



DELIVERING THE POWER  
**OF T CELLS** TO  
CANCER PATIENTS

# Immatics Corporate Presentation

June 30, 2022

# Forward-Looking Statements



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# Building a Leading TCR Therapeutics Company



## Two Clinical-Stage Modalities

Pipeline of TCR-T and TCR Bispecific product candidates in clinical & preclinical development



## Clinical PoC for Cell Therapy

Objective responses across multiple solid tumors in early TCR-T clinical development



## Differentiated Platforms

Unique technologies to identify true cancer targets and right TCRs



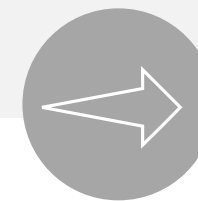
## Therapeutic Opportunity

Potential for addressing large patient populations with high prevalence targets in solid tumors



## Strategic Partnerships

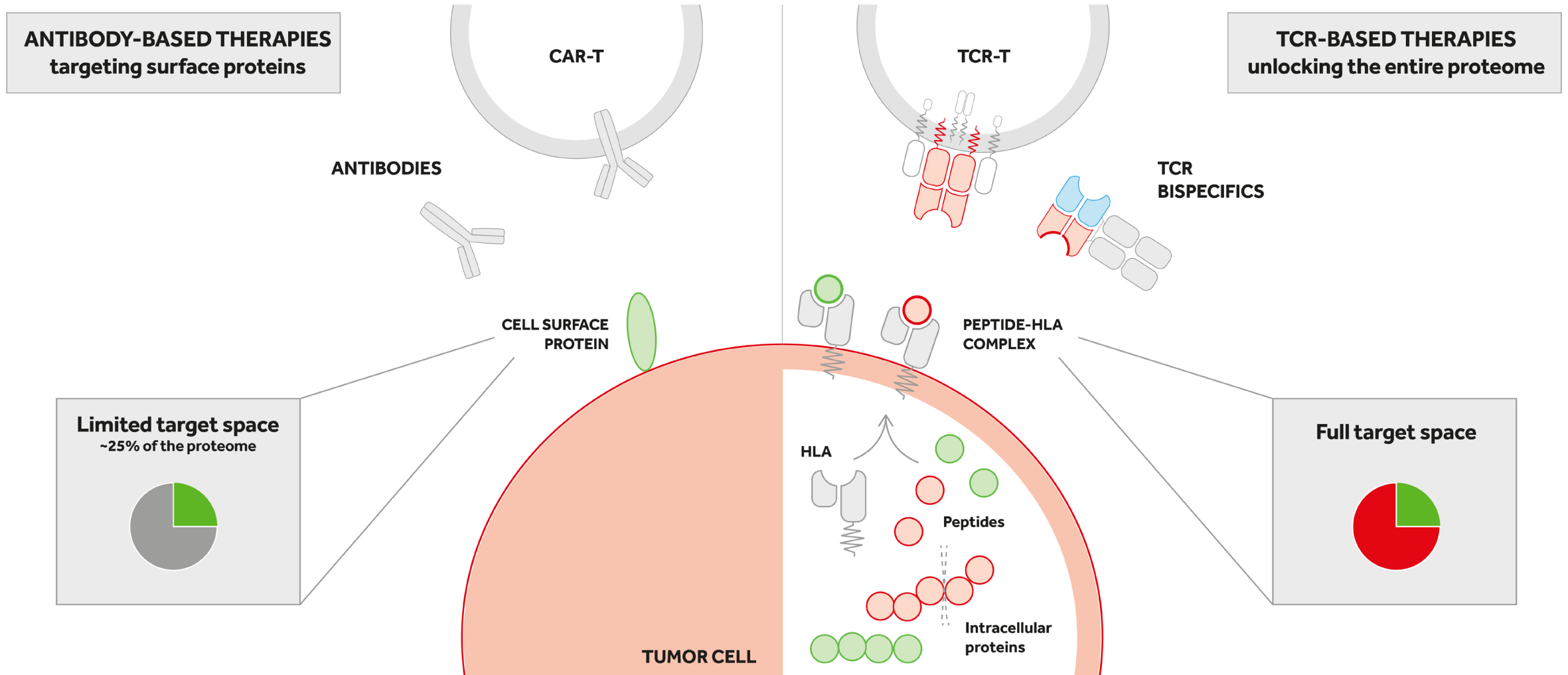
World-leading industry players with synergistic expertise



## Solid Cash Runway

To reach next value inflections points across our portfolio

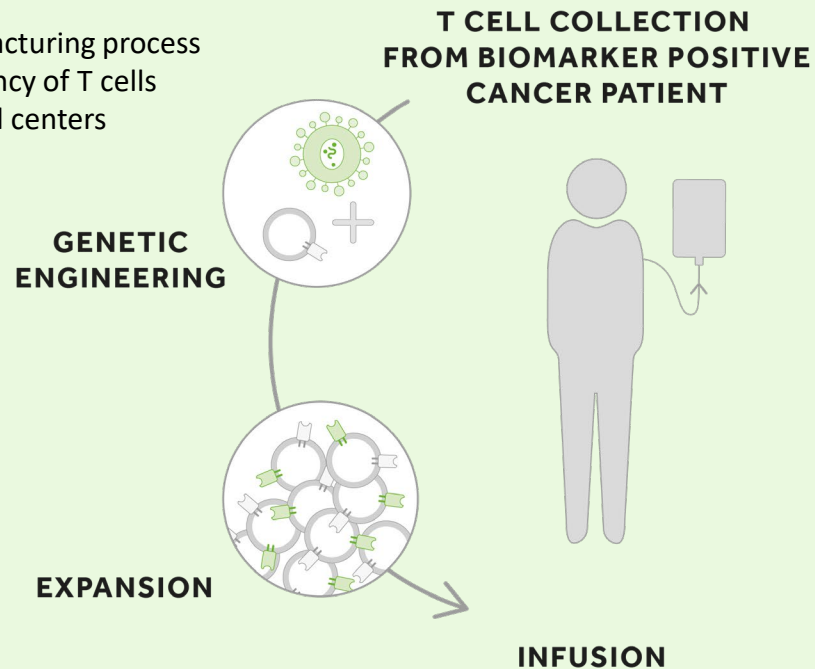
# Our TCR-based Approaches Leverage the Full Target Space beyond the Cancer Cell Surface



# Two Distinct TCR-based Therapeutic Modalities in Clinical Development

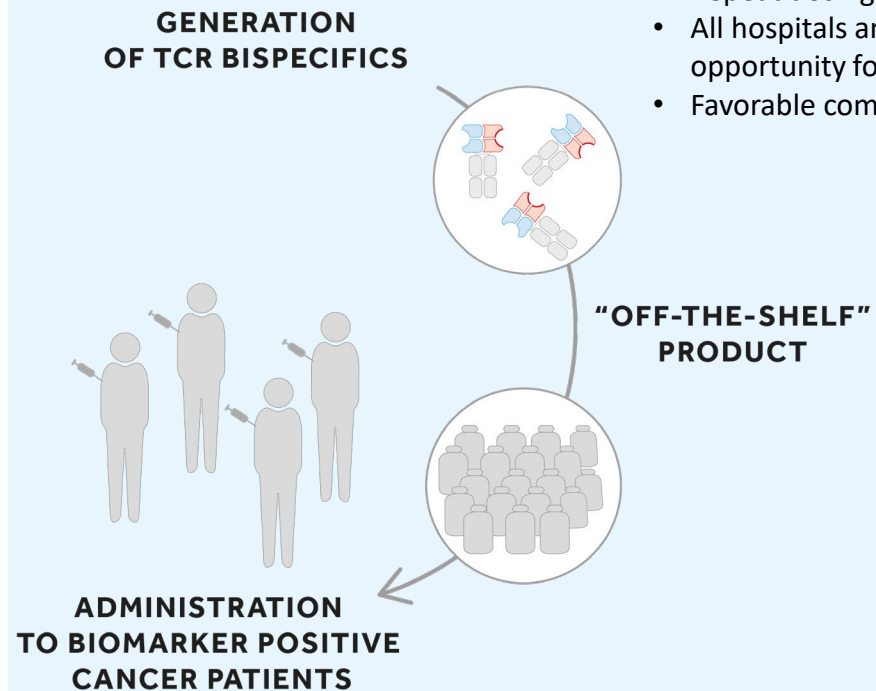
## Autologous TCR-T (ACTengine®)

- Strong clinical activity in patients with high tumor burden<sup>1</sup>
- Single dose<sup>2</sup>
- Proprietary manufacturing process for enhanced potency of T cells
- Specialized medical centers



## TCR Bispecifics (TCER®)

- Off-the-shelf biologic for immediate treatment
- Repeat dosing
- All hospitals and out-patient, opportunity for larger patient reach
- Favorable commercial characteristics



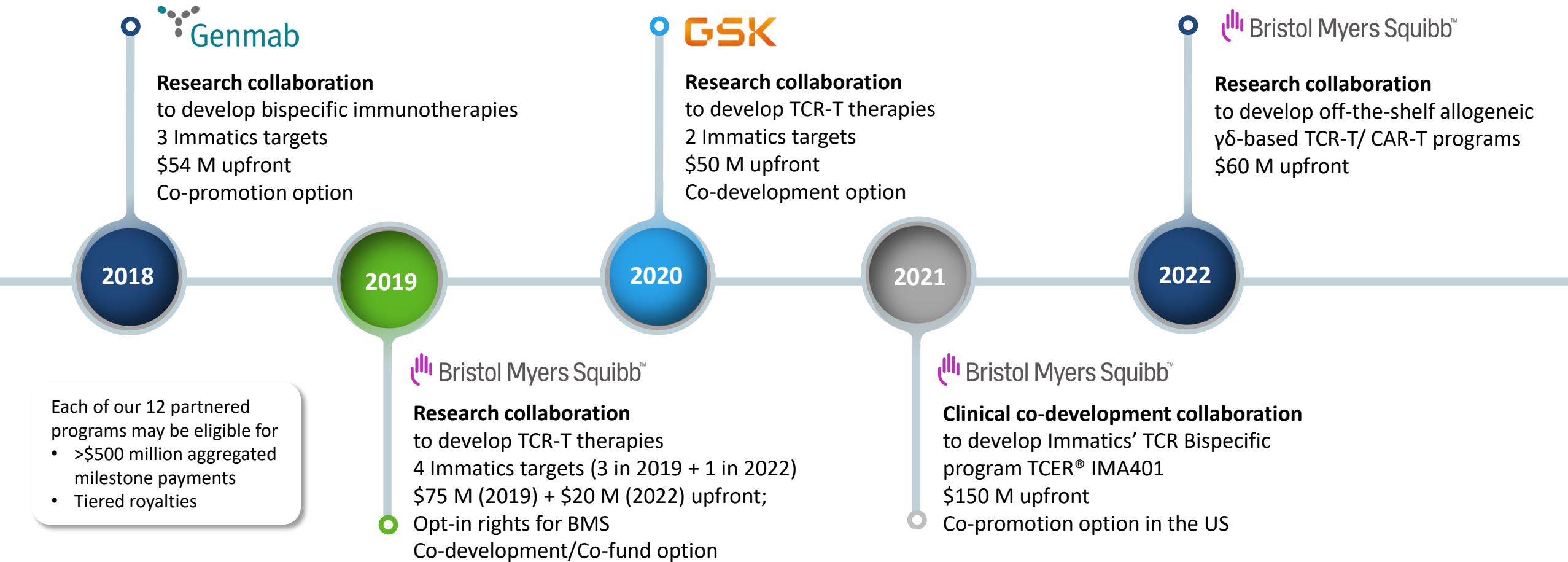
**Differentiated positioning of ACTengine® vs. TCER® based on patient population, medical need and geographical reach**

# Our Pipeline of TCR-based Adoptive Cell Therapies and Bispecifics

Modality	Product Candidate	Target		Preclinical	Phase 1a <sup>1</sup>	Phase 1b <sup>1</sup>	Phase 2/3
ACTengine® Autologous ACT	IMA203	PRAME					
				+ Checkpoint Inhibitor <sup>2</sup> 			
	IMA203CD8	PRAME					
	IMA201	MAGEA4/8					
	IMA204	COL6A3					
Autologous ACT	4 programs	Undisclosed					
	2 programs	Undisclosed					
ACTallo® Allogeneic ACT γδ T cells	IMA30x	Undisclosed					
	2 programs	Undisclosed					
TCER® Bispecifics	IMA401	MAGEA4/8					
	IMA402	PRAME					
	IMA40x	Undisclosed					
Bispecifics	3 programs	Undisclosed					

# Strategic Collaborations

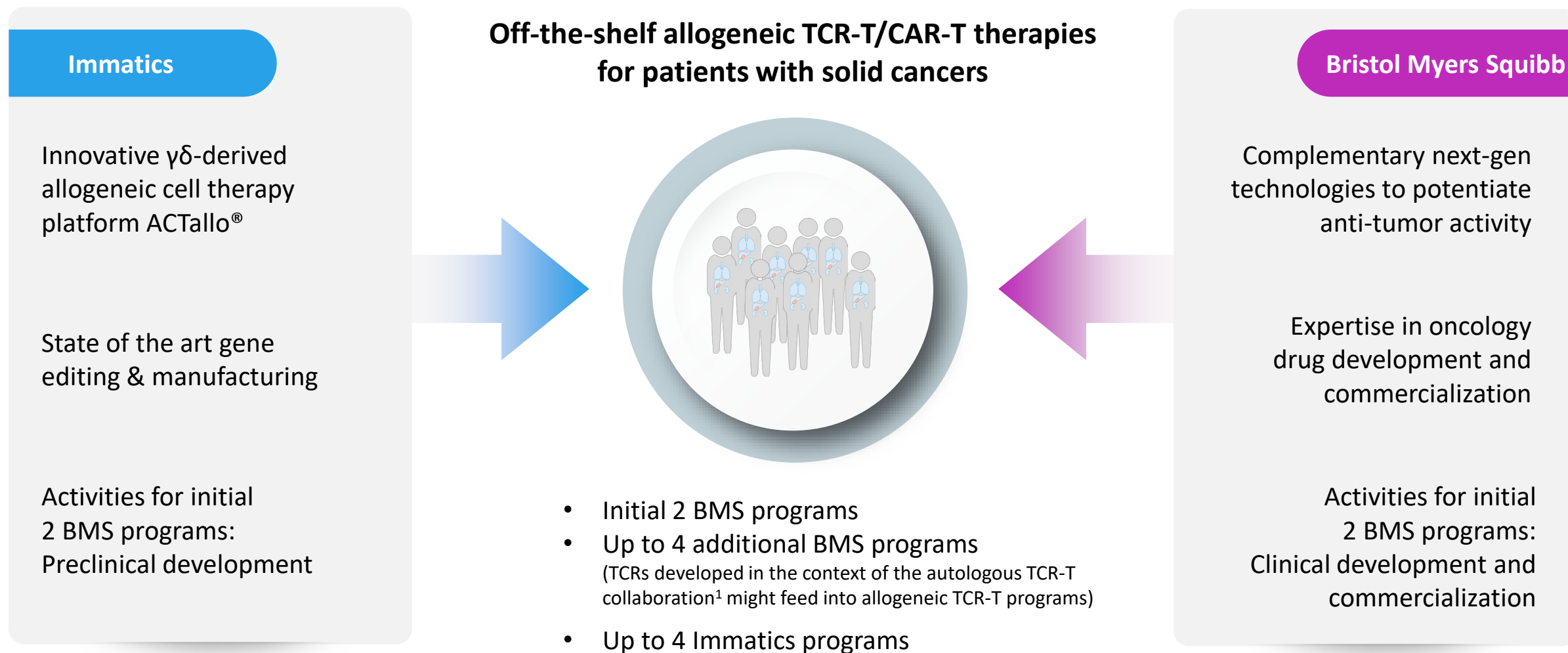
Synergistic Expertise that Can Foster Transformative Innovations for ACT and Bispecifics



Broadening the clinical framework beyond our pipeline

# Immatics and Bristol Myers Squibb – New Allogeneic Multi-program Collaboration

## Leveraging Complementary Technologies & Capabilities for the Benefit of Cancer Patients





# Potential for Large Patient Populations across Multiple Solid Cancers

	IMA201 / IMA401 MAGEA4/8	IMA203 / IMA402 PRAME	IMA204 COL6A3 exon 6
Selected solid cancer indications with significant target prevalence <sup>1</sup>	<p>Sarcoma Subtypes – up to 80%</p> <p>Squamous NSCLC – 50%</p> <p>HNSCC – 35%</p> <p>Bladder Carcinoma – 30%</p> <p>Esophageal Carcinoma – 25%</p> <p>Uterine Carcinosarcoma – 25%</p> <p>Ovarian Carcinoma – 20%</p> <p>Melanoma – 20%</p>	<p>Uterine Carcinoma – 100%</p> <p>Sarcoma Subtypes – up to 100%</p> <p>Melanoma – 95%</p> <p>Uveal Melanoma – 80%<sup>2</sup></p> <p>Ovarian Carcinoma – 80%</p> <p>Squamous NSCLC – 65%</p> <p>Kidney Carcinoma – up to 45%</p> <p>Cholangiocarcinoma – 35%</p> <p>Adeno NSCLC – 25%</p> <p>Breast Carcinoma – 25%</p> <p>HNSCC – 25%</p> <p>Esophageal Carcinoma – 20%</p> <p>HCC – 20%</p> <p>Bladder Carcinoma – 20%</p>	<p>Pancreatic Carcinoma – 80%</p> <p>Breast Carcinoma – 75%</p> <p>Stomach Carcinoma – 65%</p> <p>Sarcoma – 65%</p> <p>Esophageal Carcinoma – 60%</p> <p>Squamous NSCLC – 55%</p> <p>Adeno NSCLC – 55%</p> <p>HNSCC – 55%</p> <p>Uterine Carcinosarcoma – 55%</p> <p>Colorectal Carcinoma – 45%</p> <p>Mesothelioma – 45%</p> <p>Cholangiocarcinoma – 40%</p> <p>Ovarian Carcinoma – 40%</p> <p>Melanoma – 35%</p> <p>Bladder Carcinoma – 35%</p>

**ACTengine® and TCER® targets demonstrate high expression in multiple solid cancers**



## ACTengine® IMA203 – TCR-T Targeting PRAME

# ACTengine® IMA203 – TCR-T Targeting PRAME

Broadly Expressed Target on Multiple Solid Cancers Combined with Highly Specific TCR

## TARGET

HLA-A\*02-presented peptide derived from **PRAME**

Naturally and specifically presented on tumors at high target density<sup>1</sup>:  
**100-1,000 copies/cell**

Identified and validated by XPRESIDENT® quant. mass spectrometry platform

## TCR

High-affinity, specific TCR targeting PRAME

**Pairing-enhanced, engineered TCR** to avoid mispairing

High functional avidity<sup>2</sup>:  
**EC50 ~5 ng/ml**

Identified and characterized by XCEPTOR® TCR discovery and engineering platform

## CLINICAL DATA

N=18 pts treated in phase 1 dose escalation cohort

Manageable tolerability profile; no additional DLTs<sup>3</sup> & no CRS/ICANS ≥ grade 3

16 patients with at least one post treatment tumor assessment

**Objective responses in 50% (8/16) of patients, thereof 62% (8/13) of responses above DL1; all doses still below 1 bn cells**

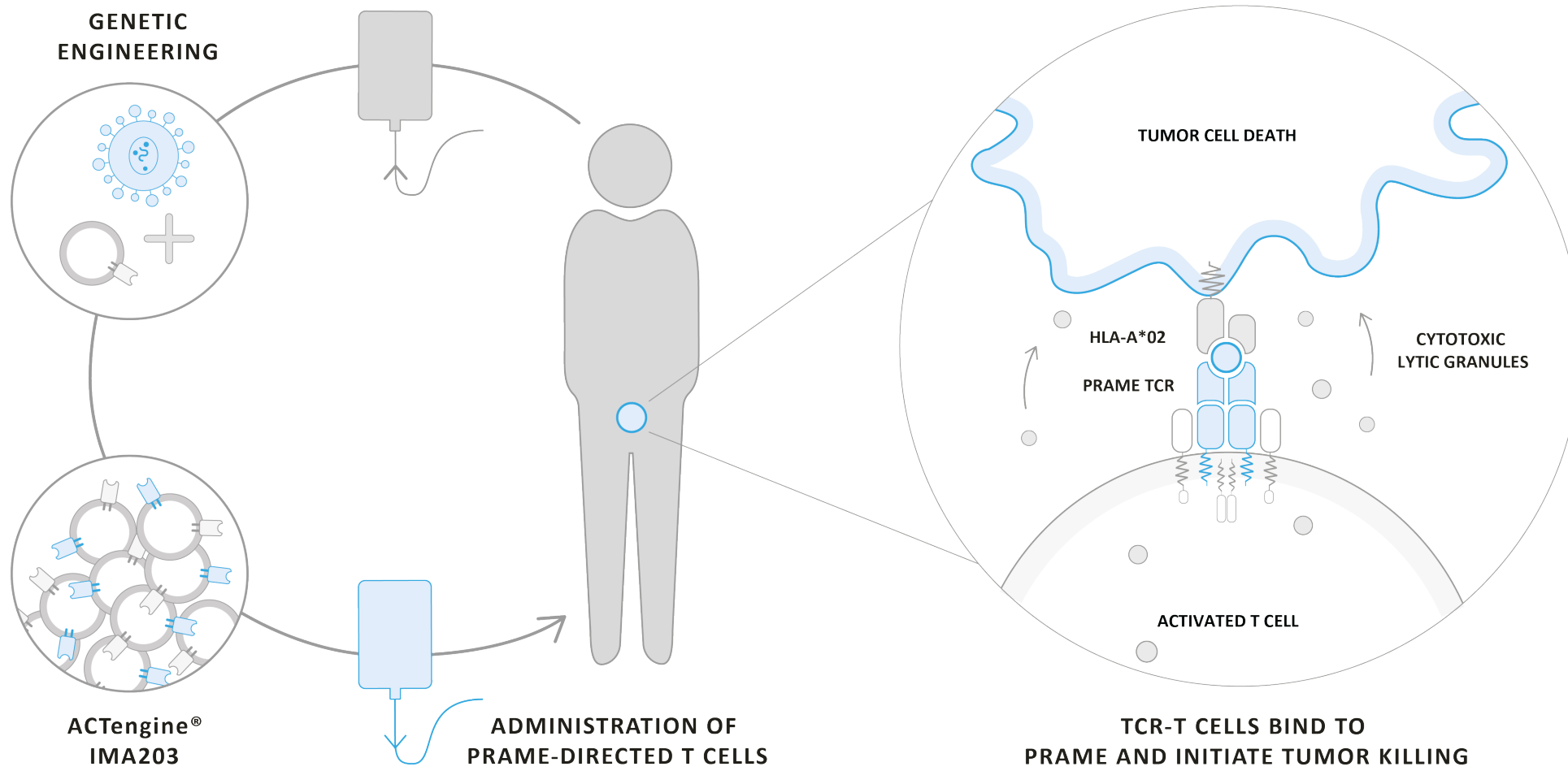
## PATIENT POPULATION<sup>4</sup>

Uterine Carcinoma – 100%  
 Sarcoma Subtypes – up to 100%  
 Melanoma – 95%  
 Uveal Melanoma – 80%<sup>5</sup>  
 Ovarian Carcinoma – 80%  
 Squamous NSCLC – 65%  
 Kidney Carcinoma – up to 45%  
 Cholangiocarcinoma – 35%  
 Adeno NSCLC – 25%  
 Breast Carcinoma – 25%  
 HNSCC – 25%  
 Esophageal Carcinoma – 20%  
 HCC – 20%  
 Bladder Carcinoma – 20%

Data cut-off – 05-Oct-2021

# ACTengine® IMA203 Targeting PRAME – Mechanism of Action

## Immatics' Leading TCR-T Approach



# Optimized Cell Therapy Products to Enhance T cell Persistence & Efficacy

## Current Proprietary Manufacturing Protocol for ACTengine® Product Candidates

### Leukapheresis



ACTengine® clinical programs: ~3 weeks

Manufacturing time (~1 week)	QC testing (Full sterility, 2 weeks)
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Commercial ACTengine® expected ~2 weeks

Manufacturing time (~1 week)	Expedited QC testing (~1 week)
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### Infusion-Ready



### Proprietary Manufacturing Process, designed to

- ✓ reduce manufacturing process to approx. 1 week<sup>1</sup>
- ✓ shorten vein-to-vein time
- ✓ generate younger T cells with increased proliferative capacity
- ✓ improve engraftment and persistence in patients while utilizing smaller doses

### In-house state-of-the-art cGMP Facility<sup>2</sup>

- ✓ Manufacturing by Immatics personnel
- ✓ Maximum capacity: 48 manufacturing runs/month
- ✓ Substantial in-house process development expertise

# ACTengine® IMA203 – Patient Flow

## Screening & Manufacturing Phase

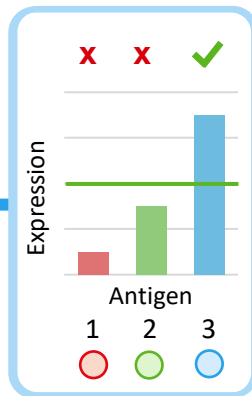
## Treatment & Observation Phase

## Long Term Follow-up

Safety and efficacy monitoring for 12 months

### HLA-A\*02 Testing

Blood sample;  
Central lab

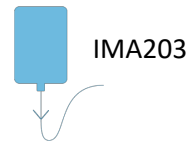


### Target Profiling

Fresh Tumor Biopsy;  
IMADetect®

### Leuka- pheresis

**ACTengine®**  
Manufacturing  
by Immatics



### Infusion of ACTengine® T cell Product

### Low dose IL-2

1m IU daily days 1-5 and  
twice daily days 6-10\*

### Lymphodepletion

30 mg/m<sup>2</sup> Fludarabine<sup>1</sup> and  
500 mg/m<sup>2</sup> Cyclophosphamide  
for 4 days

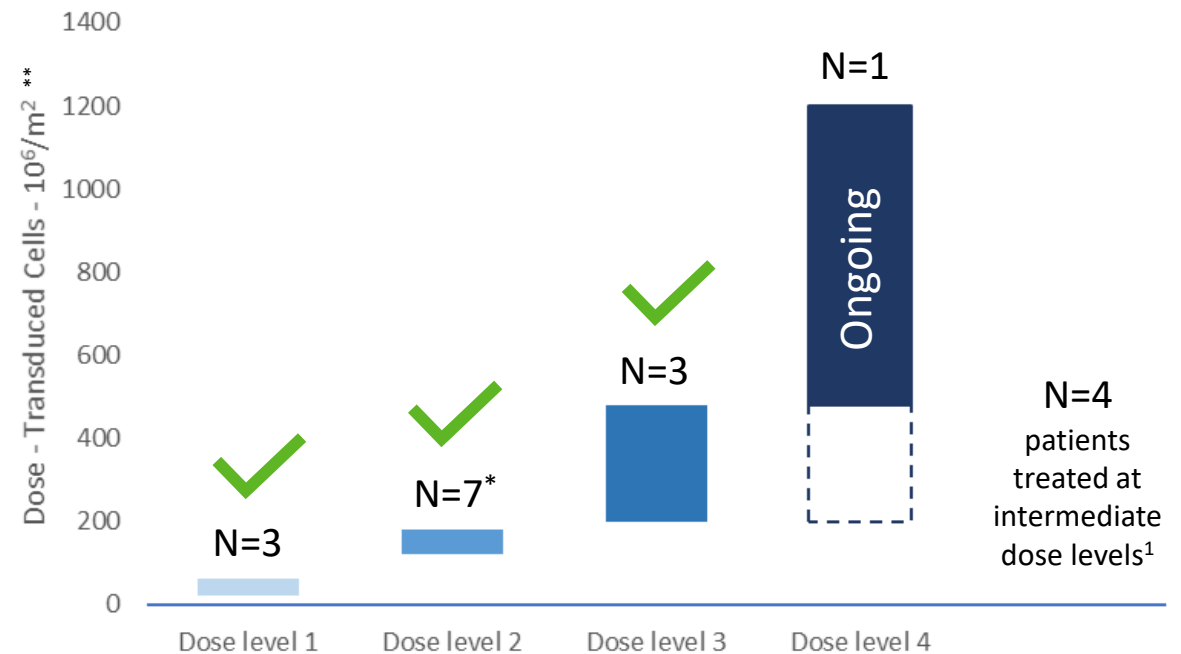
# ACTengine® IMA203 – Key Objectives & Trial Design

Presented at SITC Conference as Late-Breaking Presentation (Cut-off October 05, 2021)

## Key Study Objectives

- **Primary: Safety**
  - Investigation of Adverse Events,
  - Determination of a recommended Phase 2 dose
- **Secondary: Biological and Clinical Activity**
  - T cell engraftment and persistence
  - Objective responses as per RECIST1.1
  - Duration of response
- **Exploratory**
  - Tumor Infiltration

## Trial Design & Recruitment Status



18 patients<sup>1</sup> infused with PRAME-directed T cells at 5 clinical sites

Data cut-off – 05-Oct-2021

<sup>1</sup> Enrichment cohorts EC1 & EC2: patients infused with intermediate doses enabling infusion of patients with medical need during dose escalation observation periods, or in case of lower production yields; \* One patient infused at the same dose level as part of the enrichment cohort; \*\*Dose is shown as transduced viable CD8 T cells per m<sup>2</sup> total body surface area



# ACTengine® IMA203 – Safety Profile

## Manageable & Transient Treatment-emergent Adverse Events – No ≥ Grade 3 CRS or ICANS

TEAEs by maximum severity (N=19)<sup>1</sup>

Adverse event	All grades		≥ Grade 3		Adverse event	All grades		≥ Grade 3	
	No.	%	No.	%		No.	%	No.	%
Patients with any adverse event	19	100.0	19	100.0	table continued...				
<b>Adverse Events of Special interest</b>					<b>Cardiac or vascular disorders</b>				
Cytokine release syndrome	17	89.5	0	0.0	Hypertension	3	15.8	2	10.5
ICANS <sup>2</sup>	4	21.1	0	0.0	Atrial fibrillation	2	10.5	1 <sup>4</sup>	5.3
<b>Blood and lymphatic system disorders</b>					<b>General disorders and administration site conditions</b>				
Neutropenia*	16	84.2	15	78.9	Fatigue	7	36.8	1	5.3
Anaemia	16	84.2	9	47.4	Pyrexia	5	26.3	0	0.0
Thrombocytopenia	15	78.9	7	36.8	Oedema peripheral	3	15.8	0	0.0
Lymphopenia*	14	73.7	14	73.7	<b>Gastrointestinal disorders</b>				
Leukopenia*	12	63.2	11	57.9	Nausea	12	63.2	0	0.0
Cytopenia	1	5.3	1	5.3	Vomiting	7	36.8	0	0.0
<b>Infections and infestations</b>					Diarrhoea	7	36.8	0	0.0
Enterococcal infection	1	5.3	1	5.3	Constipation	6	31.6	0	0.0
COVID-19	1	5.3	1	5.3	<b>Investigations</b>				
Appendicitis	1	5.3	1	5.3	Aspartate aminotransferase increased	5	26.3	0	0.0
Sepsis <sup>3</sup>	1	5.3	1	5.3	Alanine aminotransferase increased	4	21.1	0	0.0
<b>Respiratory, thoracic and mediastinal disorders</b>					Blood creatinine increased	4	21.1	0	0.0
Hypoxia	2	10.5	1	5.3	<b>Other</b>				
Pleural effusion	2	10.5	1	5.3	Rash	5	26.3	0	0.0
Bronchial obstruction	1	5.3	1	5.3	Myalgia	4	21.1	0	0.0
<b>Metabolism and nutrition disorders</b>					Arthralgia	3	15.8	0	0.0
Hyponatraemia	7	36.8	1	5.3	Alopecia	3	15.8	0	0.0
Hypokalaemia	5	26.3	1	5.3	Rash maculo-papular	2	10.5	1	5.3
Decreased appetite	3	15.8	0	0.0	Orchitis	1	5.3	1	5.3
					Contrast media allergy	1	5.3	1	5.3

CRS/ICANS:  
No ≥ Grade 3 CRS  
or ICANS  
observed so far

Most Adverse  
Events were  
associated with  
lymphodepletion

DLT:  
Transient, Grade 3  
atrial fibrillation  
Onset on day 5 post  
infusion that  
resolved within 48h  
DLT triggered  
expansion of DL2

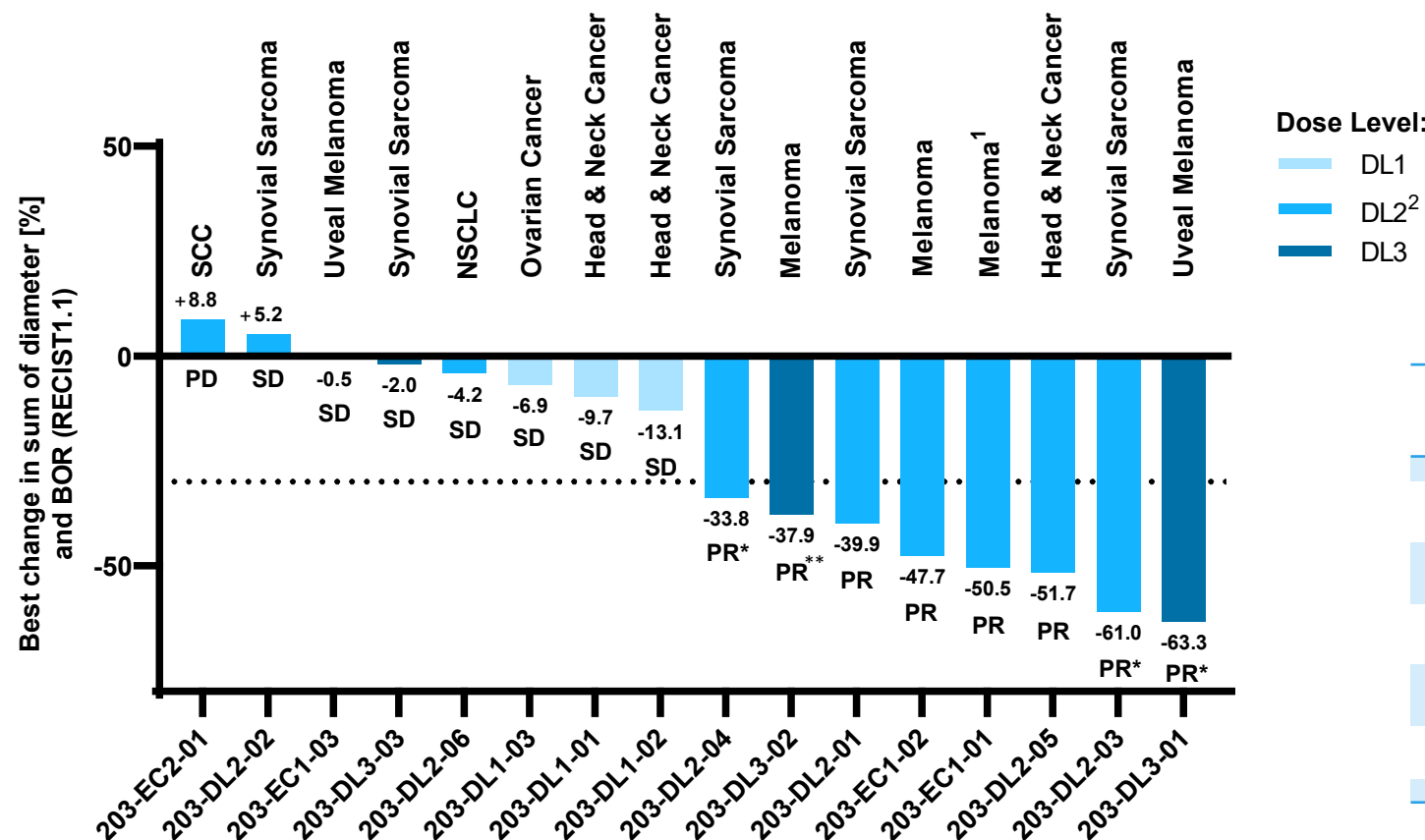
<sup>1</sup> All treatment-emergent adverse events (TEAEs) with grade 1-2 occurring in at least 3 patients (incidence ≥15.8%) and additionally all events with grade 3-5 regardless of relatedness to study treatment are presented. Data source: clinical database. Adverse events were coded using the Medical Dictionary for Regulatory Activities. Grades were determined according to National Cancer Institute Common Terminology Criteria of Adverse Events (CTCAE), version 5.0. Grades for Cytokine release syndrome and ICANS were determined according to CARTOX criteria (Neelapu *et al.*, 2018). Patients are counted only once per adverse event and severity classification; <sup>2</sup> ICANS: Immune effector cell-associated neurotoxicity syndrome; <sup>3</sup> Patient died from sepsis of unknown origin and did not receive IMA203 T cells; <sup>4</sup> DLT: Dose limiting toxicity; \*100% of patients experienced transient cytopenias ≥ Grade 3 (CTCAE v5.0)



# ACTengine® IMA203 – Change in Target Lesions

## Objective Responses across Multiple Tumor Types at Doses below 1 billion Transduced Cells

Best Overall Response (RECIST1.1)



Preliminary Objective Response Rates (RECIST1.1, confirmed and unconfirmed)

	All doses	Dosed above DL1
All comers	8/16 (50%)	8/13 (62%)
Melanoma	3/3 (100%)	3/3 (100%)
Head & Neck Cancer	1/3 (33%)	1/1 (100%)
Synovial Sarcoma	3/5 (60%)	3/5 (60%)
Uveal Melanoma	1/2 (50%)	1/2 (50%)

Data cut-off – 05-Oct-2021

<sup>1</sup> RECIST1.1 response at the timepoint of maximum change of target lesions (week 12): PD due to new lesions (leptomeningeal disease) at week 12

<sup>2</sup> Patients dosed with DL2, EC1 and EC2; \* Confirmed at subsequent scan; \*\* Confirmation pending as of data cut-off

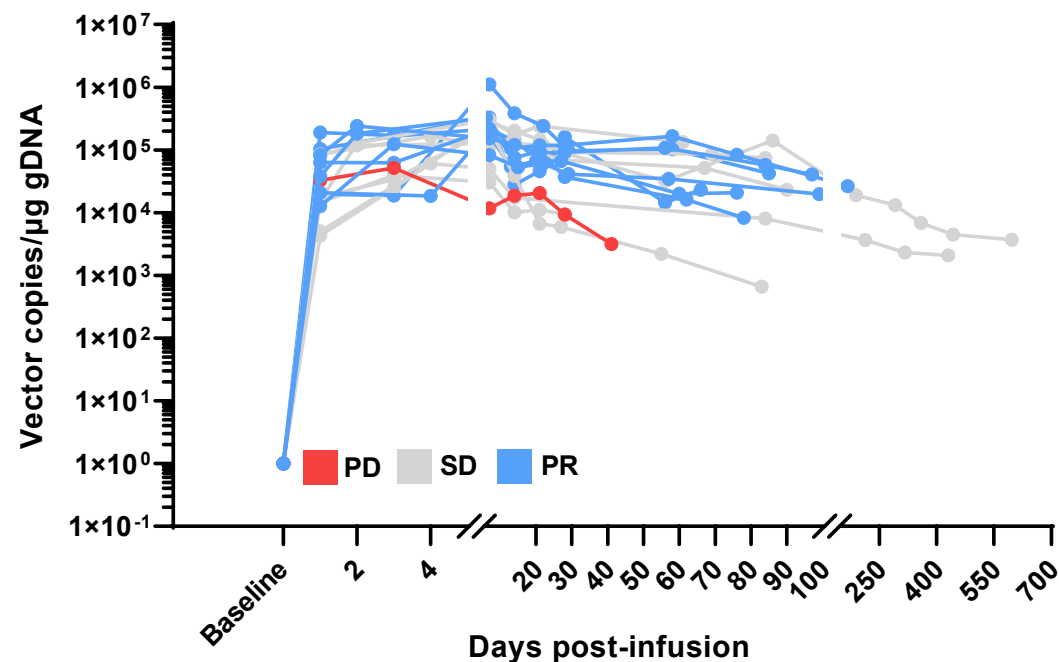
Patient ID	Indication	Dose	Week															Month							
			1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
203-DL1-01	Head & Neck Cancer	DL1																							
203-DL1-02	Head & Neck Cancer	DL1																							
203-DL1-03	Ovarian Cancer	DL1																							
203-EC1-01	Melanoma	EC1																							
203-EC1-02	Melanoma	EC1																							
203-EC1-03	Uveal Melanoma	EC1																							
203-DL2-01	Synovial Sarcoma	DL2																							
203-DL2-02	Synovial Sarcoma	DL2																							
203-DL2-03	Synovial Sarcoma	DL2																							
203-DL2-04	Synovial Sarcoma	DL2																							
203-DL2-05	Head & Neck Cancer	DL2																							
203-DL2-06	NSCLC	DL2																							
203-EC2-01	SCC	EC2																							
203-DL3-01	Uveal Melanoma	DL3																							
203-DL3-02	Melanoma	DL3																							
203-DL3-03	Synovial Sarcoma	DL3																							

Data cut-off – 05-Oct-2021

# ACTengine® IMA203 – Engraftment, Persistence & Tumor Infiltration

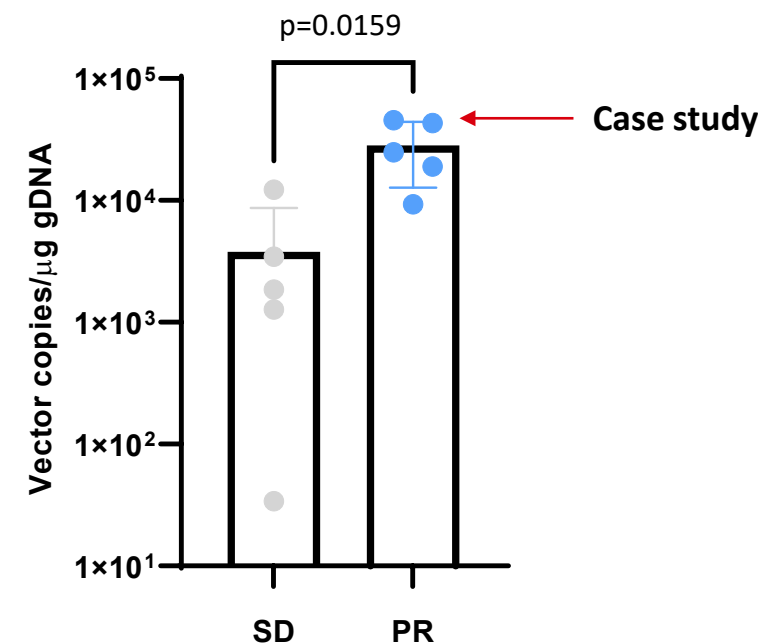
## Clinical Responses Consistent with Biological Data

### T cell Engraftment & Persistence



High T cell engraftment and persistence with trend for association of peak vector copies with clinical response<sup>1</sup>

### Tumor Infiltration post Infusion<sup>2</sup>

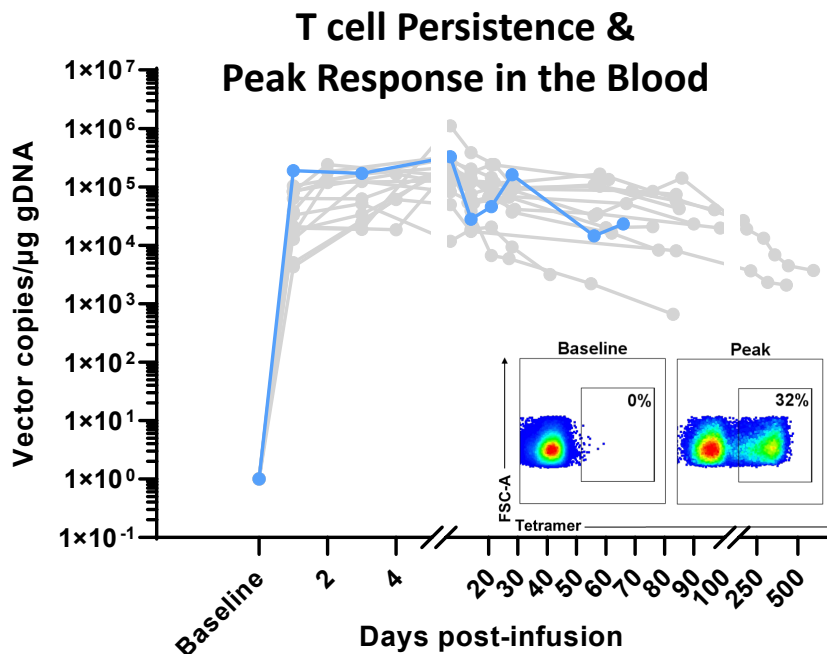


High T cell infiltration observed through serial biopsies associated with clinical response<sup>3</sup>

Data cut-off – 05-Oct-2021

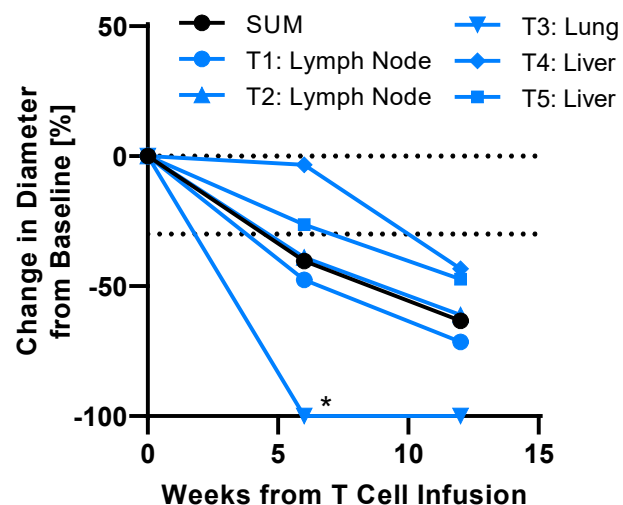
# ACTengine® IMA203 – Case Study Patient IMA203-DL3-01

## Confirmed Partial Response with Deepening Tumor Regression in Multiple Lesions

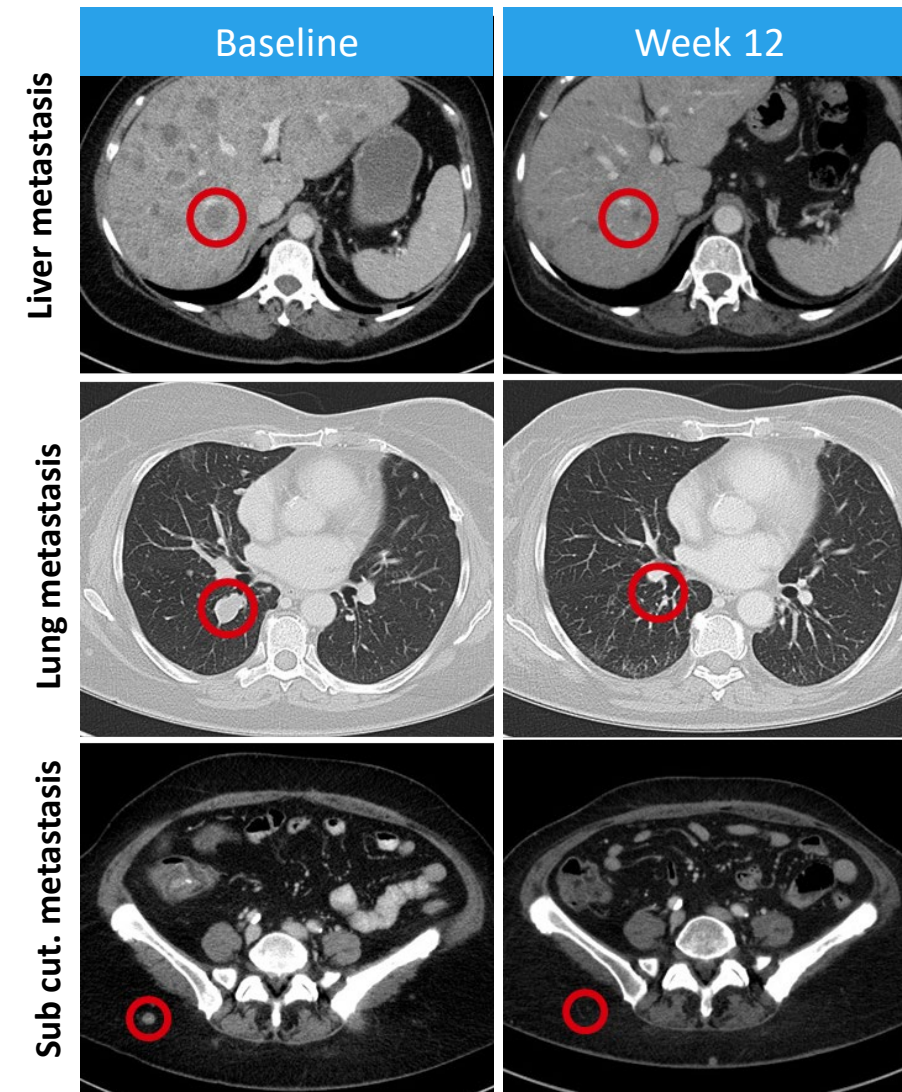


- 62-year-old female; metastatic uveal melanoma
- High tumor burden in multiple organs
- Infused at refractory disease after failing 4 prior lines of therapy including 2 lines of CPI<sup>1</sup>
- Patient received total dose of 0.59 billion transduced T cells following lymphodepletion

### Change in Target Lesions



- T cell persistence until end of observation & detection in the tumor
- All lesions decreased at week 6 - 40% decrease in target lesions response deepened at week 12 to 63% decrease
- Best Response (RECIST1.1): PR (confirmed & ongoing)



# ACTengine® IMA203 PRAME – Phase 1a Dose Escalation Interim Update

## Preliminary Findings after Completion of Dose Level 3



Objective responses observed across multiple tumor types  
at dose levels below 1 billion T cells originally presumed to be subtherapeutic

### SAFETY

- 3** Dose levels completed, all below 1 bn cells
- 0** Additional DLTs<sup>1</sup>
- 0** Grade ≥3 CRS or ICANS<sup>2</sup>

### CLINICAL ACTIVITY

- 50%** ORR<sup>3</sup> across all doses and multiple solid cancers (8/16 patients)
- 62%** ORR<sup>3</sup> at DL2\* & DL3 (8/13 patients) – all still dosed below 1 bn cells

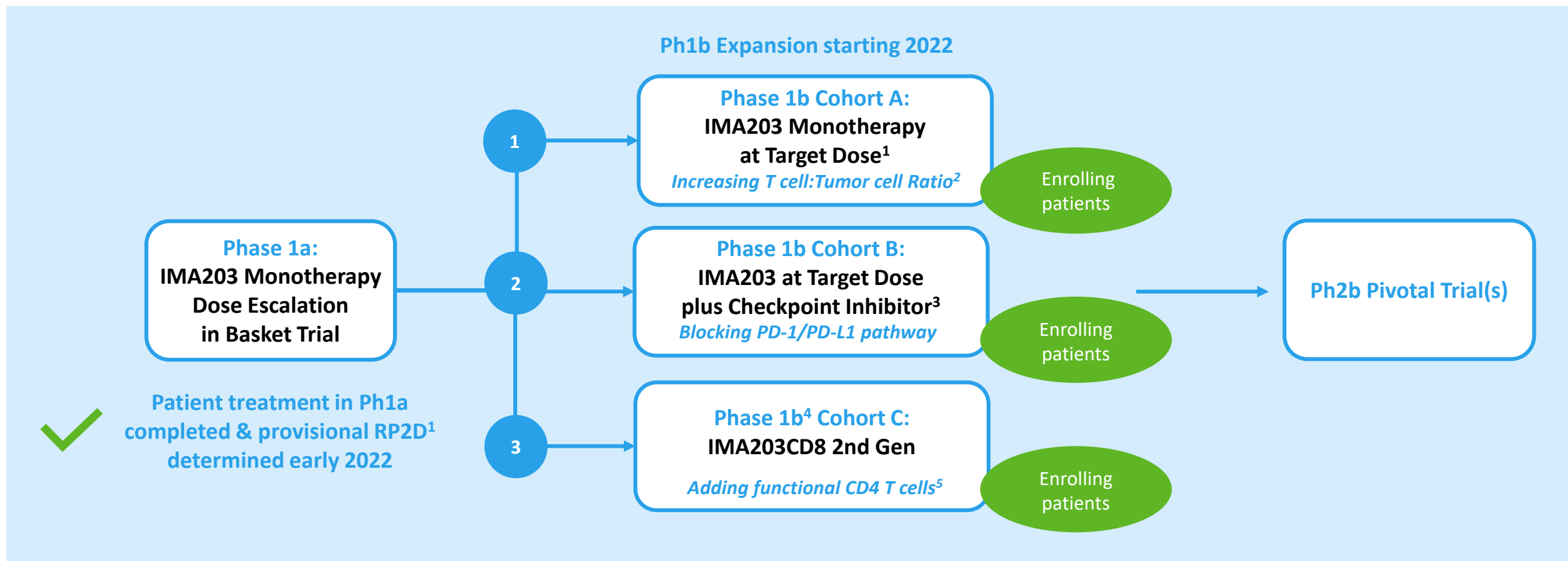
### BIOLOGICAL ACTIVITY

- Blood** High T cell engraftment and persistence
- Tumor** High T cell infiltration associated with clinical response

Data cut-off – 05-Oct-2021

# Our Plans to Achieve Long-Lasting Responses with TCR-T cells against PRAME

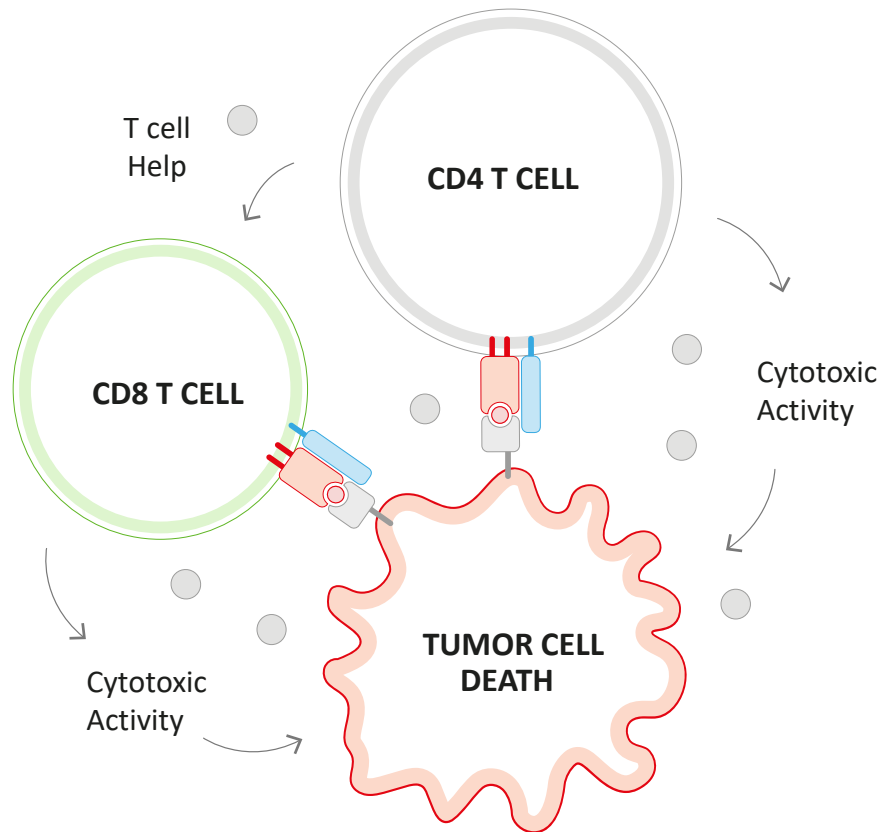
## Addressing Relevant Secondary Resistance Mechanisms to Increase Durability of Response



Each expansion cohort is designed to evaluate the observed objective response rate, demonstrate durability of response & provide the basis for entering registration trials

# ACTengine® IMA203CD8 – Next-generation TCR-T

Building on First-Gen IMA203 Success to Further Improve Anti-Tumor Activity



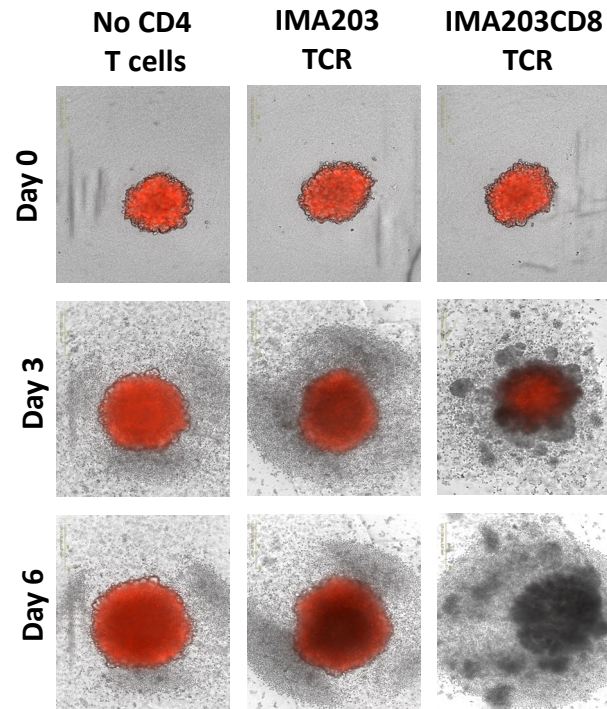
- Engagement of CD4 T cells by CD8 co-transduction reported to boost anti-tumor activity in TCR-T trials
- Recent data from leukaemia patients treated with CAR-T achieving decade-long remissions show that CD4 T cells dominate at the later time points of response<sup>1</sup>
- Functional superiority of the **CD8αβ** construct over multiple other CD8 constructs in preclinical experiments
- Proprietary 4-in-1 lentiviral vector to engineer CD4 and CD8 T cells with the PRAME-specific IMA203 TCR and CD8αβ construct (IMA203CD8)



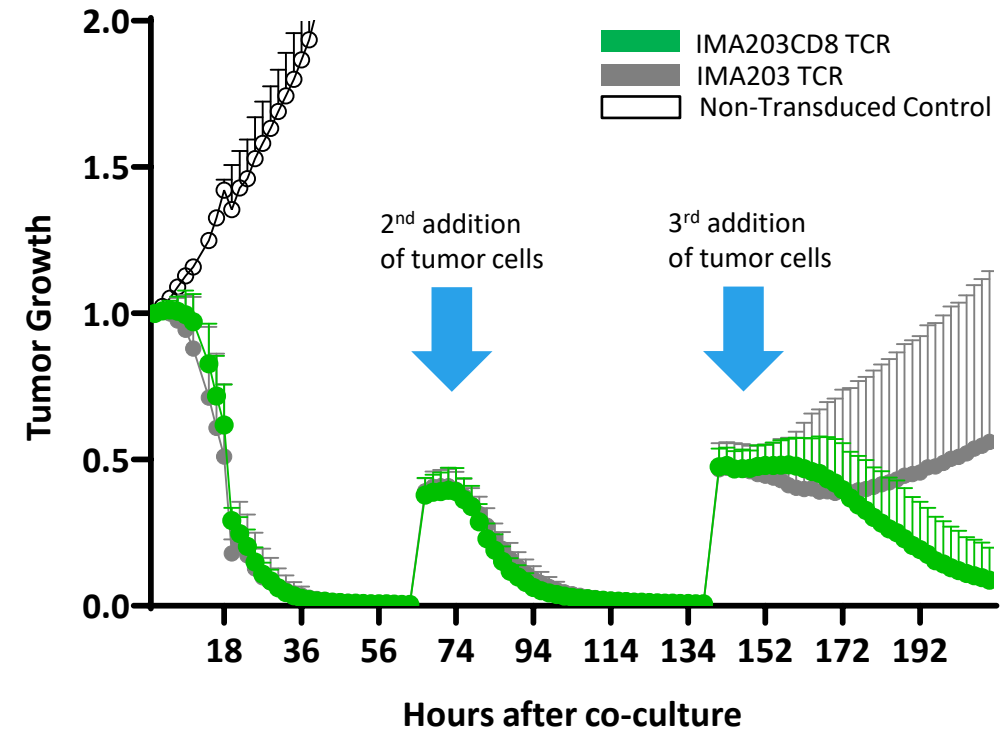
# ACTengine® IMA203CD8 – Preclinical Assessment of Anti-Tumor Efficacy

## Co-Transduction of CD8 Enhances Anti-Tumor Activity *in Vitro*

### 3D Spheroid Killing – CD4 T cells



### Serial Killing Assay – CD8 & CD4 T cells



Engagement of CD4 T cells may enhance depth and durability of anti-tumor response and clinical outcome of TCR-T in solid cancer patients





**ACTengine® IMA201 and IMA204**  
**– TCR-T Targeting MAGEA4/8 and COL6A3**

# ACTengine® IMA201 Targeting MAGEA4/8

## Key Features

### TARGET

HLA-A\*02-presented peptide derived from **MAGEA4 and/or MAGEA/8**

**>5-fold higher** peptide copy number per tumor cell than a commonly used MAGEA4 target

Naturally and specifically presented on tumors at high target density<sup>1</sup>:  
**100-1,000 copies/cell**

Identified and validated by XPRESIDENT® quant. mass spectrometry platform

### TCR

High-affinity, specific TCR targeting MAGE4/8

High functional avidity<sup>2</sup>:  
**EC50 ~10 ng/ml**

Identified and characterized by XCEPTOR® TCR discovery and engineering platform

### CLINICAL DATA

Dose escalation ongoing, target dose level to commence

Too early for assessment of safety or anti-tumor activity

### PATIENT POPULATION<sup>3</sup>

Sarcoma Subtypes – up to 80%  
Squamous NSCLC – 50%  
HNSCC – 35%  
Bladder Carcinoma – 30%  
Esophageal Carcinoma – 25%  
Uterine Carcinosarcoma – 25%  
Ovarian Carcinoma – 20%  
Melanoma – 20%

Status – 02-June-2022

# ACTengine® IMA204 First-in-Class TCR-T Targeting Tumor Stroma

## Key Features

### TARGET

HLA-A\*02-presented peptide derived from **COL6A3 exon 6**

Naturally and specifically presented on tumors at high target density<sup>1</sup>:  
**100-700 copies/cell**

Novel **tumor stroma target** identified and validated by XPRESIDENT® quant. mass spectrometry platform

### TCR

High-affinity, specific TCR targeting COL6A3 exon 6

**Affinity-maturated, CD8-independent TCR**

High functional avidity<sup>2</sup>:  
**~0.01ng/ml**

Identified and characterized by XCEPTOR® TCR discovery and engineering platform

### PRECLINICAL DATA

CD8-independent, next-generation TCR engages both, CD8 and CD4 T cells

*In vitro* anti-tumor activity against target-positive cell lines in CD8 and CD4 T cells

Complete tumor eradication in *in vivo* mouse models

### PATIENT POPULATION<sup>3</sup>

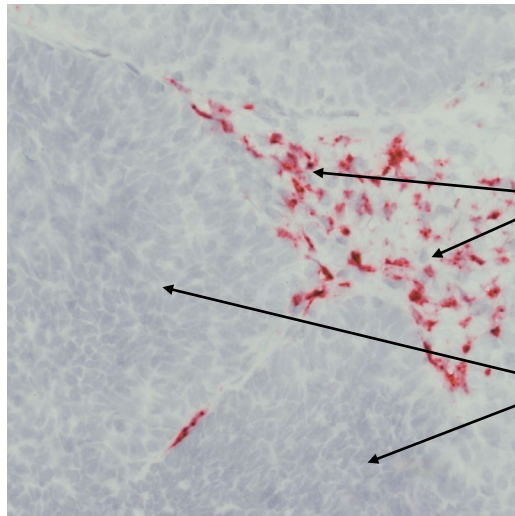
Pancreatic Carcinoma – 80%  
Breast Carcinoma – 75%  
Stomach Carcinoma – 65%  
Sarcoma – 65%  
Esophageal Carcinoma – 60%  
Squamous NSCLC – 55%  
Adeno NSCLC – 55%  
HNSCC – 55%  
Uterine Carcinosarcoma – 55%  
Colorectal Carcinoma – 45%  
Mesothelioma – 45%  
Cholangiocarcinoma – 40%  
Ovarian Carcinoma – 40%  
Melanoma – 35%  
Bladder Carcinoma – 35%

IMA204 provides a promising therapeutic opportunity for a broad patient population as monotherapy or in combination with TCR-T cells directed against tumor targets

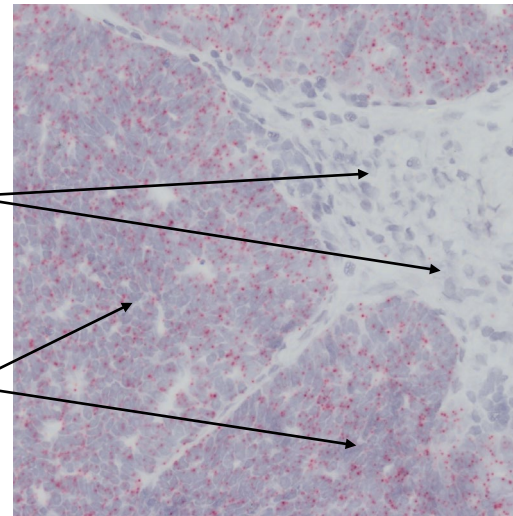
# ACTengine® IMA204 – High Affinity, CD8-independent TCR

Complete Tumor Eradication *in vitro* & *in vivo*<sup>1</sup> by Affinity-enhanced IMA204 TCR

Stroma Target (COL6A3 exon 6)  
in Ovarian Cancer sample



Example of a Tumor Target  
in same Ovarian Cancer sample

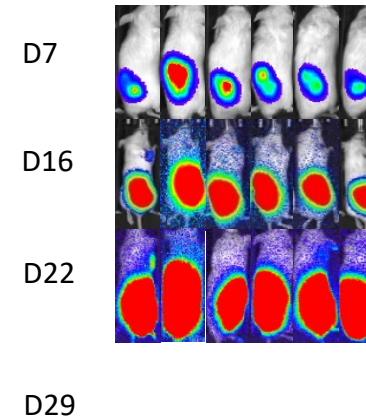


Stroma  
cells

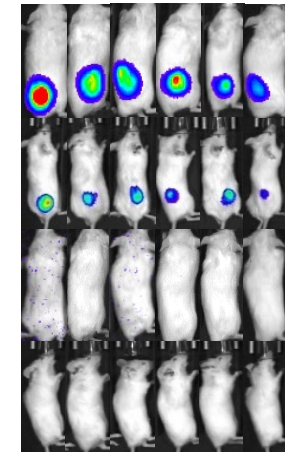
Tumor  
cells

COL6A3 exon 6 prevalently expressed at high target density  
in tumor stroma across many solid cancers

Control



IMA204 TCR



CD8-independent TCR leads to tumor eradication  
in all mice treated

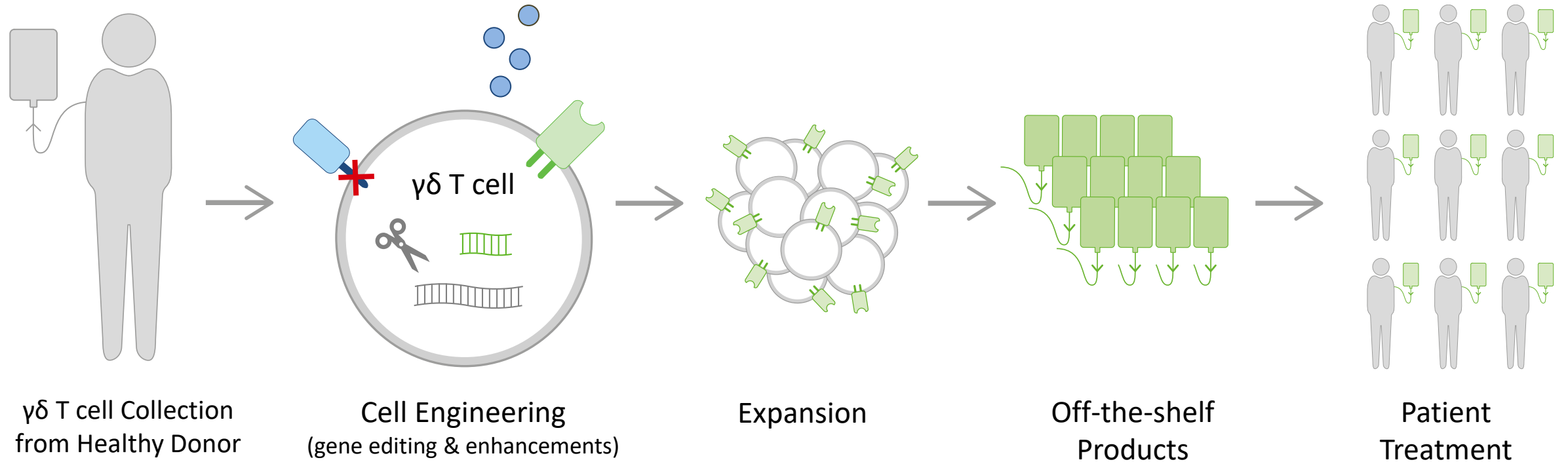
- Affinity matured CD8-independent, next-generation TCR engages both CD4 and CD8 T cells without the need of CD8 co-transduction
- IND-enabling studies are nearing completion





**ACTallo® – Our Next-generation Off-the-shelf TCR-T**

# ACTallo® – Immatics' Allogeneic Cell Therapy Approach



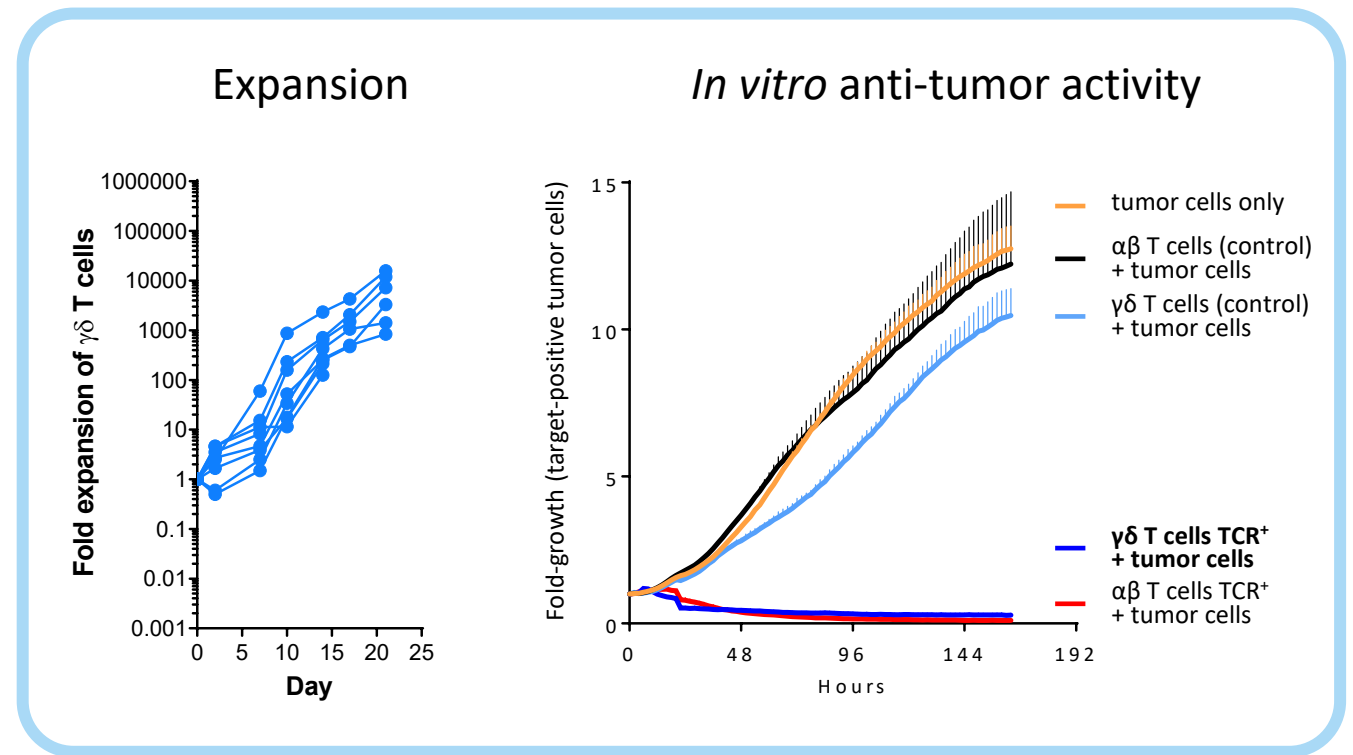
- **Off-the-shelf cell therapy**, no need for personalized manufacturing → reduced logistics and time to application
- **Potential for hundreds of doses** from one single donor leukapheresis → lower cost of goods
- **Use of healthy donor material** provides standardized quality and quantity of starting material

# Why $\gamma\delta$ T cells?

## $\gamma\delta$ T cells Are Well Suited for an Off-the-shelf Cell Therapy Approach

### $\gamma\delta$ T cells

- ✓ are abundant in the peripheral blood
- ✓ show intrinsic anti-tumor activity
- ✓ naturally infiltrate solid tumors & correlate with favorable prognosis
- ✓ are HLA-independent, thus do not cause graft-vs-host disease in allogeneic setting
- ✓ can be expanded to high numbers in a cGMP-compatible manner
- ✓ can be effectively redirected using  $\alpha\beta$  TCR or CAR constructs



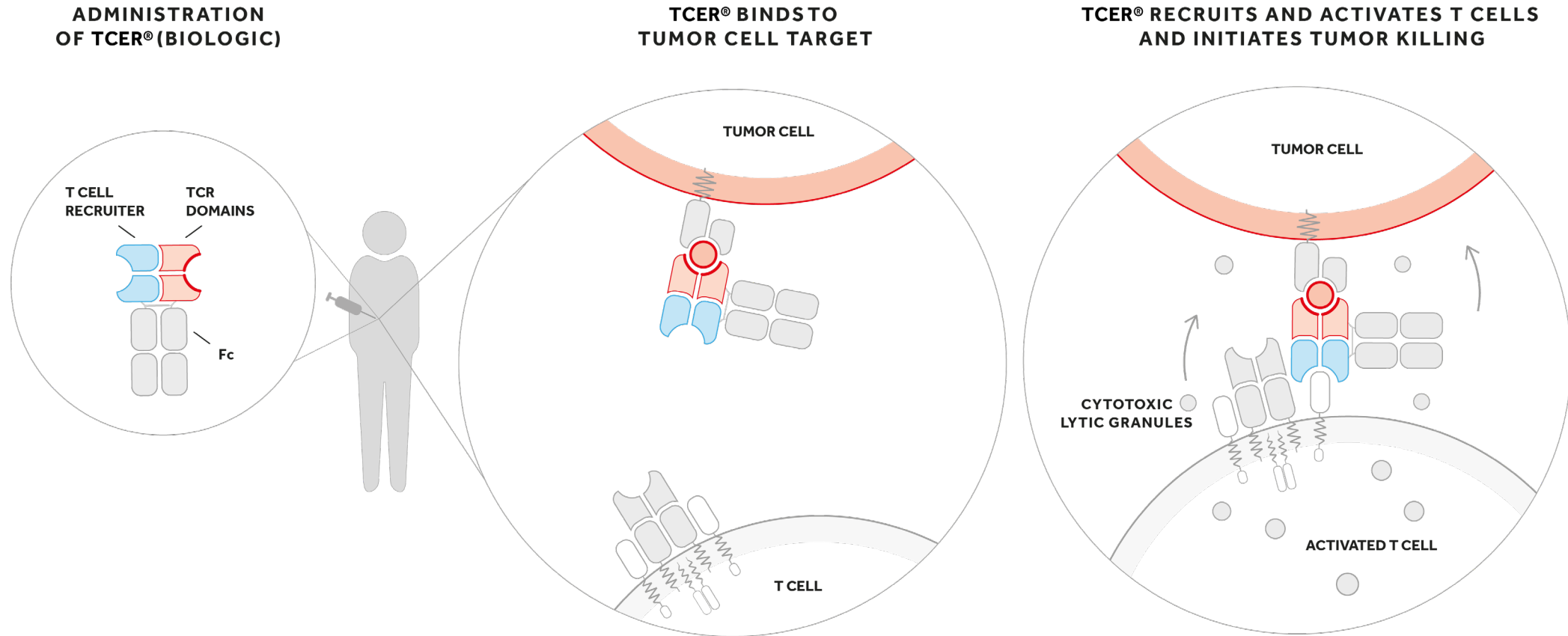


## TCER® – TCR Bispecifics

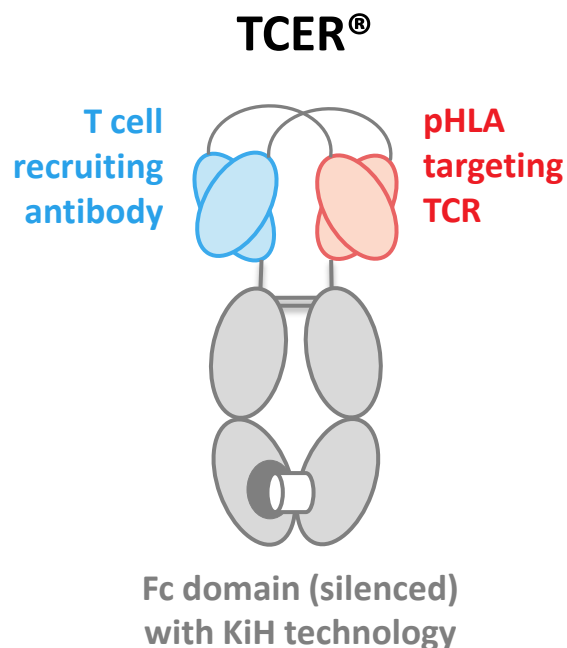


# TCER® – Mechanism of Action

## Immatics' Off-the-Shelf TCR Bispecifics Approach



# TCER® – Immatics' Half-Life Extended Bispecifics



## pHLA targeting TCR

- ✓ **High-affinity TCR** targeting HLA-restricted tumor-specific peptides
- ✓ Broad therapeutic window through **XPRESIDENT®-guided** affinity maturation (>1000x)<sup>1</sup>
- ✓ **Complete tumor eradication** in mouse xenograft models at low doses

## T cell recruiting antibody

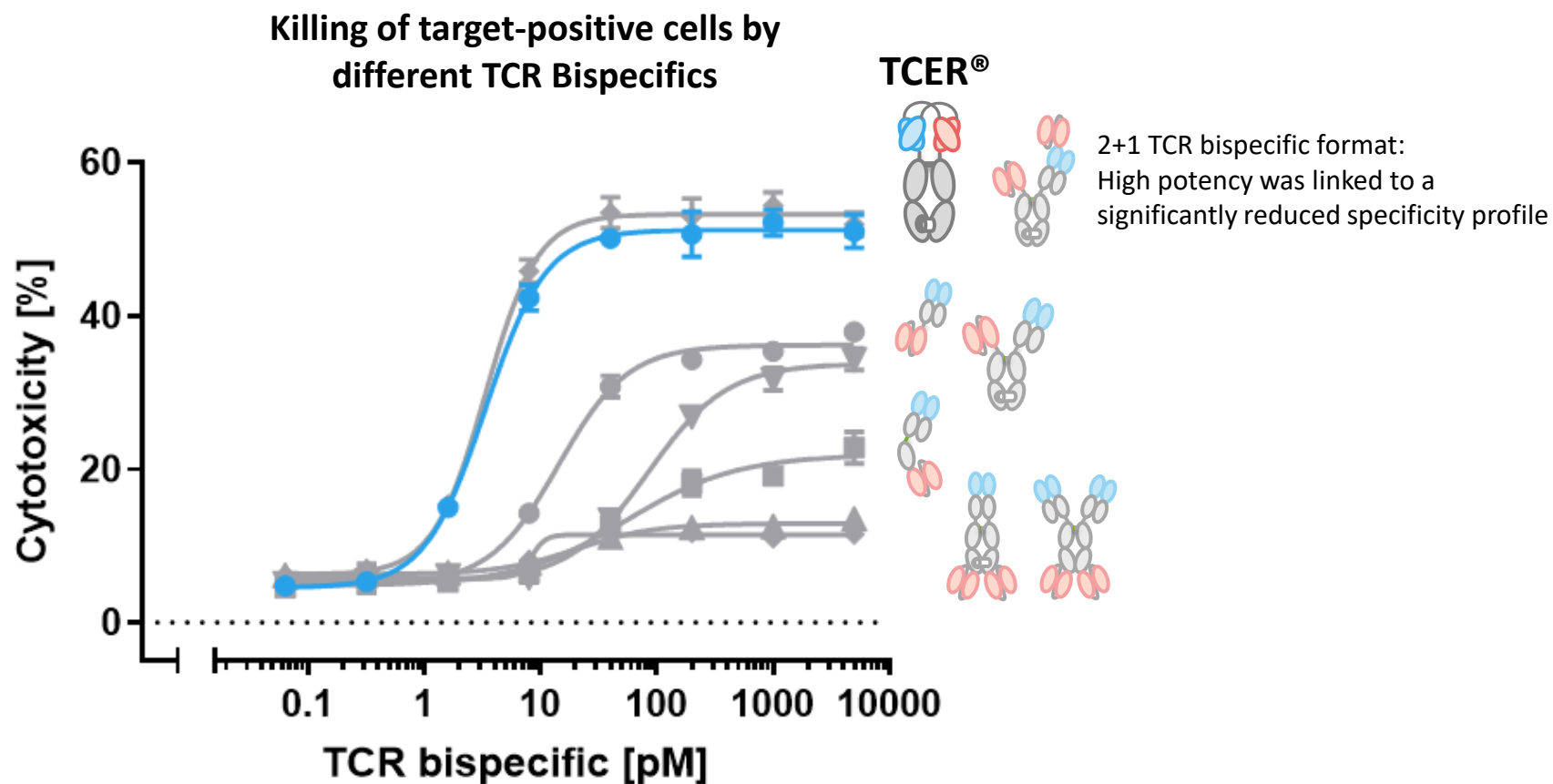
- ✓ **Low-affinity** T cell recruiter against both **TCR & CD3**
- ✓ **Optimized biodistribution** aiming for enrichment at tumor site and **prevention of CRS**<sup>2</sup>
- ✓ **Superior anti-tumor activity** in mouse models as compared to widely used CD3 recruiters

## Next-generation TCER® format

- ✓ Off-the-shelf biologic with antibody-like manufacturability<sup>3</sup> and low cost of goods
- ✓ Superior anti-tumor activity<sup>4</sup> compared to six alternative bispecific formats
- ✓ Half-life of several days expected in humans

**Our TCER® format is designed to maximize efficacy while minimizing toxicities in patients**

# Potency of Our Proprietary TCR Bispecific Format TCER®



- Seven different TCR Bispecific formats were evaluated with a pHLA targeting TCR and the identical T cell recruiting antibody
- TCER® format had higher combination of potency and specificity<sup>1</sup> than six alternative TCR Bispecific format designs evaluated

**Flexible Plug-and-play platform: TCER® format successfully validated for different TCRs & different T cell recruiting antibodies**

## Building a Pipeline of Next-Gen Half-Life Extended TCR Bispecifics

	IMA401	IMA402	IMA40X
	MAGEA4/8	PRAME	Undisclosed
Status	Start of Phase 1 trial in May 2022	Clinical GMP batch targeted in 2022 Phase 1 trial in 2023	TCER® engineering and preclinical testing ongoing
Preclinical Proof-of-concept – Efficacy / Safety	<ul style="list-style-type: none"> <li>➤ Complete remission of estab. tumors in xenograft mouse models at low doses</li> <li>➤ Very broad therapeutic window (reactivity tumor compared to normal cells)</li> </ul>		n/a
Half-life	Half-life extended to several days via effector function silenced Fc part		
Clinical Development Strategy	<ul style="list-style-type: none"> <li>➤ First-in-human basket trial</li> <li>➤ Adaptive design aiming at fast dose escalation</li> <li>➤ Development strategy includes TCER® as add on to checkpoint inhibitor-based standard of care in early lines of treatment</li> </ul>		

# Phase 1 Clinical Trial to Evaluate TCER<sup>®</sup> IMA401 Targeting MAGEA4/8

## Trial Overview

**Biomarker positive patients** with recurrent and/or refractory solid tumors

- HLA-A\*02:01
- MAGEA4/8 (Immatics' IMADetect<sup>®</sup> test)

**Basket trial** in indications with high MAGEA4/8 prevalence, e.g. sqNSCLC, SCLC, HNSCC, bladder carcinoma, esophageal carcinoma, ovarian carcinoma, melanoma, uterine carcinosarcoma, sarcoma subtypes

**Phase 1a: Dose escalation cohort**

**Phase 1b: Dose expansion cohort(s)**

**Up to N=50 patients**

**Up to 15 centers**

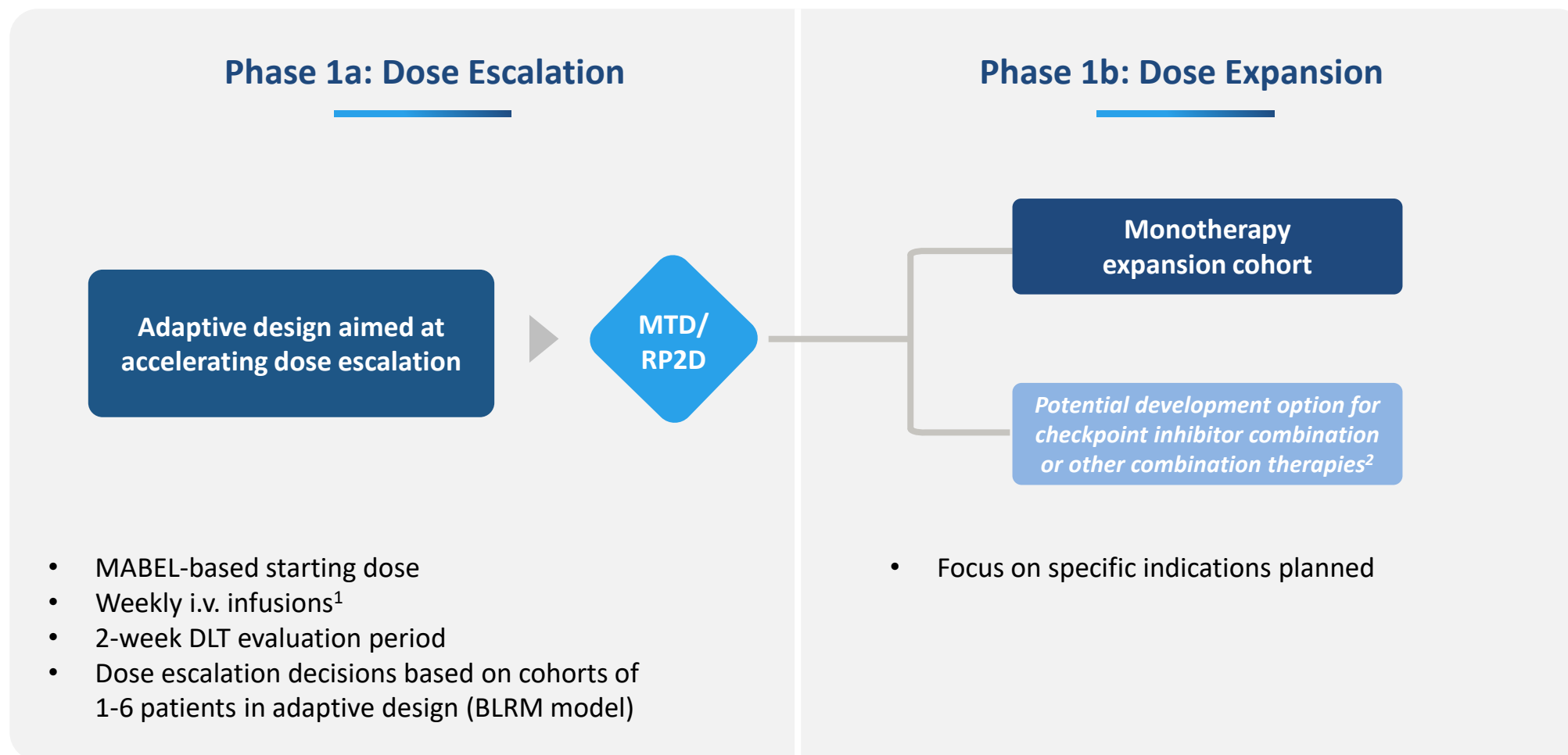
## Primary Objective

- Determine maximum tolerated dose (MTD) and/or recommended phase 2 dose (RP2D)

## Secondary Objectives

- Safety and tolerability
- Initial anti-tumor activity
- Pharmacokinetics

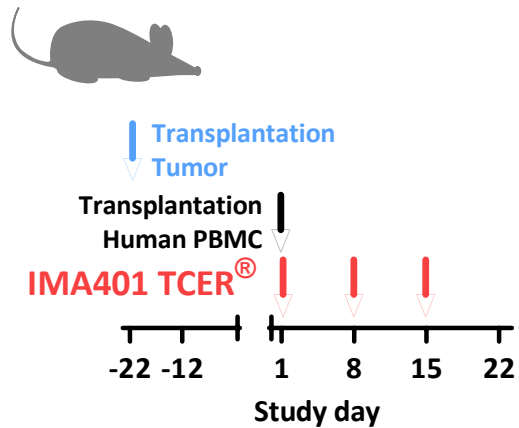
# Phase 1 Clinical Trial to Evaluate TCER<sup>®</sup> IMA401 Targeting MAGEA4/8



# TCER® IMA401 Targeting MAGEA4/8

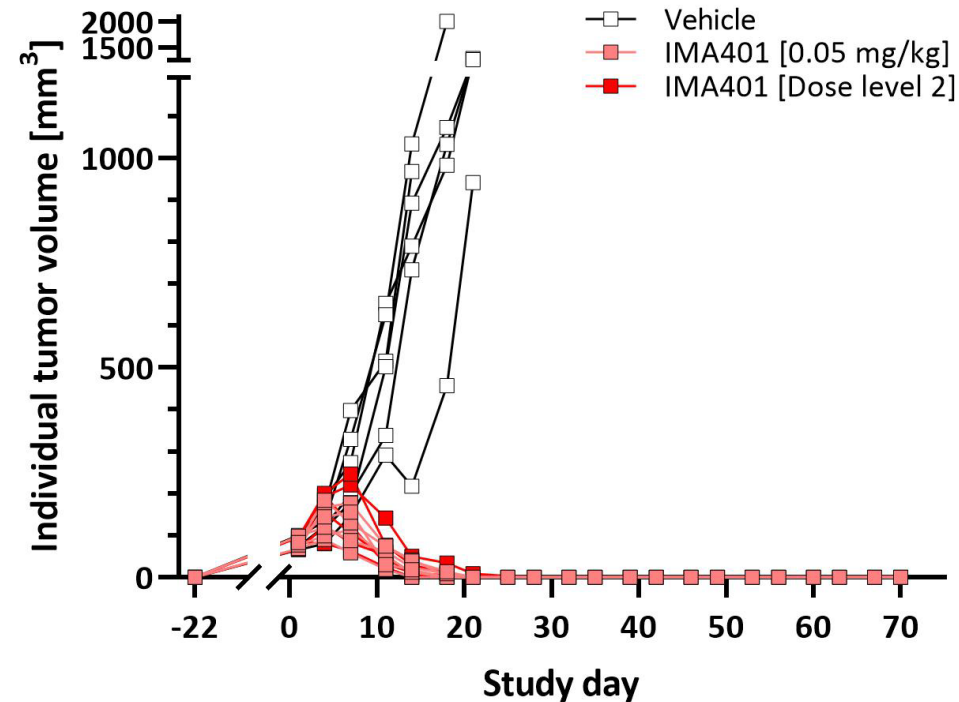
Product Candidate in Clinical Development with Bristol Myers Squibb

## Treatment schedule



N=6 mice per group, two PBMC donors  
Dose: two dose levels

## Tumor Model in Mice<sup>1</sup>



- **Complete remissions observed in all animals** even at low IMA401 dose of 0.05 mg/kg
- No detectable outgrowth of tumors during **prolonged observation period of 70 days**

# TCER® IMA402 Targeting PRAME

Preclinical-stage Product Candidate Fully Owned by Immatics

## PRAME Target Peptide

- **HLA-A\*02-restricted PRAME peptide** targeted by TCER® IMA402 is one of the most frequently expressed intracellular cancer targets for TCR-based therapies
  - Homogenously expressed at high prevalence across multiple solid tumors including melanoma, lung cancer, gynecological cancers (ovarian, breast, uterine) and others

## Preclinical Proof-of-Concept Data

- **High *in vitro* potency** in killing of tumor cells with physiological PRAME peptide levels
- Favorable safety profile with broad therapeutic window between tumor and normal cell reactivity *in vitro*
- **Consistent tumor regression** including complete responses in NOG mice treated at low doses
- **Extended serum half-life** of several days<sup>1</sup> expected in humans driven by the TCER® Fc part

## Well Progressing CMC Development

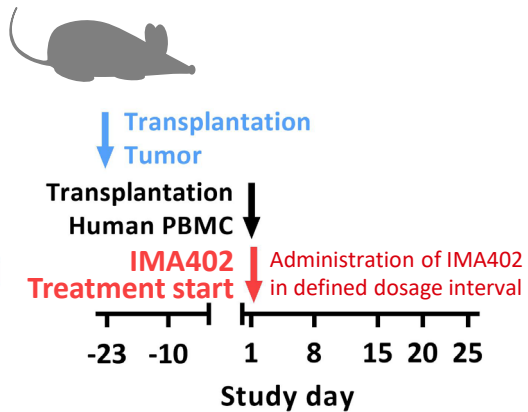
- Current data support antibody-like manufacturability and developability
- GMP process development and IND-enabling activities ongoing
- Manufacturing of the clinical batch for the Phase 1 trial expected in 2H 2022



# TCER® IMA402 – Efficacy Assessment in Tumor Model in Mice

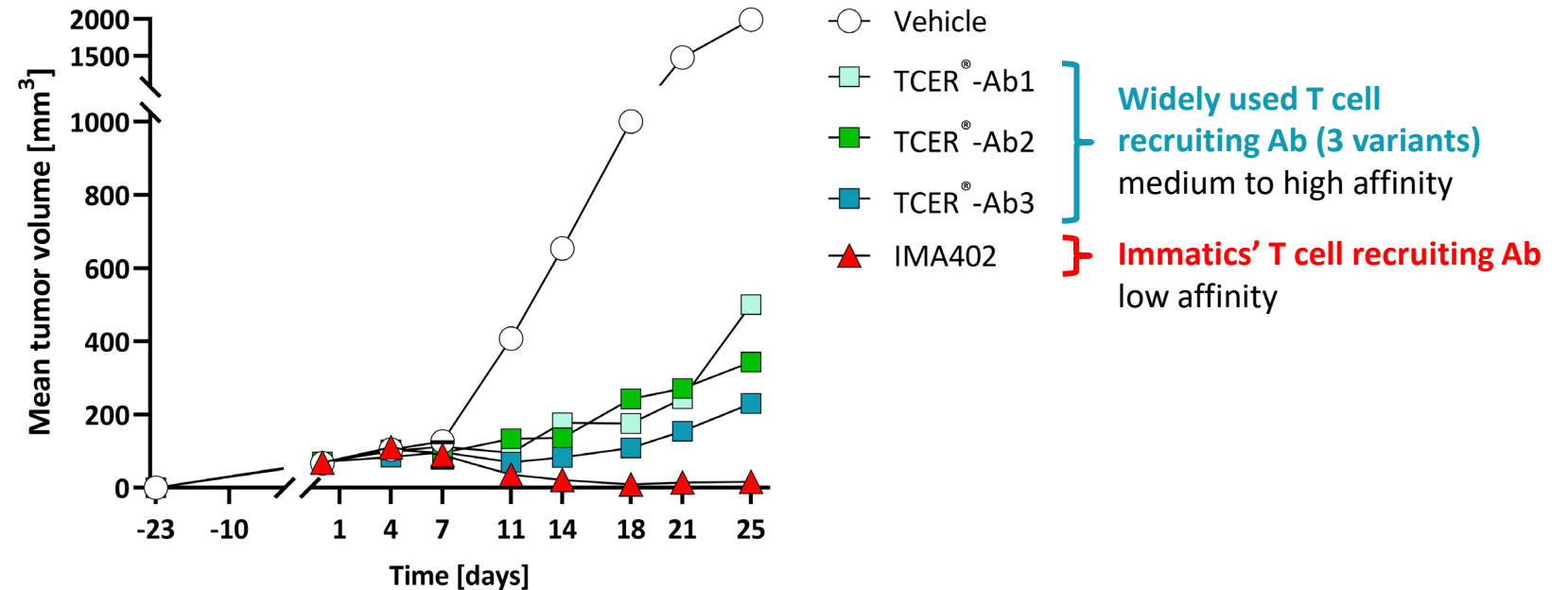
## Superior Tumor Control Using a Proprietary, Low-Affinity Recruiter

### Treatment schedule



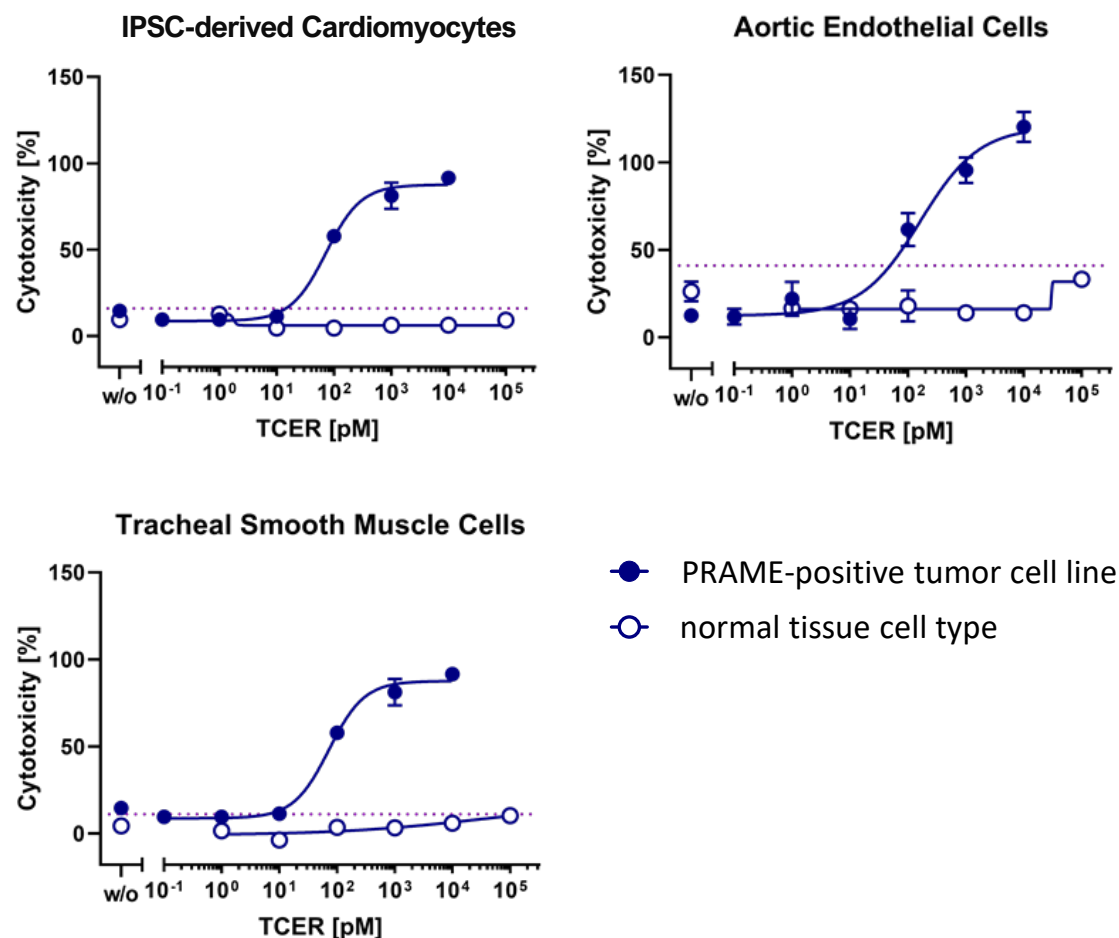
N=6 mice per group, two PBMC donors  
Dose: 0.025 mg/kg

### Tumor Model in Mice<sup>1</sup>



Proprietary, **low-affinity T cell recruiting region** demonstrates superior tumor control compared to analogous TCER® molecules designed with higher-affinity variants of a widely used recruiter

# TCER® IMA402 – *In vitro* Safety Assessment with Normal Tissue Cells



Normal Tissue Type	Therapeutic Window (x-fold)
IPSC-derived astrocytes	≥1,000
IPSC-derived GABA neurons	≥1,000
IPSC-derived cardiomyocytes	≥1,000
Human Pulmonary Fibroblasts	≥1,000
Human Cardiac Microvascular Endothelial Cells	≥1,000
Human Dermal Microvascular Endothelial Cells	≥1,000
Human Aortic Endothelial Cells	≥1,000
Human Coronary Artery Smooth Muscle Cells	≥1,000
Human Tracheal Smooth Muscle Cells	≥1,000

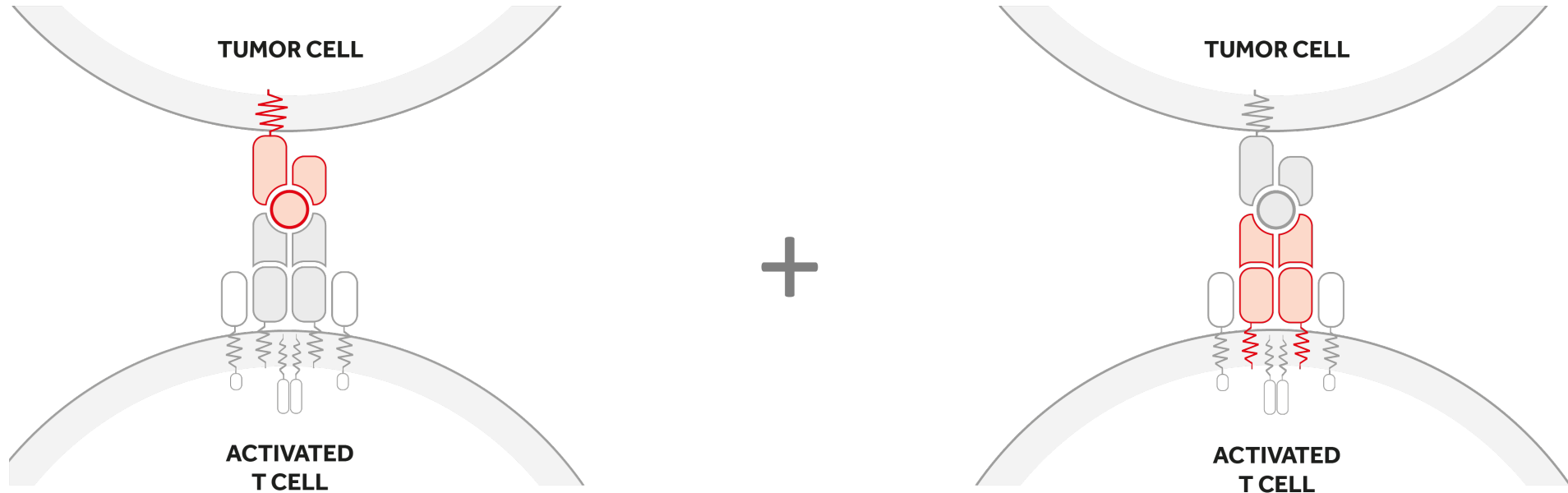
- Cytotoxicity against N≥9 different human normal tissue cell types
- TCER® IMA402 shows a **minimum of 1,000-fold therapeutic window** between normal tissue cell reactivity and tumor cell reactivity



## Immatics' Proprietary Target and TCR Discovery Platforms

# True Cancer Targets & Matching Right TCRs

Goal to Maximize Anti-Tumor Activity and Minimize Safety Risks of TCR-based Immunotherapies



## True Targets via XPRESIDENT® technology platform

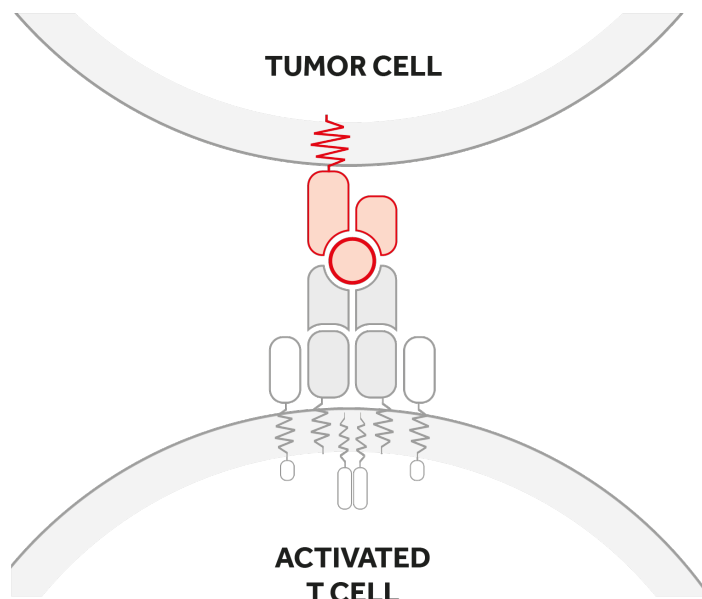
- are naturally presented on tumor tissues as identified by mass-spec
- are absent or presented at only low levels on normal tissues
- are presented at high copy numbers to trigger a pharmacological response

## Right TCRs via XCEPTOR® technology platform

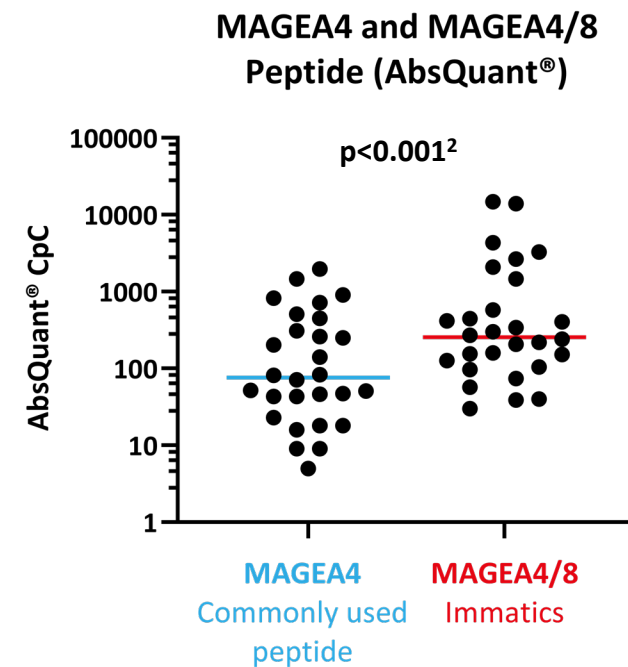
- recognize the target peptide with high affinity and specificity
- show selective killing of tumor cells
- are developed to be suitable for two different therapeutic modalities, Cell Therapies and TCR Bispecifics

# Immatics' Unique Capability – Identification of the most Relevant Target

## Example of MAGEA4/8 Peptide Target



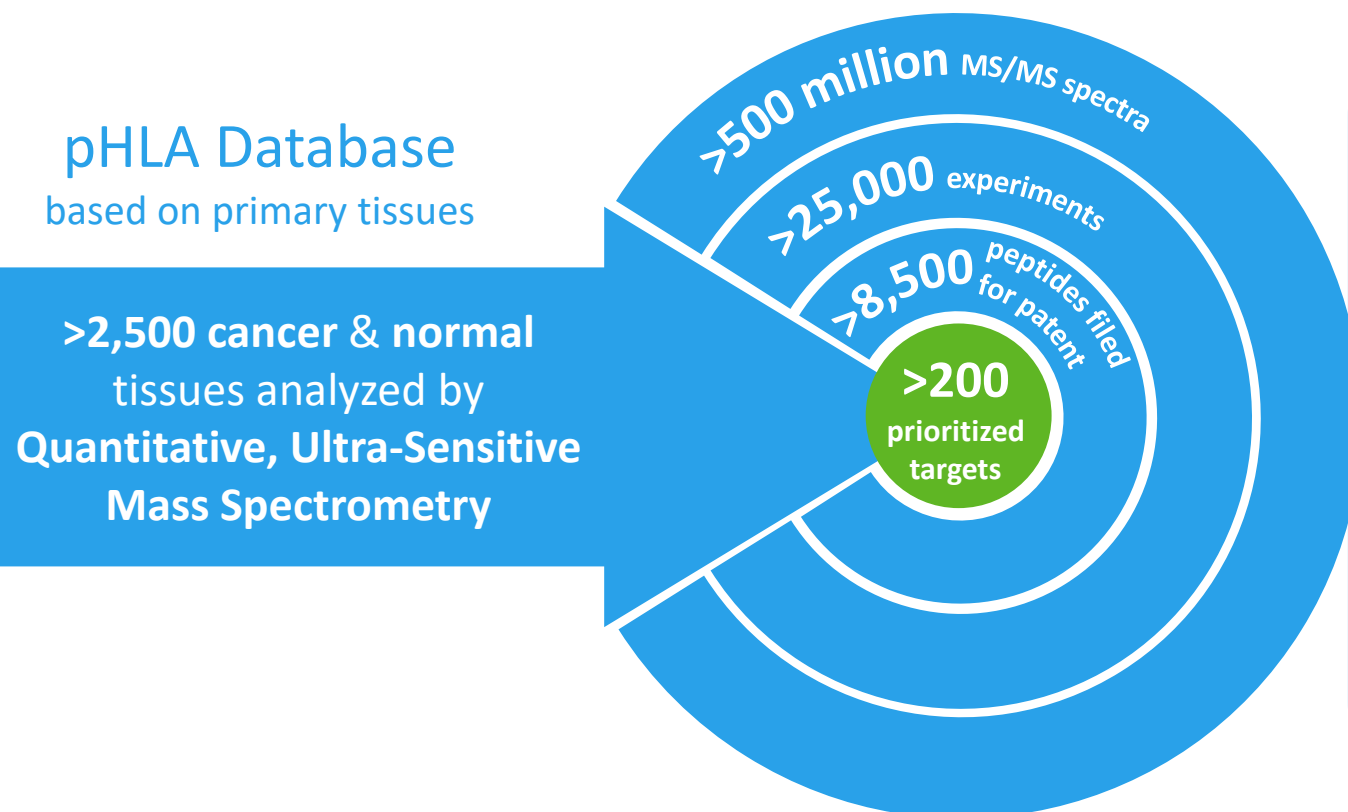
Ranking of  
pHLA targets



XPRESIDENT® quantitative information on target density<sup>1</sup> between peptides originating from the same source protein

MAGEA4/8 target is presented at >5-fold higher target density<sup>1</sup> than a commonly used MAGEA4 target peptide

# Pool of 200 Prioritized Targets as Foundation for Future Value Generation



## 200 Prioritized Targets

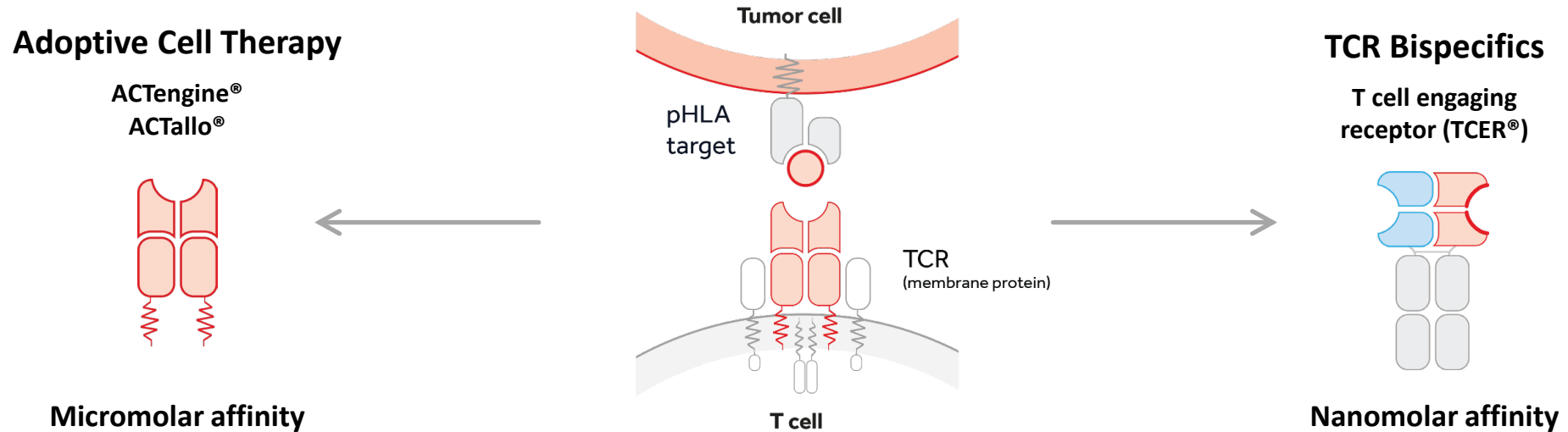
### Grouped in 3 Target Classes:

1. **Well known and characterized parent protein (20%)**  
e.g. MAGE family cancer testis antigens
2. **Unknown or poorly characterized parent protein (60%)**  
e.g. stroma target COL6A3 exon 6
3. **Crypto-targets/Neoantigens (20%)**  
Novel target class which includes RNA-edited peptides & non-classical neoantigens

~50% of our prioritized targets are non-HLA-A\*02 restricted, substantially broadening the potential patient reach

# Development of the Right TCR – XCEPTOR® Technology

## TCR Discovery and Engineering for ACT and TCR Bispecifics

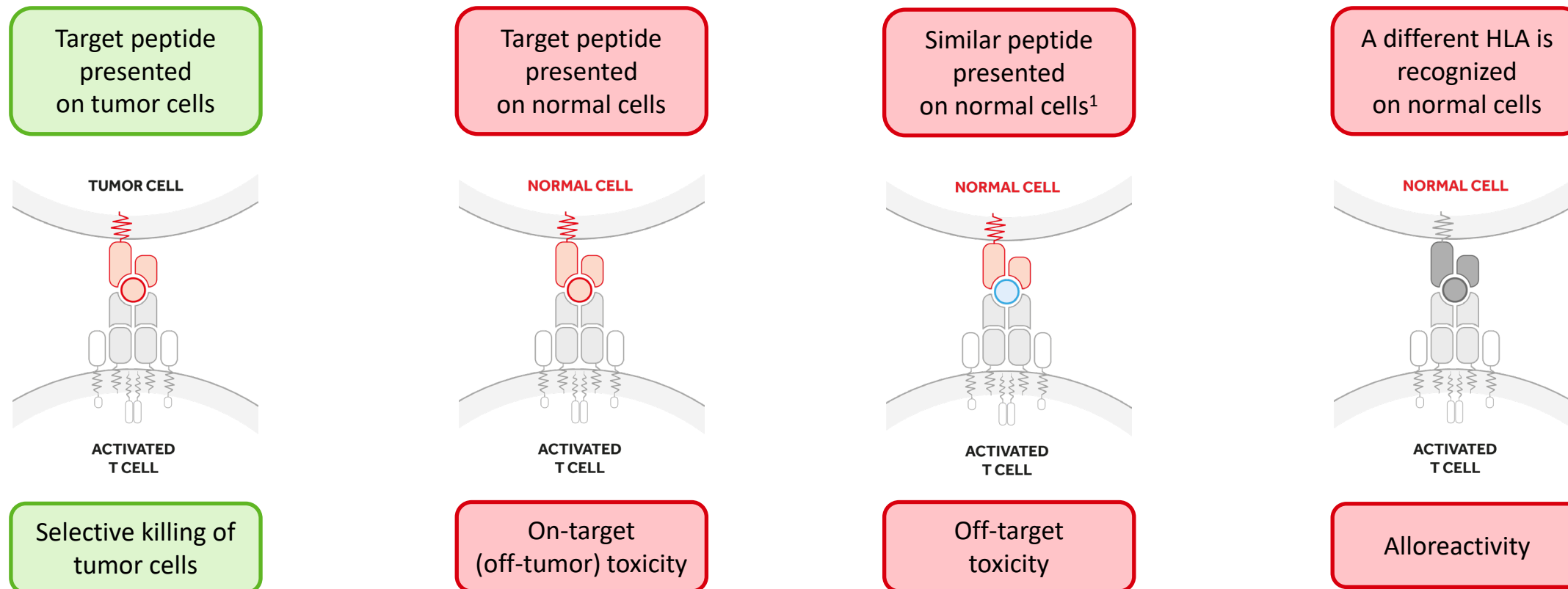


- Fast, efficient and highly sensitive discovery of highly specific, natural TCRs
- Protein engineering capabilities to design and mature TCRs with increased affinity while retaining specificity
- Early de-selection of cross-reactive TCRs by the unique interplay between Immatics' target and TCR discovery platforms XPRESIDENT® and XCEPTOR® during TCR discovery<sup>1</sup> and TCR maturation<sup>2</sup>



# Optimal Target Selection & TCR Specificity for Minimizing Safety Risks

Unique Interplay between Technology Platforms Allows Early De-risking for Clinical Development



**XPRESIDENT®-guided screening** for on- and off-target toxicities of TCRs based on the extensive database of peptides presented on normal tissues



## Corporate Information & Milestones

# Experienced Global Leadership Team Across Europe and the US



**Harpreet Singh**  
Chief Executive Officer  
Co-Founder  
>20 yrs biotech experience



**Arnd Christ**  
Chief Financial Officer  
>20 yrs biotech experience  
(Probiobrug, NovImmune, Medigene, InflaRx)



**Cedrik Britten**  
Chief Medical Officer  
>10 yrs pharma & biotech experience  
(BioNTech, GSK)



**Carsten Reinhardt**  
Chief Development Officer  
>20 yrs pharma & biotech experience  
(Micromet, Roche, Fresenius)



**Steffen Walter**  
Chief Technology Officer  
Co-Founder Immatics US  
>15 yrs biotech experience



**Toni Weinschenk**  
Chief Innovation Officer  
Co-Founder  
>15 yrs biotech experience



**Rainer Kramer**  
Chief Business Officer  
25 yrs pharma & biotech experience  
(Amgen, MorphoSys, Jerini, Shire, Signature Dx)



**Edward Sturchio**  
General Counsel  
>15 yrs pharma & biotech experience  
(Schering, Merck, Novartis, Advanced Accelerator Applications, Abeona Therapeutics)



**Jordan Silverstein**  
Head of Strategy  
>10 yrs biotech experience  
(Advanced Accelerator Applications, InflaRx)



# Strong, Focused and Highly Integrated Trans-Atlantic Organization



**Tübingen, Germany, ~195 FTEs**  
Target & TCR discovery and  
TCR Bispecifics development



**Houston, Texas, ~140 FTEs**  
Cell therapy development  
and manufacturing



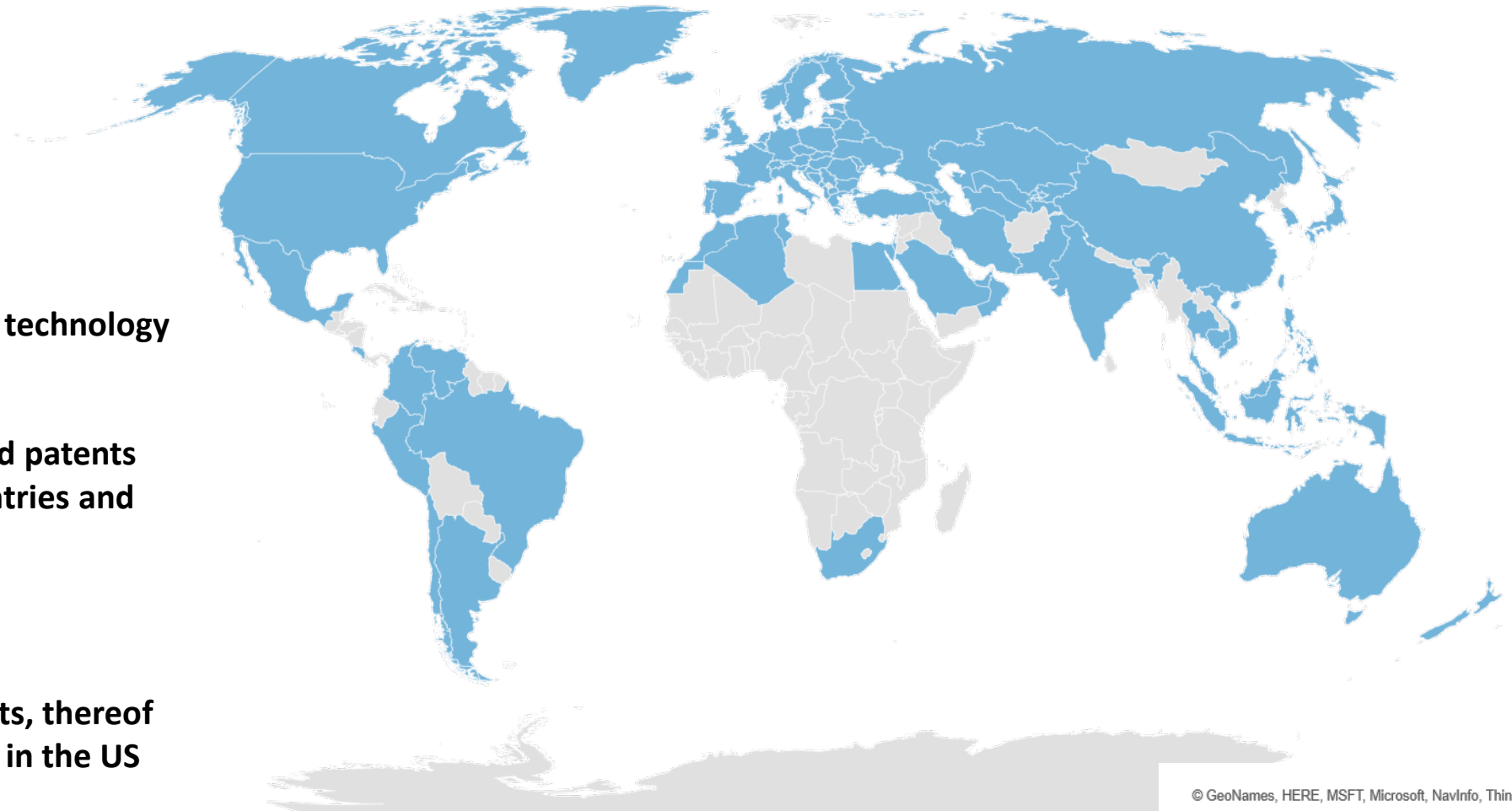
**Munich, Germany, ~55 FTEs**  
Various operating functions

# Robust IP Portfolio

## Immatics' Patent Estate – Territorial Coverage

**Cancer targets, TCRs and technology protected by:**

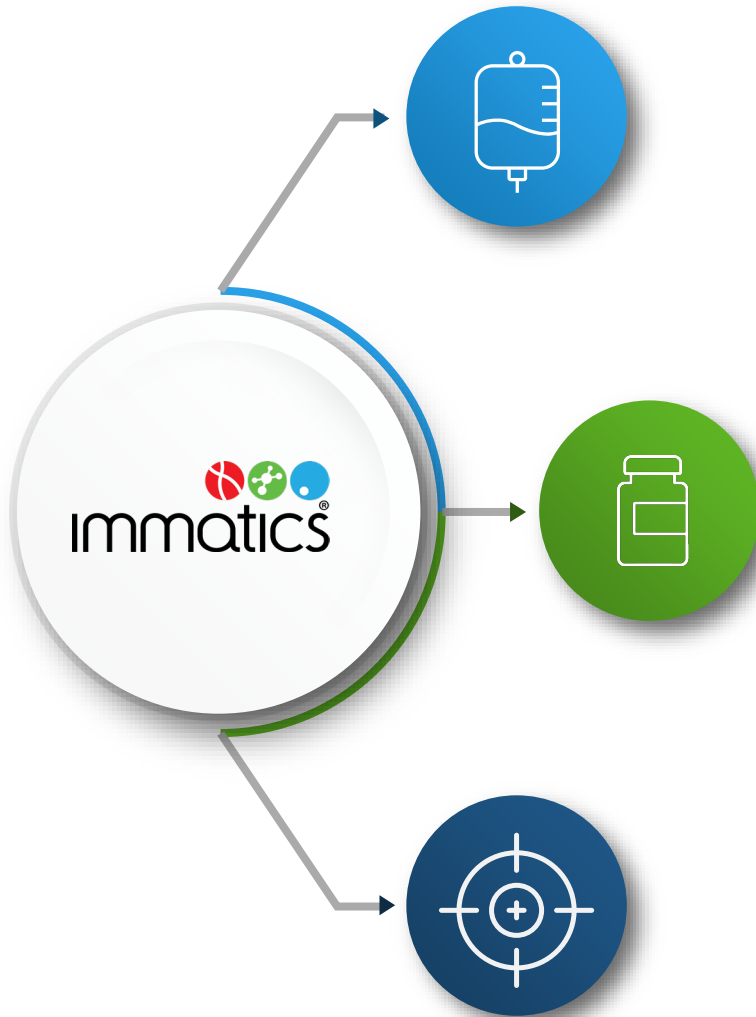
- 5,800 applications and patents filed in all major countries and regions
- >115 patent families
- >2,000 granted patents, thereof >500 granted patents in the US



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# Near-Term Value Drivers and Development Milestones

## Clinical Expansion of TCR Bispecifics and the Next-generation of TCR-T



### Advance clinical development of ACTengine® candidates

- Multiple IMA203 Ph1b expansion cohorts:  
Monotherapy, checkpoint combination, 2nd-gen approach IMA203CD8
- Next IMA203 monotherapy data read-out in 2H 2022
- Initial data read-out for checkpoint combination, IMA203CD8 YE 2022
- Advance IMA204 to the clinic, submission of IND application YE 2022

### Further clinical development of TCER® candidates

- Start of Ph1 trial for IMA401 (MAGEA4/8) in May 2022
- Manufacturing of IMA402 clinical batch in 2H 2022, clinical trial in 2023
- Innovative TCER® program(s) IMA40X in preclinical development

### Leverage full potential of targeting PRAME

- Focused & accelerated development of IMA203 expansion cohorts
- Develop IMA402, an off-the-shelf TCER®



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CANCER PATIENTS

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