

Precision Immunotherapy for Solid Tumors

Non-Confidential Corporate
Presentation
January 2025

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RIGHT TARGETS. RIGHT CELLS. RIGHT PLACE.

We target oncogenic driver mutations to deliver transformative therapies for patients with solid tumors

- **Leader in Precision Immunotherapy** - developing a deep pipeline of TCR-based therapies that have **first-in-class / best-in-class potential**
- **Focus on targeting the most frequent oncogenic driver mutations** in solid tumors; including KRAS, P53, and PIK3CA
- **Proprietary platform technologies** to build potent and persistent T cell therapies and generate bispecific T cell Engagers
- Science-driven team and founders focused on continued innovation to **develop novel therapies with curative potential**

Development Pipeline Milestones



AFNT-211

A11 KRAS G12V

- Lead KRAS targeting program
- Phase 1a data generation ongoing in 2L+ solid tumor indications
- Dose escalation proceeding on track across ~10 US sites with indication-specific expansions planned

Completion of Dose Escalation
anticipated 2H25



AFNT-212

A11 KRAS G12D

- Doubles addressable KRAS population
- Introduces THRIVE non-viral gene-editing platform to enable future product development
- IND-enabling studies complete, IND submission planned 1H25

IND clearance
anticipated 2025



AFNT-313

A2 P53 R175H

- Expands beyond KRAS to address largest P53 population
- Differentiated development candidate designed to integrate immunostimulatory signals for optimal T-cell activation
- IND-enabling studies underway

Pre-IND feedback
anticipated 2025

Lead programs offer attractive US market opportunities with blockbuster potential

AFNT-211

6.3k

Colorectal	1,800
Pancreatic	1,900
Lung	1,500
Other	1,100

A11 KRAS G12V

AFNT-212

7.7k

Colorectal	2,600
Pancreatic	2,500
Lung	1,100
Other	1,500

A11 KRAS G12D

AFNT-313

13.4k

Colorectal	4,300
Breast	2,200
Pancreatic	1,200
Other	5,700

A2 P53 R175H

Cell therapies approved for solid tumors may benefit from an attractive development path



- First TCR T cell (A2 MAGE A4)
- Single arm registrational study - 44 pts with **synovial sarcoma**

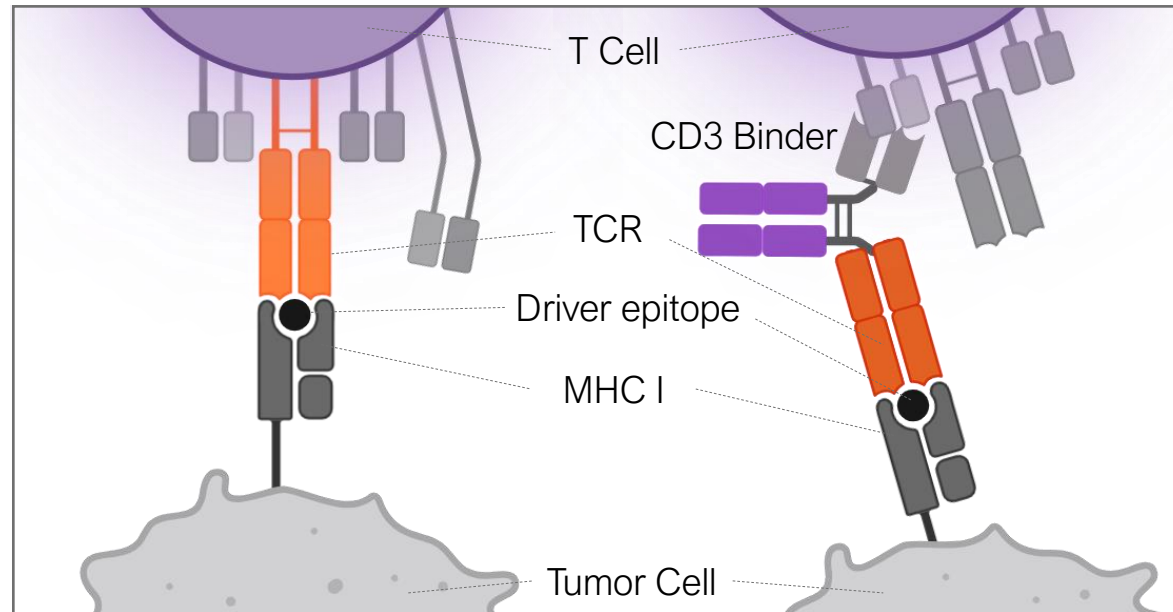


- First TIL therapy
- Single arm registrational study - 73 pts with **advanced melanoma**

Affini-T is Developing Two TCR-Based Therapeutic Modalities

TCR-T Cell Therapies

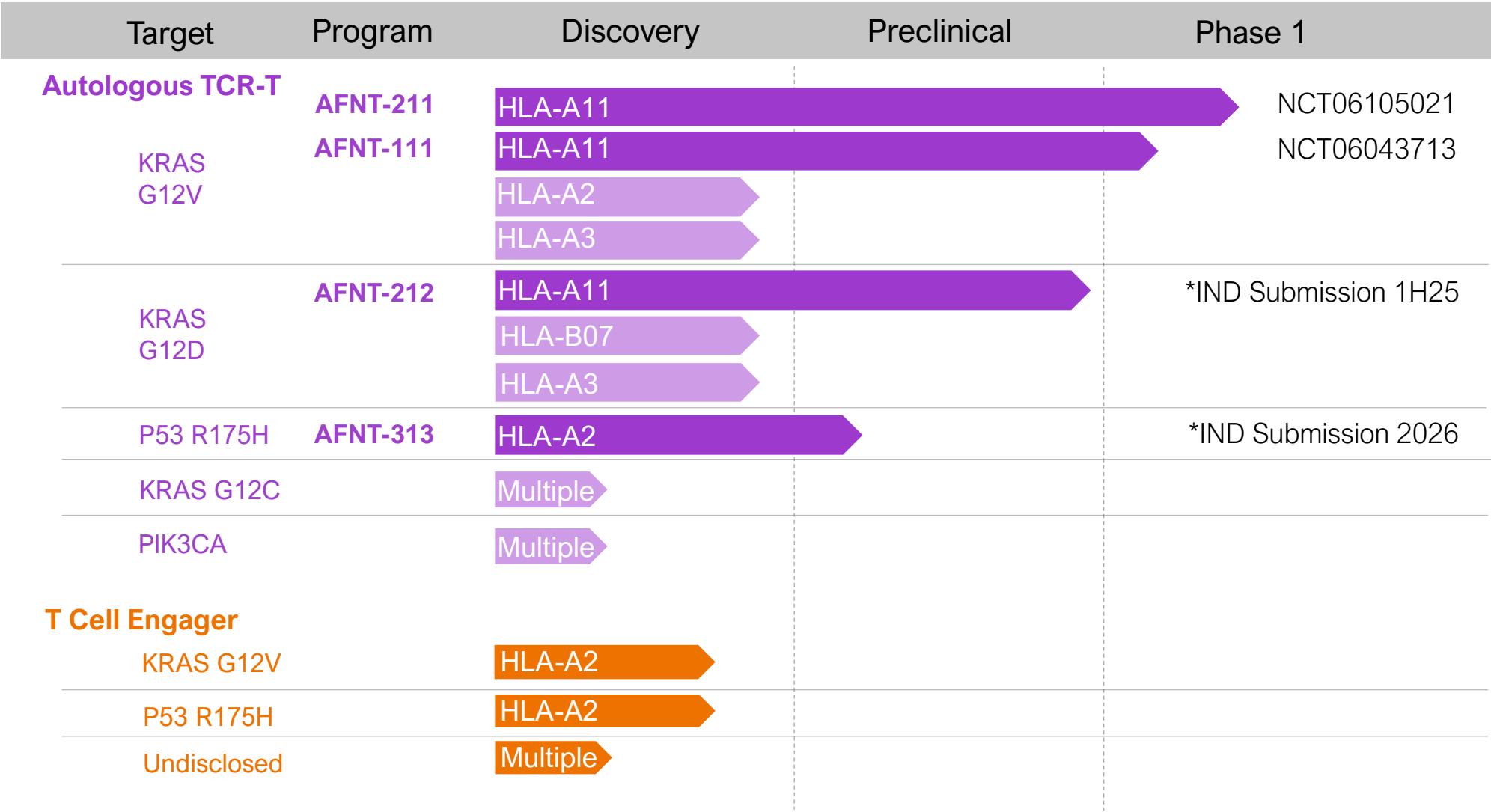
T cells engineered with a transgenic TCR that allows recognition of specific driver mutant epitopes



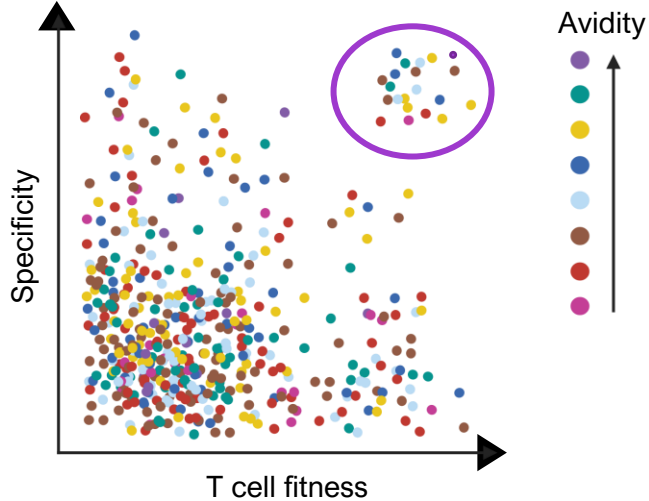
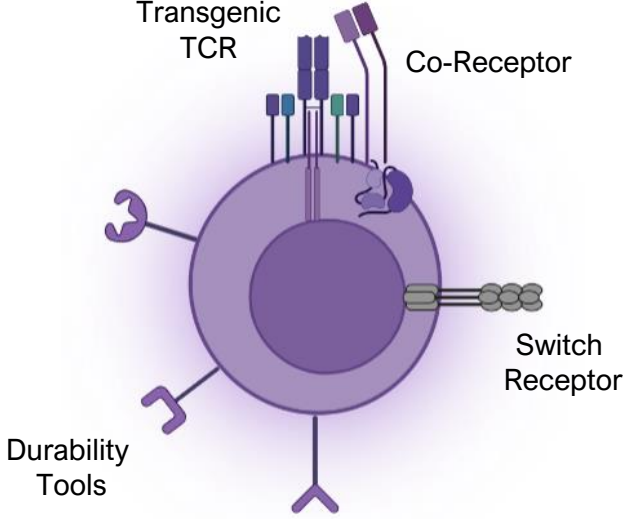
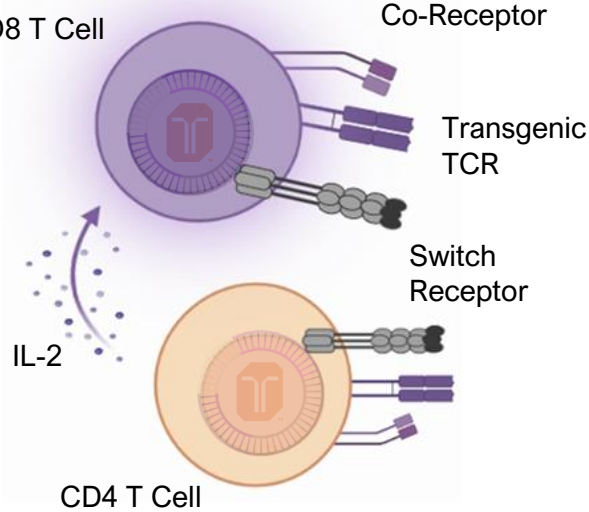
Bispecific T Cell Engagers

Bispecific biologics combining a TCR moiety to recognize the driver mutant epitope with a CD3 binding moiety to recruit endogenous T Cells

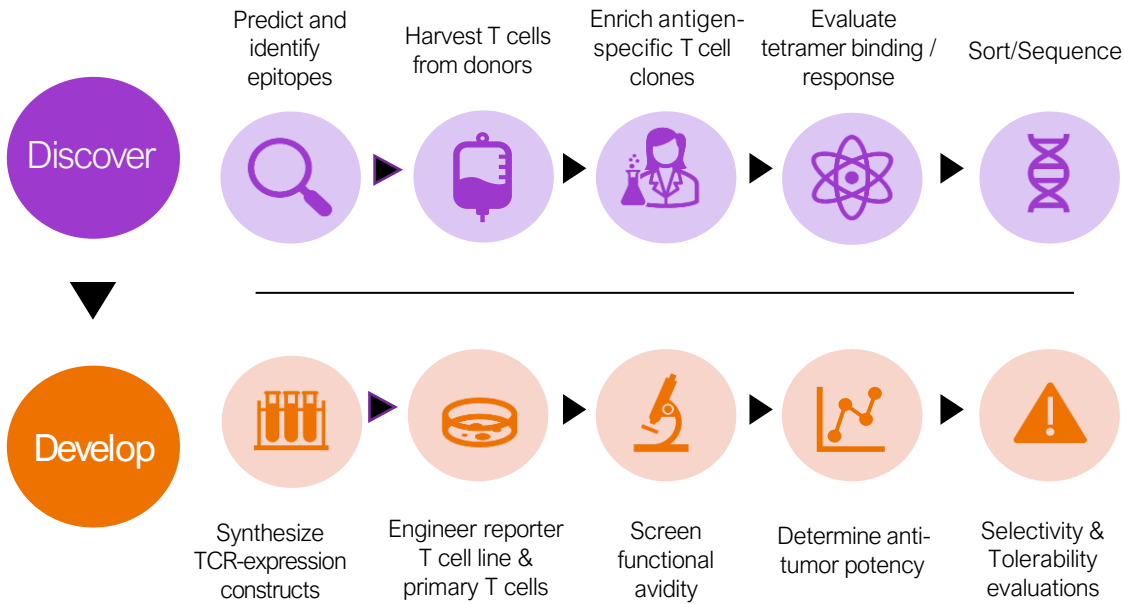
First-In-Class Potential for Multiple Products Targeting Oncogenic Drivers in Solid Tumors



Affini-T Platform Technologies Designed To Generate Potent & Tolerable TCR-T Cells

<p style="text-align: center;">TAILOR™ TCR Discovery</p>	<p style="text-align: center;">TUNE™ Synthetic Biology</p>	<p style="text-align: center;">THRIVE™ Engineering and Manufacturing</p>
 <p>A scatter plot with 'Specificity' on the y-axis and 'T cell fitness' on the x-axis. The plot contains numerous colored dots representing different TCRs. A legend on the right indicates 'Avidity' levels with a color scale from purple (high) to pink (low). A purple circle highlights a cluster of high-specificity, high-fitness TCRs.</p>	 <p>A diagram of a purple T cell. It features a 'Transgenic TCR' (purple) and a 'Co-Receptor' (purple) on its surface. 'Durability Tools' (purple) are shown as various receptors on the cell. A 'Switch Receptor' (grey) is also present. The cell is shown in a cross-section view.</p>	 <p>A diagram showing two T cells. The top one is a 'CD8 T Cell' (purple) and the bottom one is a 'CD4 T Cell' (orange). Both have a 'Transgenic TCR' (purple) and a 'Co-Receptor' (purple). The CD4 T cell also has a 'Switch Receptor' (grey). 'IL-2' (represented by small blue dots) is shown being released from the CD4 T cell towards the CD8 T cell.</p>
<p>Optimized system that integrates decades of learning with predictive algorithms to identify highly functional & specific TCRs against diverse targets</p>	<p>Program cell persistence in the TME to enhance durability with switch receptors (eg Fas-41BB), co-receptor and other armoring technologies</p>	<p>Scalable manufacturing with gene editing technologies to generate a robust engineered product with high stemness phenotype</p>

TAILOR™ TCR Discovery Platform Validated Across Multiple Programs

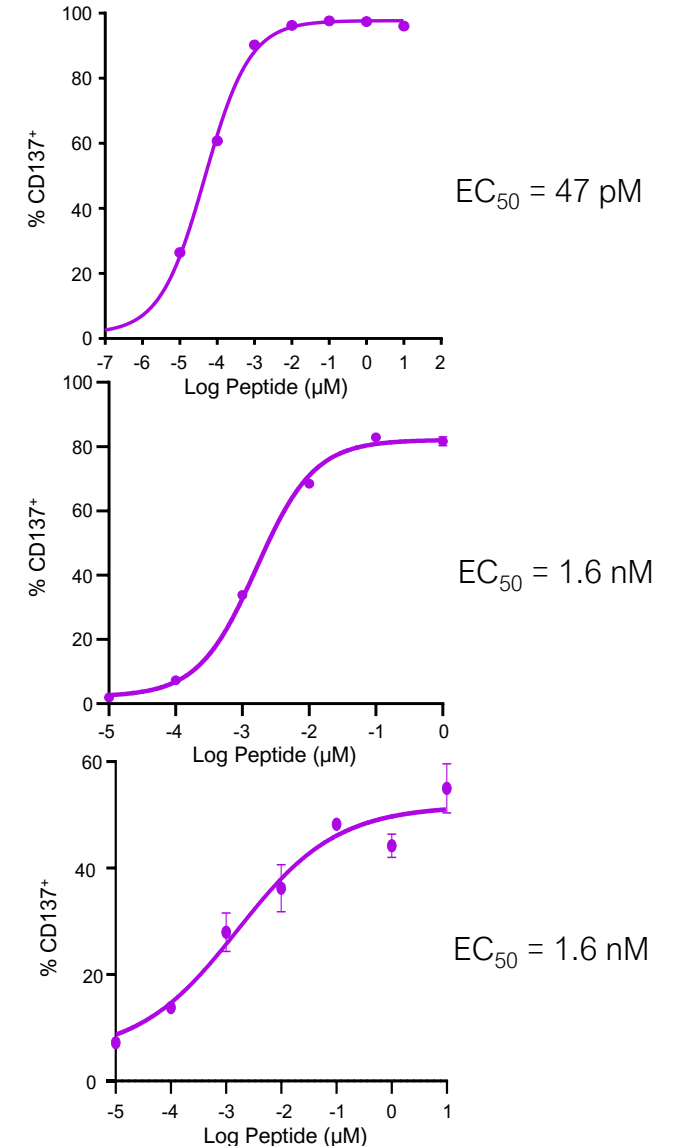


- High-throughput method identifies rare, high-affinity, thymus-vetted TCRs for clinical translation
- Selection based on avidity and preclinical tolerability profile: No cross-reactive self-peptides by XScan, and no allo-reactive responses by B-LCL screening


KRAS A11 G12V TCR
AFNT-111
AFNT-211

KRAS A11 G12D TCR
AFNT-212

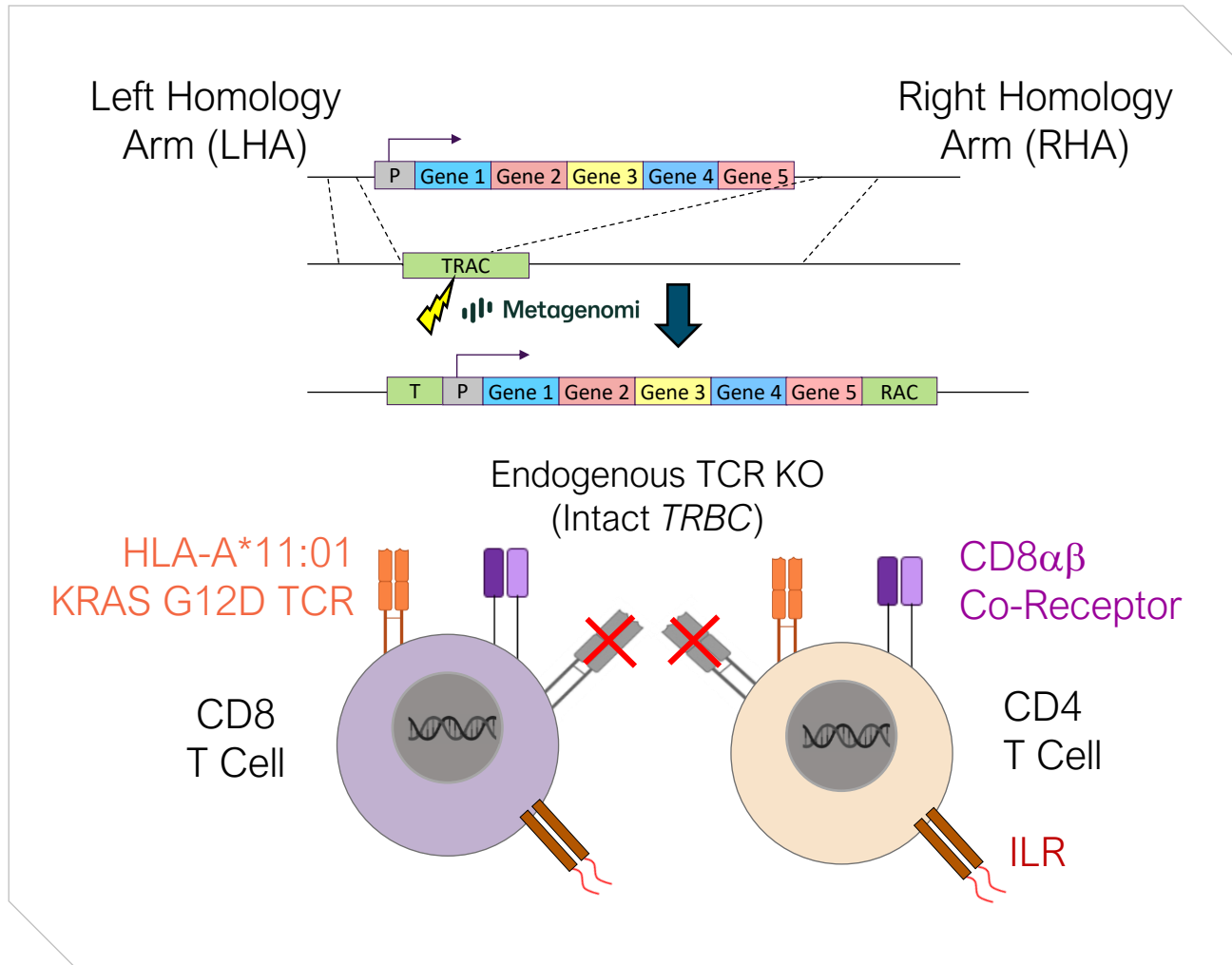
P53 A2 R175H TCR
AFNT-313



TUNE™ Enhancing T Cell Fitness and Persistence with Novel Synthetic Biology

Syn-Bio Component	Purpose	Product	Development Stage		
			Discovery	Preclinical	Clinical
CD8 α / β coreceptor 	<ul style="list-style-type: none"> • Signal 1 (TCR targeting) in CD4+ T cells • Coordinated immune response 	AFNT-111 AFNT-211 AFNT-212 AFNT-313			
FAS-41BB 	<ul style="list-style-type: none"> • Signal 2 (co-stimulation) upon FASL binding: proliferation, memory, and enhanced metabolic function • Prevention of T cell death mediated by apoptosis in the TME 	AFNT-211 AFNT-313			
ILR (undisclosed) 	<ul style="list-style-type: none"> • Signal 3: Constitutive cytokine support • Proliferation, survival in the TME 	AFNT-212 AFNT-313			

THRIVE™ Non-Viral Knock-In (KI) Provides Advantages over Vector Based Platforms



Larger cargo size

Enables engineering with multiple syn-bio components (up to 10kb)

Targeted integration

Reduces potential risk compared to random virus integration

Consistent expression

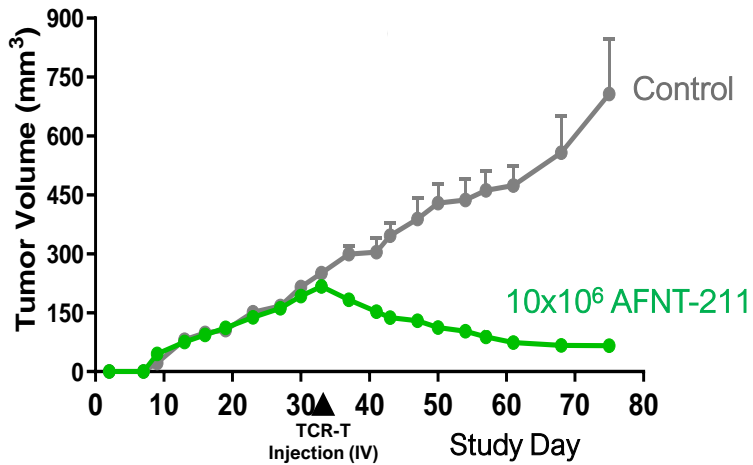
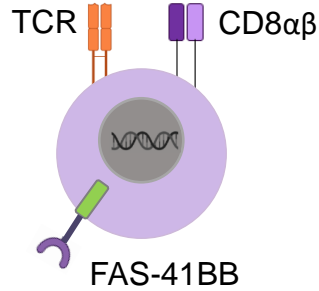
Defined vs. variable copy number

Lower COGS

And faster development timelines

Enhanced survival in TME

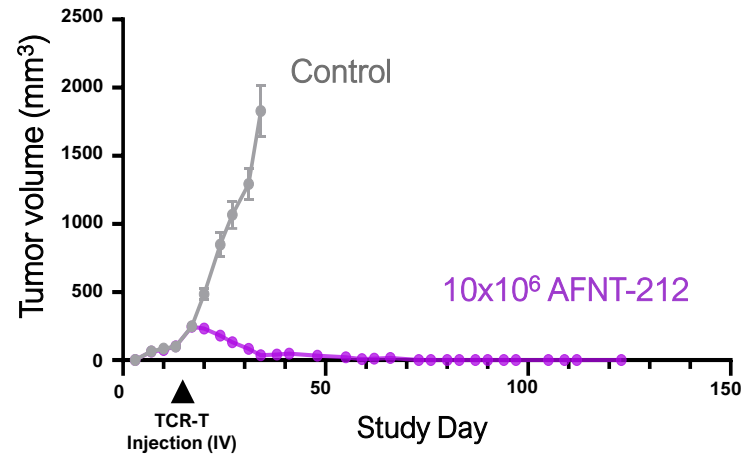
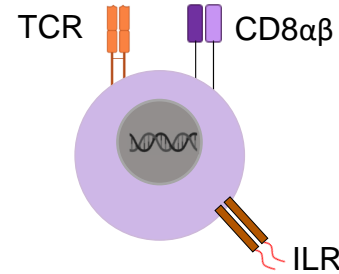
AFNT-211
Integrates
Signal 1+2



AFNT-211 induced tumor regression in established Breast (SW527) tumors with mut. KRAS G12V

Enhanced proliferation in periphery

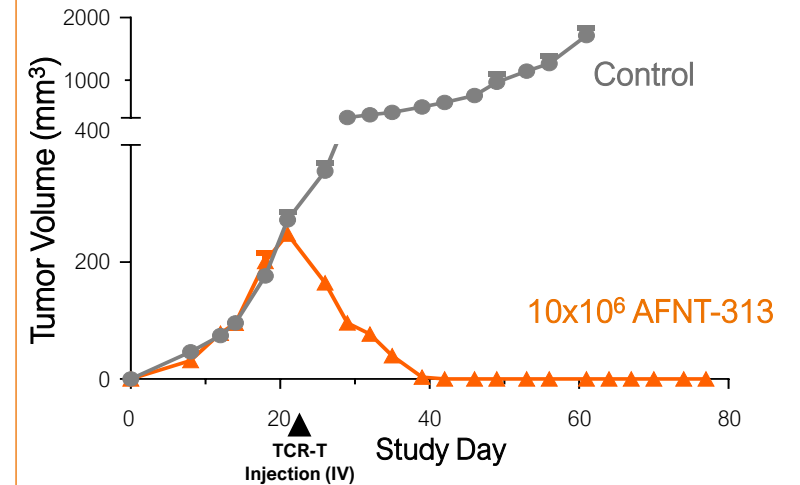
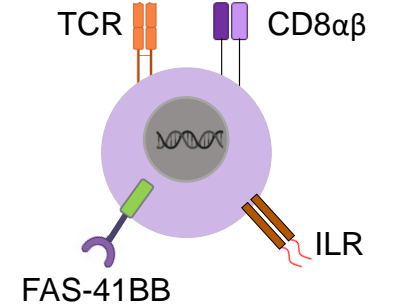
AFNT-212
Integrates
Signal 1+3



AFNT-212 induced tumor eradication in established Colorectal (CL40) tumors with mut. KRAS G12D

Support in TME & Periphery

AFNT-313
Integrates
Signal 1+2+3



AFNT-313 induced tumor eradication in established Ovarian (TYK-nu) tumors with mut. p53 R175H

AFNT-211: A11 KRAS G12V TCR Engineered T Cells + FAS-41BB Durability Switch Receptor

TAILOR™

KRAS G12V TCR



CD8αβ Co-Receptor

Synthetic Biology
(FAS-41BB)

TUNE™

Cell Engineering
THRIVE™

Activated
CD8 T Cell

Activated
CD4 T Cell

IL-2

CD8 Co-Receptor

KRAS TCR

FAS-41BB FAS-L

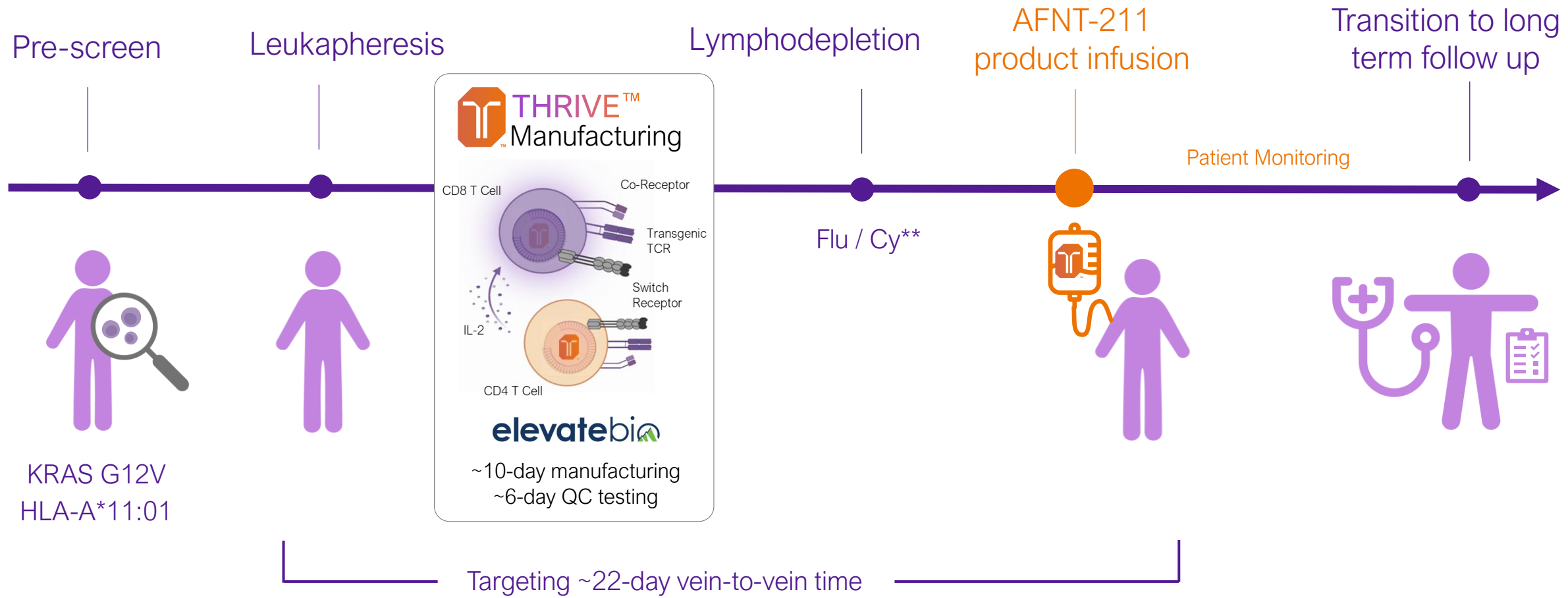
Peptide / MHC
Complex

Perforin + Granzyme
IFN-γ / TNF-α

Tumor Cell Death

IFN-γ / TNF-α

AFNT-211: Patient Journey



**Lymphodepleting chemotherapy (LDC) with cyclophosphamide 500mg/m2/day and fludarabine 30mg/m2/day intravenously (I.V.) on Days -6 to -3, (4 days),

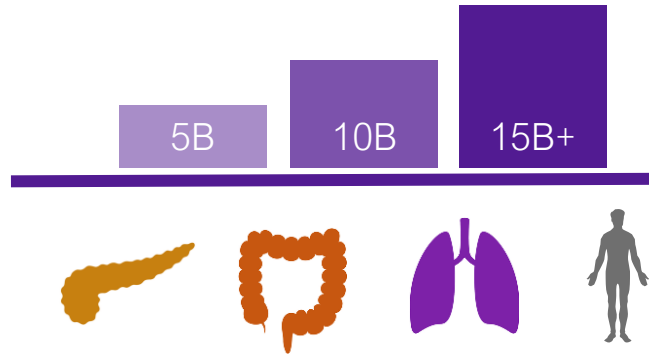
AFNT-211: Clinical Development Plan

Phase 1a Basket Trial Dose Finding

Phase 1b/2 Expansion Cohorts

Registration Study

KRAS G12V-mutated tumors
& HLA-A*11:01 allele
2nd+ Line



Sample size N=15-20
~10 US clinical trial sites

**Optimal Biological Dose /
Proof of Clinical Concept**

PDAC → 2nd/3rd line

CRC → 2nd/3rd line

NSCLC → 2nd/3rd line

Tissue-agnostic → 2nd/3rd line

Sample size up to N=20 per indication

**Interim
Analysis**

Registrational trial design
based on data and FDA
interaction (e.g. single-arm)

Target N=~80 for potential indication
Expand to 35-40 sites US/EU5/CAN

**Aim for approval on
ORR & DoR**

*Excluding primary brain tumors

AFNT-211: Patient Selection & Biomarker Strategy

I. Patient Selection

- **KRAS G12V** mutation – routinely reported by PCR, NGS, and CGP; by tumor or liquid biopsy (ctDNA)
- **HLA A*11:01** – via standard typing assays (Histogenetics – ASHI accredited) or CGP
- **2L+**, Upside: frontline consolidation

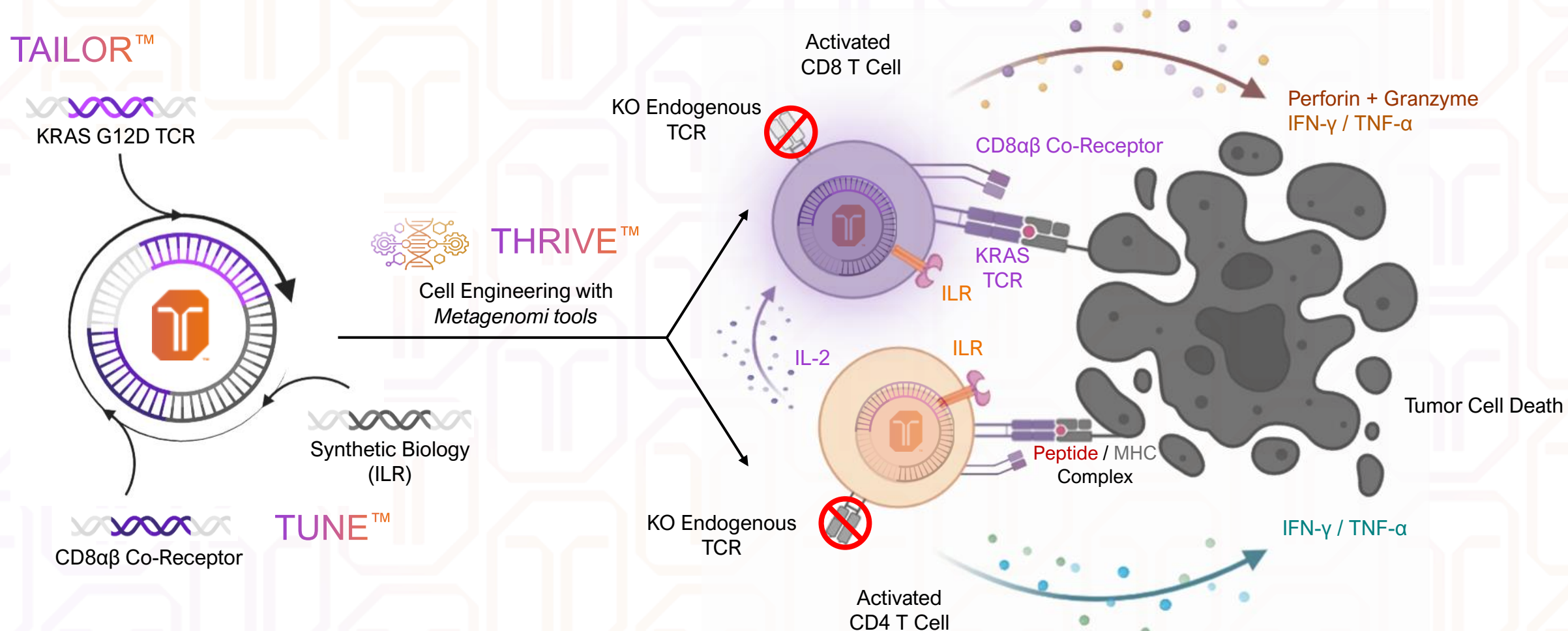
II. Monitoring - Peripheral Blood

- **PK:** TCR-T expansion (VCN and/or CK), C_{max} , T_{last} , AUC
- **PD:** TBNK depletion and reconstitution; cytokines, e.g. IL7, IL15, IFN γ
- **MRD:** ctDNA
- **TCR-T phenotyping:** TCR-T cell differentiation, activation, and exhaustion
- **Safety:** Replication-competent lentivirus, insertion site analysis

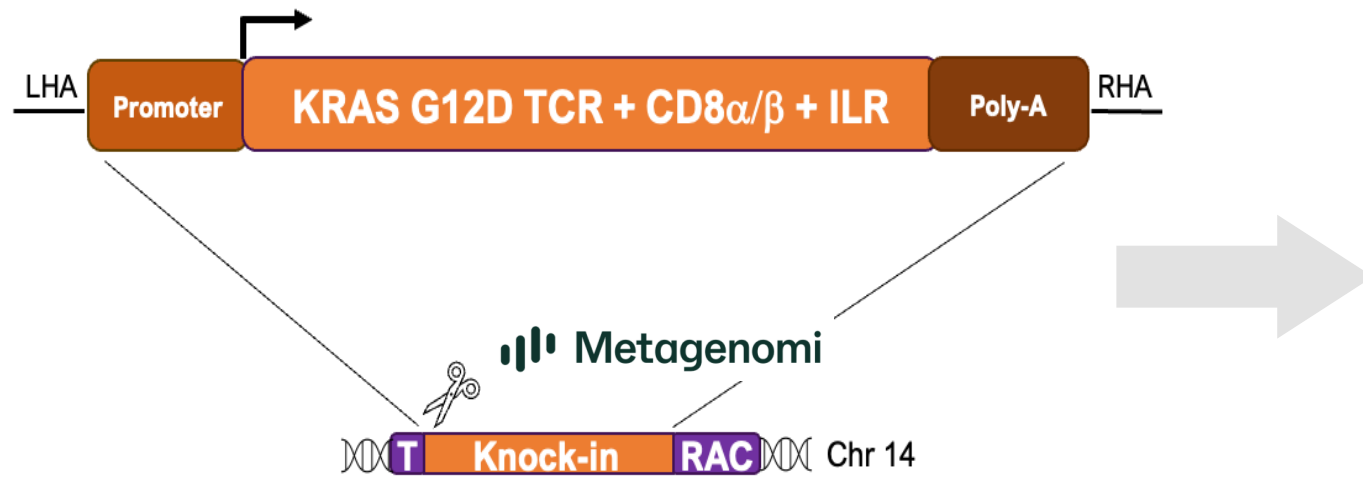
III. Phenotyping - Tumor

- **RECIST:** Imaging response assessment
- **TME:** AFNT-211 TCR-T cell infiltration and phenotyping, Host immune infiltration (including CD4 and CD8)
- **Tumor characterization:** TMB, MSI, PD1, FasL, IFN γ and APM

AFNT-212: A11 KRAS G12D TCR Engineered T Cells + Durability Switch Receptor + Gene Editing



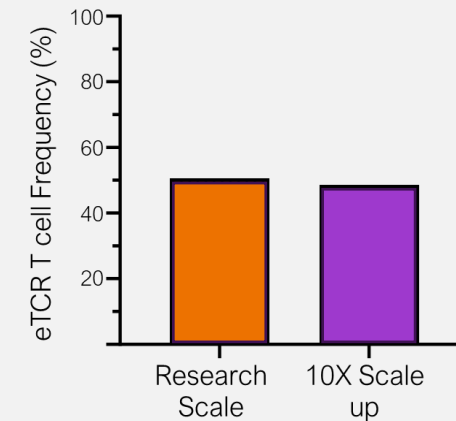
THRIVE™ High Efficiency Non-viral Delivery of Large Transgenes at cGMP Scale



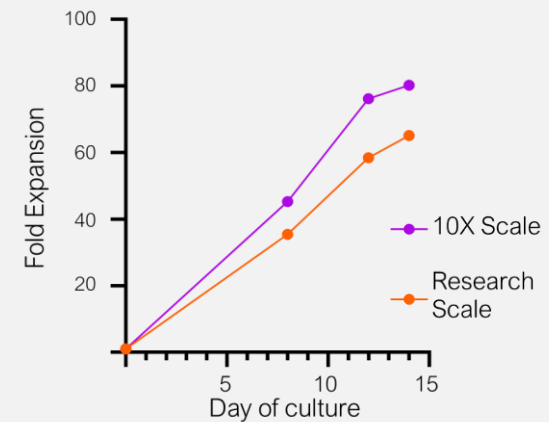
Transgenes inserted within the endogenous TRAC gene via CRISPR/Cas driven homology mediated repair

TRAC-inserted knock-in of 6.3kb 5 gene cassette

Transgene Integration Frequency

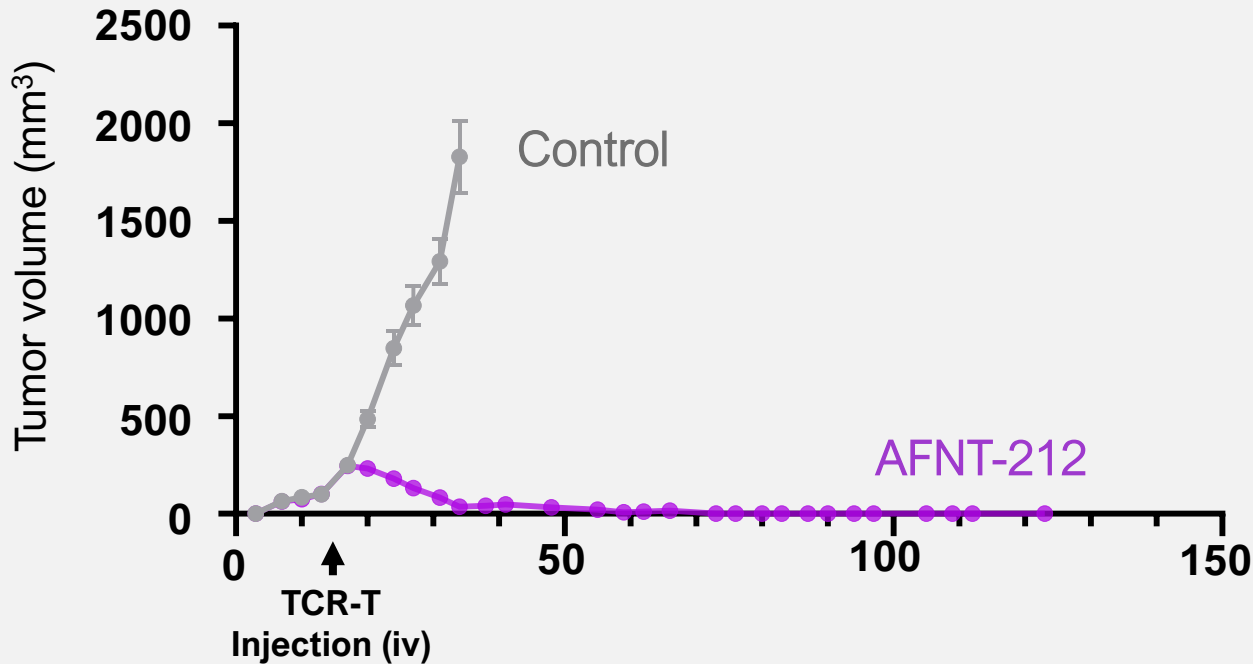


Expansion Kinetics

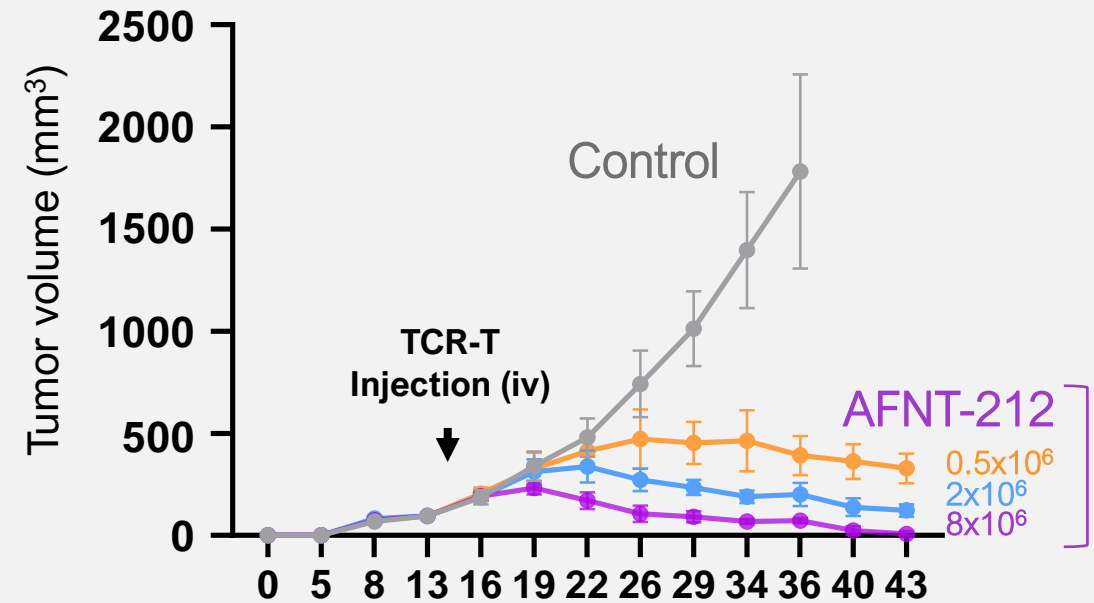


AFNT-212 Showed Robust Anti-tumor Activity in Established Tumor Mouse Models *in vivo*

AFNT-212 T cells induced complete remission for over 120 days

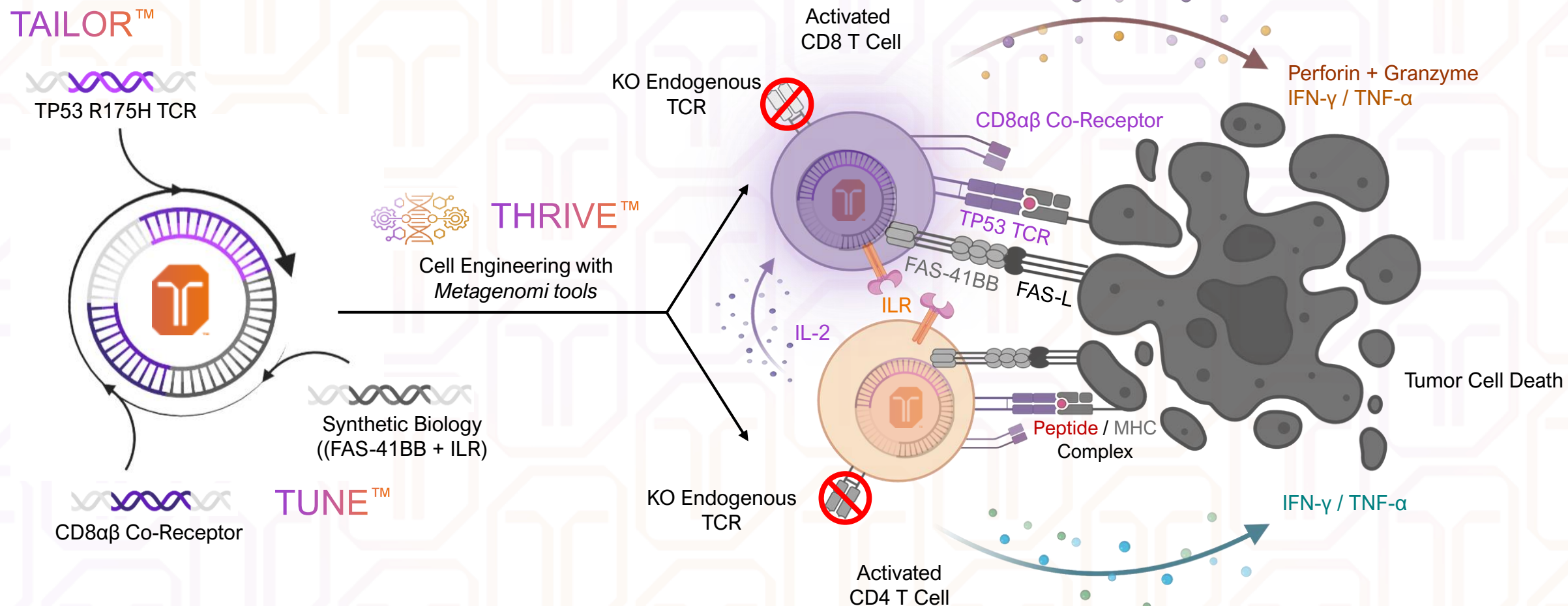


AFNT-212 T cells drove a dose-dependent anti-tumor response

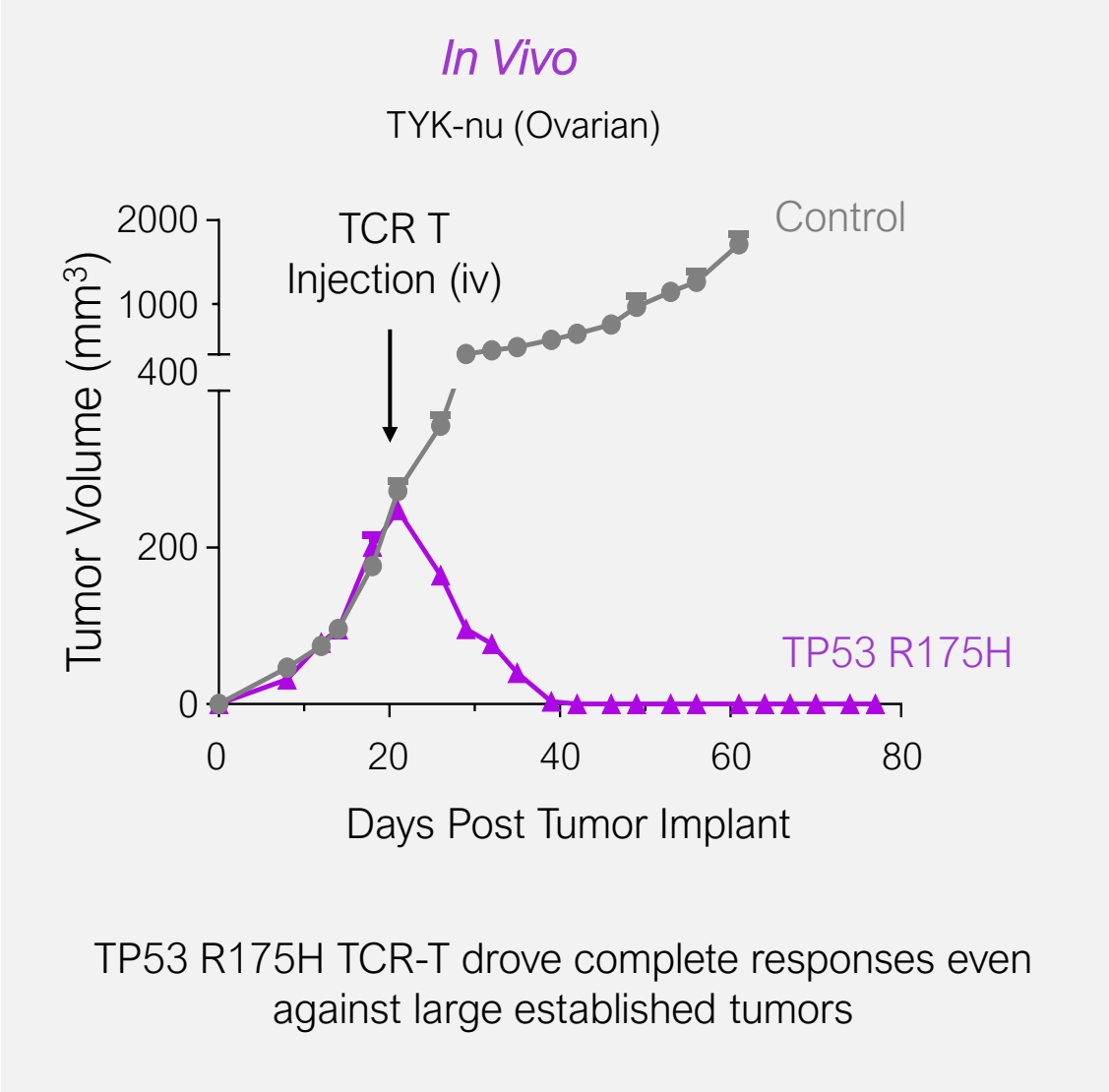
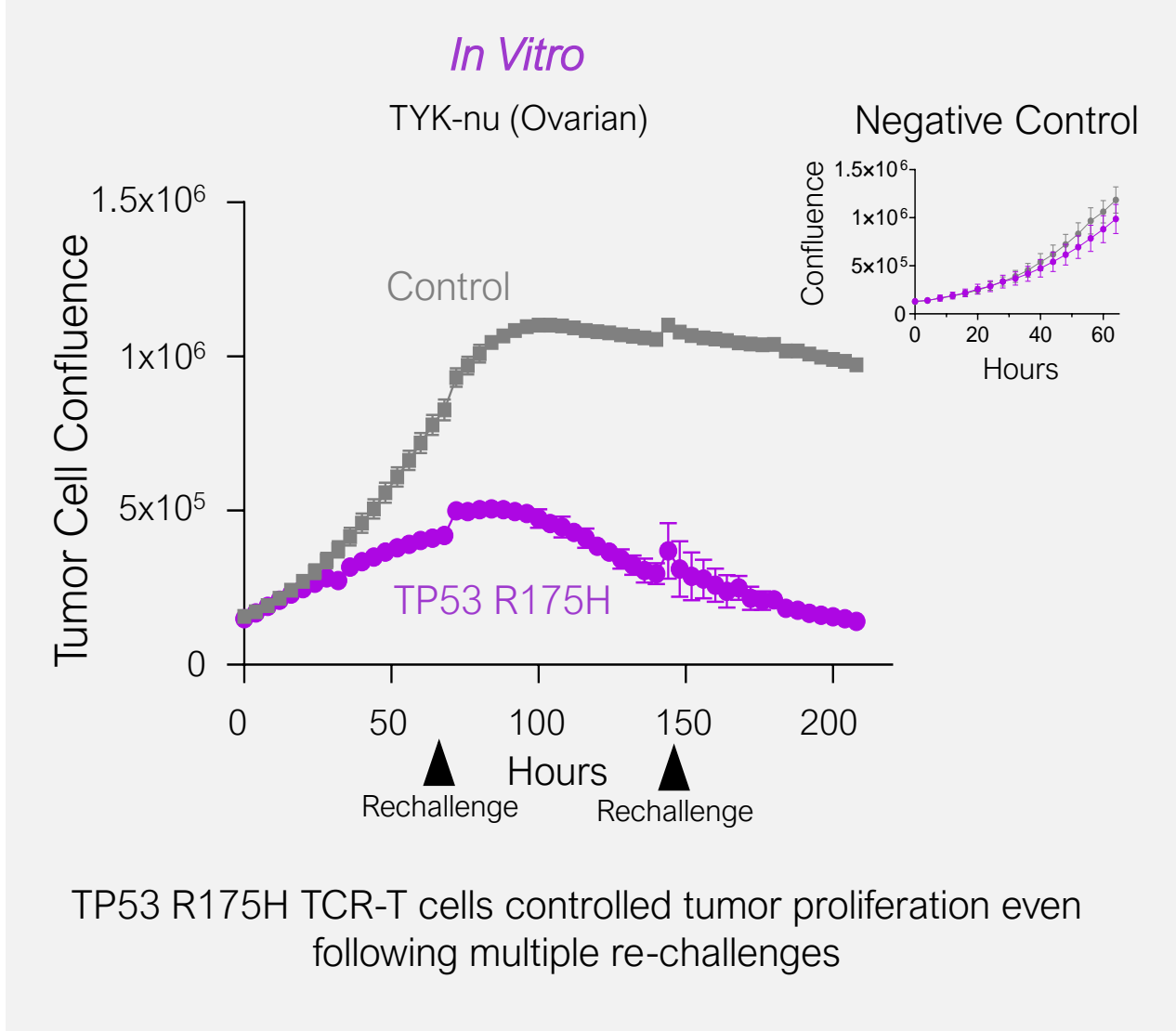


(CL-40 colorectal model)

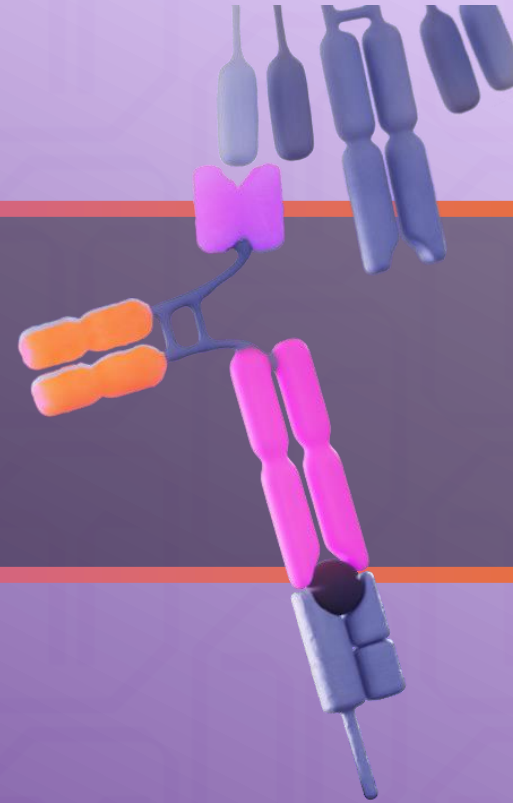
AFNT-313: A2 TP53 R175H TCR Engineered T Cells + 2 Durability Switch Receptors + Gene Editing



AFNT-313 TCR-T Showed Robust Preclinical Tumor Cell Control *In Vitro* and *In Vivo*



TETHER™ T Cell Engagers Highlights

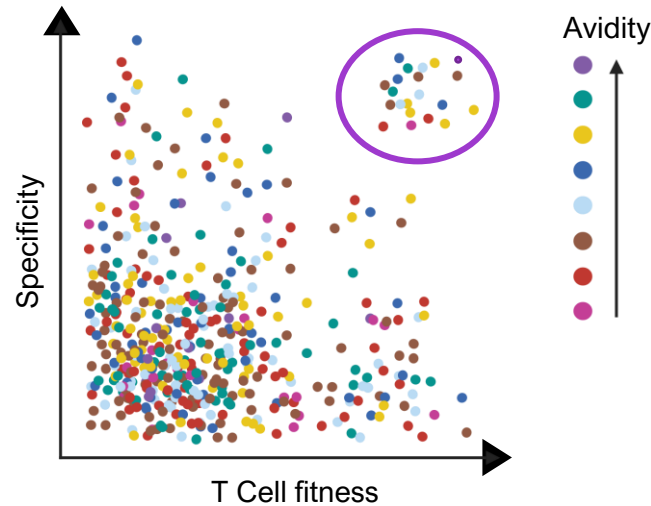


Affini-T Platform Technologies Designed to Generate Highly Specific & Active T Cell Engagers

1

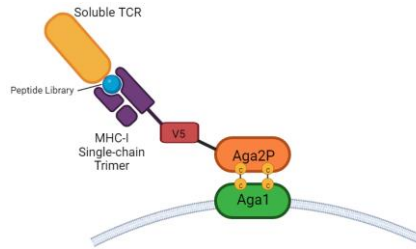
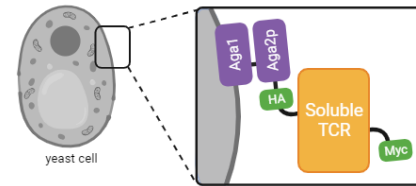
TAILOR™ TCR Discovery

- High throughput screening, predictive algorithms, and decades of learning
- Generate highly functional and tolerable TCRs against diverse targets



2

Affinity Maturation



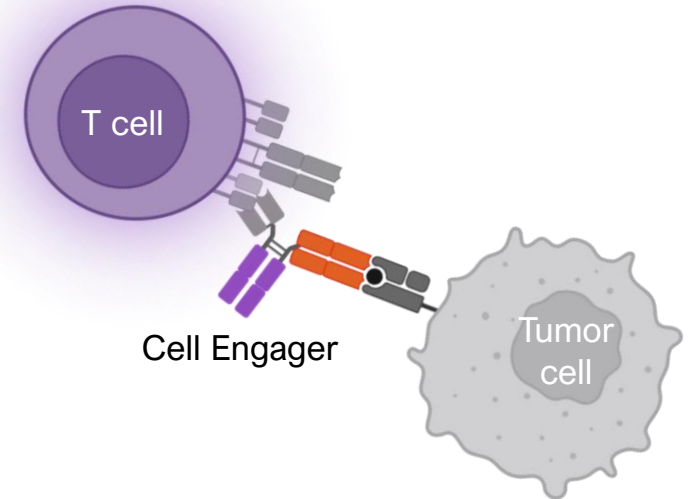
Yeast Display Modalities

- Libraries to identify high affinity TCRs
- Libraries for specificity screenings

3

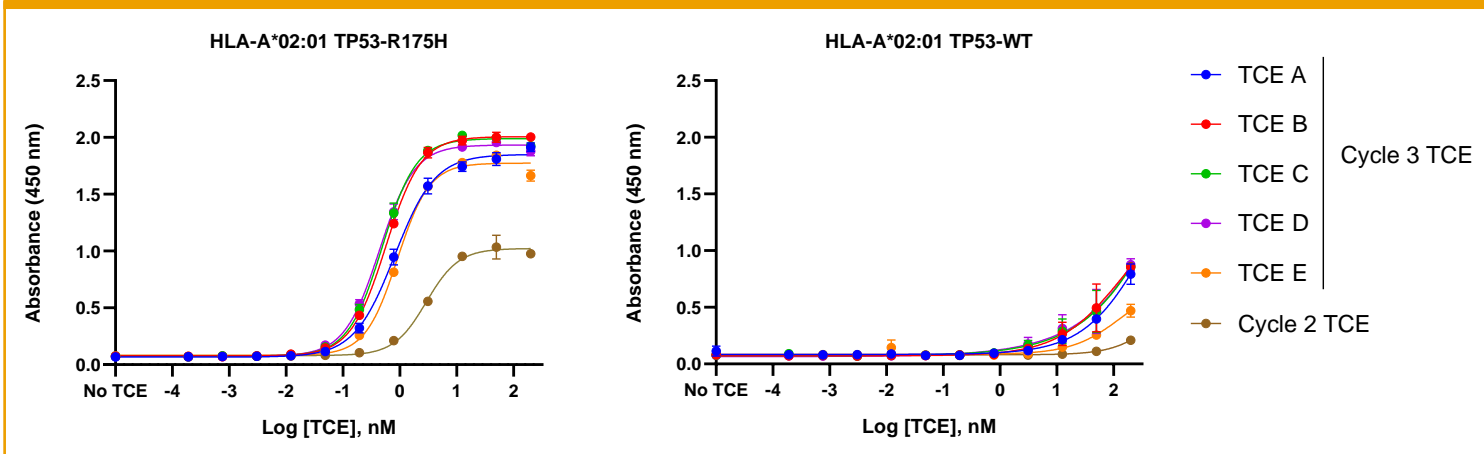
TETHER™ T Cell Engagers

- Affinity matured TAILOR™ TCRs with high specificity and affinity
- Balanced CD3 binders for optimal T cell engagement
- Bispecific T cell engager format with long half-life

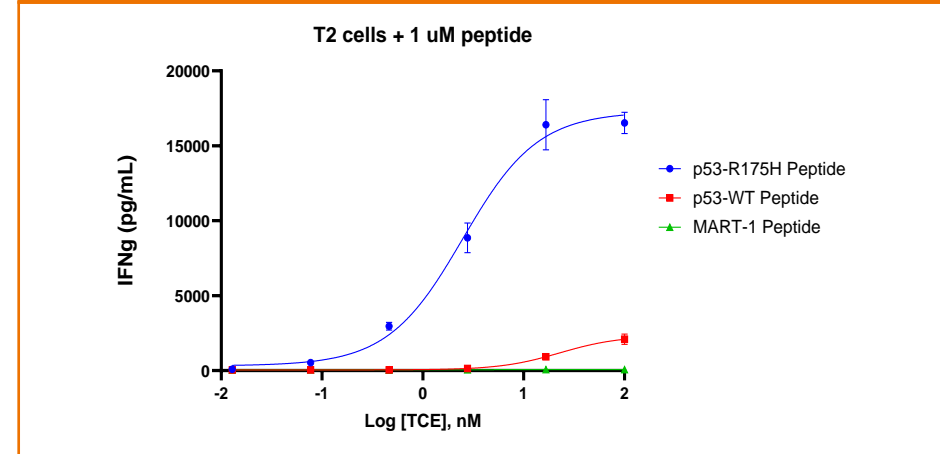


A2 TP53 R175H TETHER TCEs Showed Robust & Specific Tumor Killing & T Cell Activation *In Vitro*

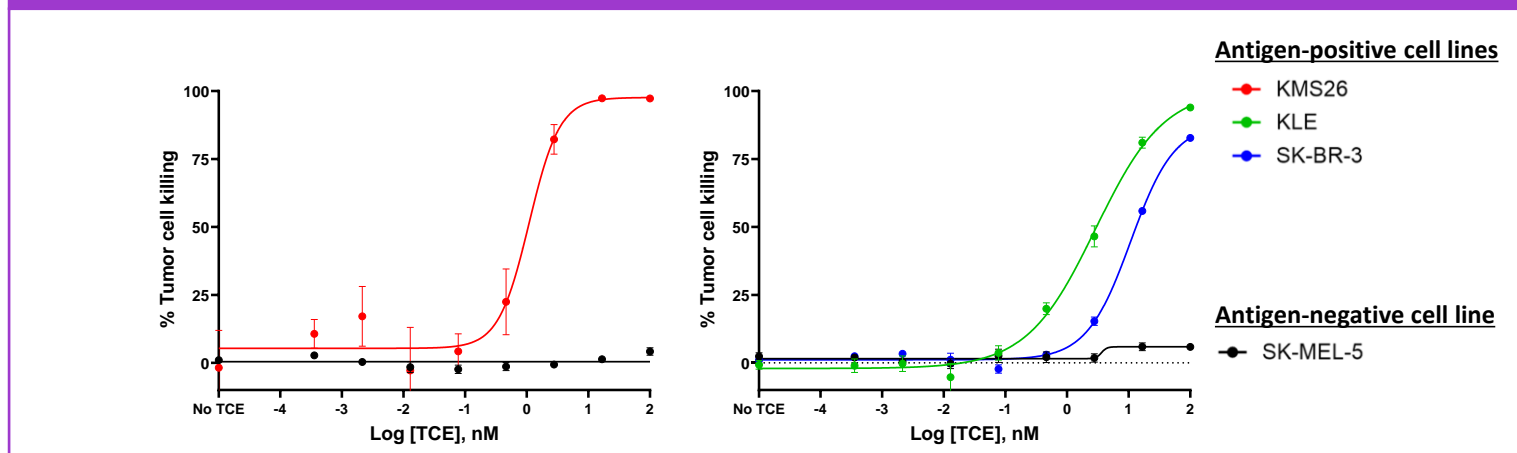
Specific binding



Robust activation

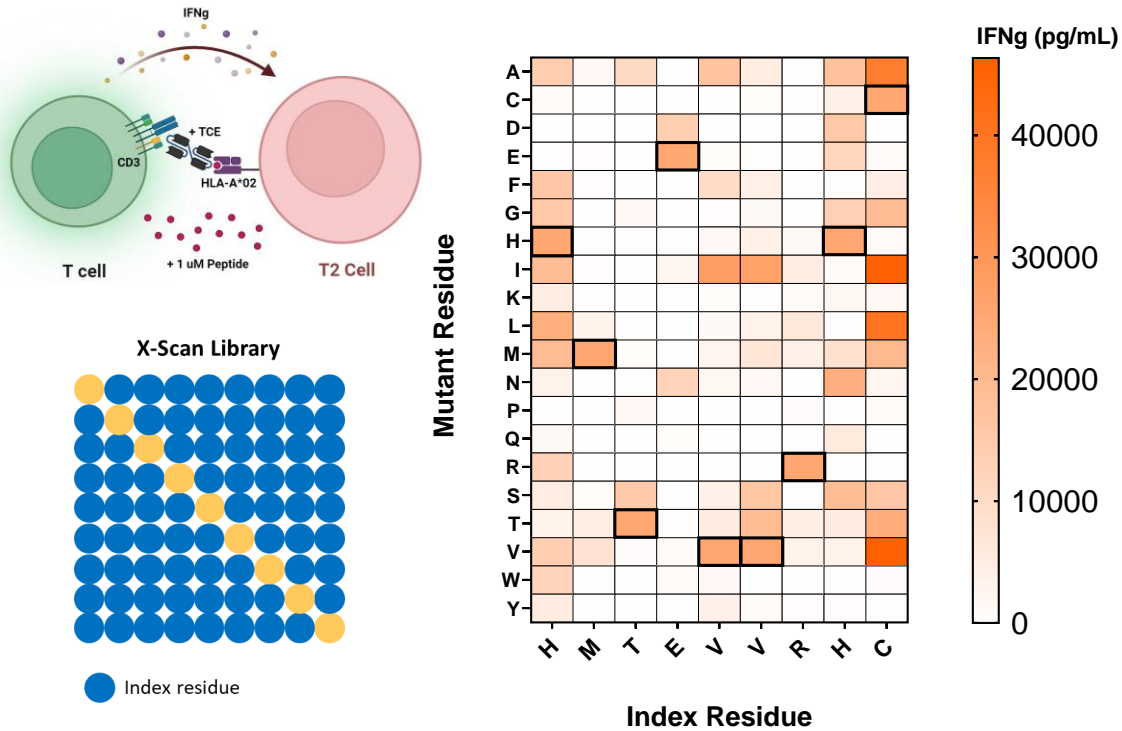


Potent and specific tumor cell killing

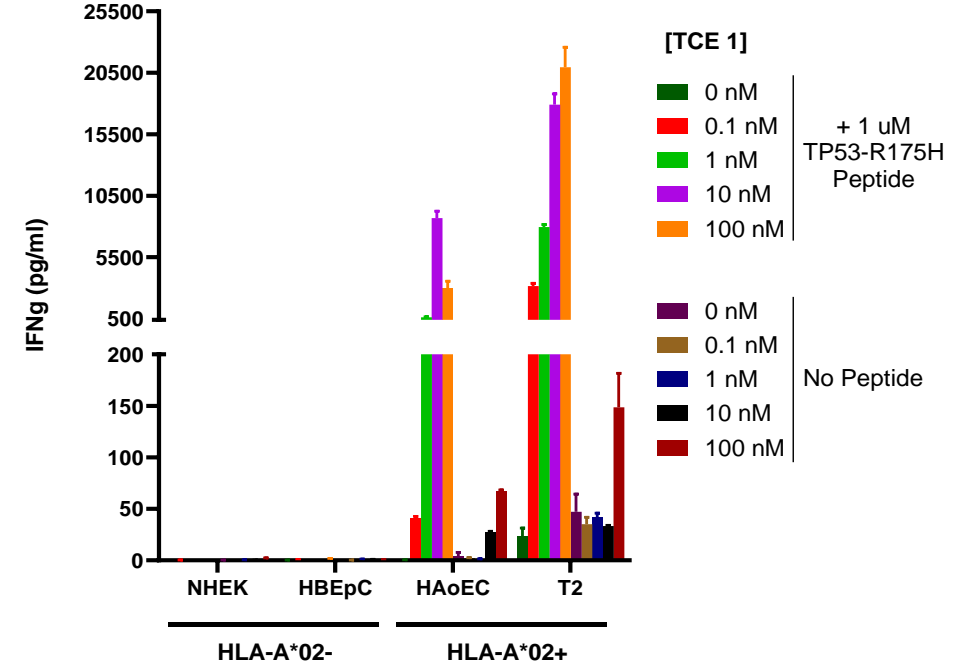


Preliminary Data for A2 TP53 R175H TCE Suggested Favorable Tolerability & Specificity Profiles

X-scan off-target binding profile



Did not mediate activation toward select normal tissues



Normal Tissue	HLA-A Alleles
Normal Human Epidermal Keratinocytes (NHEK)	A*11:01
Human Bronchial Epithelial Cells (HBEpC)	A*11:01
Human Aortic Endothelial Cells (HAoEC)	A*02:01/A*11:01
T2 Cells (Positive Control)	A*02:01

Experienced Management Team Supported by Blue-Chip Investor Syndicate

Executive Leadership



Jak Knowles, MD
Co-Founder and CEO



Kathy Bergsteinsson, MBA
Chief Financial Officer

Morgan Stanley



Dirk Nagorsen, MD
Chief Medical Officer



Kim Nguyen, PhD
Chief Technical Officer



Loïc Vincent, PhD
Chief Scientific Officer



Kathy Yi, MBA
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AnaptysBio



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Independent



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Scientific Advisors



Jim Allison, PhD



Pam Sharma, MD



Strategic Partners



Upcoming Catalysts

AFNT-211

A11 KRAS G12V

- Phase 1a data generation ongoing in 2L+ solid tumor indications
- Completion of dose escalation anticipated 2H25

AFNT-212

A11 KRAS G12D

- IND enabling studies complete
- IND clearance anticipated 2025
- THRIVE™ non-viral gene-edited FiH

AFNT-313

A2 P53 R175H

- IND enabling studies underway
- Pre-IND feedback anticipated 2025
- 2026 IND



Precision Immunotherapy
targeting oncogenic driver
mutations to develop potentially
curative therapies for patients
with solid tumors

