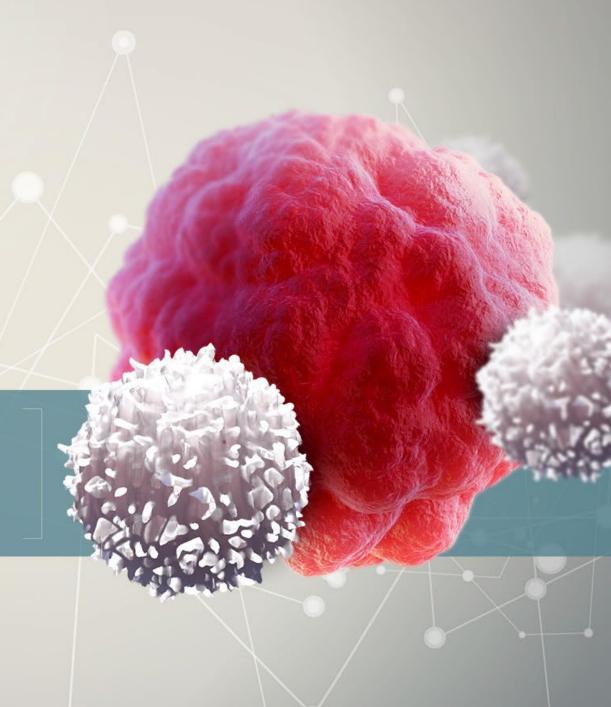


CORPORATE PRESENTATION

GT-EPIC™: EXQUISITELY PERSONALIZED IMMUNOTHERAPIES FOR CANCER



Q3, 2023

## GENEOS OVERVIEW



Mission to develop Personalized Therapeutic Cancer Vaccines (PTCV's) to treat cancer



## Leveraged DNA Medicines IP from INOVIO

- Exclusive license in the FIELD of patient specific cancer prevention or therapy
- Additional PTCV-related IP developed internally



## Institutional VC backed company

- \$50M raised to date
- Key investors Santé Ventures, INO, KIP-VC, Flerie Invest, 3B Future Health Fund, Shanghai Healthcare Capital



# GT-30 Phase 1b/2a POC study in 2L Advanced HCC

- Fully enrolled, near completion



## **EXPERIENCED LEADERSHIP TEAM**



NIRANJAN Y. SARDESAI, PH.D President & CEO, BOD Founder



JOANN PETERS, MHA Chief Operating Officer



ILDIKO CSIKI, MD, PH.D Consulting Chief Medical Officer



**FEDERICA F. O'BRIEN**Consulting CFO



**DAVID WURTMAN, MD MBA**Consulting Chief Business Officer



JIAN YAN, PH.D VP, Research & Discovery



MYRNA THOMAS VP, QA Manufacturing



**BETH JUNKER, PH.D**Consultant CMC / Manufacturing

### BOARD OF DIRECTORS

**DR. SAMUEL BRODER** Independent; Former NCI Director

**DR. CASEY CUNNINGHAM**Santé Ventures

DR. JAMES EADIE Santé Ventures

DR. LAURENT HUMEAU
Inovio Pharmaceuticals

MR. SANGWOO LEE KIP-VC, USA

**DR. TED FJÄLLMAN** Flerie Invest

DR. ROBERTO DE PONTI (BOD OBSERVER)
3B Future Health Fund

**DR. JING BAO (BOD OBSERVER)**Shanghai Healthcare Capital

### **ADVISORS**

**DR. DAVID B. WEINER** Wistar Institute

DR. CHI VAN DANG
Ludwig Institute, Johns Hopkins University

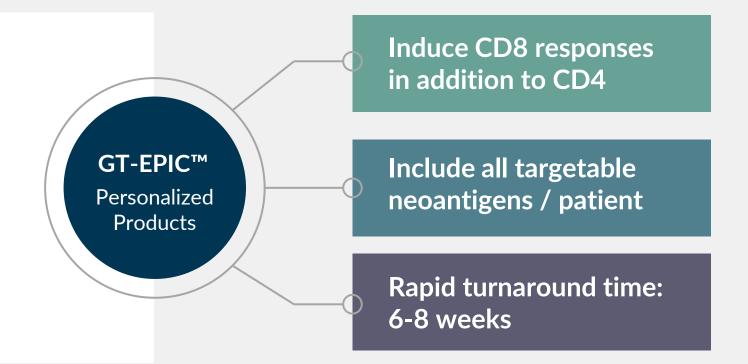
MS. SHAWN TOMASELLO Independent; Formerly Kite, Celgene



# GENEOS PERSONALIZED CANCER VACCINES (PCV) ARE VERSATILE & POTENT

### **GENEOS DIFFERENTIATION:**

- Deliver cancer neoantigens & shared antigens in patient specific product
- Leverage patient's own immune system to in vivo select neoantigens that induce T cells and drive clinical responses



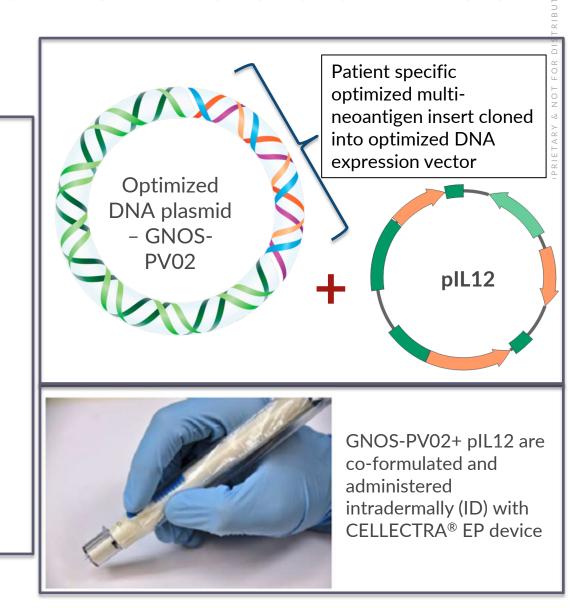
Targeting all neoantigens can address potential issues with tumor immune escape & polyclonal, multi-focal tumors



## GENEOS PERSONALIZED TREATMENT DRIVES IMMUNE RESPONSES BY DESIGN

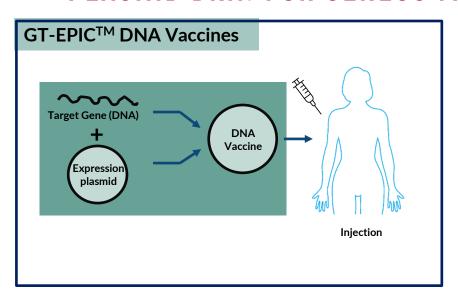
# OPTIMIZED DNA NEOANTIGENS + plL12 + CELLECTRA® ELECTROPORATION (EP)

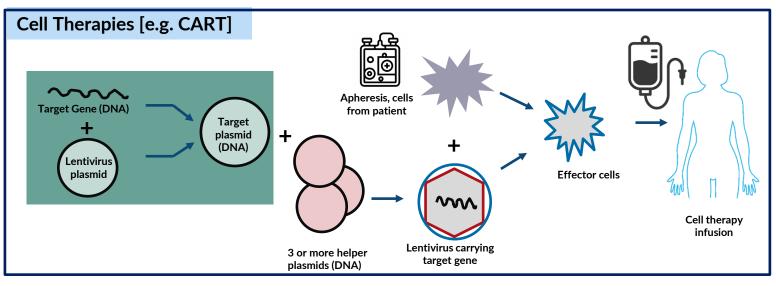
- Personalized product has three components
  - Optimized DNA plasmid encoding neoantigens
  - IL-12 (pIL12): Cytokine immune-modulator; Boosts T cells
  - CELLECTRA® delivery device (in vivo electroporation;
     EP): Efficient plasmid uptake for optimal antigen production
- Combination activates robust functional antigen specific
   CD4+ & CD8+ killer T cells

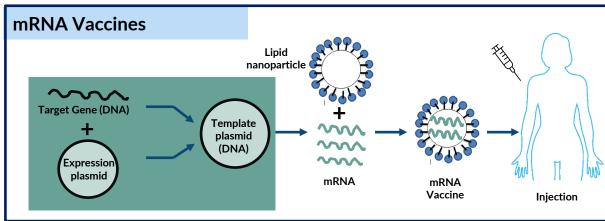


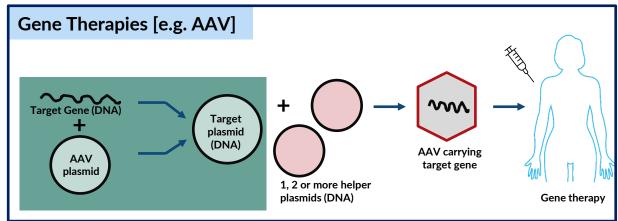


# ALL CELL & GENE THERAPY (CGT) PRODUCTS START WITH THE MANUFACTURING OF PLASMID DNA: FOR GENEOS IT IS ALSO THE FINAL DRUG PRODUCT





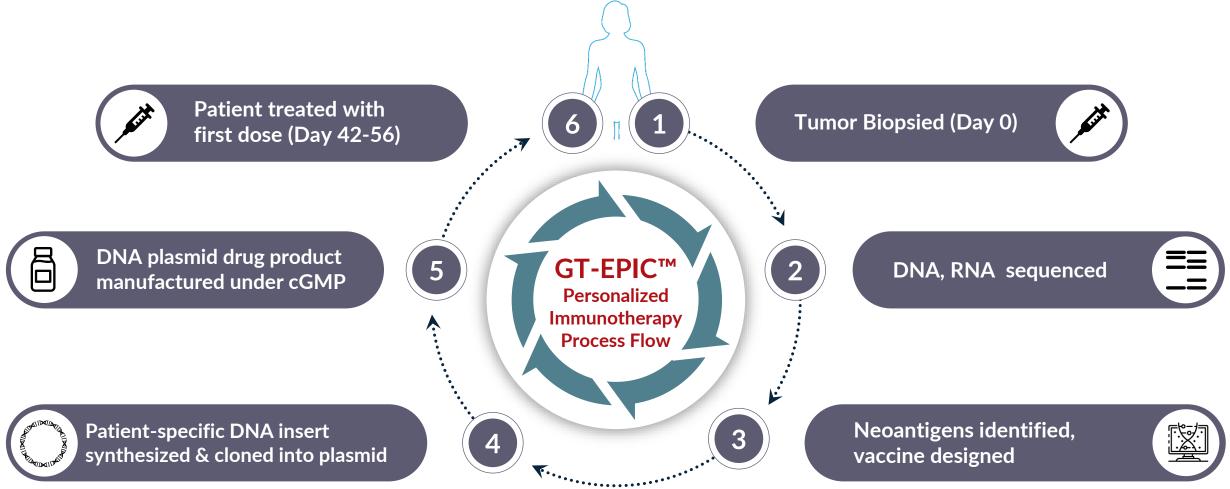




- Manufacturing represents a key competitive advantage for Geneos' GT-EPIC<sup>TM</sup> DNA platform
- · Higher process complexity drives larger manufacturing TAT and COGS for the other CGT platforms. 🏯



# GT-EPIC™ PERSONALIZED IMMUNOTHERAPY PROCESS HAS BEEN SUCCESSFULLY TRANSLATED TO THE CLINIC





## HEPATOCELLULAR CARCINOMA

#### LARGE UNMET CLINICAL NEED

US cases per year:

29,000

EU: 34,000/year WW: 800,000/year



5-year survival rate, 2nd behind pancreatic cancer

#### **CPI IN ADVANCED HCC**

14-17% 4 mo

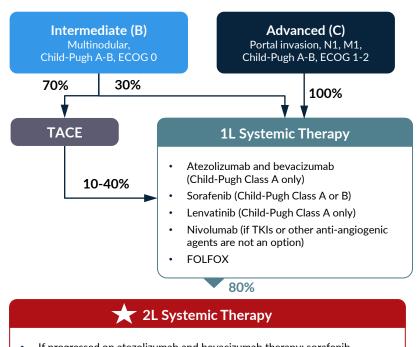
Respond to CPI **Immunotherapy** 

**Median PFS** 

14 mo

**Median OS**  ★ GENEOS-targeted segment in HCC could benefit from CD8+ inducing therapy

#### **Treatment Overview**



- If progressed on atezolizumab and bevacizumab therapy: sorafenib,
- If not previously treated with checkpoint inhibitor: nivolumab, nivolumab with ipilimumab, pembrolizumab
- Other options: regorafinib, cabozantinib, ramucirumab, lenvatinib



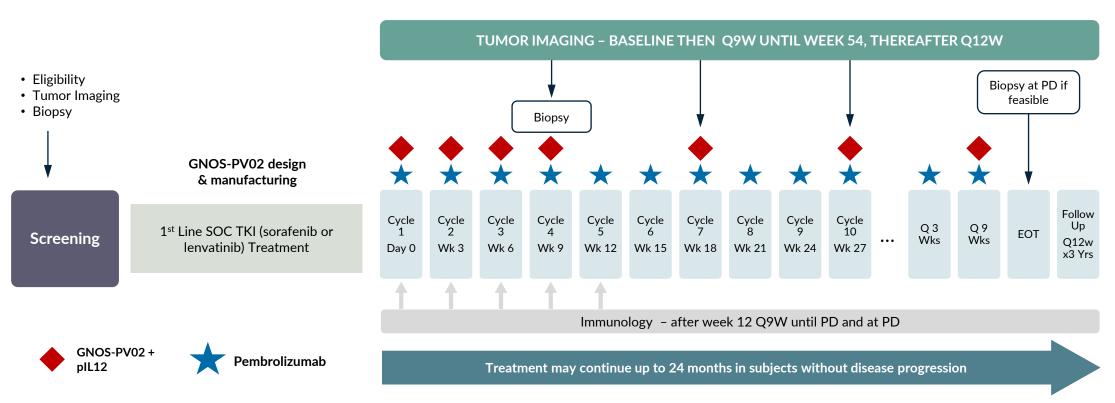
3L+ Systemic Therapy



## GT-30 CLINICAL TRIAL IN 2ND LINE ADVANCED HCC

Advanced HCC patients who progress during or are intolerant to 1st Line TKI treatment (sorafenib or lenvatinib)

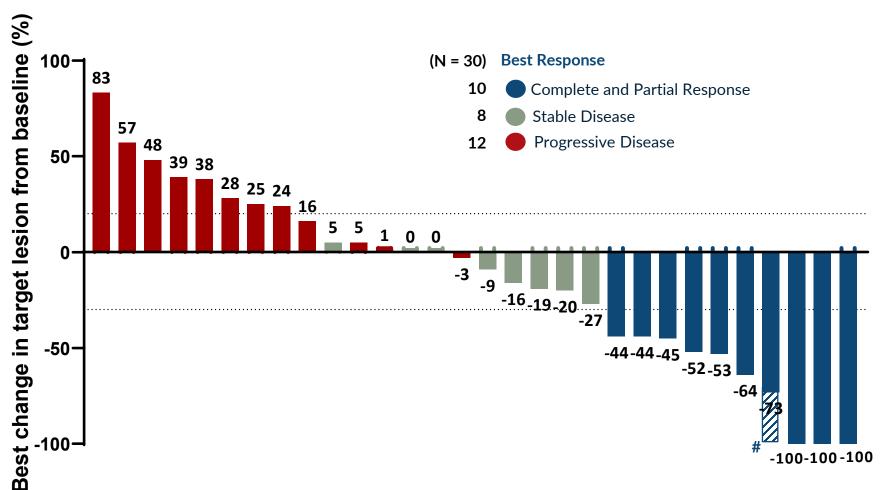
- Goal is to demonstrate safety, immune responses, and enhanced efficacy (ORR, PFS, OS) compared to single agent anti-PD1 therapy
- N = 24 patients (Johns Hopkins University, Mount Sinai, New Zealand Clinical Research)
- Study expanded to n = 36 patients based on current promising data





## GT-30 ITT: 32.3% ORR IN THE FIRST 31 PATIENTS (3 CR, 7 PR)

Best Response in 30 evaluable patients (as of Feb 10, 2023)\*



Disease Control Rate (DCR) per RECIST 1.1: 58.1% (CR/PR/SD, 18/31)

Target Lesion Control Rate: 67.7% (21/31)

# Pt 2105, a confirmed PR, achieved secondary resectability



## GT-30-2915: COMPLETE RESPONSE

73 yo, white male
HCC (Aug2019)
Microwave ablation (Aug2019)
TACE (Dec2020 and Jan2021)
Lenvatinib (Jun2021; BOR SD)
GNOS-PV02 (Sep2021)
Etiology: non-viral
T2N0M0 (II) BCLC B
Beta-catenin mutation
(CTNNB1 S45F)

### **Significant Medical History:**

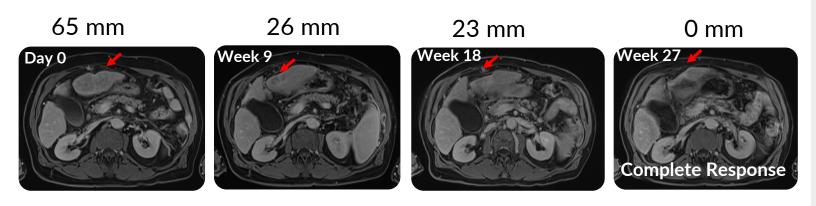
Hepatitis C, Cirrhosis, heavy alcohol intake

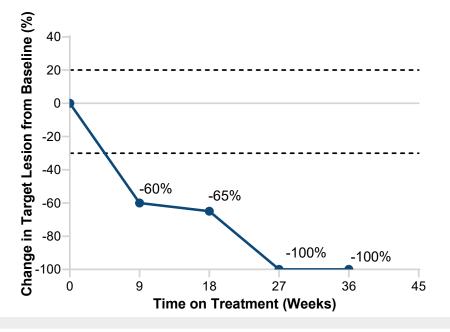
Neos: 40

PTCV doses: 9 Status: On study

**SAEs:** 2, unrelated to study treatments **AEs related to PTCV/pIL12/EP:** 2, both

Grade 1







# GT-30-2105: Reduction in Liver Primary & Two Lung Mets Creates Path to Resection Resulting in Cancer Free Status

68 yo, white female HCC (May2019) Lenvatinib (Jun2019) Radiotherapy (Jul2020) GNOS-PV02 (Nov2021) Etiology: non-viral T4N0M1 (IVB) BCLC C

### **Significant Medical History:**

- Hypertension, diverticulitis, GERD
- DVT, Meningioma

Neos: 40

PTCV Doses: 5
Status: Follow up

**Hepatectomy Surgery: 20Apr22** 

Pembro Q6W: 17May22

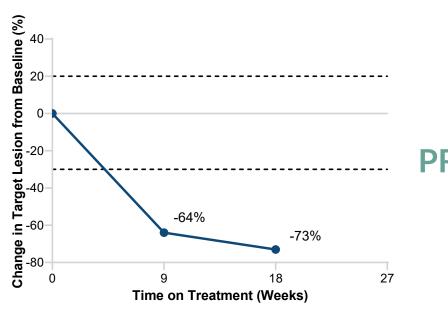
No SAEs

No AEs related to PTCV/pIL12/EP

Target lesions (mm)			
Location	d0	w9	w18
Lung LUL	11	4	2
Lung RML	22	8	7

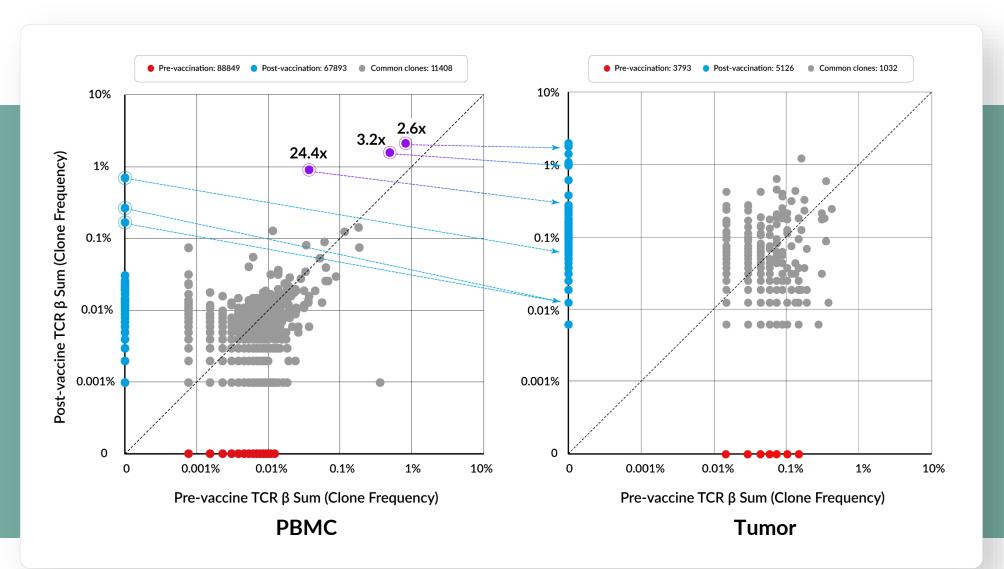
Patient with liver primary and two lung mets; liver biopsied to determine neos for design of PTCV but lung lesions used as target lesions for RECIST1.1

- After 5 PTCV+pIL12 doses, liver lesion shrank to point of secondary resectability and lung mets shrank to point of being amenable to XRT
- Lesions responded fully to XRT and surgery
- Patient now disease and recurrence free





# GT-30 CASE STUDY: POST-VACCINATION INCREASE IN PERIPHERAL T CELL CLONES THAT INFILTRATE THE TUMOR



# PBMC (Blood) and Tumor:

Expansion of several pre-vac clones (purple) and detection of multiple new T cell clones (blue) post-vaccination

Arrows highlight infiltration of high frequency clones from blood into the tumor post-vaccination



### **Clinical Efficacy**

- 3 CR & 7PRs detected by RECIST 1.1 out of first 31 patients on treatment and 30 evaluable as of February 2023
- Overall, 16/31 patients with some level of tumor reduction; 18/31 with CR/PR/SD on treatment (DCR)
- Next data readouts: 36 Pts in Q3 2023

### Safety Data

- No treatment related SAEs noted to-date
- 200+ doses of GNOS-PV02+pIL12 and 350+ doses of pembrolizumab across 31 subjects treated

### **Treatment Feasibility**

- Personalized cancer treatments can be designed, manufactured, & administered successfully
- Delivered up to 40 neoantigens/patient
- 6-8 week turnaround time feasible



# GT-10 CASE STUDY: EFFICACY FROM A SECOND TUMOR TYPE EXPANDS CLINICAL APPLICABILITY

PT # 10-101: 21y Female, Anaplastic Astrocytoma/GBM

### **Diagnosis & Primary Treatment**

- IDH positive, MGMT methylated
- Two surgeries; Radiation; Temozolomide

#### **GT-10 Treatment**

Monotherapy with GNOS-PV + plL12

### Single Patient Compassionate Use IND

- PCV contains both MHC Class I and II antigens
- 30 antigens (27 neoantigens; 3 shared antigens)

Patient treated only with PCV monotherapy

## GT-10 Treatment & Outcome (as of June 30<sup>th</sup>, 2022; Treatment ongoing):

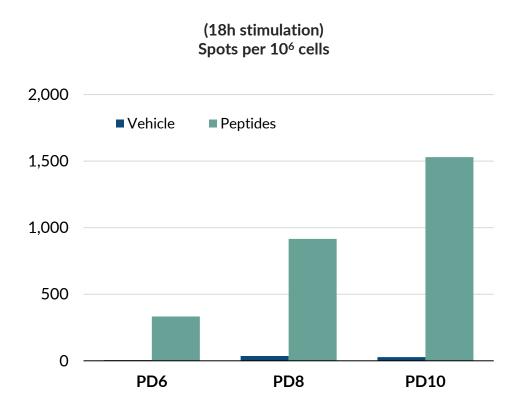
- Patient is recurrence free 60+ mo since 1° surgery & 48+ mo on GT-10 treatment
- No related SAEs

Extended RFS/PFS, OS - illustrates the persistence of response to GT-EPIC™ immunotherapy



# GT-10 IFN $\gamma$ ELISPOT ANALYSIS: RESPONSES DETECTED TO 93.3% ENCODED ANTIGENS

# Cumulative ELISpot data from vaccine antigens show induction of a robust response



### Responses detected to:

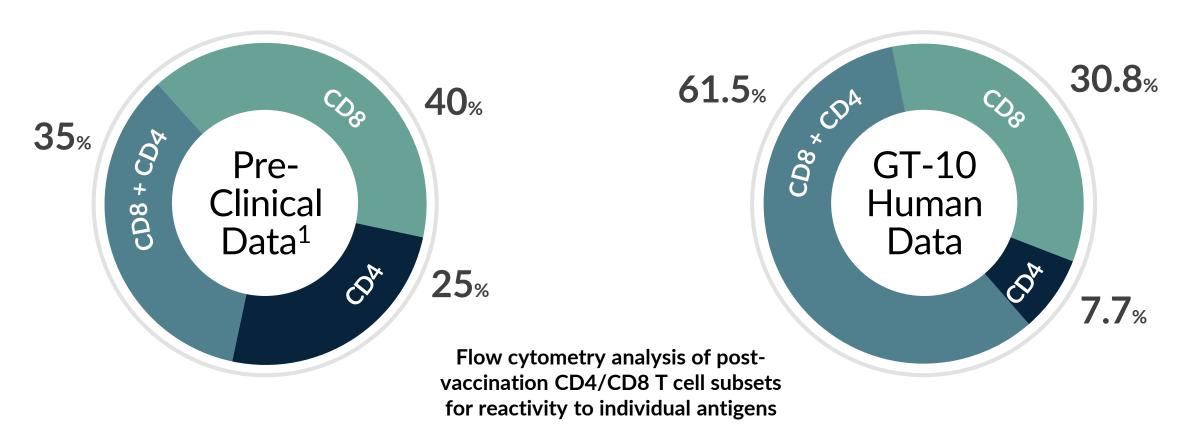
3/3 (100%) shared tumor antigens 25/27 (92.6%) neoantigens

(30 antigens)





# GT-10 IMMUNE PHENOTYPING: GT-EPIC™ VACCINES INDUCE CD8 T CELL RESPONSES TO HIGH PROPORTION OF ENCODED ANTIGENS



- Humans: 92% of the epitopes tested individually yield CD8 T cells
- Pre-clinical (mouse) models: 75%



# GT-EPIC™ CLINICAL DATA SUPPORTS A MOA BASED ON TUMOR-SPECIFIC IMMUNE-ACTIVATION

### Strong induction of antigen-specific T cells

- IFNγ-producing T cells detected by ELISpot analysis
- Polyfunctional T cells TNF, IFNγ, multiple cytokines, activation markers detected in antigen-specific manner

### CD4 & CD8 T cell responses

• CTL phenotype

### **Tumor Infiltration by Lymphocytes (TILs)**

- Pre- vs. on-treatment TCR sequence analysis demonstrates expansion of new clones in both blood and tumor tissue
- Infiltration of newly expanded clones into the tumor

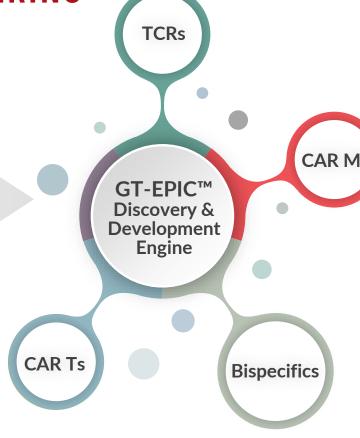


VALIDATED MOA DRIVES GT-EPIC™ DISCOVERY ENGINE FOR NOVEL TCRs TO HIGH VALUE TARGETS FOR PARTNERING **→** 

# **GT Personalized Vaccine Administration** geneos THERAPEUTICS MANUFACTURE PATIENT SPECIFIC PRODUCTS IN LESS THAN HALF THE TIME OF **OUR COMPETITORS** shipped for DNA **On-Treatment Immune Monitoring** 3. Proprietary neoa ELISPOT - 200311 Sample - Individual epitopes DNA insert design optimization

#### **Discovery Engine Highlights:**

- Vaccine induced CD8 T cells
- Over 800+ (neo) antigens encoded in PCVs to date
- Select Mutations/Neos include: p53, β-catenin
- Select shared cancer antigens: Survivin, gp100



Validated TCRs, BCRs to novel antigens provide additional assets for partnering in the cell therapy space - CARTs, CAR Macrophages, TCRs, Bispecifics



## GENEOS SUMMARY



**Compelling Clinical Data** 



High Neoantigen Payloads
Drive CD8s & TILs



**Multiple Upcoming Catalysts** 



**Strong Discovery Platform** 





BRINGING PATIENT SPECIFIC TUMOR TARGETED IMMUNOTHERAPIES TO THE WORLD, ONE PATIENT AT A TIME.

Dr. Niranjan Y. Sardesai

