



affini



Unlocking the power of T cells
against oncogenic driver mutations

40th Annual J.P. Morgan Healthcare Conference
January 2022

Forward-Looking Statements

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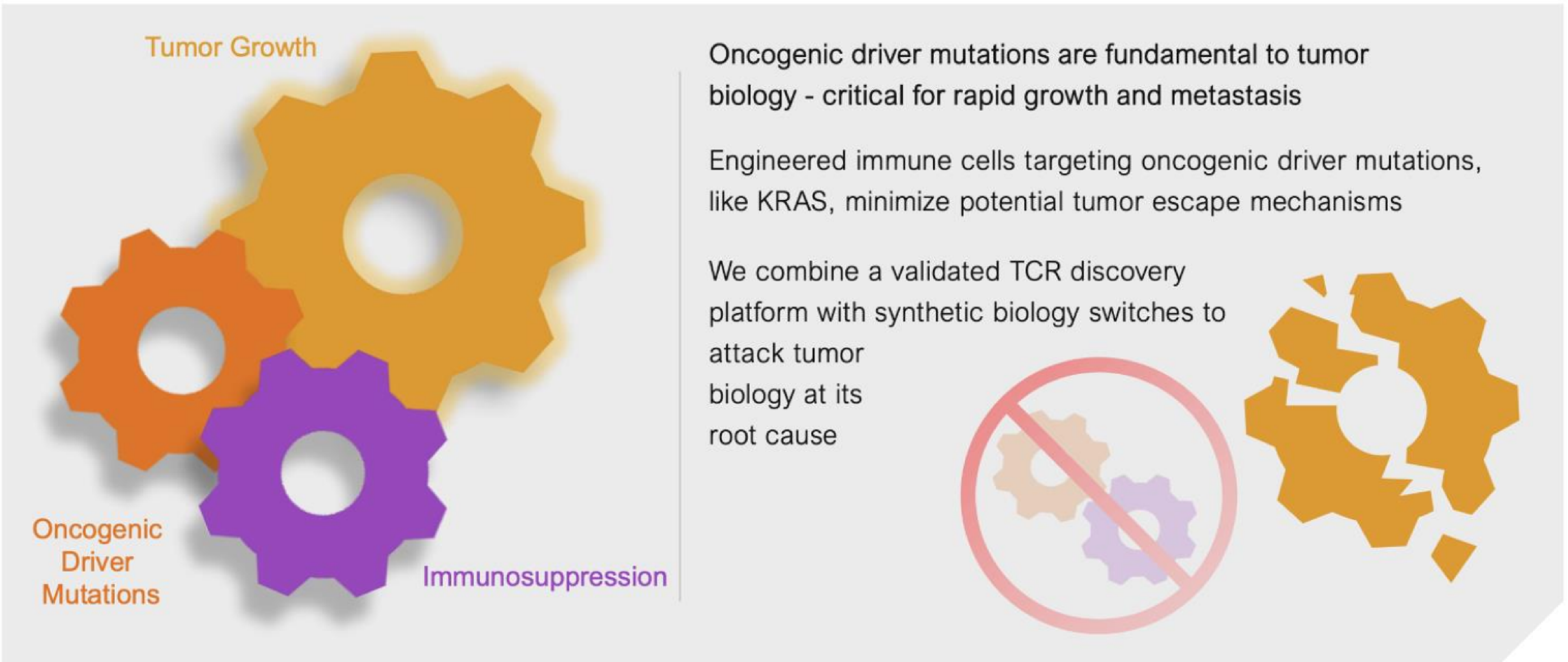
This presentation also includes statistical and other industry and market data that we obtained from industry publications and research, surveys and studies conducted by third parties as well as our own estimates of potential market opportunities. All of the market data used in this presentation involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data. Industry publications and third party research, surveys and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information. Our estimates of the potential market opportunities for our product candidates include several key assumptions based on our industry knowledge, industry publications, third-party research and other surveys, which may be based on a small sample size and may fail to accurately reflect market opportunities. While we believe that our internal assumptions are reasonable, no independent source has verified such assumptions.



Right Targets. Right Cells. Right Place.

We orchestrate the immune system to target oncogenic driver mutations to deliver transformative therapies intended to cure patients

Tumor growth powered by oncogenic driver mutations and immune cell dysfunction



Engineering T cells with the potential to cure hard-to-treat solid tumors

Right Targets

- Solid tumors are dependent on oncogene driver mutations like KRAS
- Driver mutations are present in every cancer cell, not in healthy tissues, and cannot be easily lost

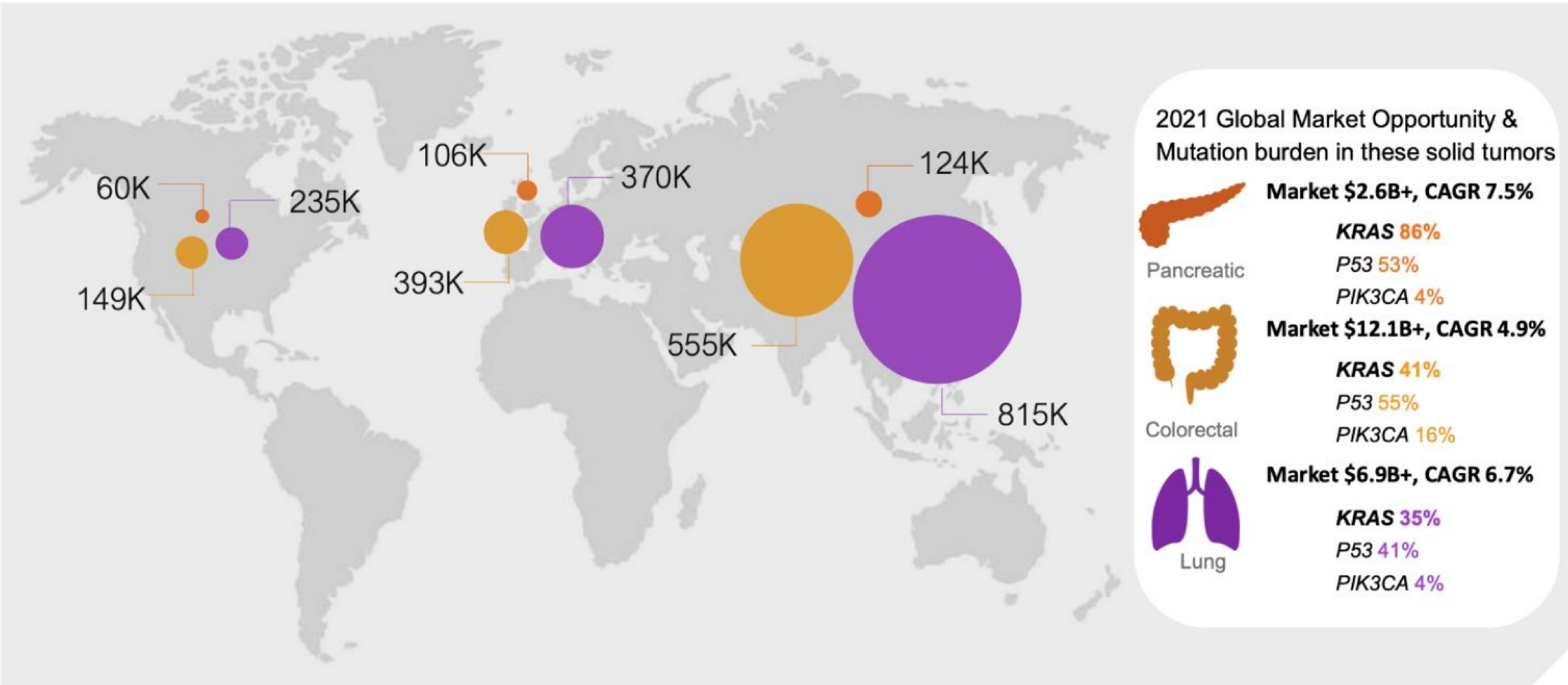
Right Cells

- Coordinate a CD4⁺ and CD8⁺ T cell response to sustain anti-tumor activity/response
- Select immune cells for stemness and central memory phenotype
- Manufacture to maintain cell naïveté and reduce terminal differentiation

Right Place

- Direct T cells to traffic into the solid tumor bed using proprietary TCRs
- Prevent deletion of T cells by converting Fas/FasL death signal to 41BB survival signal
- Convert immunosuppressive tumor microenvironment (TME) signals into stimulatory cues using additional synthetic biology switches

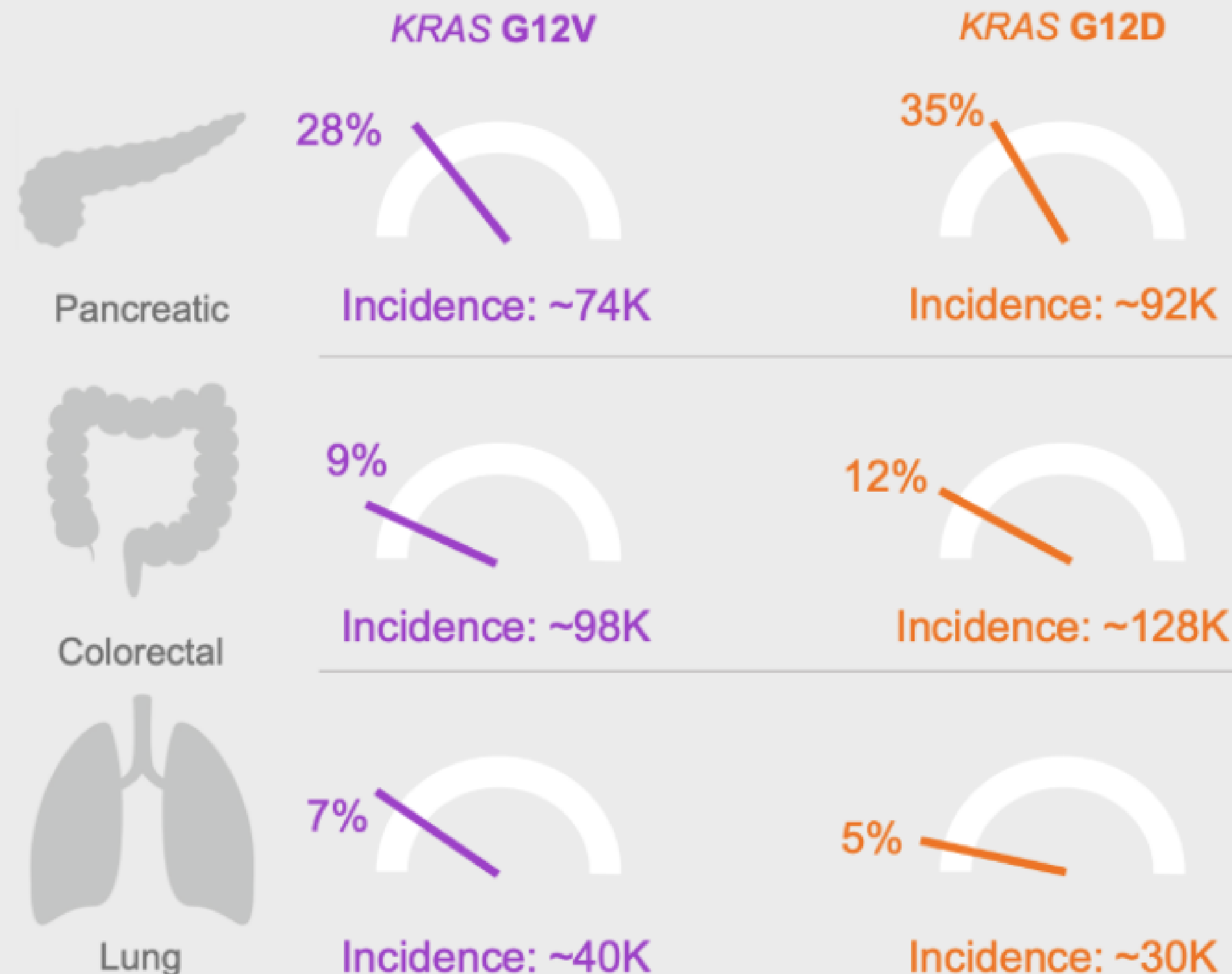
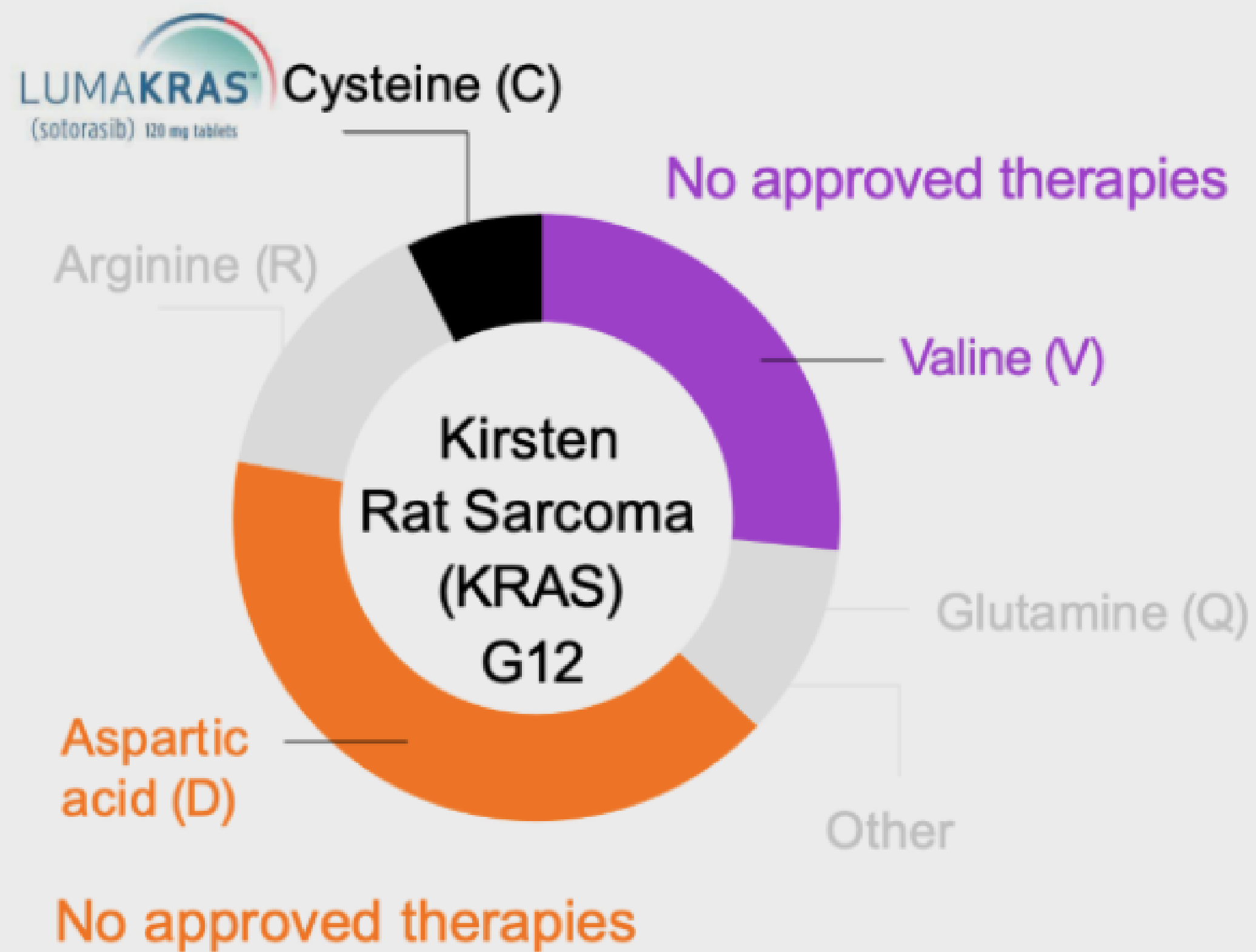
Solid tumors dependent on oncogenic drivers represent a large global market



Attacking KRAS: the most prevalent oncogenic driver mutation in solid tumors

Relative frequency of G12 mutations

Incidence of G12V and G12D in certain adenocarcinomas



Exceptional Science Leaders and Experienced Team



Fred Hutch co-founders among the most prolific in the industry for generating T cell clinical programs



Phil Greenberg, MD



Aude Chapuis, MD



Tom Schmitt, PhD

nature medicine

T cell receptor gene therapy targeting WT1 prevents acute myeloid leukemia relapse post-transplant

Aude G. Chapuis, Daniel N. Egan, Merav Bar, Thomas M. Schmitt, Megan S. McAfee, Kelly G. Paulson, Valentin Voillet, Raphael Gottardo, Gunnar B. Ragnarsson, Marie Bleakley, Cecilia C. Yeung, Petri Muhlhauser, Hieu N. Nguyen, Lara A. Kropp, Luca Castelli, Felecia Wagener, Daniel Hunter, Marcus Lindberg, Kristen Cohen, Aaron Seese, M. Juliana McElrath, Natalie Duerkopp, Ted A. Gooley & Philip D. Greenberg 

Cancer Cell

T Cells Engineered against a Native Antigen Can Surmount Immunologic and Physical Barriers to Treat Pancreatic Ductal Adenocarcinoma

Ingunn M. Stromnes,^{1,3} Thomas M. Schmitt,¹ Ayaka Hulbert,¹ J. Scott Brockenbrough,¹ Hieu N. Nguyen,¹ Carlos Cuevas,⁴ Ashley M. Dotson,¹ Xiaoxia Tan,³ Jennifer L. Hotes,¹ Philip D. Greenberg,^{1,3,5,6,*} and Sunil R. Hingorani^{1,2,5,6,*}

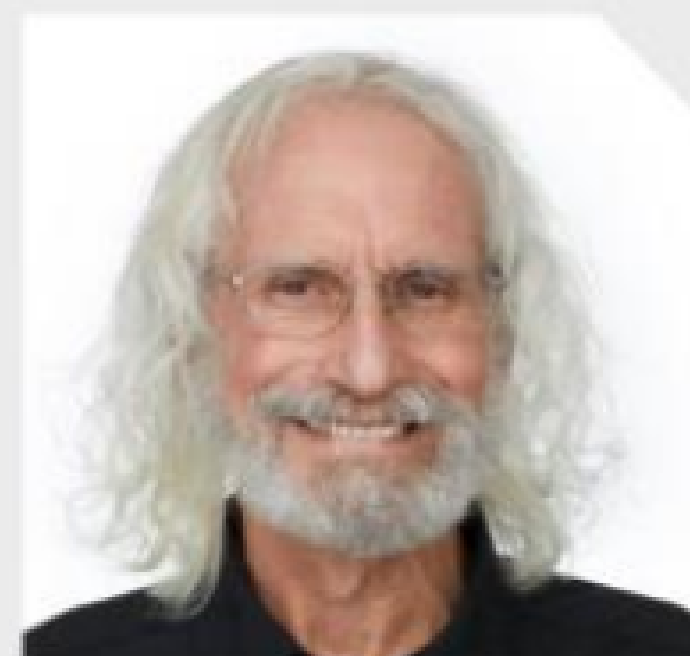
Advanced over a dozen T cell therapeutic candidates



Undisclosed
industry
partners

Guided by a world class SAB specialized in T cell biology and immunology

Scientific Advisory Board



Phil Greenberg, MD
Scientific Co-Founder



Aude Chapuis, MD
Scientific Co-Founder



Tom Schmitt, PhD
Scientific Co-Founder



Jim Allison, PhD



Pam Sharma, MD, PhD



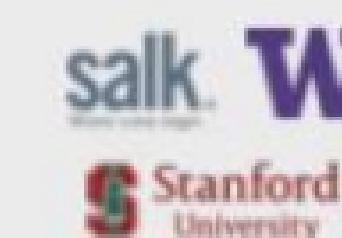
Rafi Ahmed, PhD



David Kranz, PhD



Susan Kaech, PhD



Experienced management poised to advance multiple T cell therapies

C-Suite



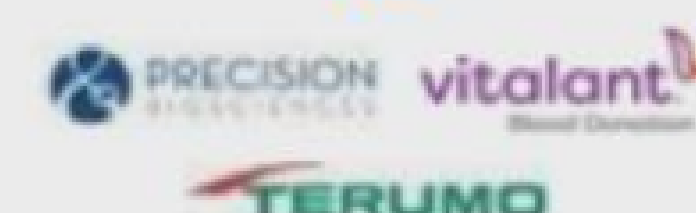
Jak Knowles, MD
CEO



Loïc Vincent, PhD
CSO



Kim Nguyen, PhD
CTO



Kathy Yi, MBA
COO/CFO



Seed Investors



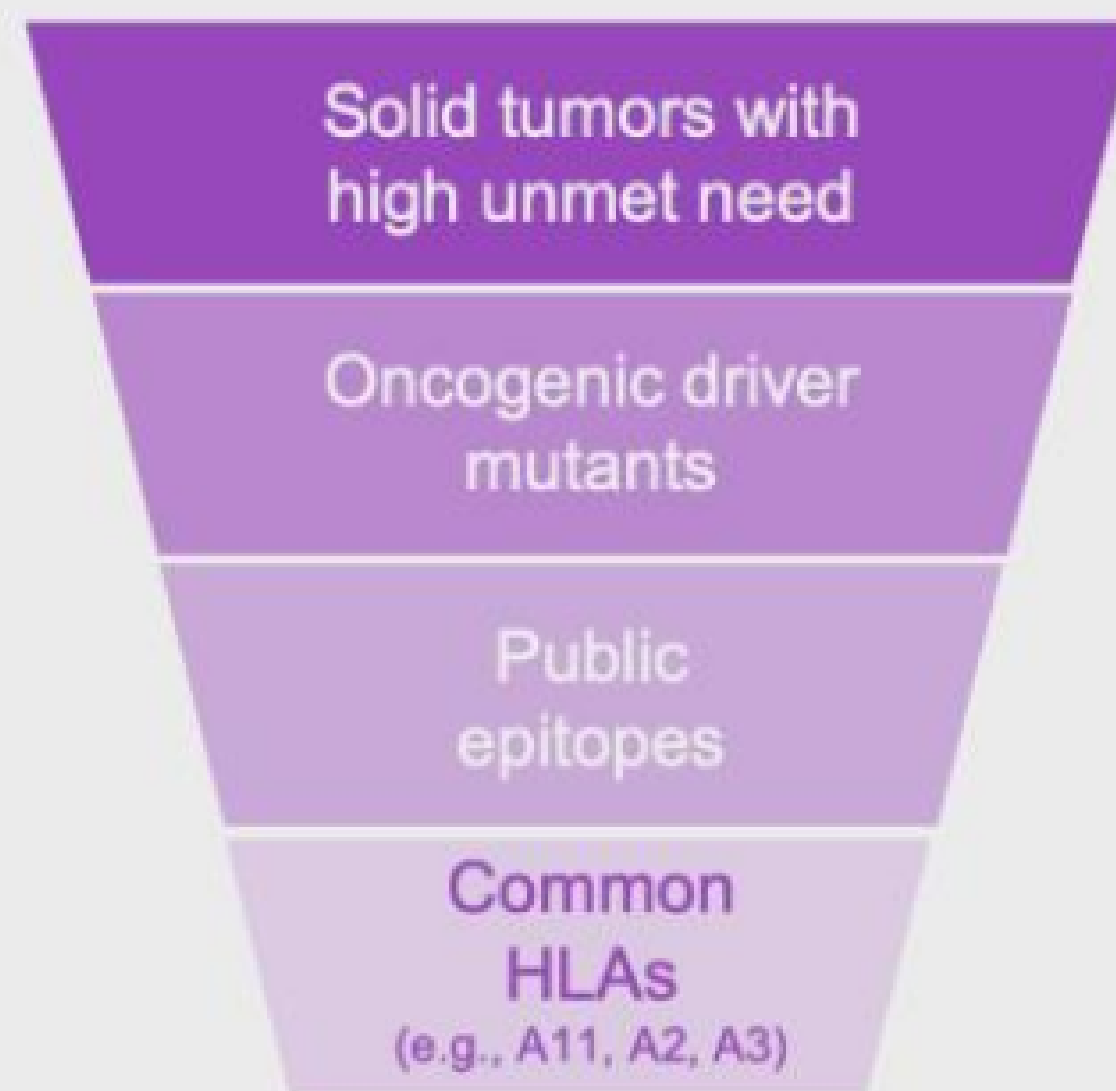
Solving Hard-to-Treat Solid Tumors



Affini-T's discovery platform has identified multiple TCR clinical candidates

Prioritize

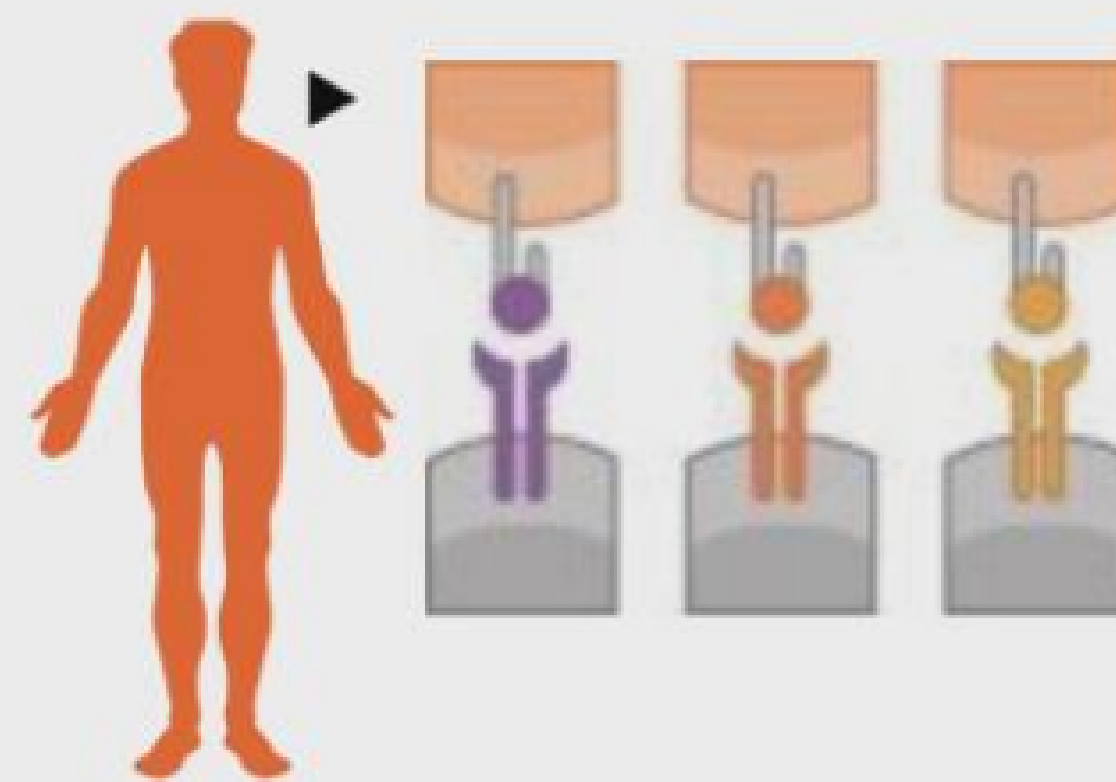
Streamline top targets to prosecute



Candidate targets

Discover

Isolate high affinity TCRs from panel of healthy donors



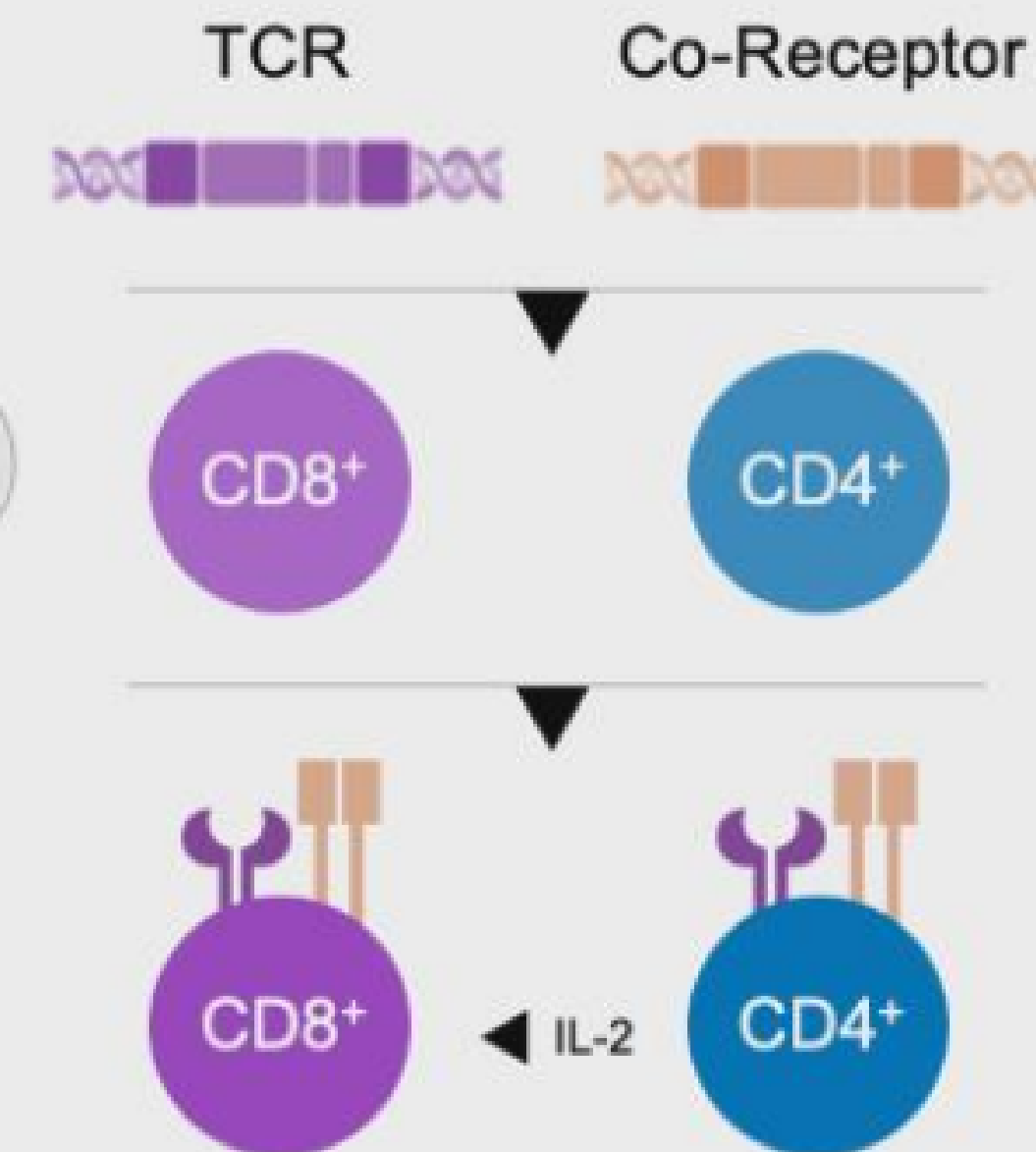
TCRs with de-risked safety profile that have cleared thymic selection



Robust and proven TCR discovery platform

Engineer

Lentiviral engineering of patient T cells



Leverage coordinated CD4/CD8 T cell response for improved efficacy and durability

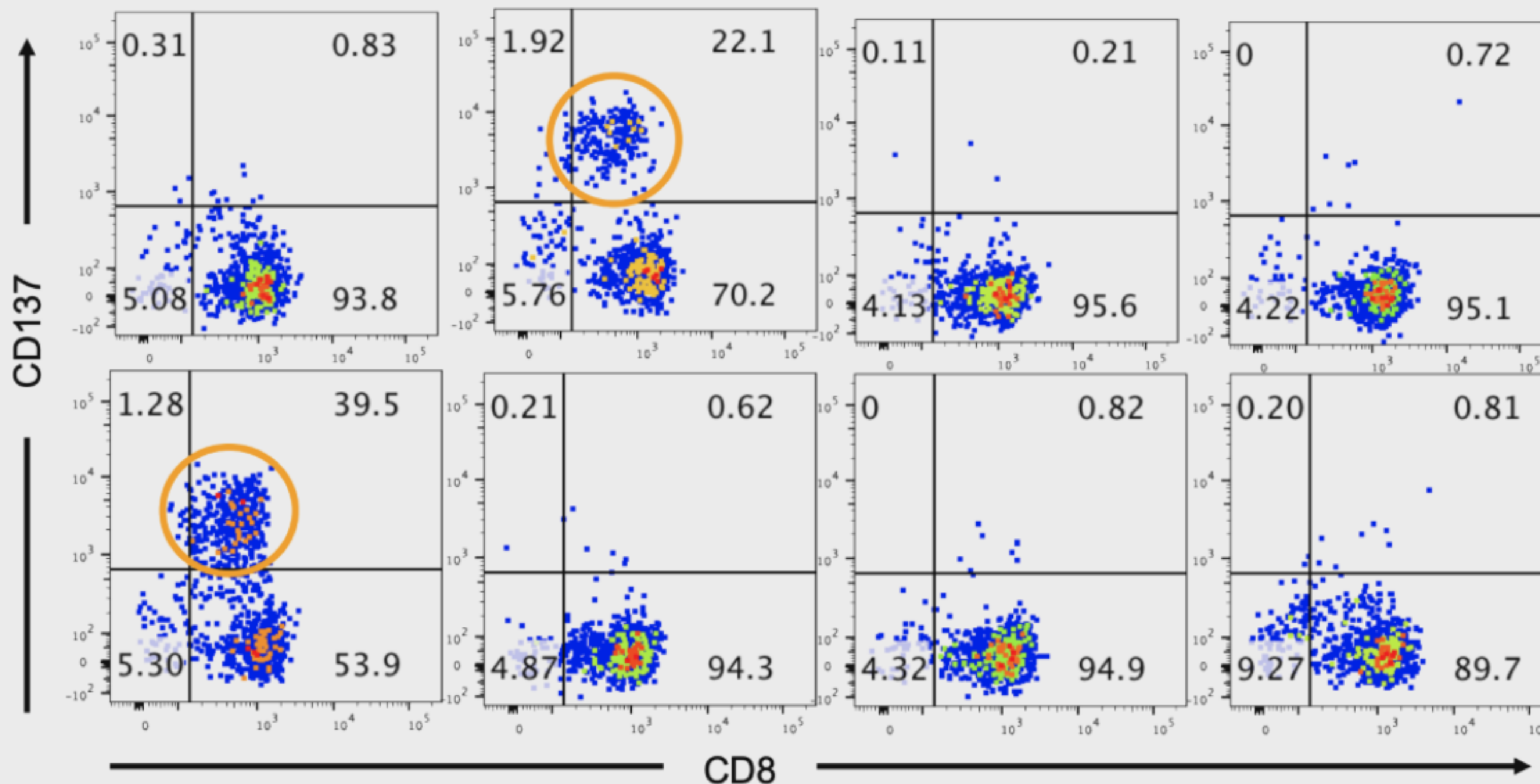
Treat

Infuse engineered T cell therapeutic

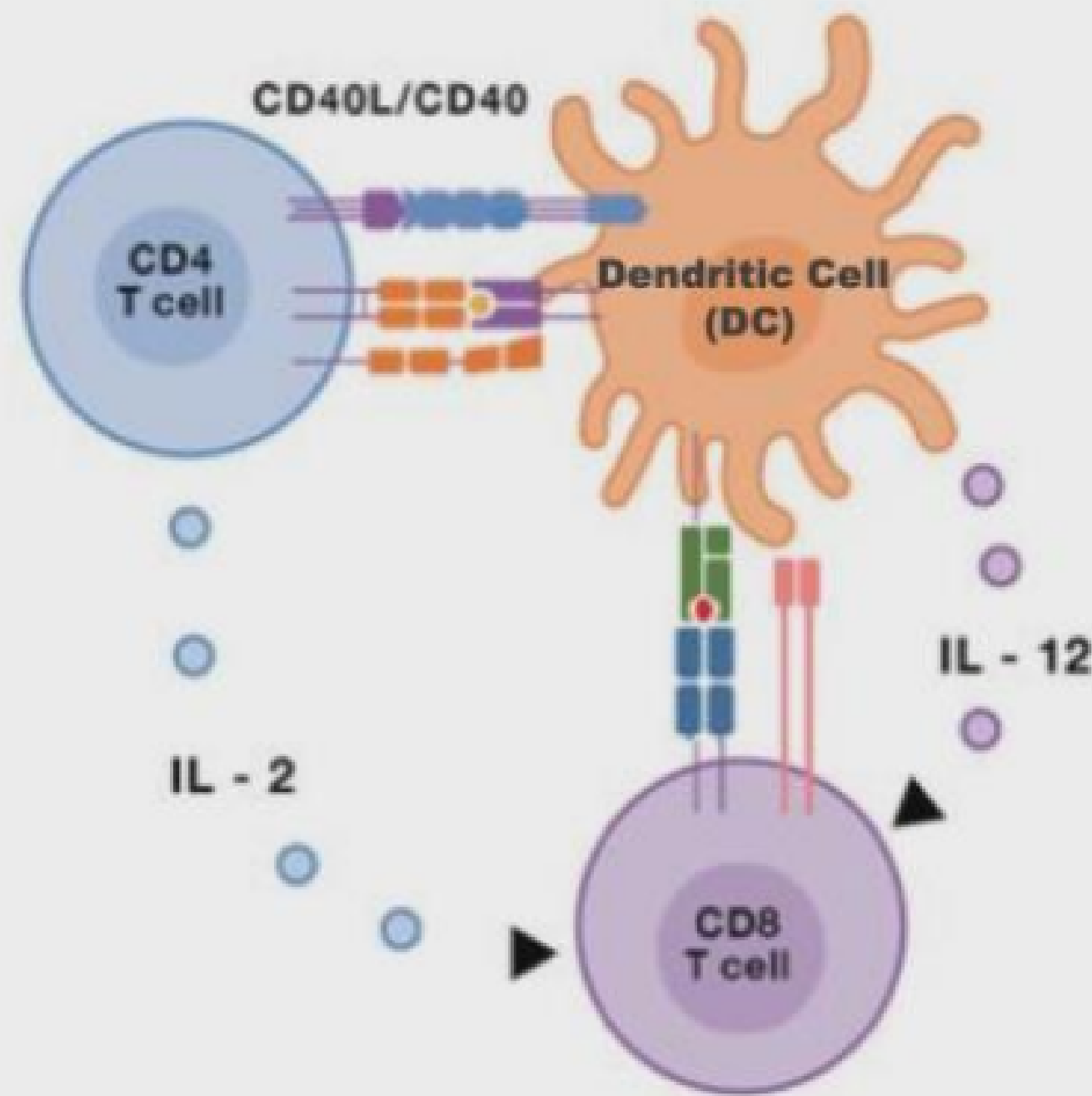


KRAS G12V and G12D TCRs are highly selective and do not react to Wild Type peptide

A11/KRAS:
(G12D-specific)

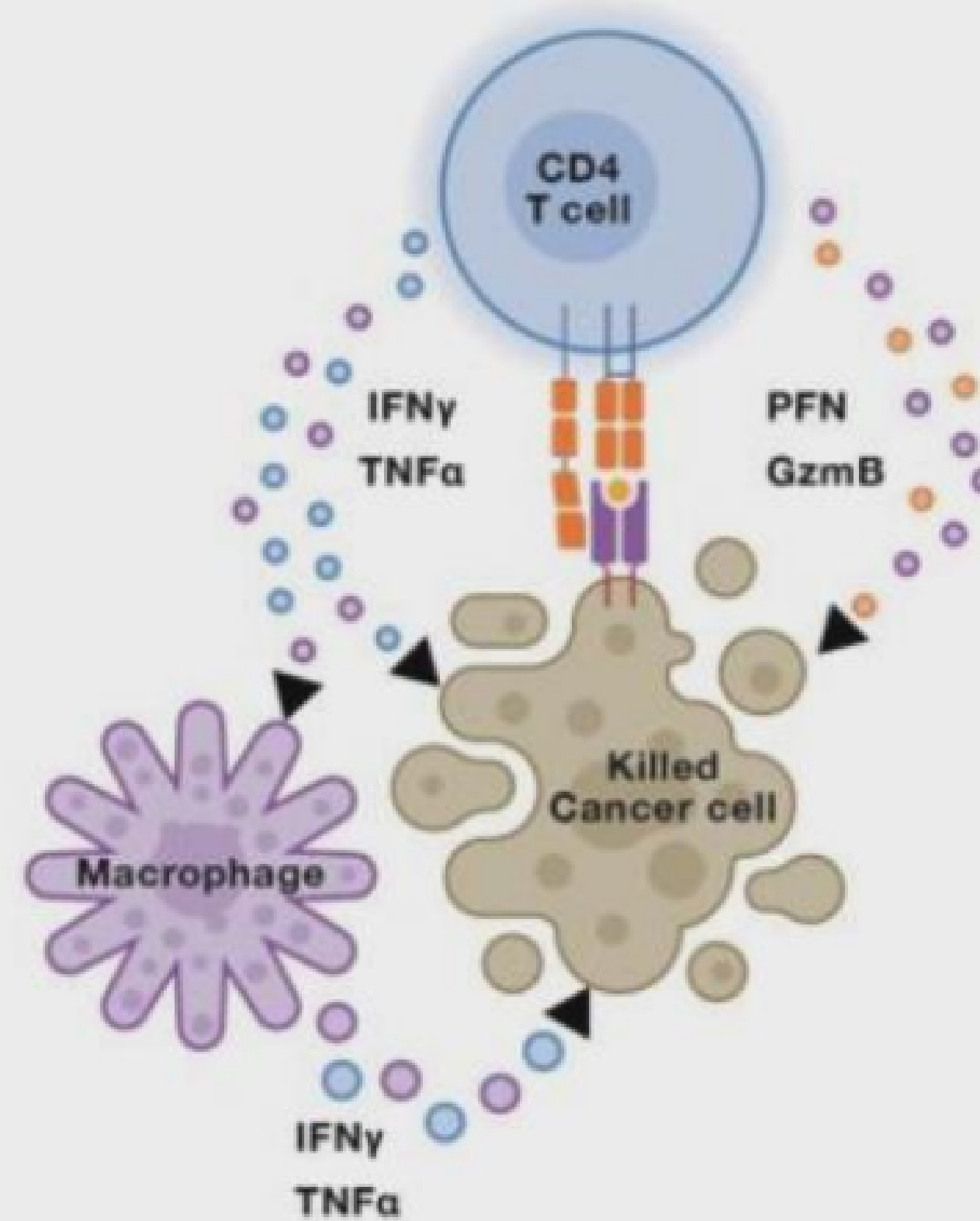


Fundamental immunology: CD4 T cells can orchestrate durable anti-tumor responses



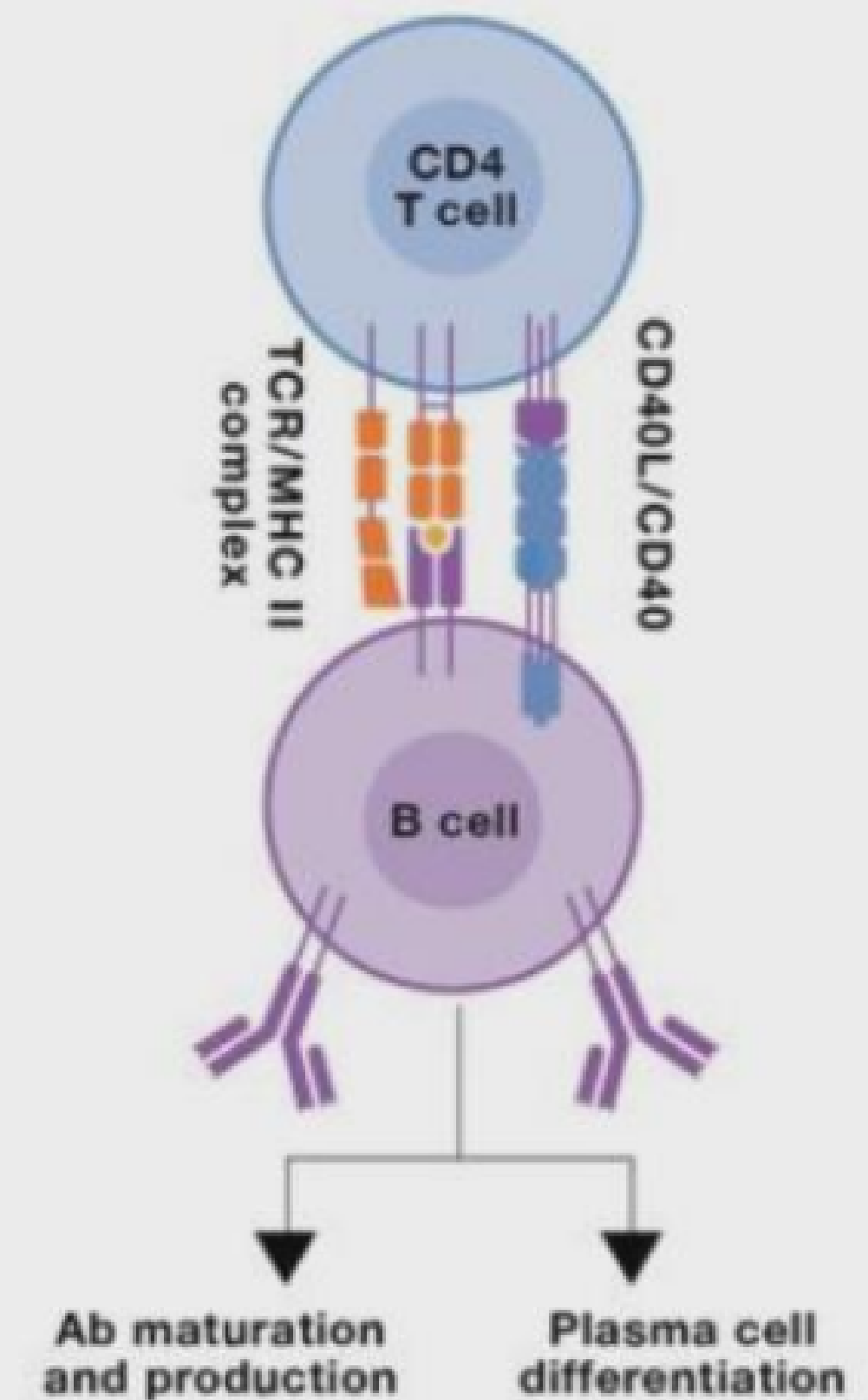
CD4 Helper

Promotes CD8 T cell function, proliferation and survival



CD4 Effector

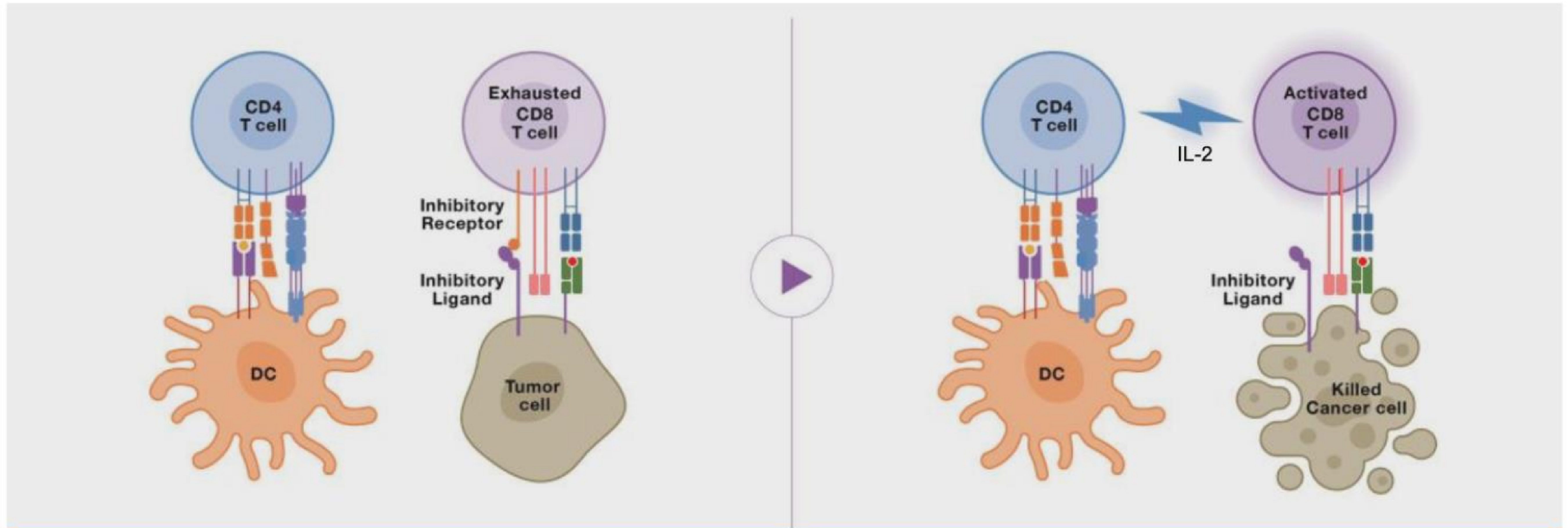
Direct and indirect killing of tumor cells



CD4 Activator

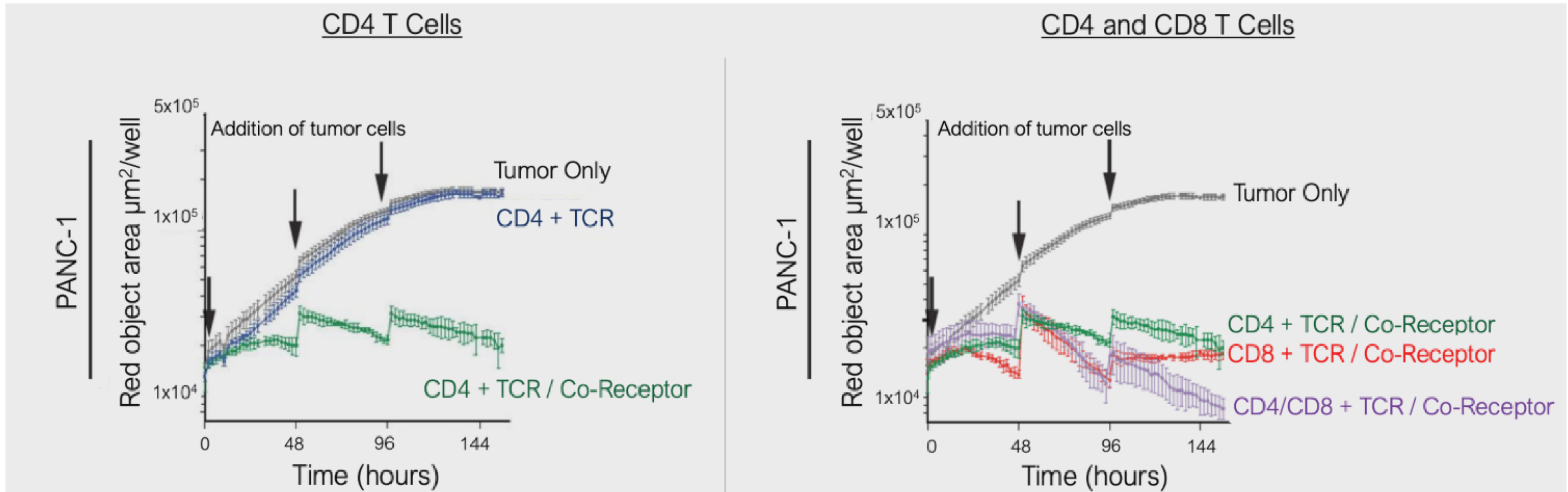
Drives B cell maturation and antibody production

CD4 T cells prevent CD8 T cell exhaustion and prolong persistence in the hostile TME



CD4 T cells retain helper phenotype, engage CD8 T cells in the TME to promote effector activation and prevent CD8 T cell exhaustion leading to enhanced CD8 persistence

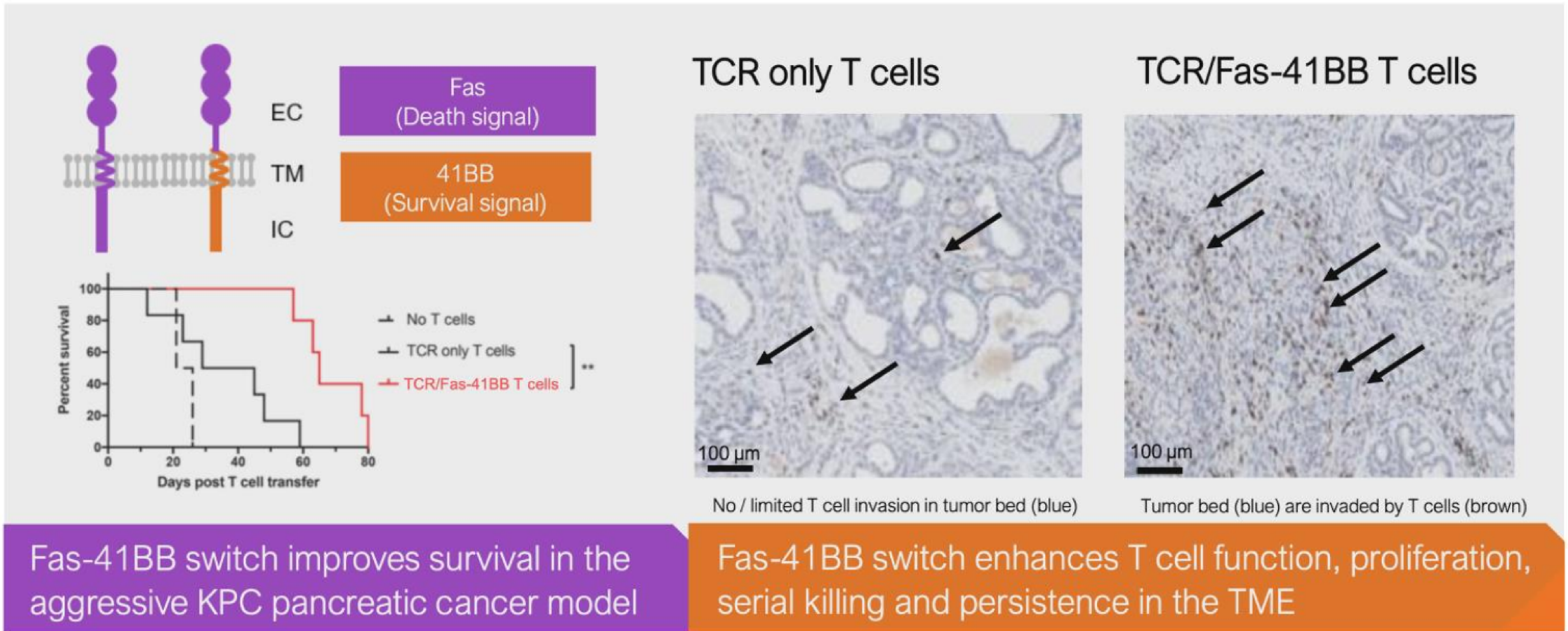
CD4/CD8 coordinated T cell response induces deep and durable anti-tumor activity



Addition of co-receptor to CD4 T cells creates additional effector function for tumor killing

CD4 T cells synergize with CD8 T cells to enhance anti-tumor activity

Enhancing T cell survival and function in the TME to create durable responses



Differentiated Pipeline of TCR T Cell Therapies



Deep and differentiated pipeline of TCR T cell therapies

Target	Approach	Discovery	Preclinical	Clinical
Oncogenic Driver Programs				
KRAS G12V	Lentiviral	HLA-A11		
	Lentiviral + Switch	HLA-A2		
	Lentiviral + Switch	HLA-A3		
KRAS G12D	Lentiviral + Switch	HLA-A11		
Mutant P53	Lentiviral + Switch	HLA-A2		
2 nd Gen (Multiple)	Gene edits			
3 rd Gen (Multiple)	Synthetic bio enhancements			
4 th Gen (Multiple)	Allo, <i>In situ</i>			
Viral Driver Programs				
MCPyV	Lentiviral	HLA-A2		
Undisclosed	Lentiviral + Switch			

Affini-T technologies positioned to advance a robust pipeline of immune innovation

First-Generation

- Coordinated CD4/CD8 response
- Fas-41BB durability receptor
- Switch receptors (e.g., inhibitory receptors, cytokine receptors)
- Boost MHC expression
- Disrupt endogenous TCR

Next-Generation

- CD28 costimulatory chimeric receptors (e.g., PD-1)
- Cytokines to increase immune response
- Enzymes to break down fibrotic tissue
- Epigenetic modification
- Immune adjuvants
- Chemokine receptors to improve trafficking

Building infrastructure to support rapid bi-coastal company growth



Seattle Research Lab

- TCR Discovery
- Manufacturing Support
- Early-Stage Process and Analytical Development
- Alliance Management



Boston Interim Lab

- Gene Editing
- Next-Gen Engineering
- Synthetic Biology
- CMC



Boston HQ

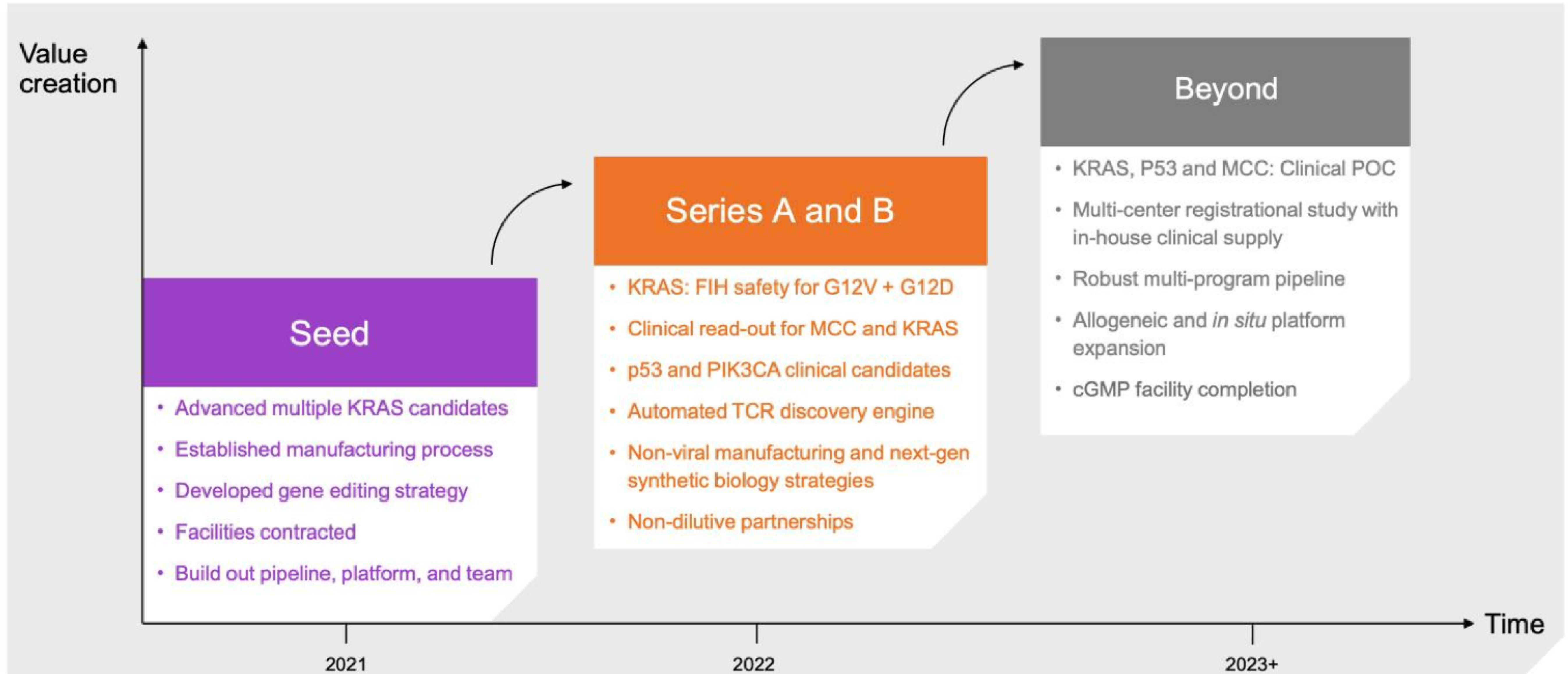
- Executive Management
- Discovery Research
- Regulatory and Quality
- Clinical Operations
- CMC
- Operations and Finance
- Business Development
- HR and Legal



Boston cGMP Clinical and Commercial Manufacturing

- 95,000 sq.ft.
- Clinical and Commercial Production
- Quality Assurance and Quality Control
- Facilities and IT
- Warehousing

Multi-generational pipeline will create meaningful value







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