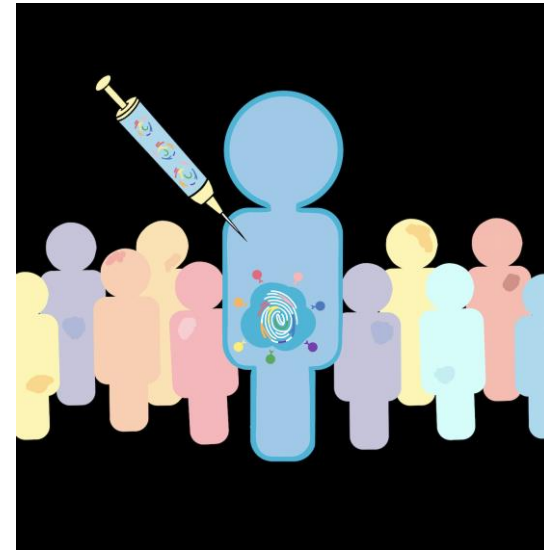


- Personalized Vaccines
- Adoptive T cell Therapy

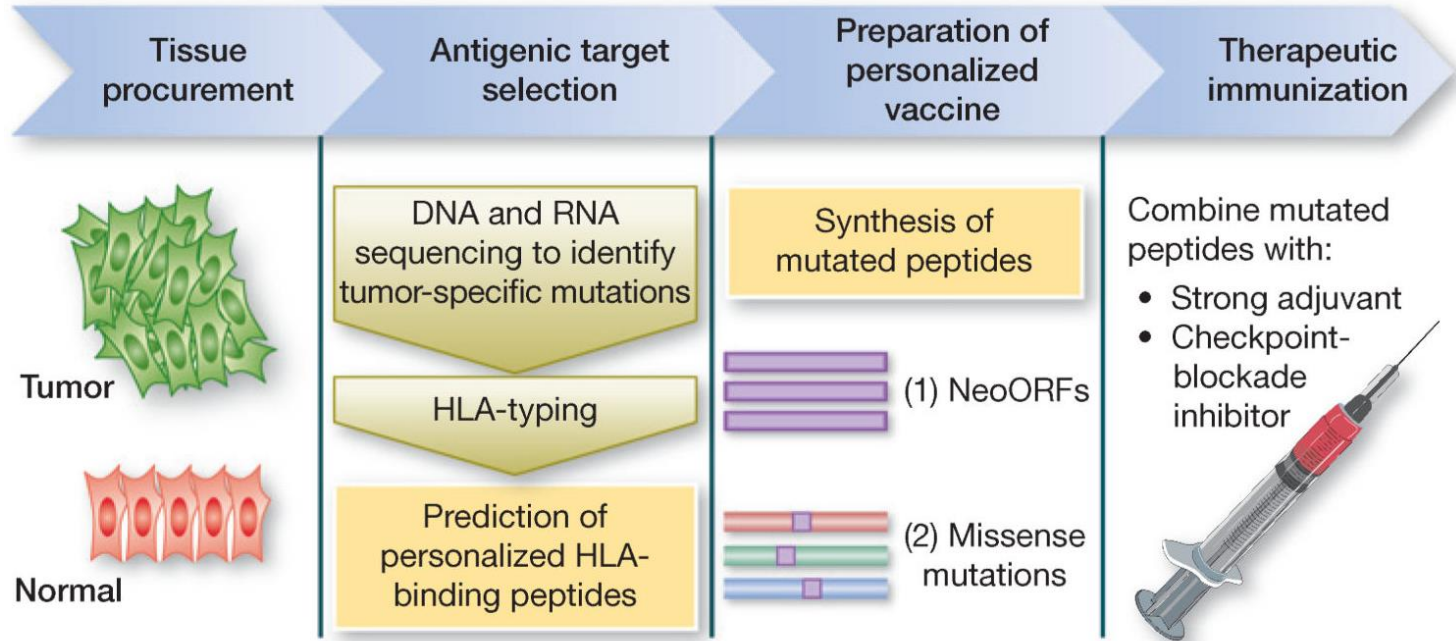
Neoantigens as Cancer Immunotherapy Targets: A paradigm shift



A Personal Approach



Generation of a personalized neoantigen vaccine



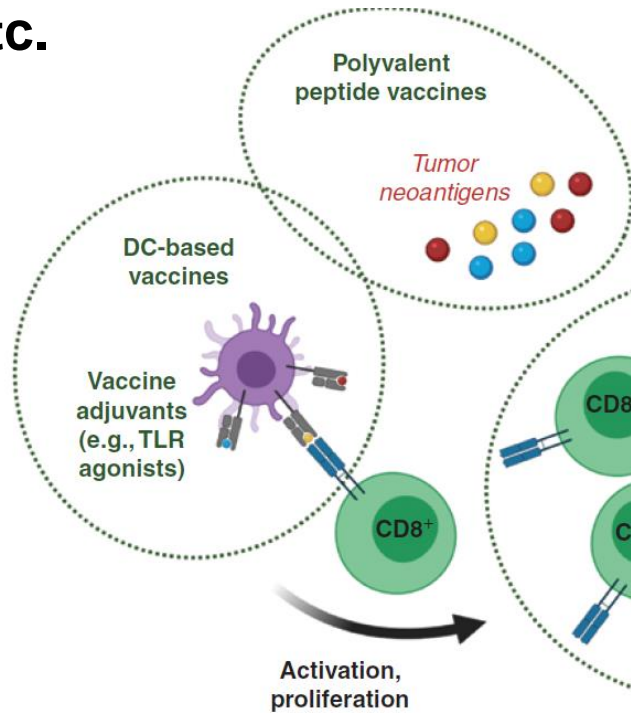
How can Neoantigens be Targeted
Therapeutically?

VACCINES

Peptide
RNA/DNA
DC
Viral etc.



Priming Phase:
Lymph nodes

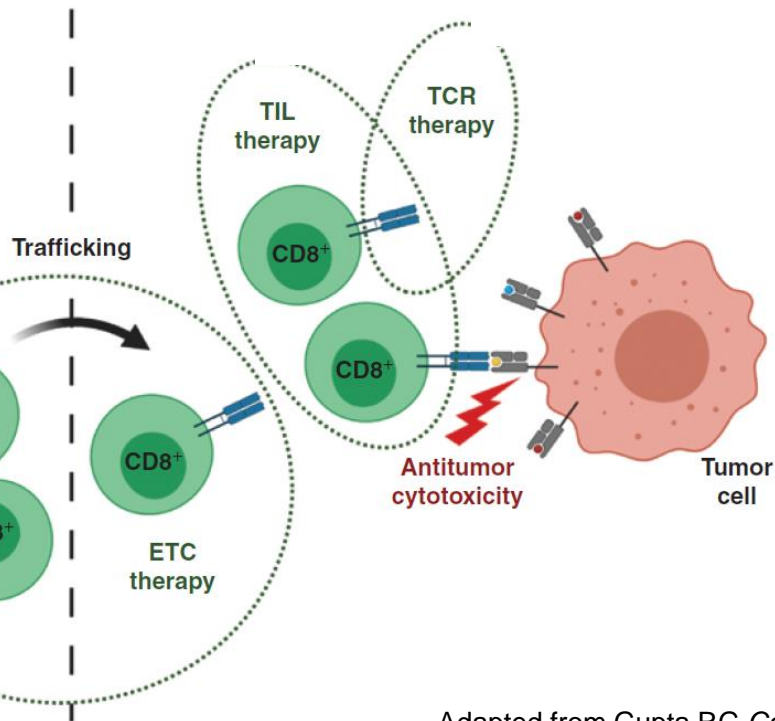


Adoptive Cell Transfer

TIL
neoTCR



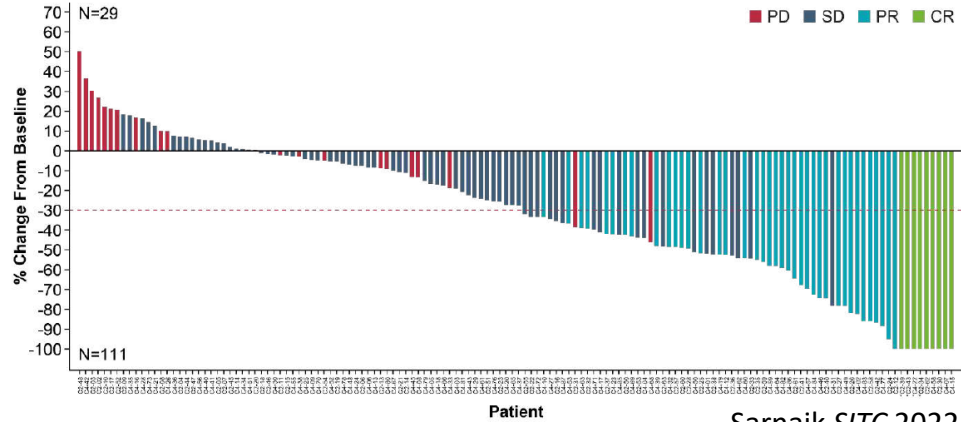
Effector Phase:
Tumor



TILs



Lifileucel

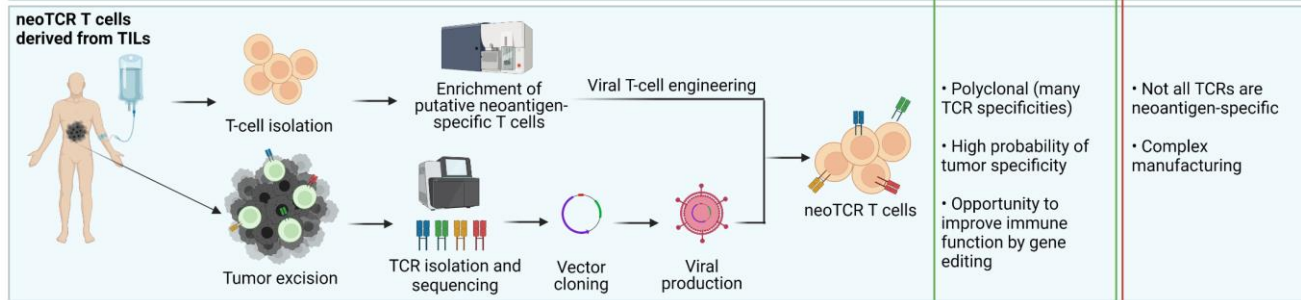


Sarnaik *SITC* 2022

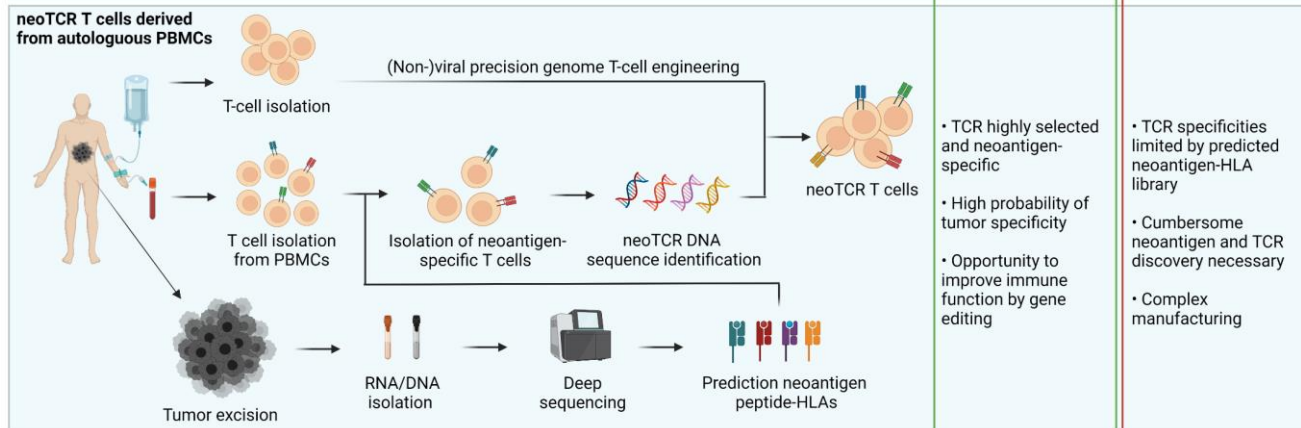
TILs

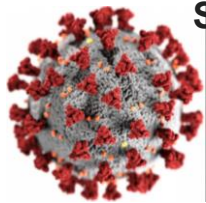


neoTCR T cells derived from TILs



neoTCR T cells derived from autologous PBMC



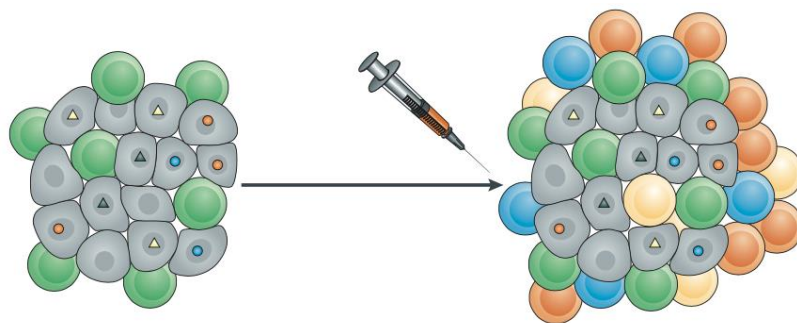


SARS-CoV-2

Pathogens — Prophylactic vaccines

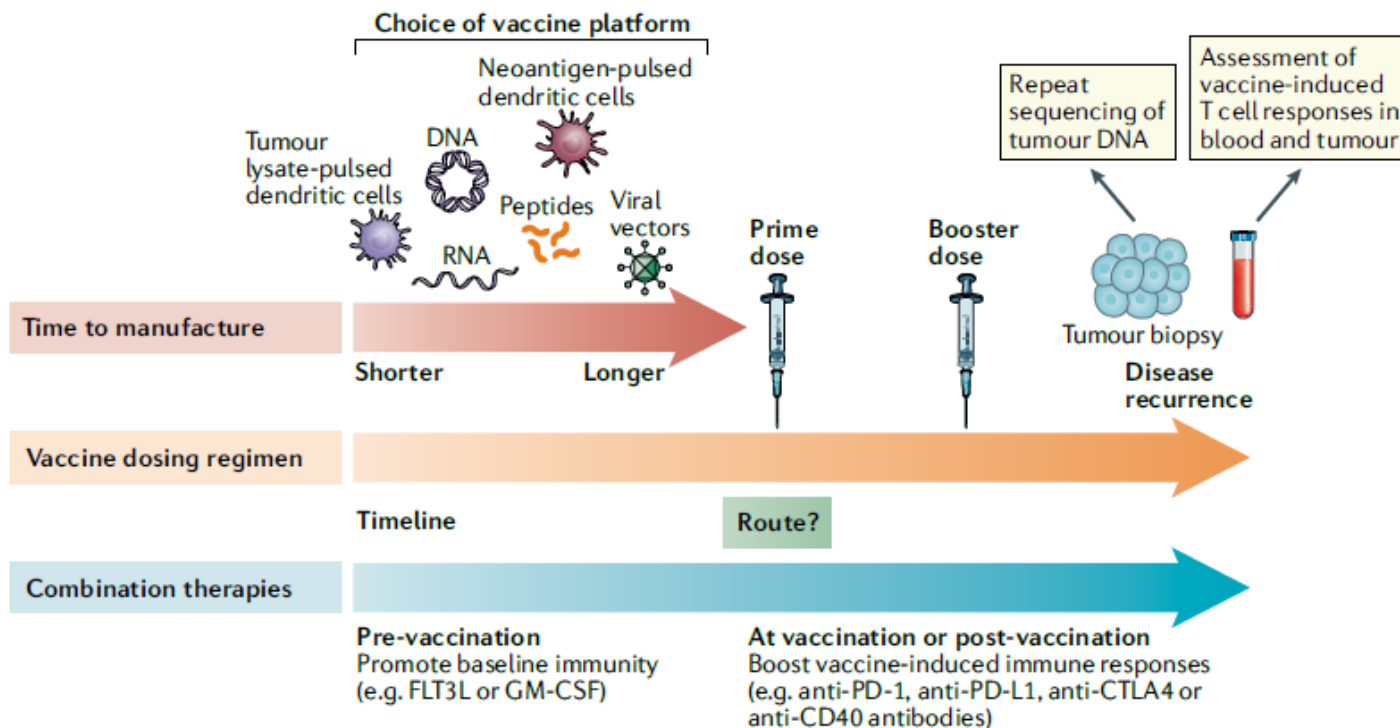
Antibody generation
? T cell responses

Cancer — Therapeutic vaccines



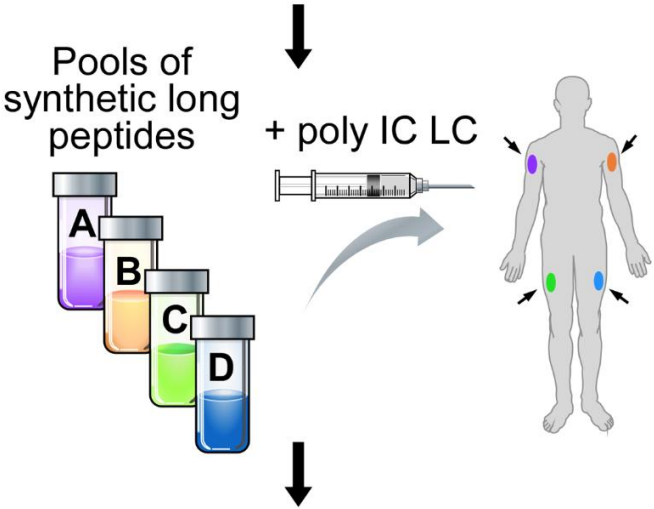
Increased breadth
and diversity of
tumour-specific
T cell response

Therapeutic Cancer Vaccines: Considerations

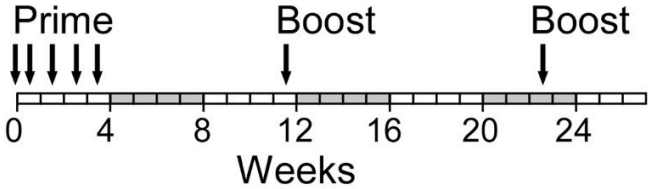


NeoVax in High risk melanoma patients: Study Design

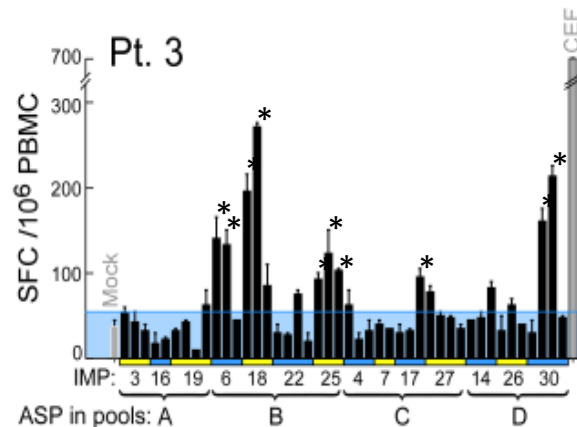
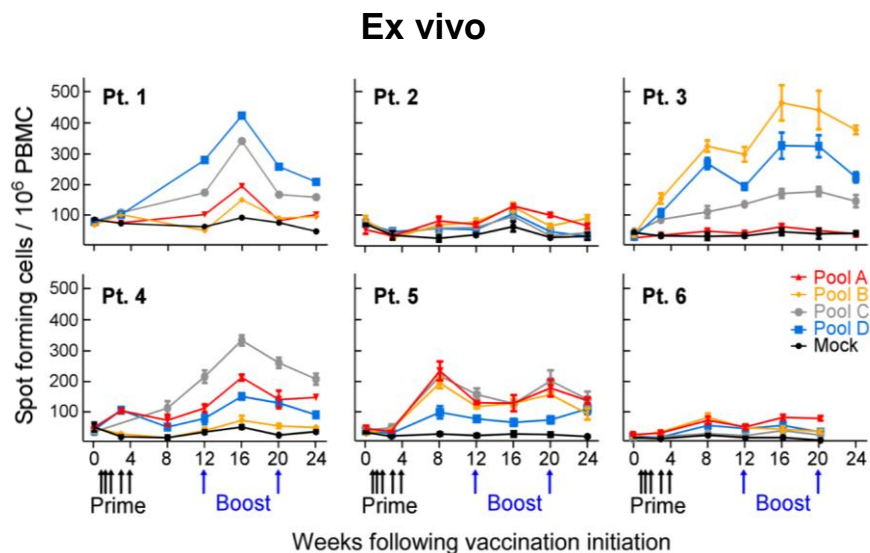
Personal vaccine manufacture



Vaccine administration



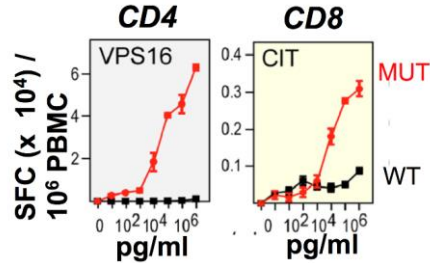
Neoantigen-specific T cell responses



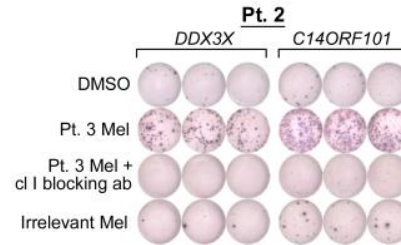
	CD4	CD8
<i>ex vivo</i>	18%	0%
1 stimulation	60%	16%

Vaccine-induced T cell responses:

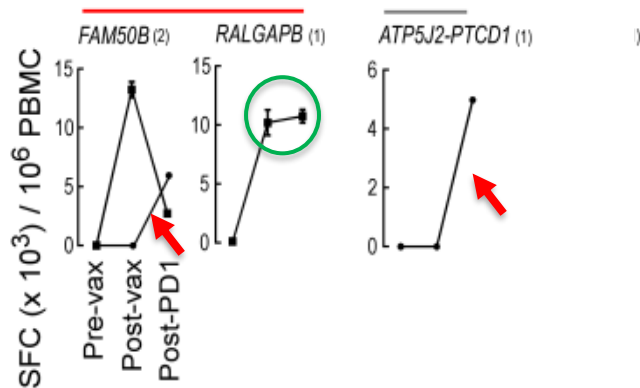
1) Specific for the mutant epitope



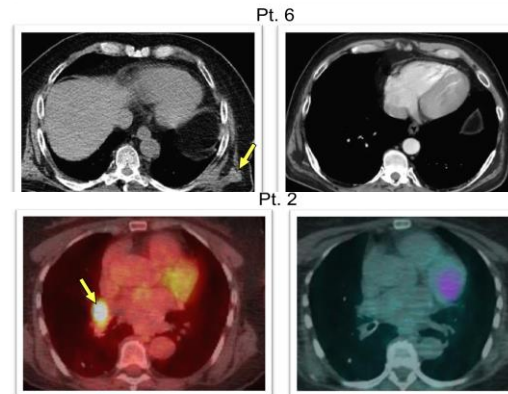
2) Reactive against autologous tumor



4) Broaden after PD-1 inhibition

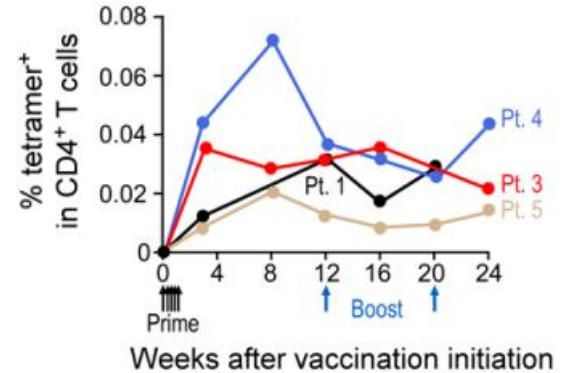
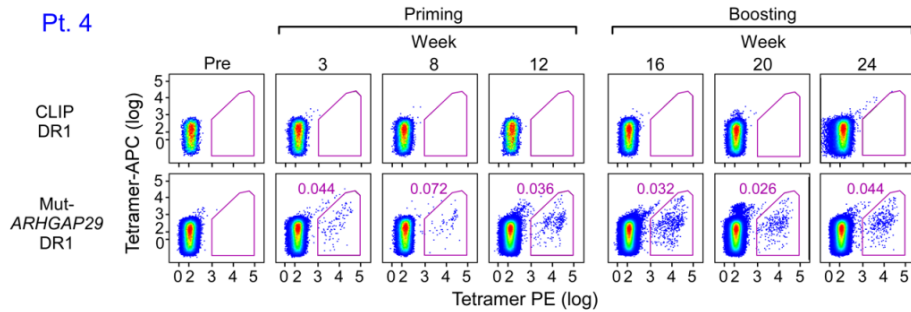


4) CR after Vax + anti-PD-1



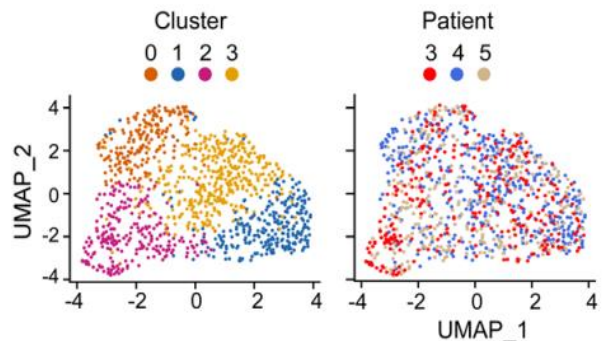
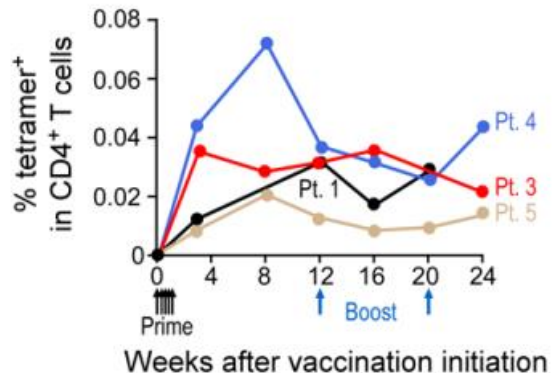
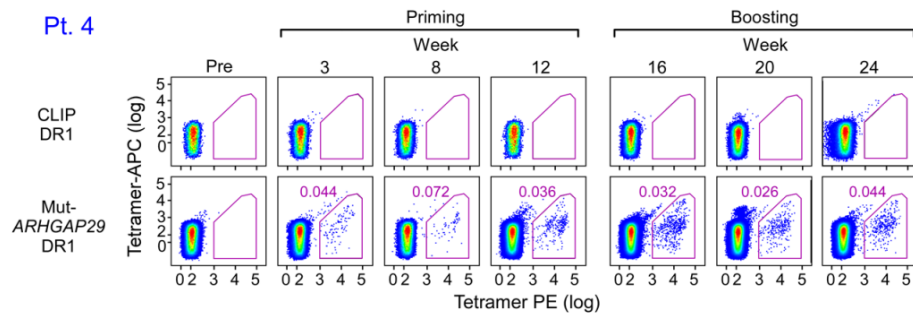
Transcriptional profile of neoantigen-specific T cells over the course of vaccination

Pt. 4



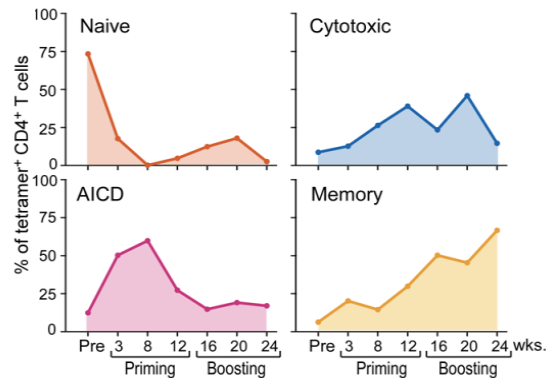
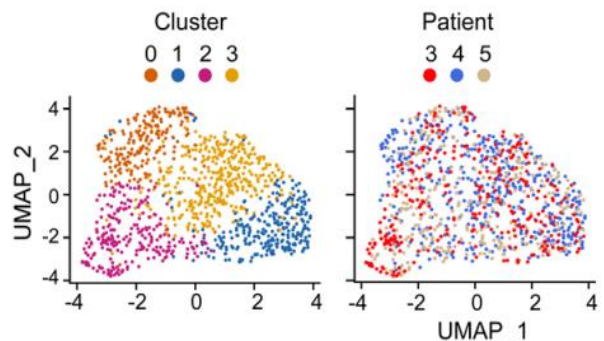
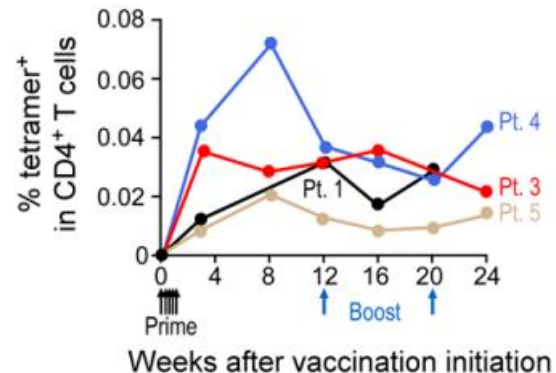
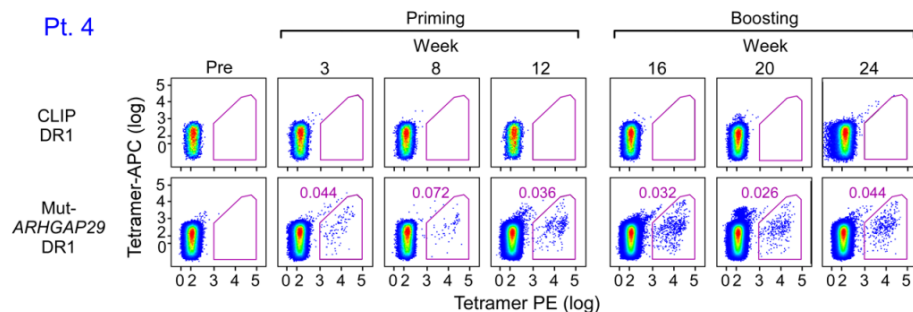
Transcriptional profile of neoantigen-specific T cells over the course of vaccination

Pt. 4

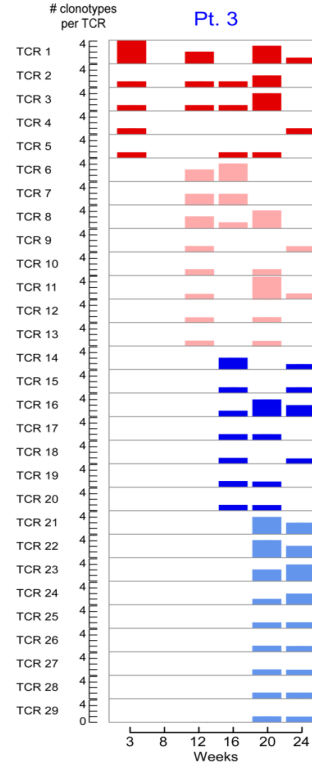
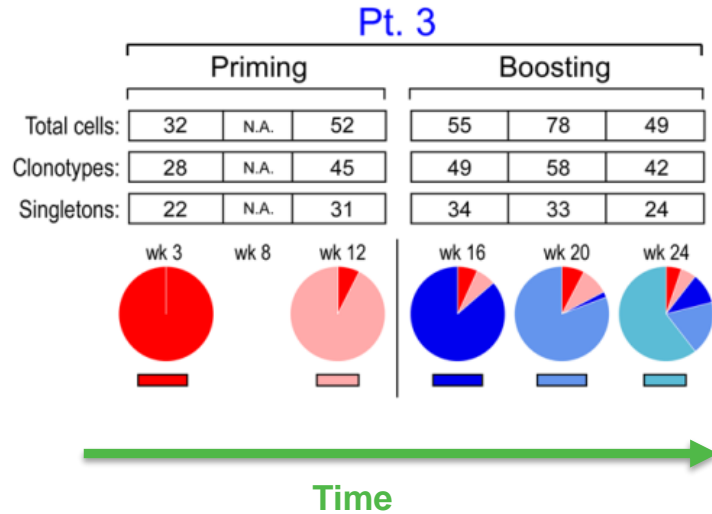


Transcriptional profile of neoantigen-specific T cells over the course of vaccination

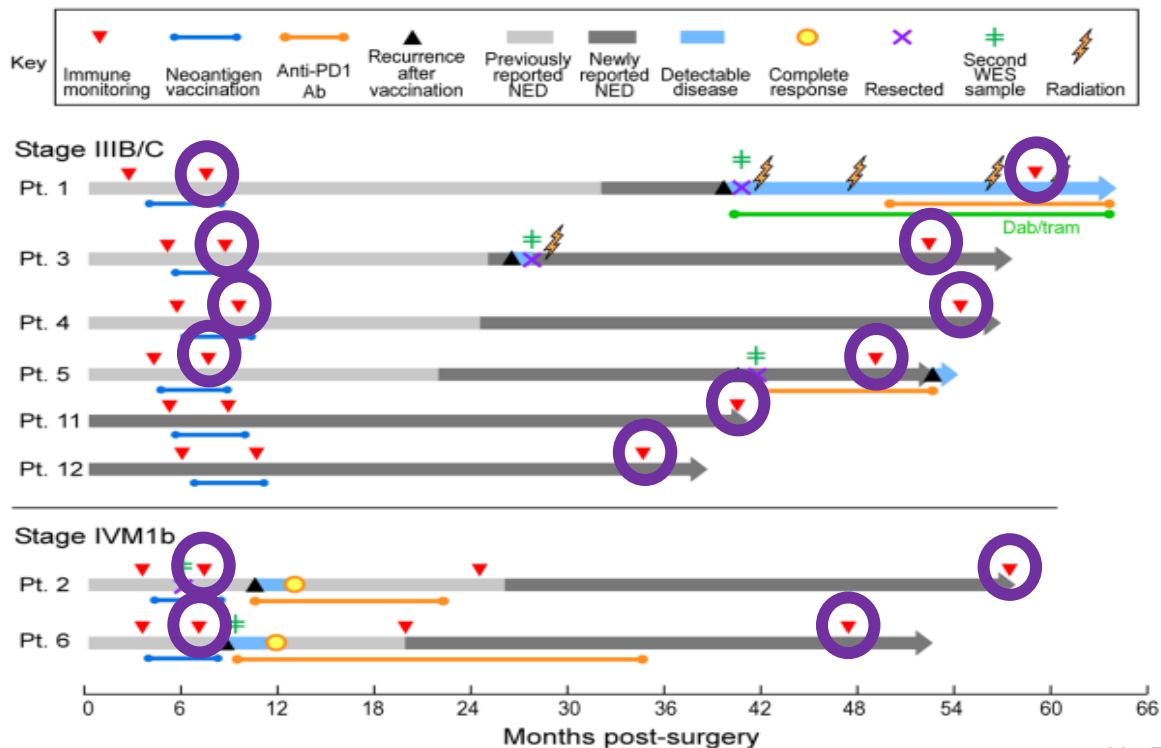
Pt. 4



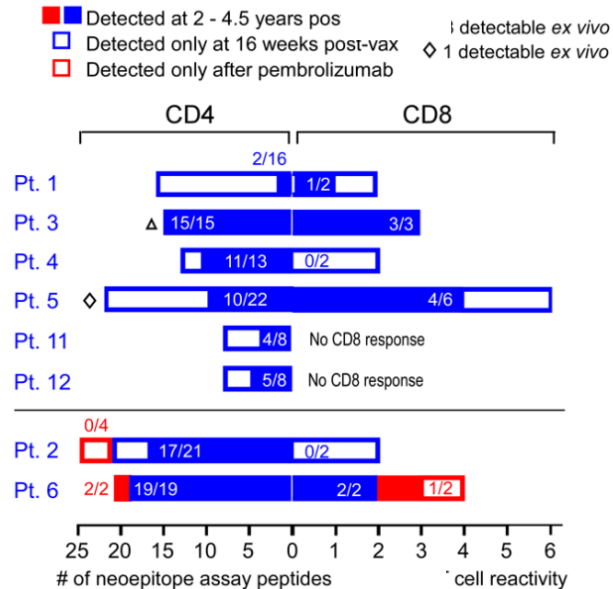
The TCR repertoire diversifies over time after vaccination



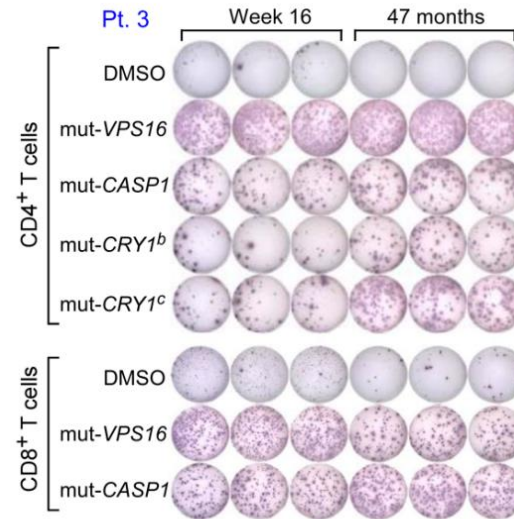
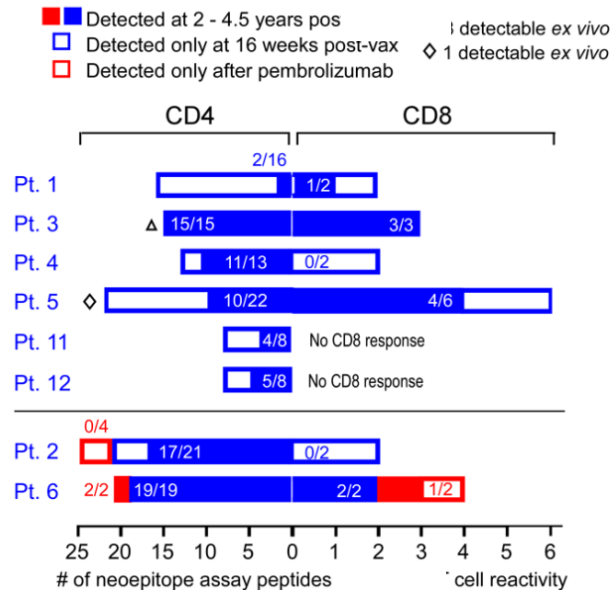
Clinical Course of Patients with High Risk Melanoma (Long-term)



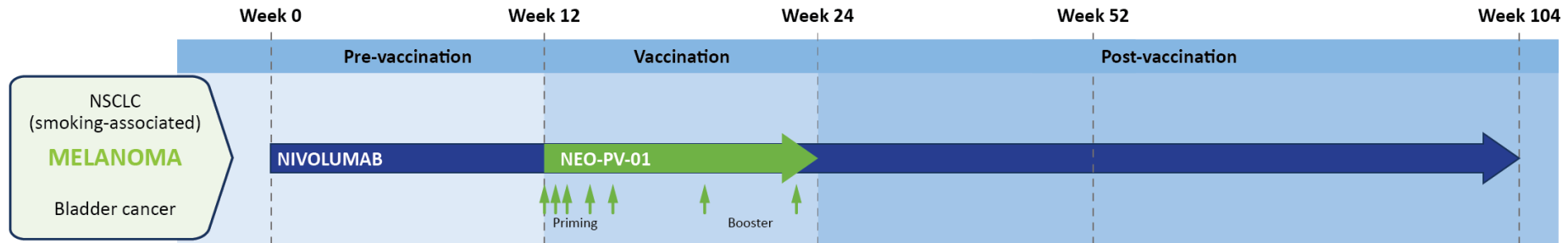
Vaccine-induced neoantigen specific T cells persist over several years



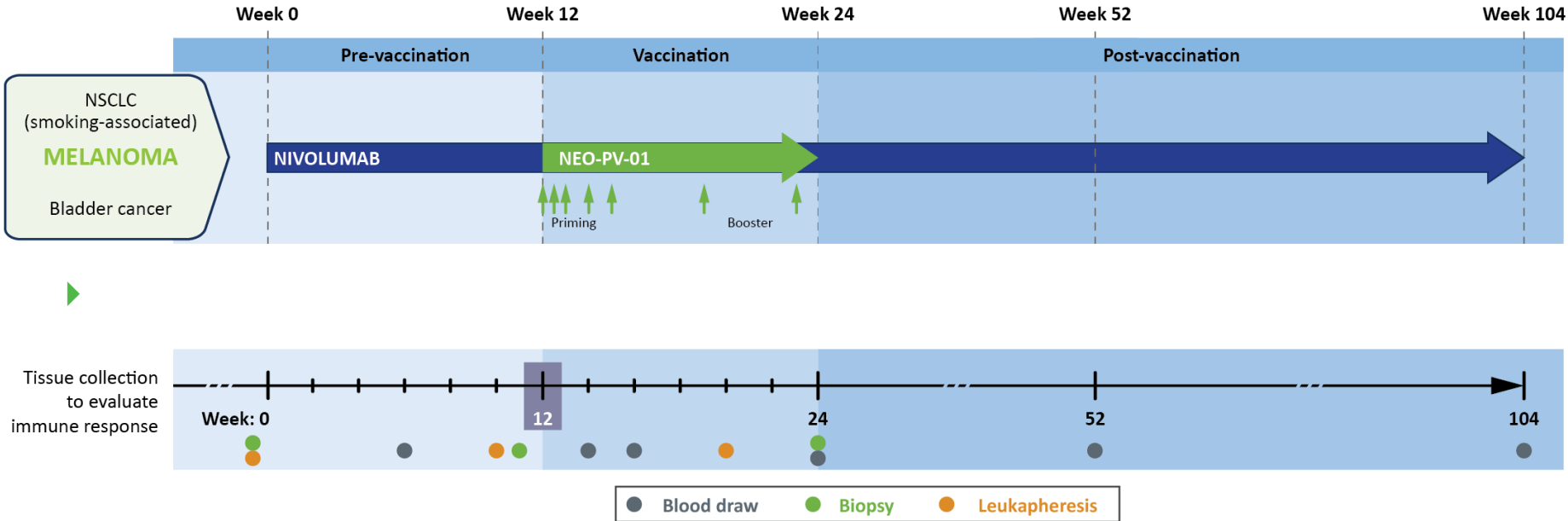
Vaccine-induced neoantigen specific T cells persist over several years



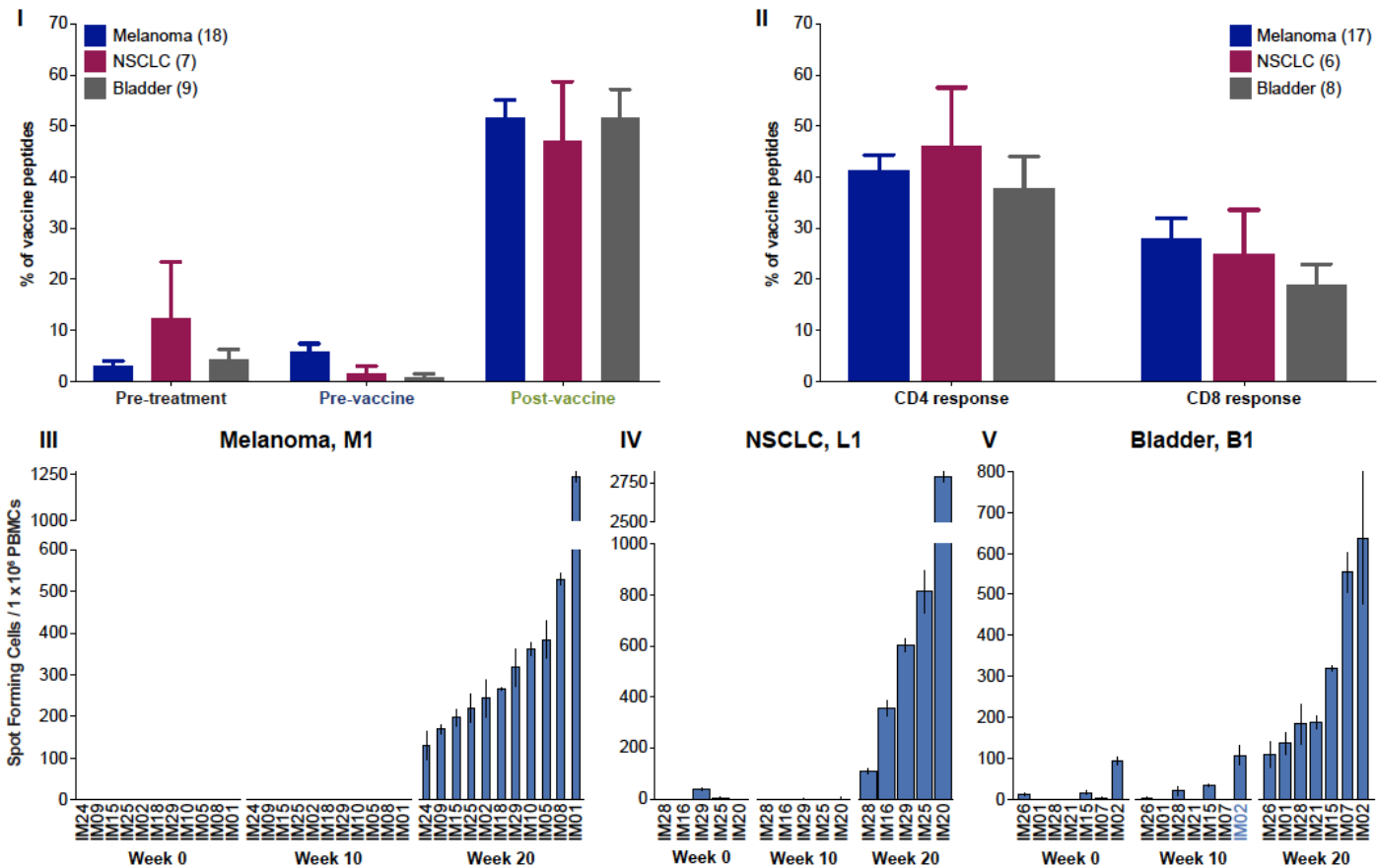
NT-001: Personalized peptide vaccine (PV-01) + Nivolumab in metastatic patients (melanoma, NSCLC, and urothelial cancer)



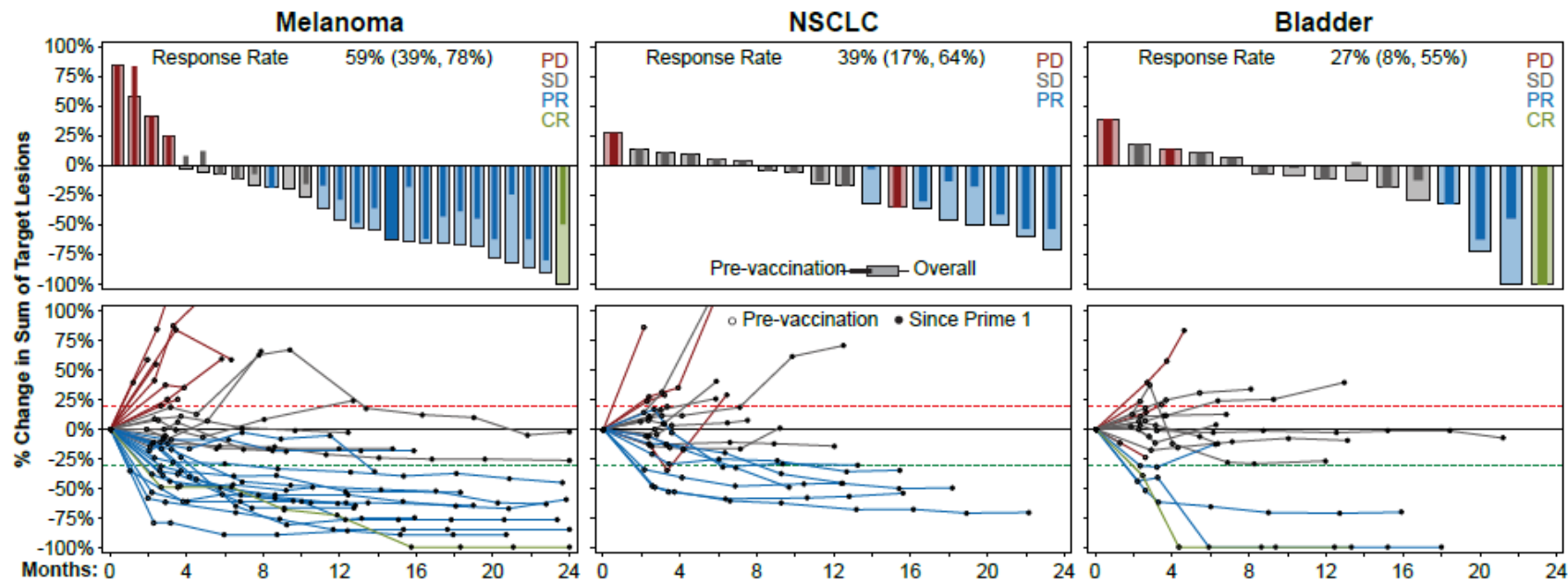
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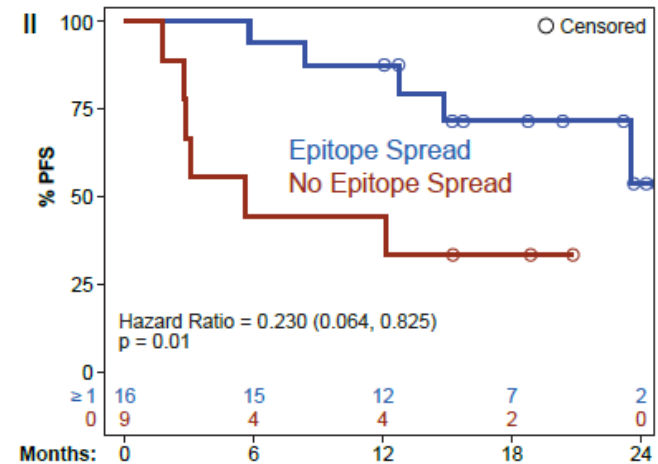
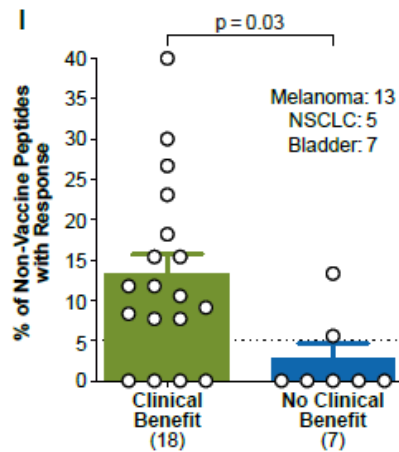
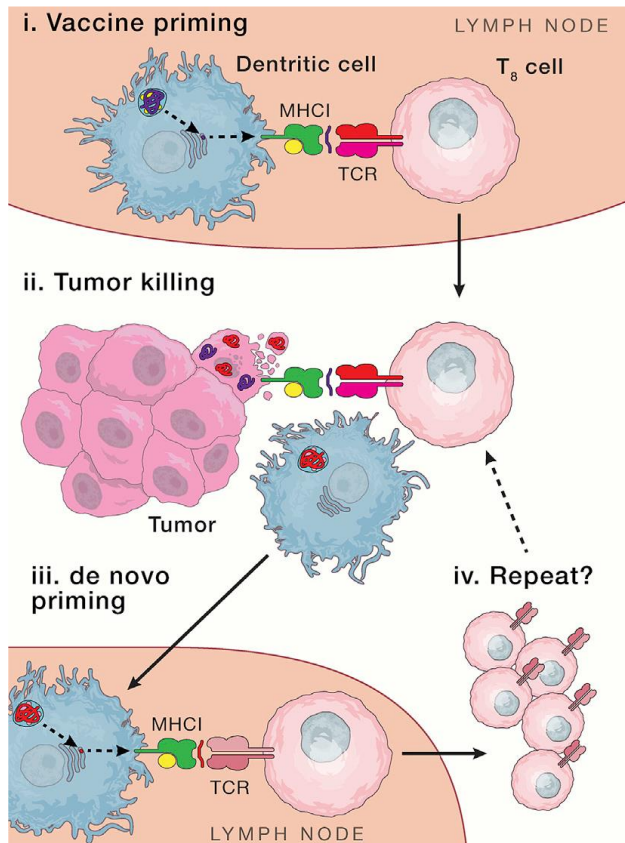
NEO-PV-01 + Nivolumab Induces Neoantigen-Specific Immune Responses



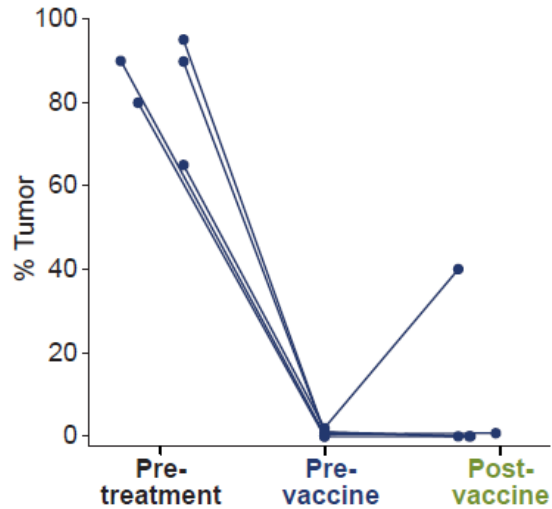
Anti-tumor activity



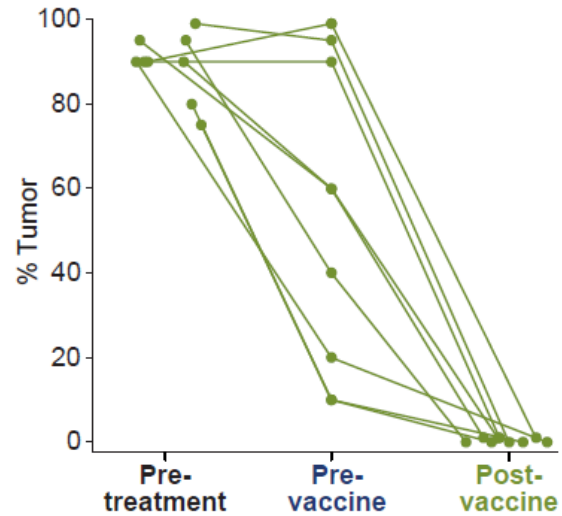
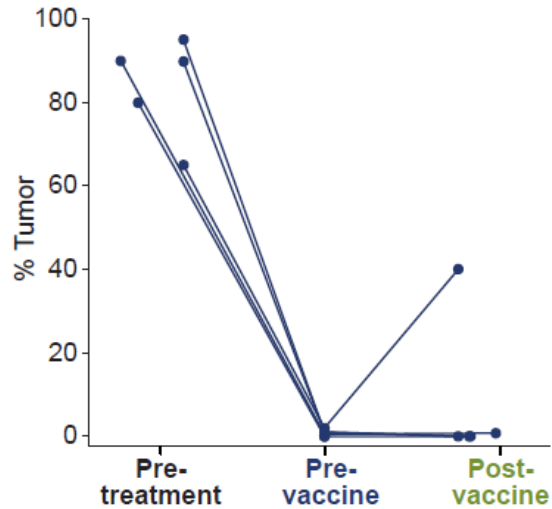
Epitope spreading correlates with durable progression-free survival



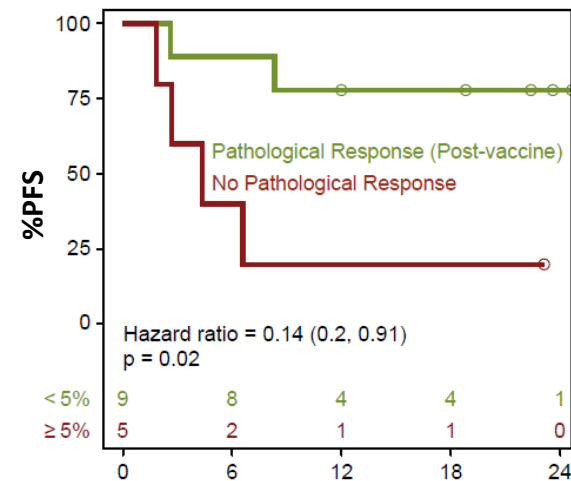
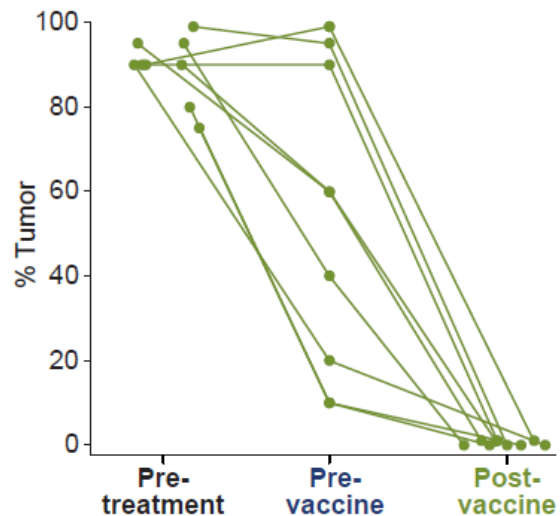
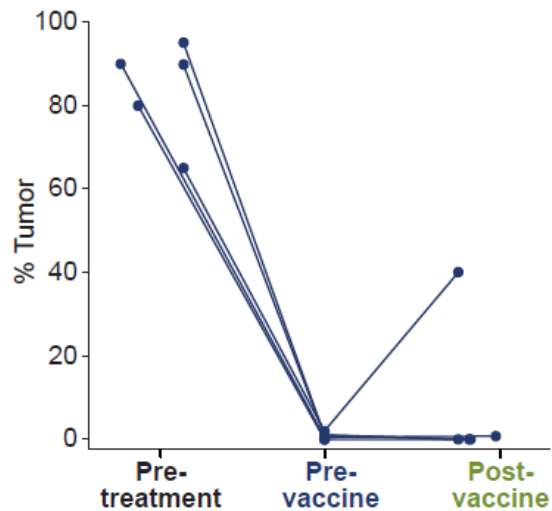
Pathologic response post vaccine is associated with clinical benefit



Pathologic response post vaccine is associated with clinical benefit

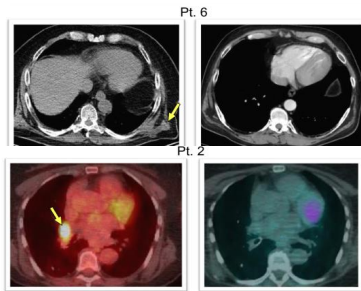


Pathologic response post vaccine is associated with clinical benefit



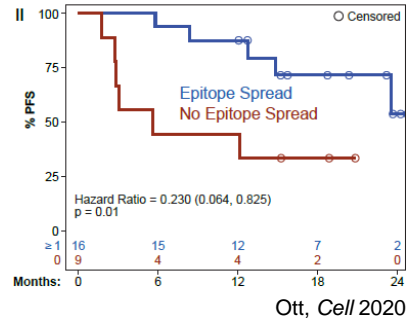
Encouraging signals for efficacy, however no definitive data ... until recently

CRs with α -PD-1 post long peptide Vax in Melanoma

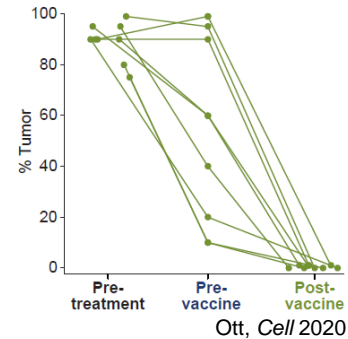


Ott & Hu, *Nature* 2017

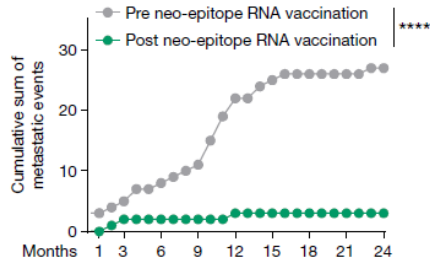
Epitope Spreading post long-peptide Vax in mel, NSCLC, bladder Ca



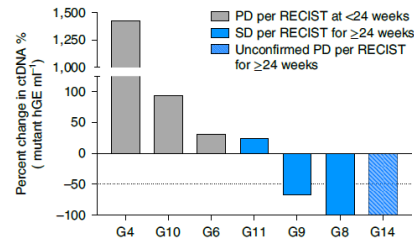
Path CR post long-peptide Vax in melanoma



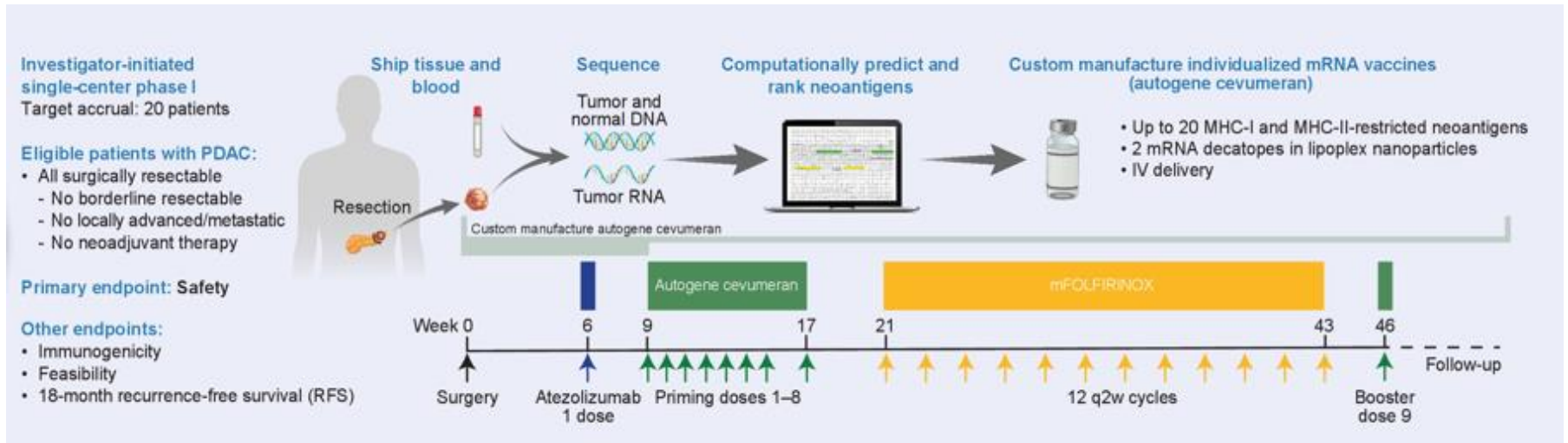
Decreased Recurrences post RNA Vax in Melanoma



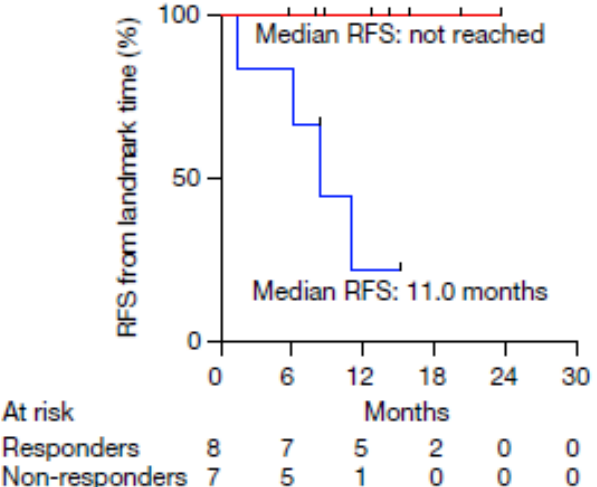
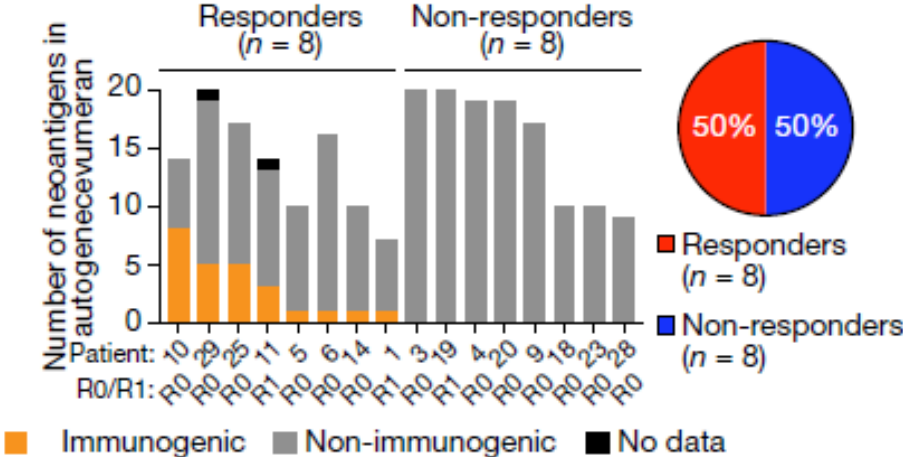
CRs by ctDNA post Vax in colorectal ca



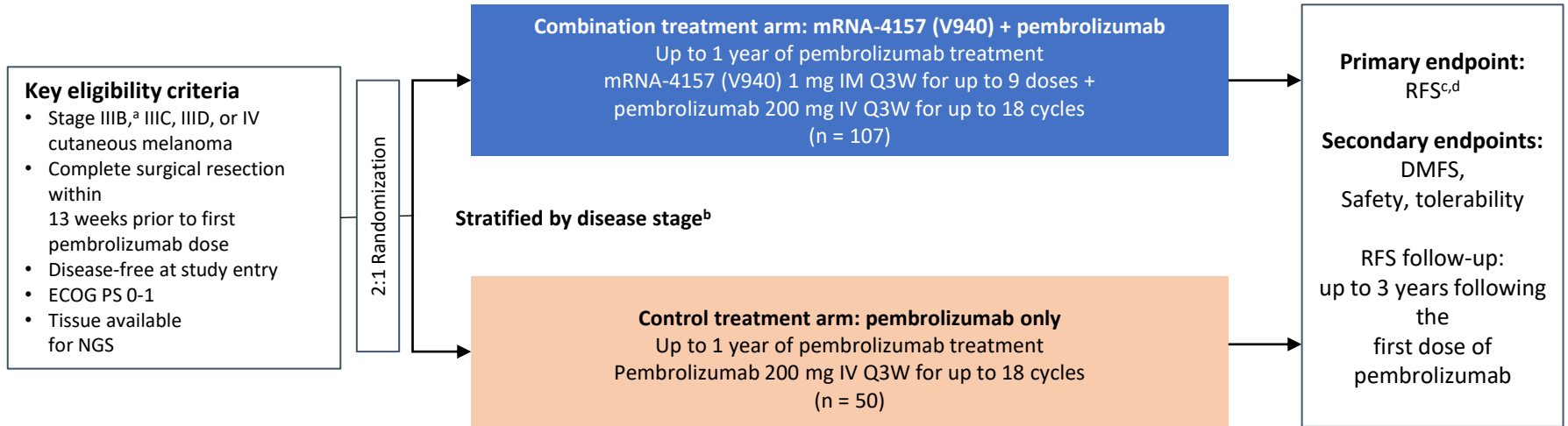
Personalized Neoantigen RNA vaccine autogene cevumeran in patients with resectable pancreatic ductal carcinoma (PDAC)



T cell immunity induced with a Personalized RNA vaccine + Atezolizumab + mFOLFORINOX correlates with delayed recurrence



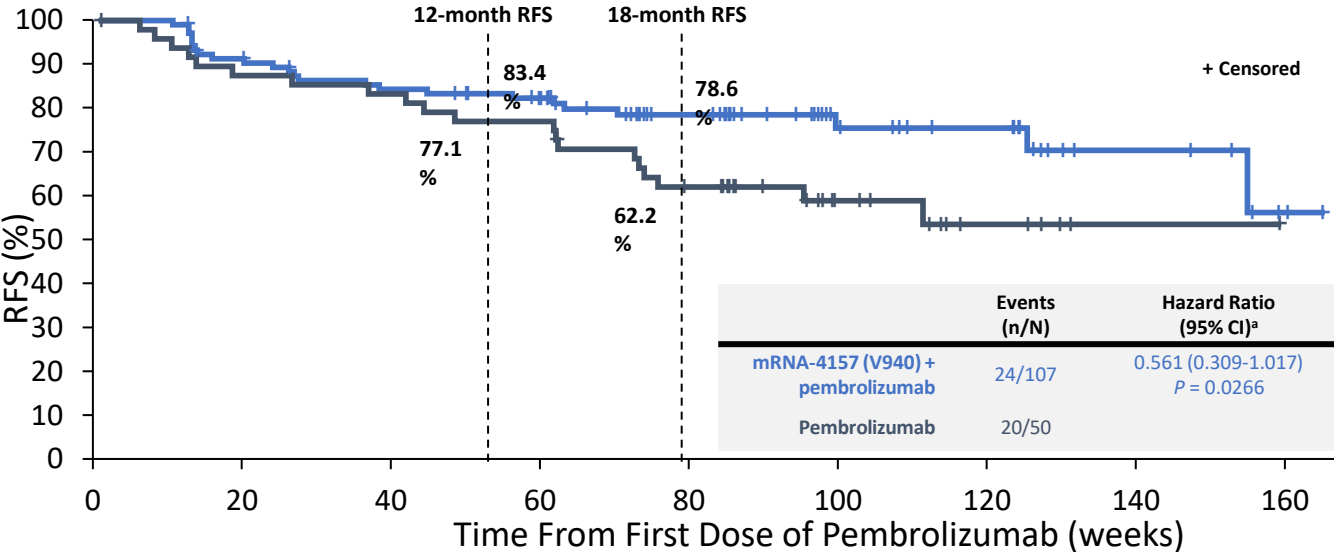
And now a randomized Phase 2 trial: mRNA-4157 (V940) + pembrolizumab vs. Pembrolizumab alone



Designed with 80% power to detect an HR of 0.5 with ≥ 40 RFS events (with 1-sided alpha of 0.1)

Median follow-up^e: 23 months for mRNA-4157 (V940) + pembrolizumab
24 months for pembrolizumab only

mRNA-4157 (V940) and pembrolizumab combination treatment demonstrated a statistically significant and clinically meaningful RFS improvement



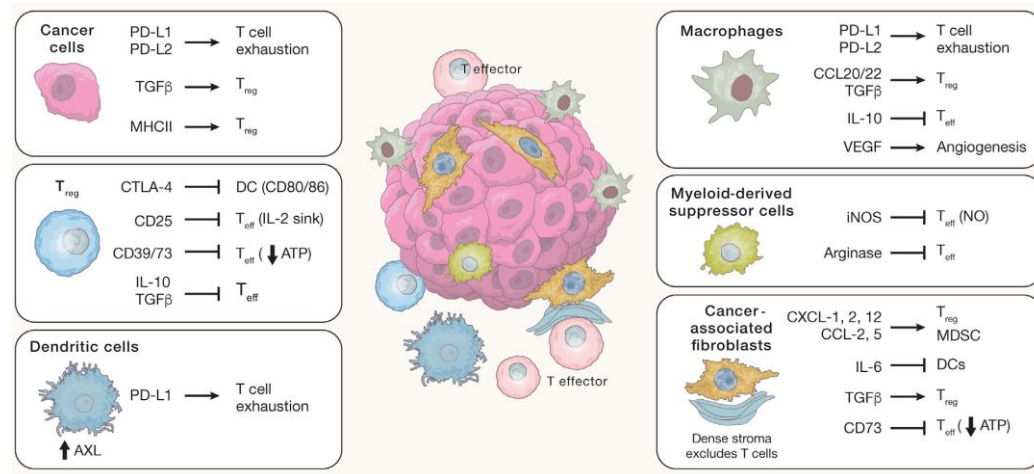
	Number at Risk								
	0	20	40	60	80	100	120	140	160
mRNA-4157 (V940) + Pembrolizumab	107	92	85	73	49	24	20	8	1
Pembrolizumab	50	42	40	37	28	13	6	1	0

Challenges/opportunities

- Time and Cost
- Magnitude and Quality of Vaccine-induced Immune responses
 - Improve vaccine formulation/immune adjuvant
 - Enhance T cell priming

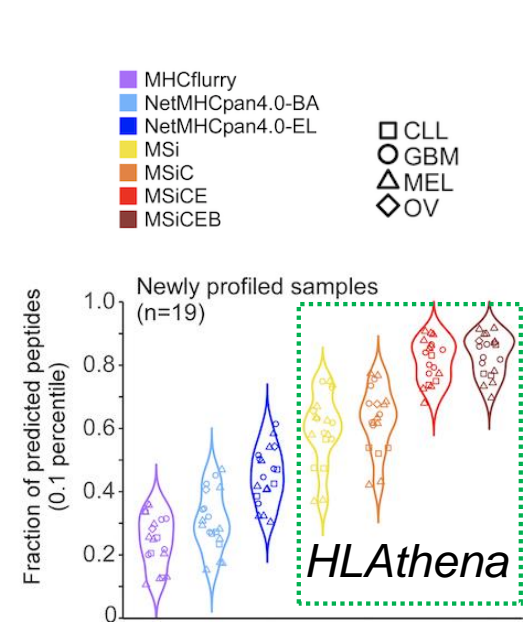
Challenges/opportunities

- Time and Cost
- Magnitude and Quality of Vaccine-induced Immune responses
 - Improve vaccine formulation/immune adjuvant
 - Enhance T cell priming
- “Manage” the TME



Challenges/opportunities

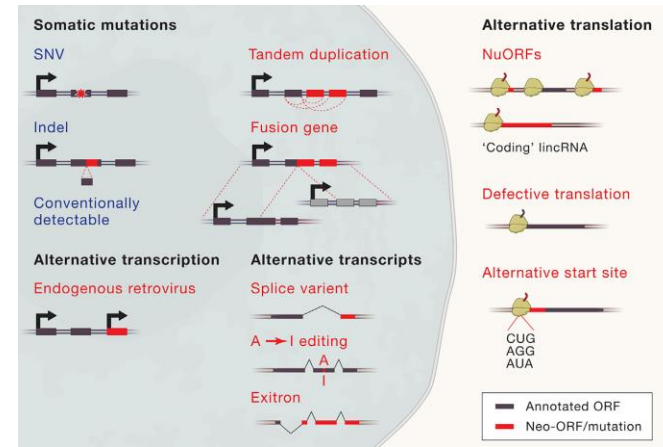
- Neoantigen discovery
- What are the most immunogenic tumor neoantigens?



9-38% increase in peptide identifications with MSEC

Challenges/opportunities

- Neoantigen discovery
- What are the most immunogenic tumor neoantigens?
- Can we increase the neoantigen discovery space?





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Immuno-Oncology & Cancer Vaccine Programs

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CSO IMUNON

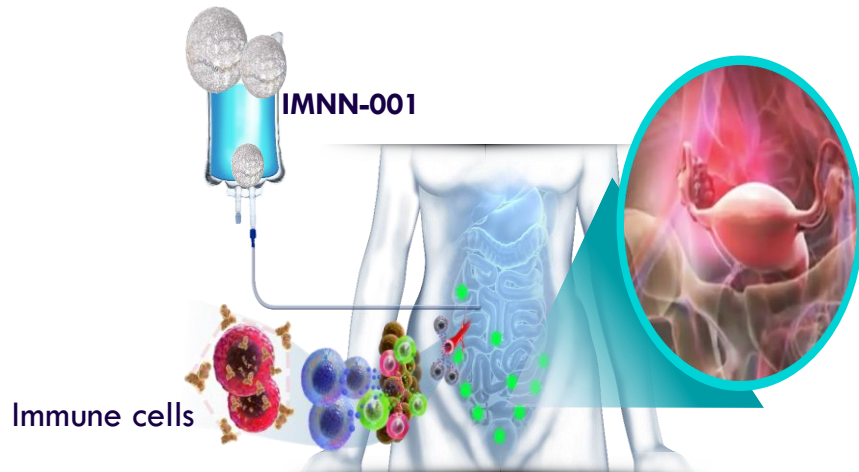
IMUNON's Immuno-Oncology & Cancer Vaccine Programs

DNA-based Technology Independent of Viruses or Devices for Delivery

TheraPlas

Immune Modulation by
Cytokine Therapy

IMNN-001



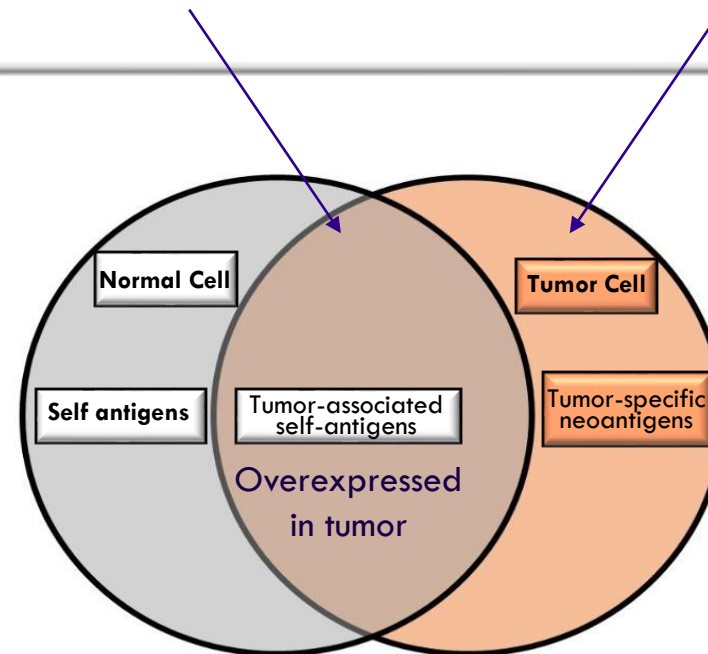
Persistent Local Delivery of IL-12
for Modulating the TME

FixPlas

Tumor-Associated Antigen
Vaccines

IndiPlas

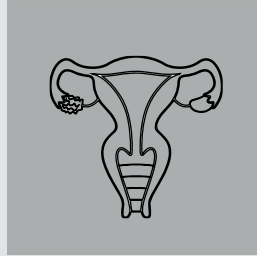
Neoantigens
(personalized vaccines)



IMNN-001 – Persistent Local Delivery of IL-12 with Formulated Plasmid

First Target: Epithelial Ovarian Cancer

- Insidious disease – late-stage diagnosis
- High recurrence rate – poor survival
- Novel approaches warranted



IMNN-001

- A gene therapy product for safe and effective delivery of IL-12
- IL-12, a powerful immune agent, a “Master Switch” to the body’s innate and adaptive immune system
- A safe alternative to rIL-12 therapy that is short-lived and exerts serious systemic toxicity

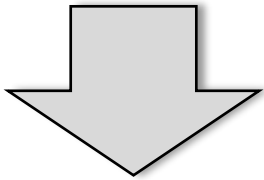
IMNN-001 Clinical Development

- Five completed clinical trials- newly diagnosed/recurrent disease
- Safety, clinical activity and biological activity in Phase 1 studies
- A Phase 2 trial (OVATION-2) offer hopes for OC patients; data read 2Q 2024
- One recently activated trial explores IMNN-001 combination with antiangiogenic agents

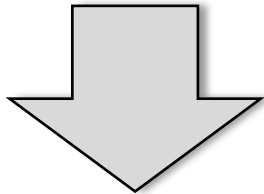
FixPlas: Cancer Vaccines

Targeting Tumor-Associated Antigens (TAAs) – Monovalent and Bivalent

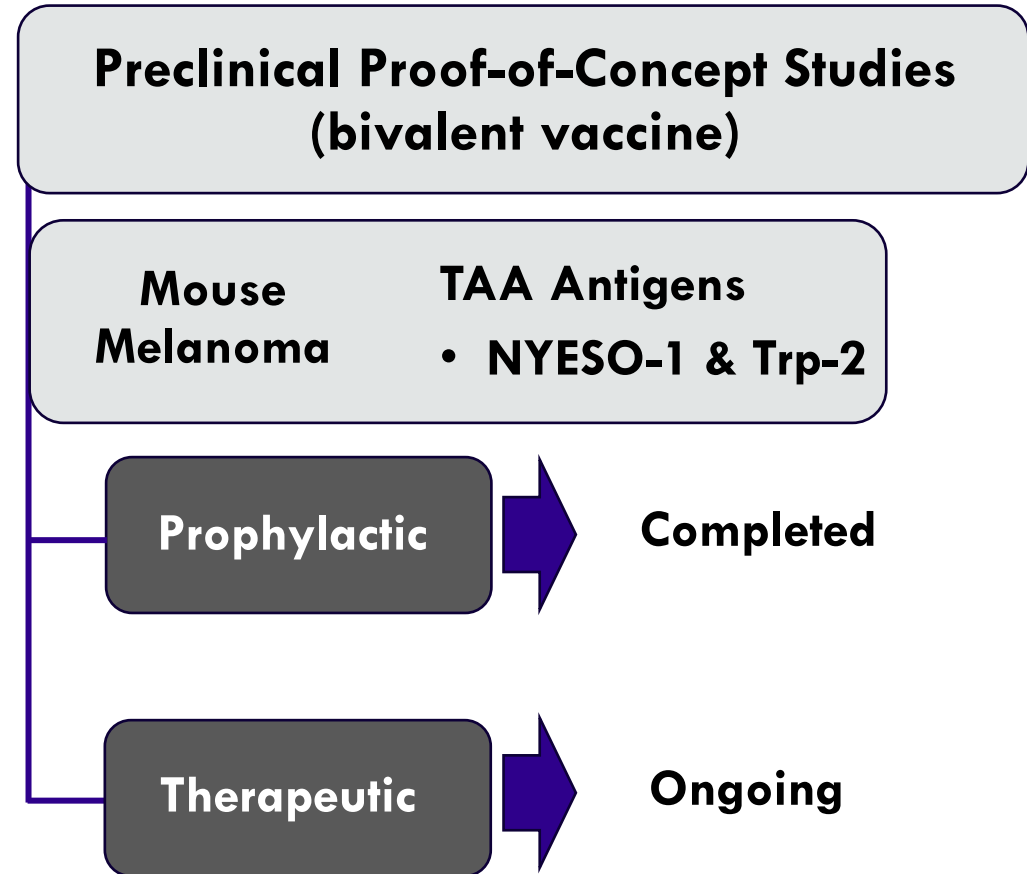
DNA Vaccines



Strong Cellular Responses
(Advantage over protein/mRNA)



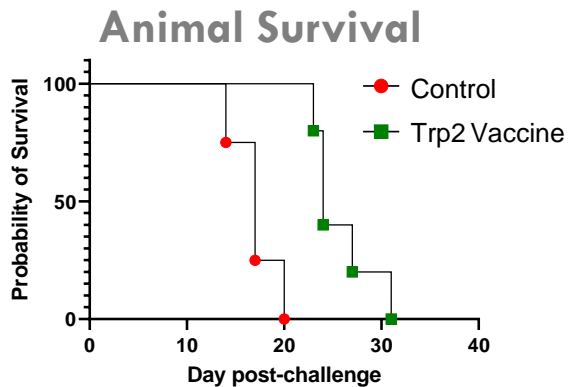
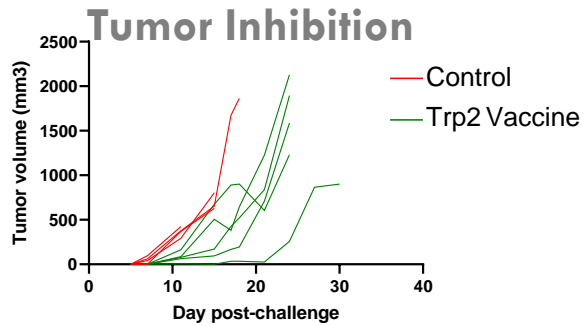
Well Suited for Cancer Therapy



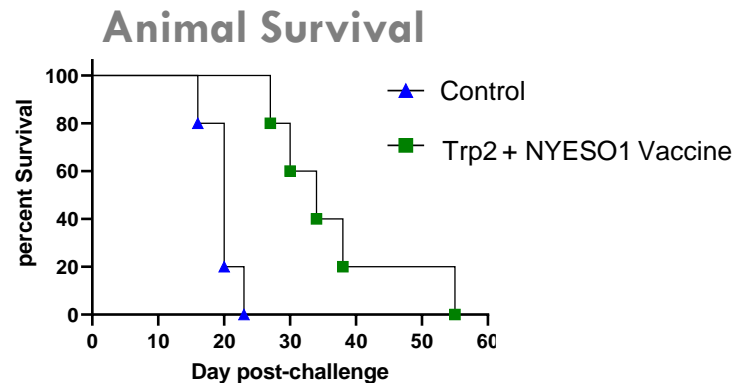
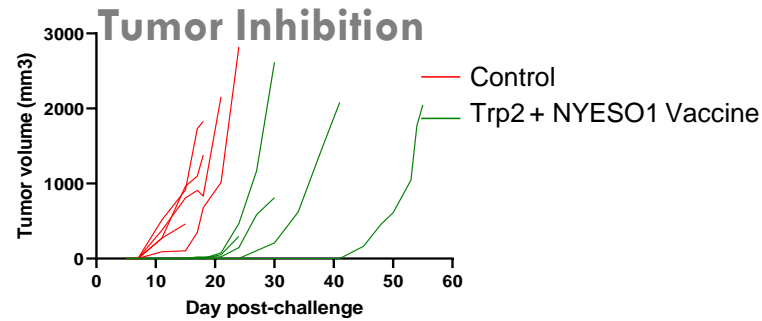
FixPlas Prophylactic-

Tumor Inhibition & Survival Improvement – Trp2/NYESO1, B16F10 Melanoma

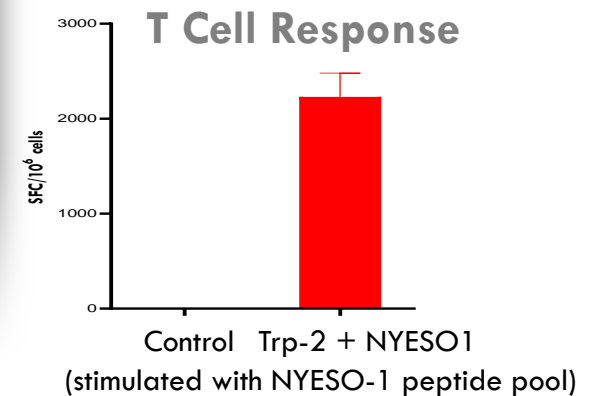
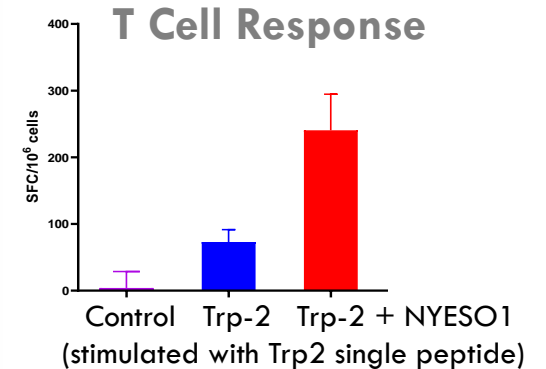
Monovalent Trp2 Vaccine (Formulated Trp2 plasmid)



Bivalent Trp2 + NYESO1 Vaccine (Formulated Trp2 + NYESO1 plasmid)



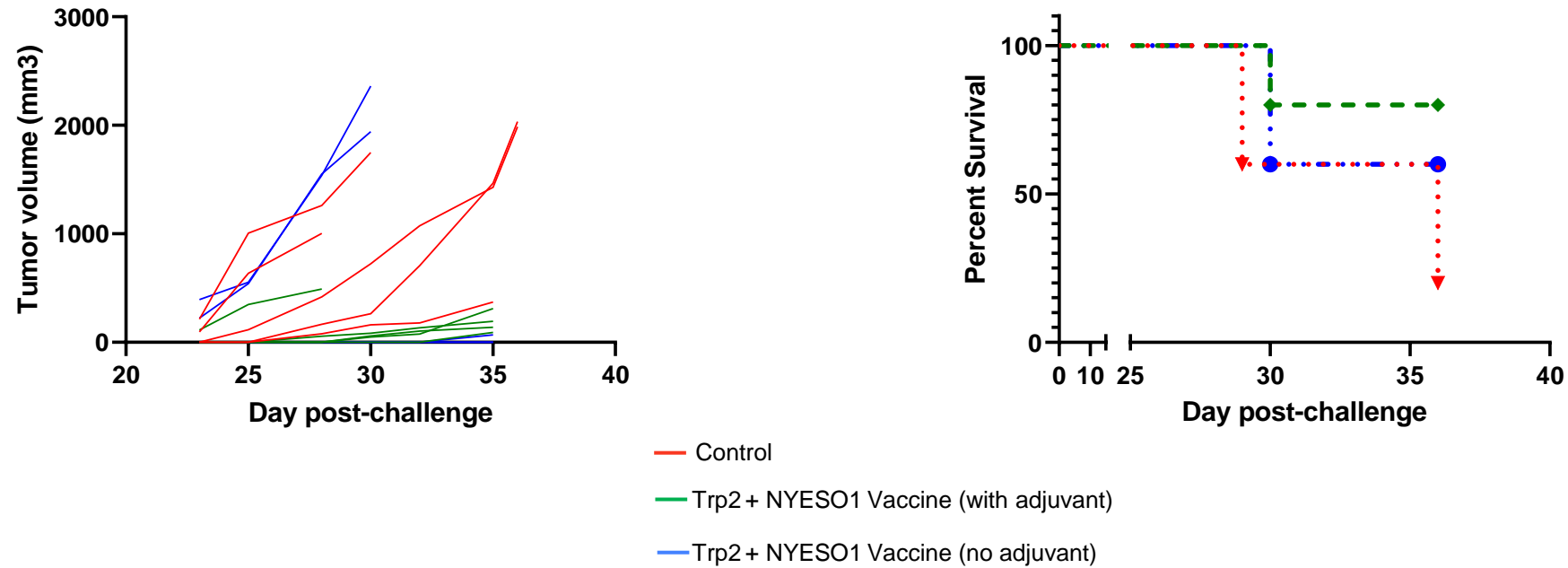
Monovalent & Bivalent (Formulated plasmid)



Vaccination Followed by Trp2 and NYESO1 Expressing B16F10 Tumors

FixPlas Therapeutic- Tumor Inhibition & Survival Improvement

Bivalent Trp2 + NYESO1 Vaccine Formulation



B16F10-Trp2 and NYESO1 Antigens Followed by Vaccination

FixPlas Development Plan

Preliminary Studies Warrant a Product Development Plan

- Selection of the disease target
- Selection of the antigen target(s)
- Optimization of the antigen
- Demonstration of robust anticancer activity
- IND-enabling studies
- Phase 1 clinical trial