

JP Morgan Healthcare Conference 2023



Disclaimer

This presentation contains "forward-looking statements," as that term is defined under the Private Securities Litigation Reform Act of 1995 (PSLRA), which statements may be identified by words such as "believe," "may," "will," "estimate," "continue," "anticipate," "intend," "expect" and other words of similar meaning. These forward-looking statements involve certain risks and uncertainties. Such risks and uncertainties could cause our actual results to differ materially from those indicated by such forward-looking statements, and include, without limitation: the success, cost and timing of our product development activities and clinical trials; our ability to submit an IND and successfully advance our technology platform to improve the safety and effectiveness of our existing TCR therapeutic candidates; the rate and degree of market acceptance of T-cell therapy generally and of our TCR therapeutic candidates; government regulation and approval, including, but not limited to, the expected regulatory approval timelines for TCR therapeutic candidates; and our ability to protect our proprietary technology and enforce our intellectual property rights; amongst others. For a further description of the risks and uncertainties that could cause our actual results to differ materially from those expressed in these forward-looking statements, as well as risks relating to our business in general, we refer you to our Annual Report on Form 10-K filed with the Securities and Exchange Commission filed for the year ended December 31, 2021, our Quarterly Reports on Form 10-Q, Current Reports on Form 8-K and other filings with the Securities and Exchange Commission. The forward-looking statements contained in this presentation speak only as of the date the statements were made and we do not undertake any obligation to update such forward-looking statements to reflect subsequent events or circumstances.

We urge you to consider these factors carefully in evaluating the forward-looking statements herein and you are cautioned not to place undue reliance on such forward-looking statements, which are qualified in their entirety by this cautionary statement. The forward-looking statements contained in this presentation speak only as of the date the statements were made and we do not undertake any obligation to update such forward-looking statements to reflect subsequent events or circumstances.

We intend that all forward-looking statements be subject to the safe-harbor provisions of the PSLRA.



Our mission and vision are clear:

To transform the lives of people with cancer by designing, developing, and delivering cell therapies

Arming cells.
Against cancer.
For good.



Leading The Cancer Revolution



Adaptimmune - an integrated cell therapy company







Focused investment to deliver near- and long-term value

Clinical programs targeting MAGE-A4



AFAMI-CEL (synovial sarcoma)

First-gen MAGE-A4 product with potential to be first approved engineered T-cell therapy for a solid tumor.

- Pivotal trial (SPEARHEAD-1) has met primary endpoint with ORR of ~39%.
- BLA submission initiated with target completion by mid-2023; eligible for priority review by FDA with RMAT



ADP-A2M4CD8 (SURPASS trials)

Next-gen MAGE-A4 product designed to be more potent and more effectively engage broader immune system.

- Ongoing Phase 1 SURPASS trial
 - Combo. arm with checkpoint inhibitor initiated
 - Initiating new cohorts in urothelial and head & neck cancers in earlier lines and in combination with SOC
- Phase 2 SURPASS-3 trial for platinum resistant ovarian cancer; supported by RMAT status, to initiate early 2023



2023-2025

Preclinical development



PRAME

Unique opportunity in a broad range of tumors; potential synergy with MAGE-A4

- IND-ready by 2023
- Target indications to be based on prevalence and high antigen expression



ALLOGENEIC



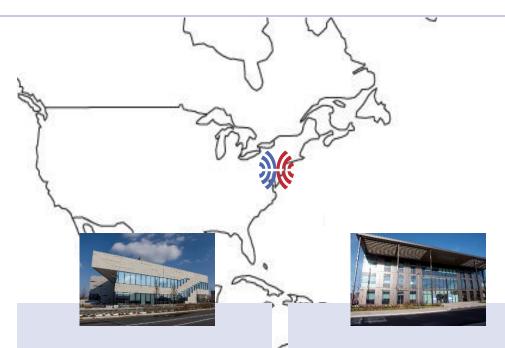
"Off-the-shelf" platform utilizing stem cell lines (iPSCs) from single donor cells to generate functional

- Progress wholly owned & partnered programs
- Astellas: HiT targeting mesothelin (target 1) and target 2 nominated
- · Genentech: off-the shelf TCR therapies and personalized off-the-shelf therapies





Best in class capabilities to design and deliver cell therapies



Philadelphia, PA, USA

- Autologous manufacturing
- Clinical development
- Commercial
- Corporate

Milton Park, UK

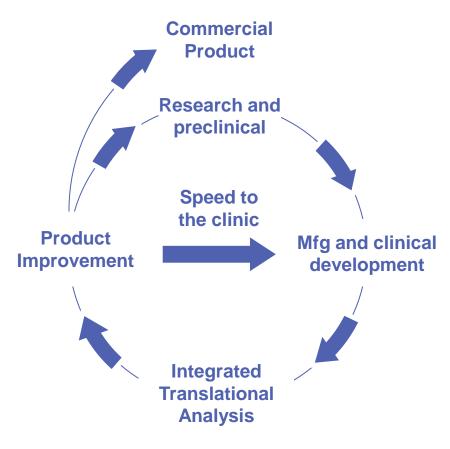
- Dedicated allogeneic manufacturing facility
- Pipeline research
- Allogeneic research
- Process and analytical development
- Corporate



Stevenage, UK

 GMP lentiviral vector manufacturing

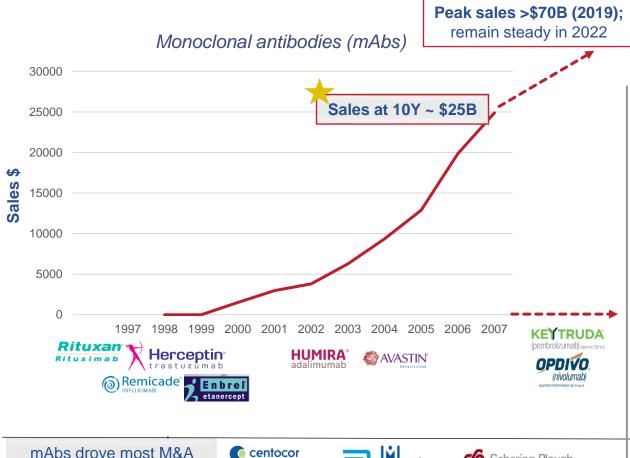
Product innovation cycle

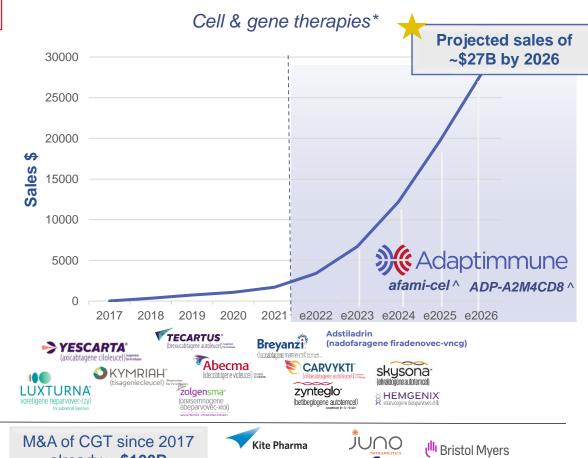


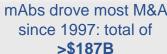


Cell and gene therapies set to transform the biopharma landscape

Adaptimmune poised to be an early leader



















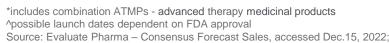






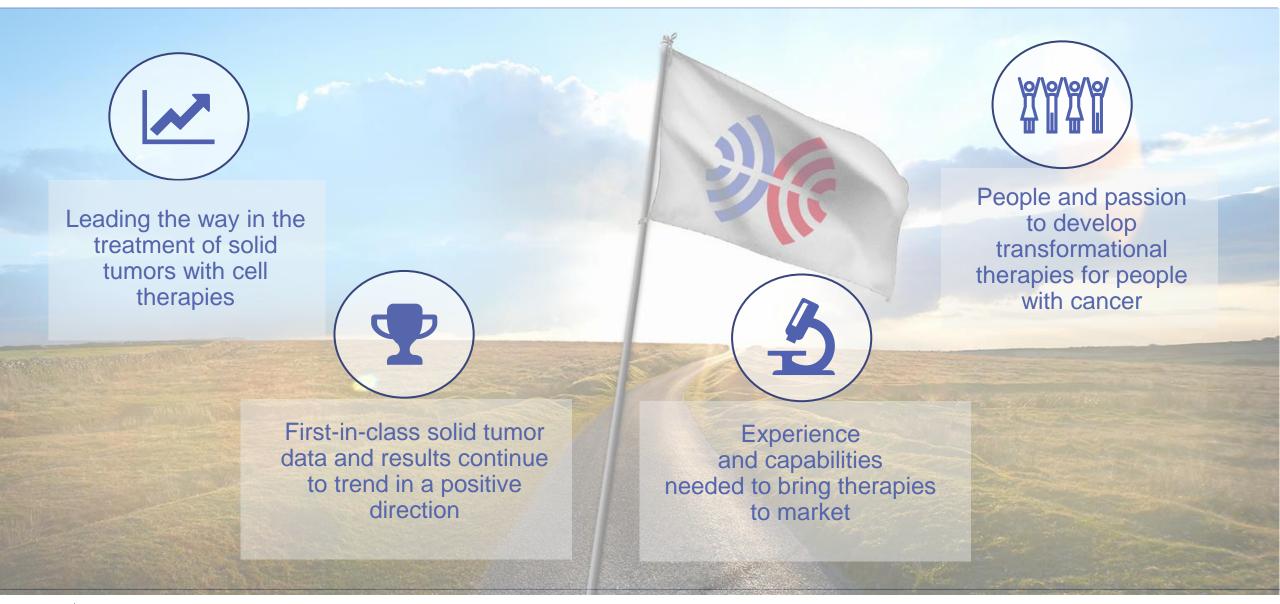








Positioned to lead and succeed



Advanced autologous engineered TCR program targeting MAGE-A4

Validated target with annual mortality of >82,000¹ patients (US and EU) with MAGE-A4+ tumors

- Clinically validated "clean" target; member of cancer testis antigen family
- **Expression across broad range of** solid tumors confirmed by screening protocol
- In early- and late-phase clinical trials with acceptable safety profile, to date, and responses in multiple solid tumor indications

- Expression levels ranging from ~15% to ~70%² across tumors
- **Encouraging responses in:**
 - Synovial sarcoma
 - Ovarian
 - Head & neck
 - Bladder
 - Gastroesophageal

- NSCLCsquamous
- Melanoma
- **MRCLS**

MAGE-A4 target for both first-gen afami-cel and next-gen (ADP-A2M4CD8) programs



Responses in multiple solid tumor indications expressing MAGE-A4

Responses reported with	Indication	Mortality US, UK & EU4*	MAGE-A4 Expression**	Potential MAGE-A4 +ve Patients	Potential MAGE-A4 +ve Patients Factored for HLA***	
afami-cel ADP-A2M4CD8	Synovial sarcoma	1,804+	% 67	1,209	496	
afami-cel	MRCLS	2,000+	% 34	680	279	
ADP-A2M4CD8	Gastroesophageal (esophageal, EGJ, and gastric)	83,384	% 20	16,677	7,388	
afami-cel ADP-A2M4CD8	Head and neck	41,409	% 22	9,110	4,036	
ADP-A2M4CD8	Urothelial	52,568	% 32	16,822	7,452	
afami-cel	NSCLC - squamous	76,875	% 35	26,906	11,919	
afami-cel	Melanoma	19,037	% 16	3,046	1,349	
ADP-A2M4CD8	Ovarian	31,558	% 24	7,574	3,355	
				TOTAL MAGE-A4: 82.024	TOTAL MAGE-A4 HLA A2: 36.274	

Significant potential for SPEAR T-cell franchise targeting MAGE-A4



Preclinical autologous engineered TCR program targeting PRAME

Validated target with annual mortality of >160,000¹ patients (US and EU) with PRAME+ tumors

- Clinically validated "clean" target; member of cancer testis antigen family
- Unique opportunity in a broader range of tumors than other targets
- First-gen in preclinical development to be IND-ready in 2023
- Considering next-gen approaches and potential synergy with MAGE-A4

- Highly expressed across a broad range of solid tumors including:
 - Breast
 - NSCLC
 - Kidney
 - Gastroesopriagear

- Melanoma
- Endometrial
- Ovarian
- Gastroesophageal Head & neck

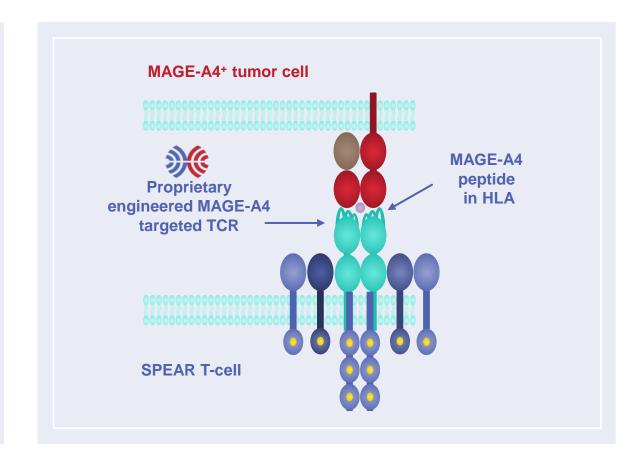




Afami-cel

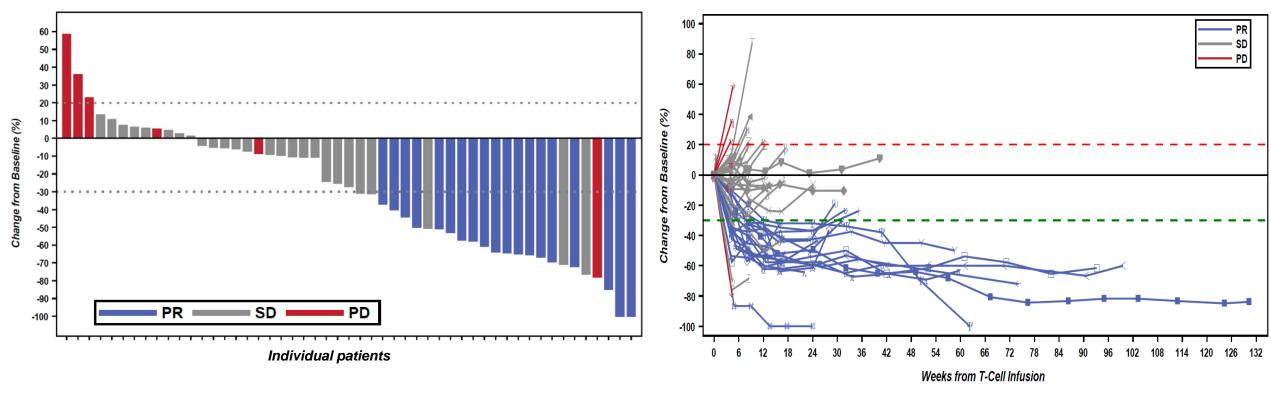
Potential to be the first commercially available engineered T-cell therapy for a solid tumor

- 1st gen TCR targeting MAGE-A4
 - Rolling BLA submission initiated for treatment of synovial sarcoma, target completion mid-2023
 - RMAT and PRIME designations
- Single dose of cells resulting in 38.6% ORR in synovial sarcoma; median duration of response 50.2 weeks
- ~300 eligible patients per year in U.S. with synovial sarcoma expressing MAGE-A4 and HLA-A2*





Response rate of 38.6% in synovial sarcoma

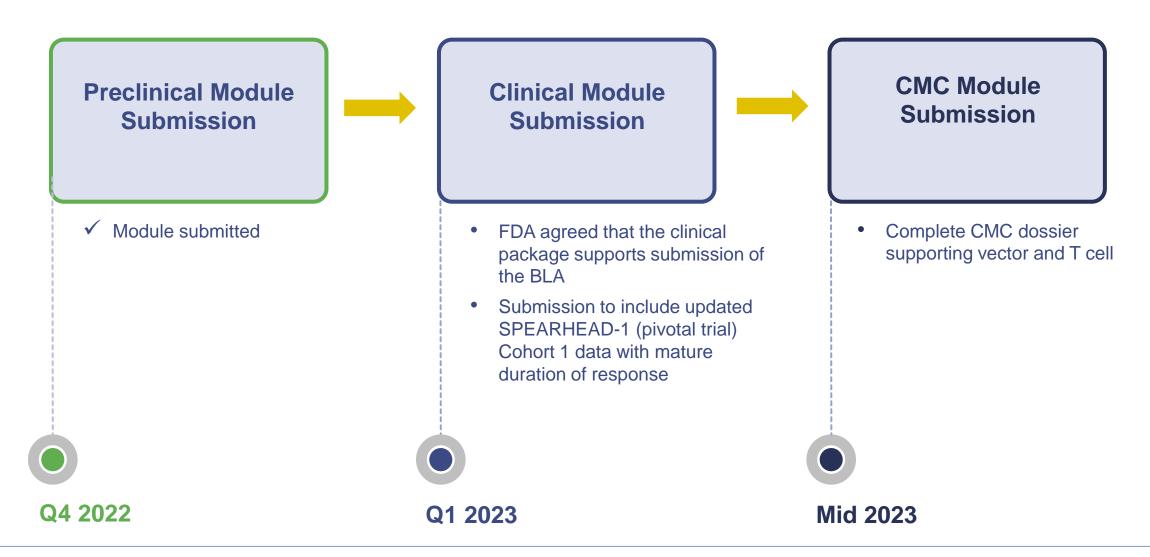




- · Afami-cel is efficacious in heavily pre-treated patients with synovial sarcoma
- Median duration of response in synovial sarcoma: 50.3 weeks (range: 11.7–122.0+)
- 8 responses ongoing as of data cut-off



Rolling BLA submission to be completed by mid-2023





Proud to receive SFA's Vision of Hope award





"Adaptimmune's contribution to improving sarcoma patient outcomes is extraordinary and is well-deserved for recognition. The company's work is taking us many steps closer to finding a cure for sarcoma."

Sarcoma Foundation of America (SFA) CEO, Brandi Felser



Bringing the potential of afami-cel to patients with synovial sarcoma

Afami-cel is a highly anticipated new treatment option for synovial sarcoma

- Currently very limited options and poor prognosis for advanced patients after 1L: <20% 5-yr survival, <15% ORR
- Adaptimmune has the opportunity to transform treatment, delivering unprecedented ~39% ORR and ~>50 weeks DOR with single dose, in heavily pre-treated patients with advanced disease

Concentrated care, few specialists, and strong experience with afami-cel through engagement in our trials

- About 80 adult Sarcoma Centers of Excellence and 18 afami-cel clinical trial sites in the US
- More than 5 years of clinical experience with afami-cel and fully owned manufacturing, supply, and customer service





Opportunity to establish focused commercialization capabilities

Ultra-targeted approach and highly specialized team, sized and customized to cell therapy in rare disease

Focused

15-20 experienced treatment sites and integrated referral networks across CoEs

Access

Strong value proposition in rare, biomarker targeted population with low budget impact to payers

Reliable

Robust processes, customized to cell therapy, delivering excellent customer experience

Simple

Straightforward testing solution and patient journey from identification to infusion

Caring Boldly

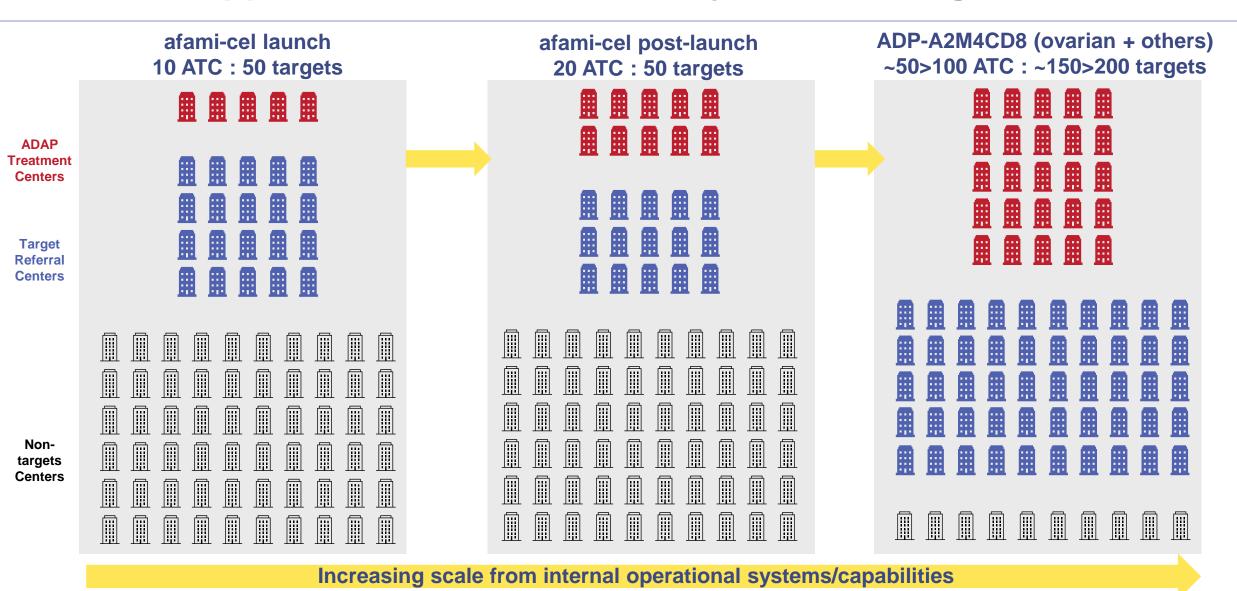
Nimble, specialized, integrated team, passionate to find solutions for patients



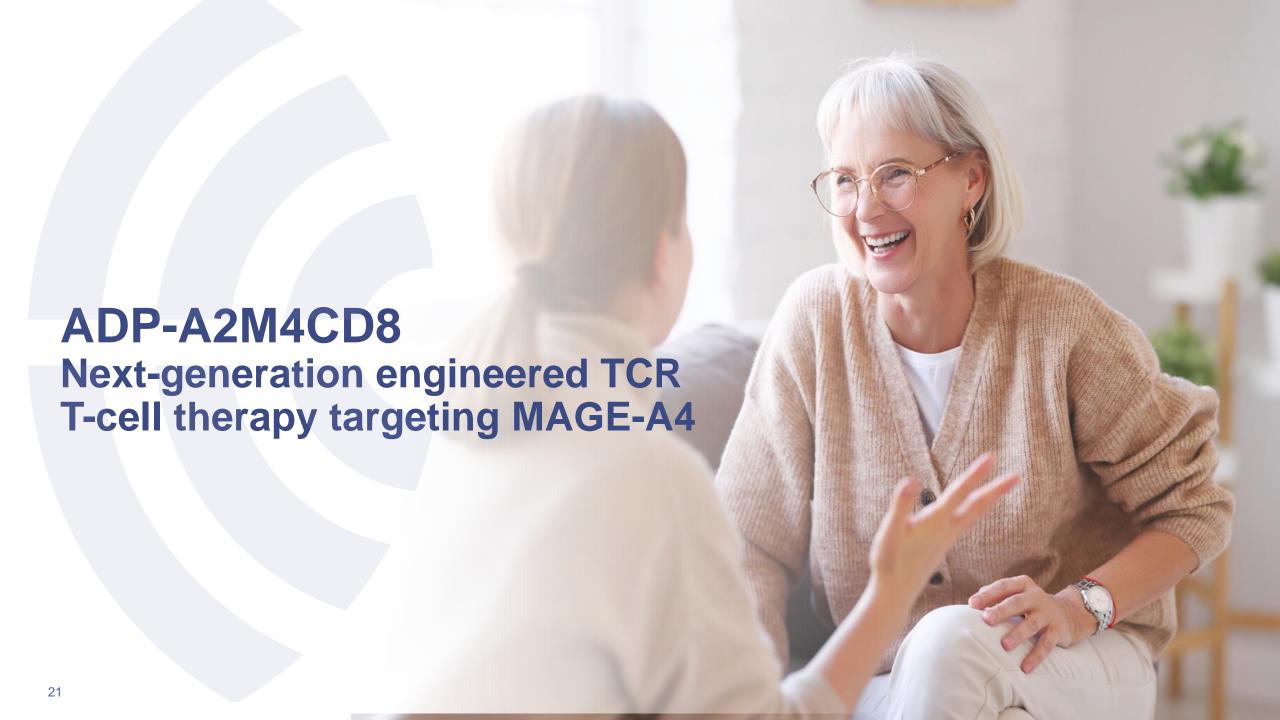
- ✓ Afami-cel provides outstanding opportunity to develop the commercial capabilities we will need as we grow to reach more patients across more indications and achieve long-term business sustainability
 - ✓ Opportunity to build capabilities and establish trust as a fully integrated cell therapy organization
 - Nimble to deliver contribution with afami-cel, and scalable to grow for pipeline
 - Enabling faster adoption, lower costs, and excellent experience for next launches, driving towards long-term business sustainability



Focused approach to ensure efficiency and enable growth









"This is an important and exciting time in oncology with TCR T-cell therapies...and their potential to help clinicians manage aggressive and difficult-to-treat cancers."

-David Hong, M.D.

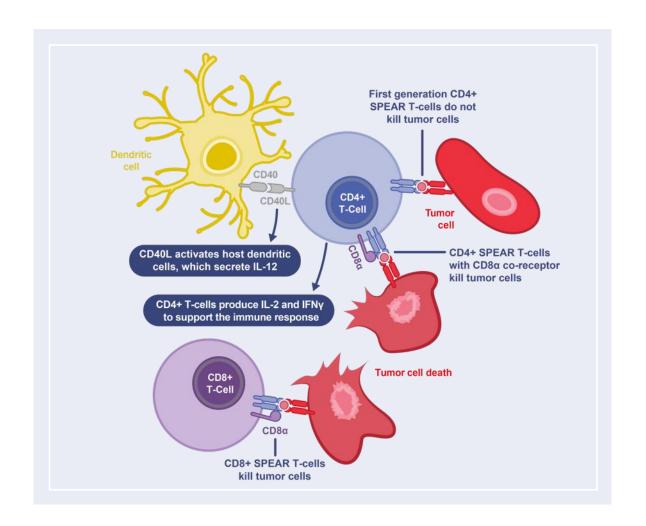
Professor and Deputy Chair of Investigational Cancer Therapeutics at The University of Texas MD Anderson Cancer Center Adaptimmune <u>ESMO 2022 press release</u>



ADP-A2M4CD8 – SURPASS family of trials

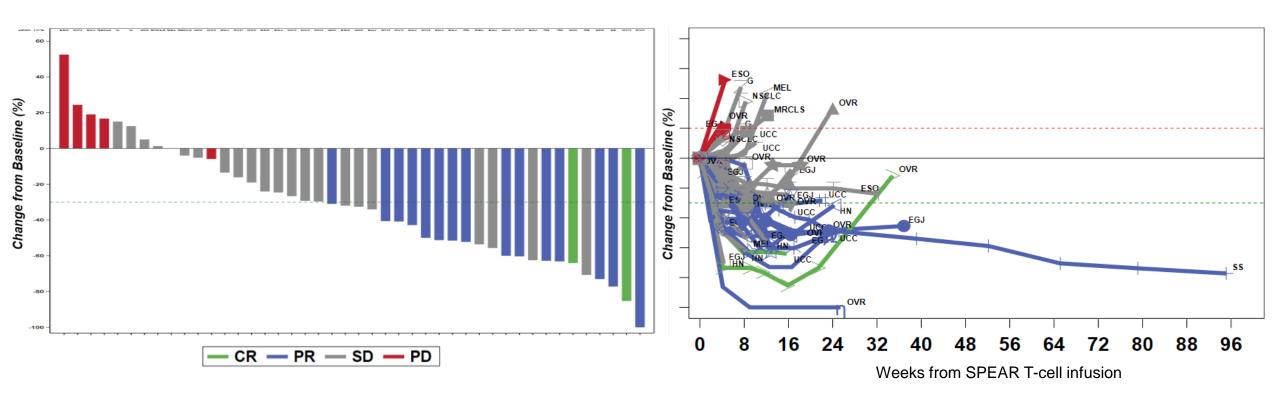
Next-gen product targeting MAGE-A4 designed to be more potent

- ✓ Same MAGE-A4 targeted TCR as afami-cel with the addition of CD8α co-receptor
- ✓ Designed to be more potent and to more effectively engage the broader immune system compared to first-gen
- ✓ Single dose of cells
- Based on results to date, focusing on ovarian, urothelial and H&N cancers
 - ✓ ORR of 52% across the three tumor types
 - √ ~ 15,000 eligible patients per year (with these three tumors) in the US and EU expressing MAGE-A4 and HLA-A2*





Results consistent: 37% response rate in SURPASS Ph 1 trial



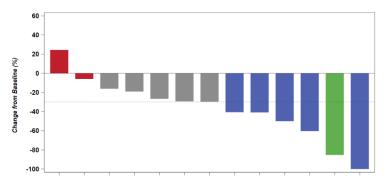
- 52% response rate in focus areas of ovarian, urothelial, and head & neck cancers (13/25)
- 75% response rate in focus areas of ovarian, urothelial, and head & neck cancers in patients with 3 or fewer prior lines of therapy (9/12)



52% ORR in Ovarian, Urothelial and H&N

Ovarian

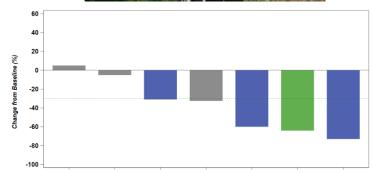




- 1 confirmed CR and 5 confirmed PRs; ORR 43% (6/14)
- Current platinum-resistant SOC ~13% response rate
- Median PFS 3-4 months

Urothelial

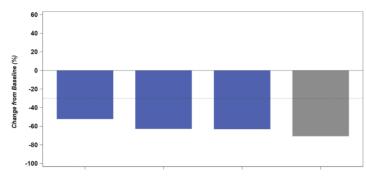




- 1 confirmed CR and 3 confirmed PRs; ORR 57% (4/7)
- Current SOC in 2nd line ~20% response rate with checkpoint inhibitor
- PFS of only ~2 months with pembrolizumab

Head and Neck*





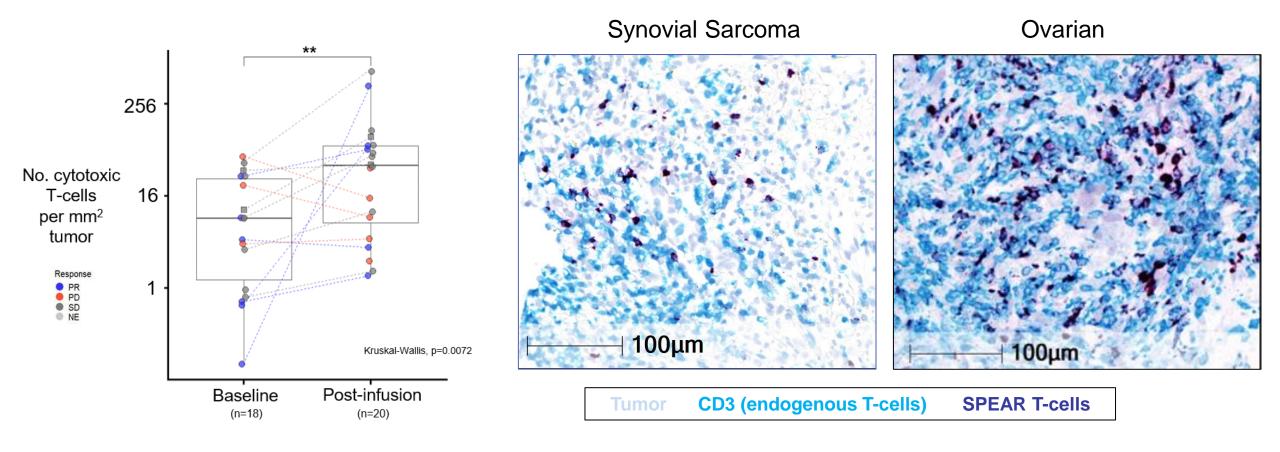
- Deep antitumor responses;
 3/4 confirmed PRs
- Current SOC 1st line pembrolizumab (CPS≥1): 19% response rate and median PFS 3-5 months





After a single dose: engineered T-cells are present in solid tumors

Other immune cells are also recruited, indicating engagement of the broader immune system in antitumor activity

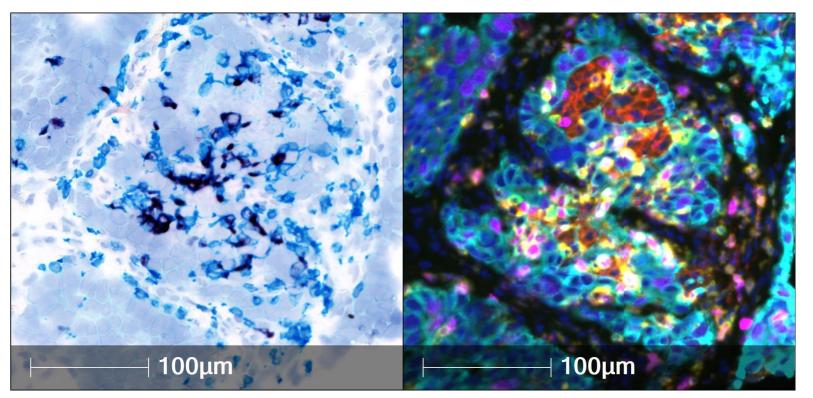


Translational data supports use of checkpoint inhibitor combination

Analysis of tumor biopsy shows upregulation of PD-L1 and other inhibitory markers in non-responding patient

Significant **engineered T-cell** infiltration into tumor

Further staining demonstrates high levels of T-cell activation alongside PD-L1+ tumor cells



CD3 - T-cell Marker
CD4 - Helper T-cell marker
CD8 - Cytotoxic T-cell marker
Ki67 - Proliferation marker
PD-L1 - Immune evasion marker
Granzyme B - Activation marker
FoxP3 - Regulatory T-cell
marker
PanCK- Tumour marker

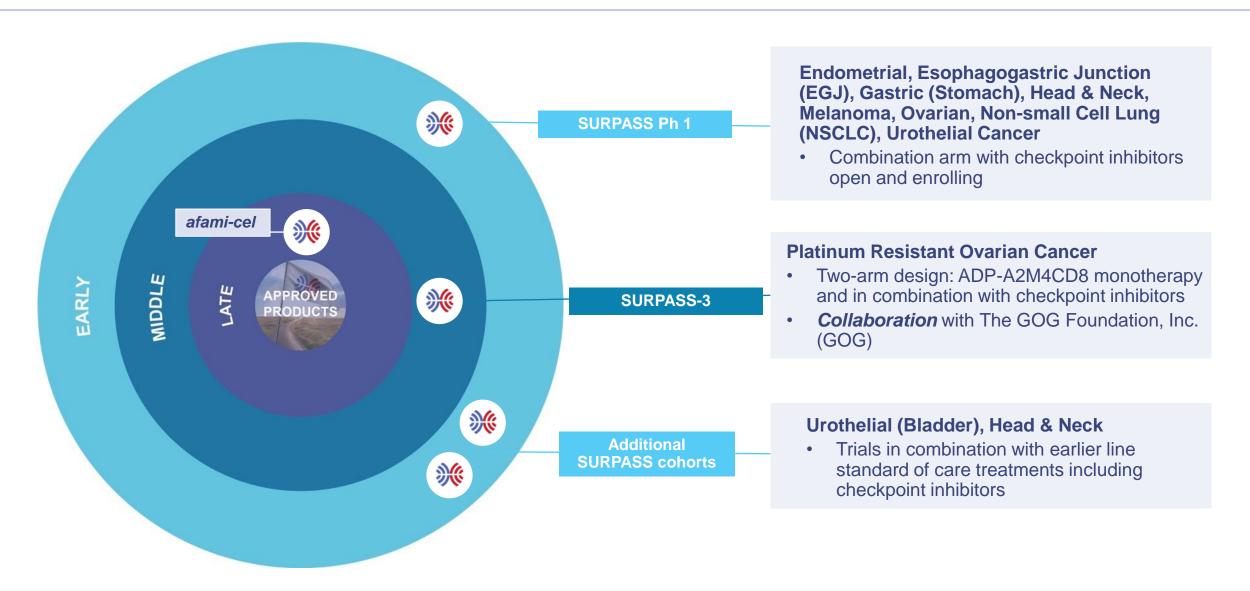
Tumor CD3 (endogenous T-cells) SPEAR T-cells



Acceptable safety profile for both afami-cel* and ADP-A2M4CD8**

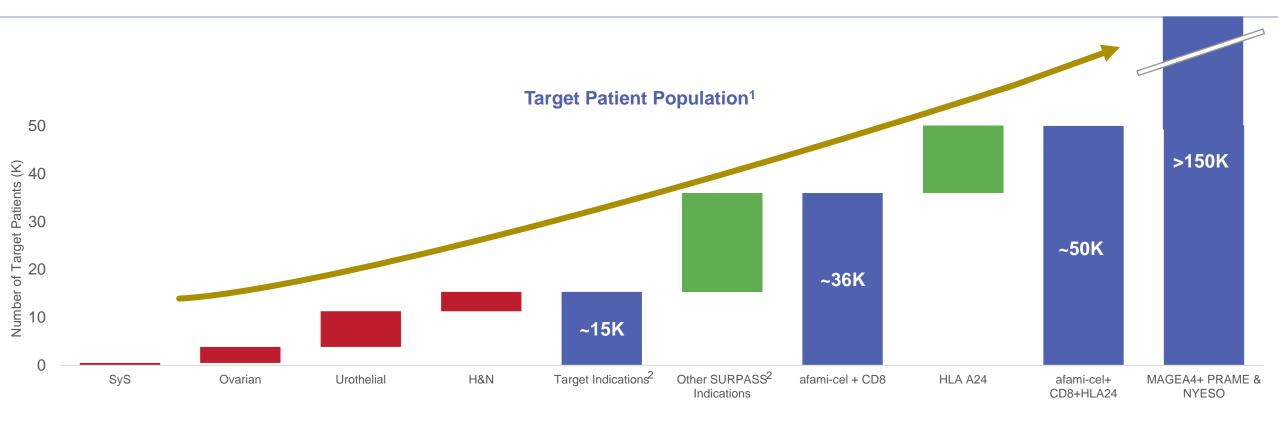
- Adverse events reported are consistent with those experienced by people with cancer undergoing chemotherapy, immuno-oncology therapy and/or adoptive cell therapy
- AEs of special interest (listed below) are generally manageable with supportive care and mitigation strategies
 - Cytokine release syndrome (CRS)
 - Neurotoxicity including immune effector cell-associated neurotoxicity syndrome (ICANS)
 - Prolonged cytopenia following lymphodepletion and SPEAR T-cell therapy

Clinical development strategy





Building towards making cell therapies mainstream in solid tumors, step-by-step



Afami-cel: Building the Base

- Set-up treatment sites
- · Establish referral networks
- Embed testing
- Operationalize commercial supply
- Build trust and connection

CD8: Broadening Indications

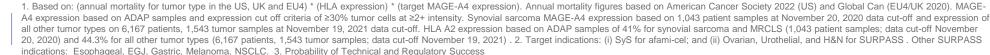
- Target indications with ~50% ORR
- >22% high MAGE-A4 expression
- Expand beyond Ovarian, Urothelial and H&N to Esophageal, EGJ, Gastric, Melanoma, and NSCLC
- Building upon established site footprint, testing, operations, connections

Other HLAs: Expanding Reach

- High PTRS ³, expanding from clinically proven platform
- ~60% coverage of population with HLA -A2 + HLA -A24, opportunity to go beyond
- Scale and reach: mainstream treatment

Other Targets: Going Beyond

- Adding indications with high PRAME & NY-ESO expression (e.g., breast, liver, testicular, thyroid, uterine)
- · Opportunity to combine and sequence
- Customize treatment based on patient target expression

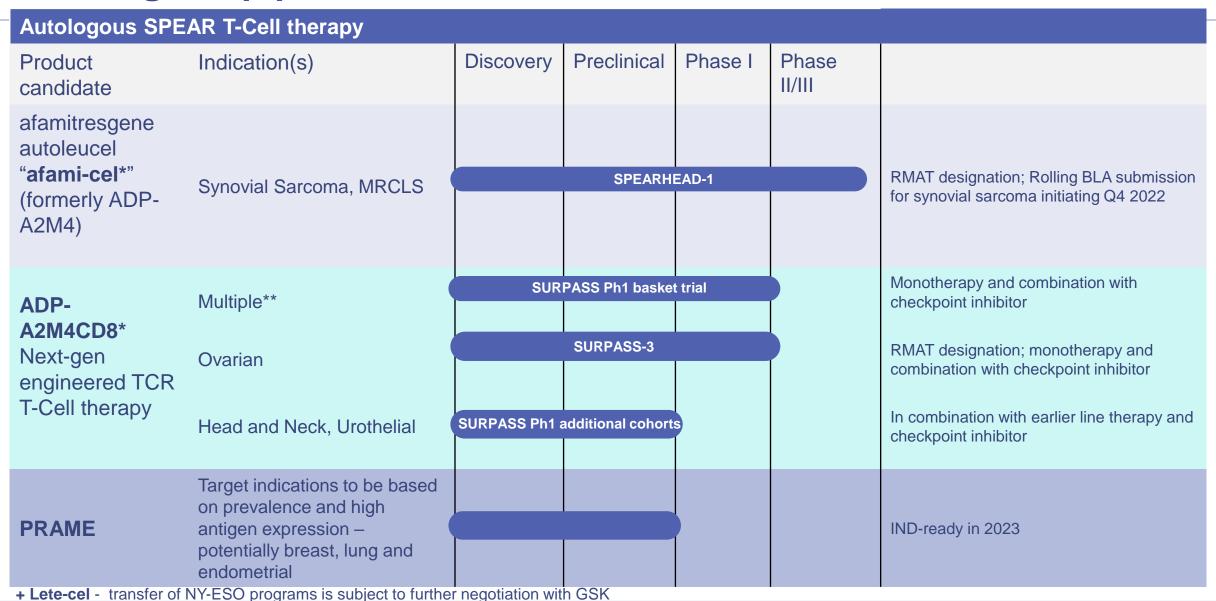




Our pipeline



Autologous pipeline





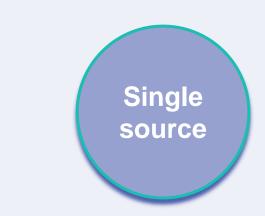
Allogeneic pipeline

Independent and in partnership with Astellas and Genentech

Allogeneic Cell Therapy Programs								
Product candidate	Indication	Discovery	Preclinical	Phase I	Phase II/III			
MAGE-A4	TBA							
HiT Mesothelin *** astellas	TBA							
Target 2 ** astellas	TBA							
Off-the-shelf" TCR therapy target 1 Genentech A Manufacraf the Earlie Group	TBA							
Personalized cell therapy platform Genentech A Manufact of the Broke Group	TBA							



Future "off-the-shelf" therapies with allogeneic platform



- iPSCs from single donor stem cells
- High proliferative potential
- Reproducible starting material



- Overcomes lentivector capacity limit
- Flexibility to add multiple next-gens or edits



- Single cell line for characterization
- Defined media composition
- No serum or feeder lines



Cell therapy manufacturing and supply



Manufacturing and supply as a key integrated capability

Experienced team

- Successfully manufactured 100s of autologous batches for patients across multiple tumor types
- Deep expertise in process and analytical development, manufacturing and supply

Secured supply

- Internal suspension lentiviral vector manufacturing
- Internal autologous drug product manufacturing
- Building internal allogeneic drug product manufacturing

Scalable infrastructure "People, Processes, and Technology"

- "Out" for more trials and processes
- "Up" for commercial GMP and increased patient numbers

Ready to supply commercial scale therapies and deliver results for patients

Faster Processing Times + Reliable Manufacturing + Reduced costs

Improved patient experience and wider access



The patient cell journey for autologous products

Current GMP manufacturing time of ~10 to 14 days



Identification and enrollment in the trial

Collection to cryopreservation ~2 days







- ✓ WBC collection (apheresis)
- ✓ Courier to manufacturing facility
- Cryopreservation



GMP manufacturing ~10 to 14 days











To clinical site for infusion





- √ Thaw WBCs and isolate T-cells
- ✓ Lentiviral transduction of SPEAR TCR
- √ T-cell expansion
- ✓ Cryopreserve dose prior to release testing

Apheresis to product release ~30 days



Adaptimmune financials



Total Liquidity at end of Q3 2022 was ~\$200M*

Runway anticipated to early 2025 with restructuring and cost savings

