

# Immatics Corporate Presentation

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March 21, 2024



*Delivering the Power of T cells to Cancer Patients*

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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

Anti-tumor activity and durability of response across multiple solid tumors in early TCR-T clinical development



## Differentiated Platforms

Unique technologies to identify true cancer targets and right TCRs

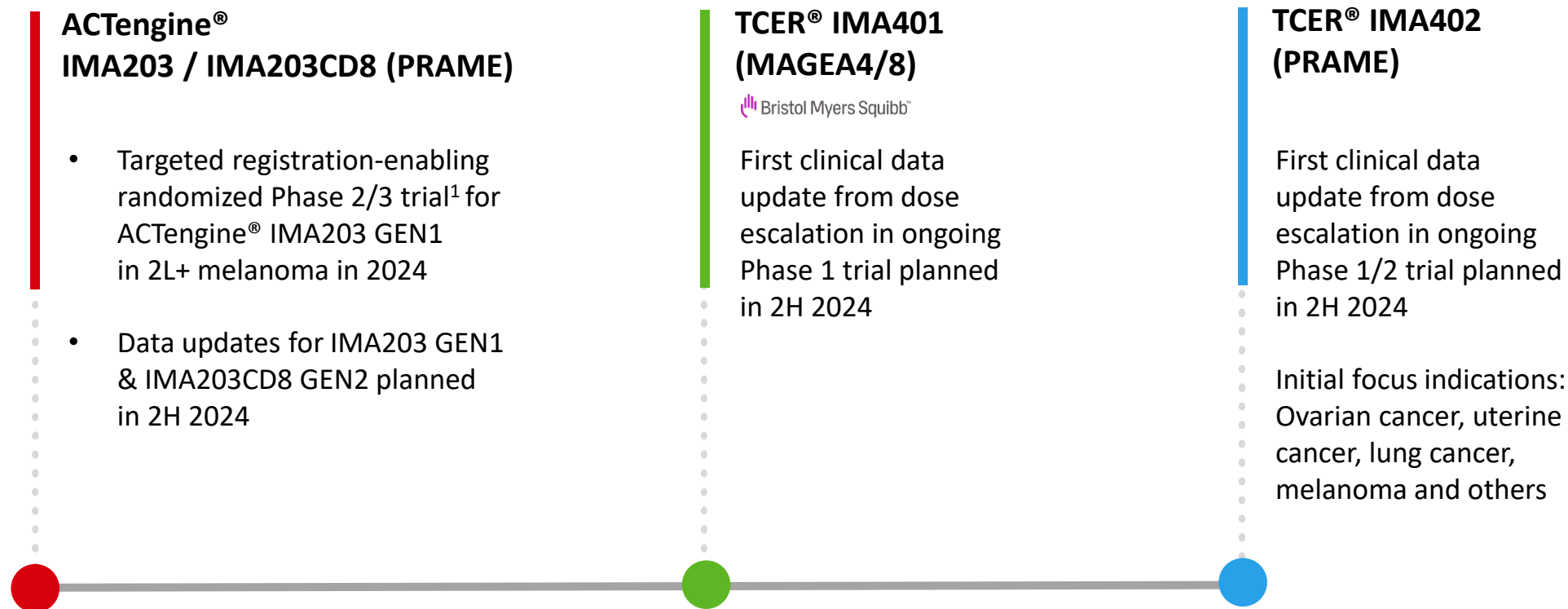


## Therapeutic Opportunity

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

# Upcoming 2024 Catalysts for ACTengine® and TCER® Clinical Lead Assets

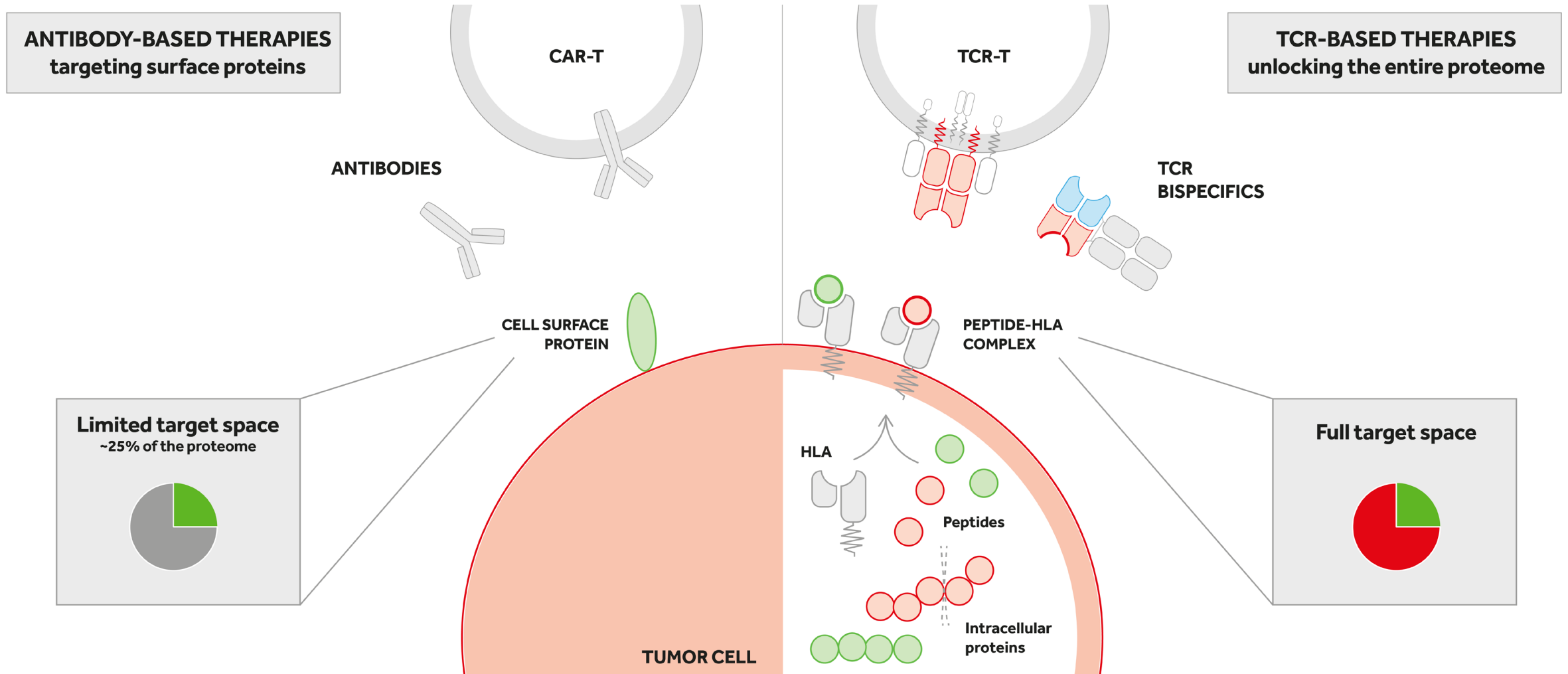
Projected Cash Runway into 2027 to Reach Multiple Value Inflections Points



Updates planned across the entire clinical portfolio throughout 2024



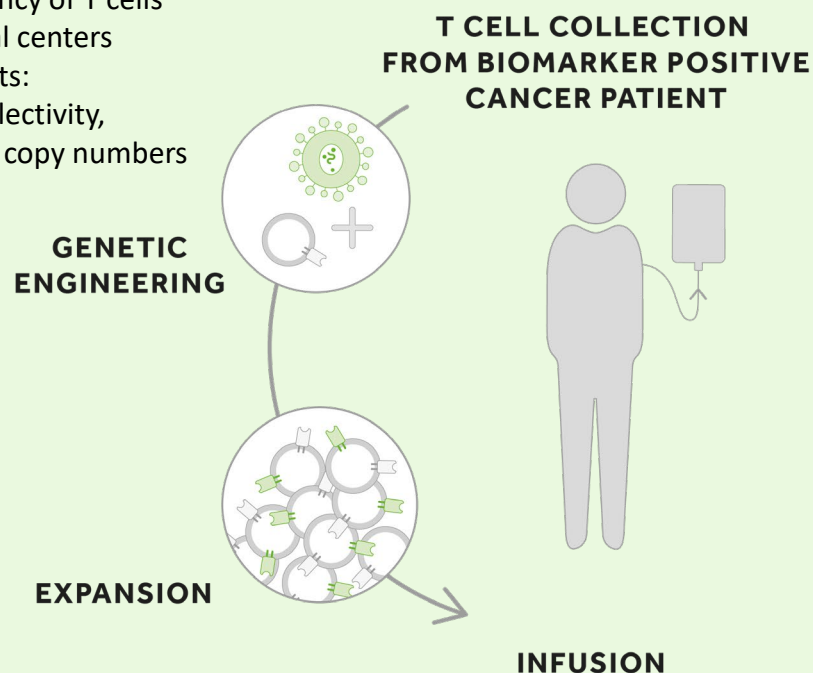
# Our TCR-based Approaches Leverage the Full Target Space beyond the Cancer Cell Surface



# Two Distinct TCR-based Therapeutic Modalities in Clinical Development

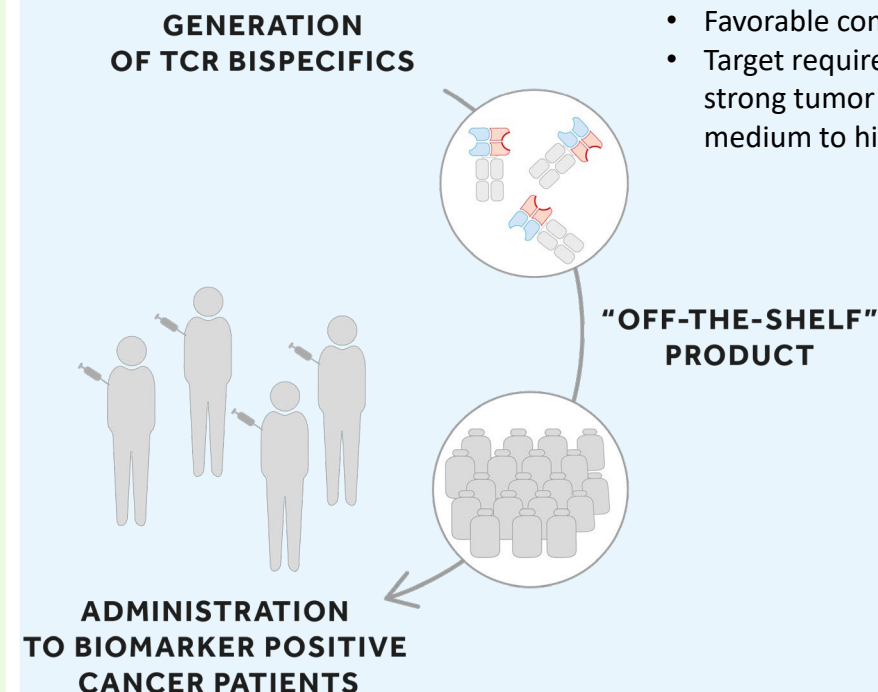
## Autologous TCR-T (ACTengine®)

- 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
- Target requirements: stringent tumor selectivity, low, medium, high copy numbers














## 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
- Target requirements: strong tumor association, medium to high copy numbers



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

# 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	Phase 3
Autologous ACT	ACTengine® IMA203	PRAME		<div></div>	<div></div>	<div></div>		
	ACTengine® IMA203CD8	PRAME		<div></div>	<div></div>	<div></div>		
	ACTengine® IMA204	COL6A3		<div></div>				
	Multiple programs	Undisclosed		<div></div>				
Allogeneic ACT γδ T cells	ACTallo® IMA30x	Undisclosed	  <sup>2</sup>	<div></div>				
	Multiple programs	Undisclosed		<div></div>				
Bispecifics	TCER® IMA401	MAGEA4/8		<div></div>	<div></div>			
	TCER® IMA402	PRAME		<div></div>	<div></div>			
	TCER® IMA40x	Undisclosed		<div></div>				
	Multiple programs <sup>3</sup>	Undisclosed		<div></div>				

# Potential for Large Patient Populations across Multiple Solid Cancers

## IMA203 / IMA402 PRAME

Uterine Carcinoma – 97%  
Uterine Carcinosarcoma – 100%  
Sarcoma Subtypes – up to 100%  
Cut. Melanoma ≥ 95%  
Uveal Melanoma<sup>1</sup> ≥ 91%  
Ovarian Carcinoma – 84%  
Squamous NSCLC – 68%  
TNBC – 63%  
Small Cell Lung Cancer – 45%  
Kidney Carcinoma – up to 40%  
Cholangiocarcinoma – 33%  
HNSCC – 27%  
Esophageal Carcinoma – 27%  
Breast Carcinoma – 26%  
Adeno NSCLC – 25%  
HCC – 18%  
Bladder Carcinoma – 18%

## IMA401 MAGEA4/8

Squamous NSCLC – 52%  
Sarcoma Subtypes – up to 60%  
HNSCC – 36%  
Bladder Carcinoma – 29%  
Uterine Carcinosarcoma – 29%  
Esophageal Carcinoma – 23%  
Ovarian Carcinoma – 23%  
Melanoma – 18%

## IMA204 COL6A3 Exon 6

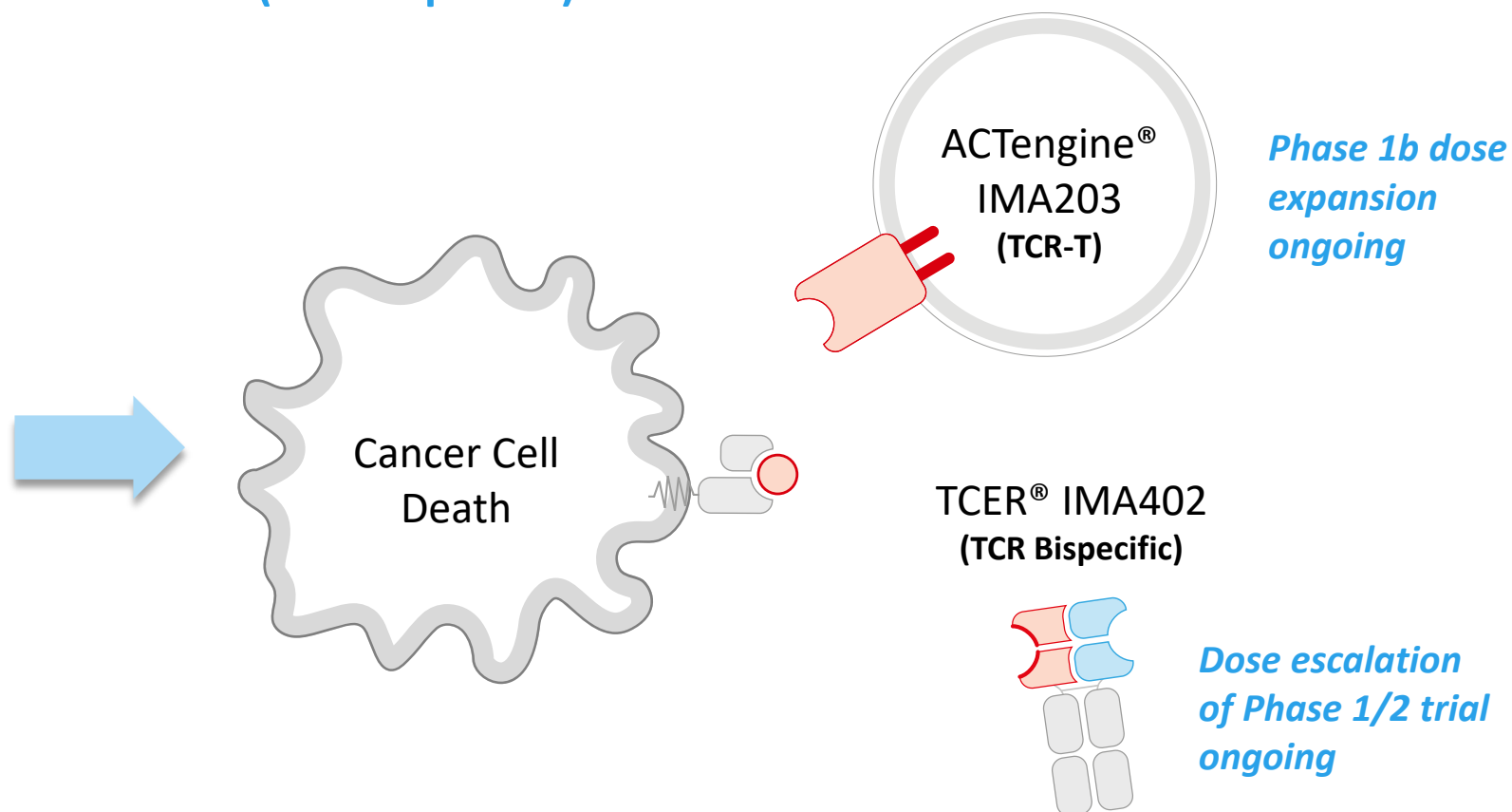
Pancreatic Carcinoma – 76%  
Breast Carcinoma – 77%  
Stomach Carcinoma – 67%  
Sarcoma – 63%  
Colorectal Carcinoma – 60%  
Esophageal Carcinoma – 60%  
Squamous NSCLC – 55%  
Adeno NSCLC – 57%  
HNSCC – 56%  
Uterine Carcinosarcoma – 50%  
Mesothelioma – 44%  
Cholangiocarcinoma – 36%  
Melanoma – 35%  
Bladder Carcinoma – 34%  
Ovarian Carcinoma – 31%

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

# Realizing the Full Multi-Cancer Opportunity of PRAME

## ACTengine® IMA203 (TCR-T) and TCER® IMA402 (TCR Bispecific)

Indication	% PRAME positive patients <sup>1</sup>
Uterine Carcinoma	97%
Uterine Carcinosarcoma	100%
Sarcoma Subtypes	up to 100%
Cut. Melanoma	≥95%
Uveal Melanoma <sup>2</sup>	≥91%
Ovarian Carcinoma	84%
Squamous NSCLC	68%
TNBC	63%
Small Cell Lung Cancer	45%
Kidney Carcinoma	up to 40%
Cholangiocarcinoma	33%
HNSCC	27%
Esophageal Carcinoma	27%
Breast Carcinoma	26%
Adeno NSCLC	25%
HCC	18%
Bladder Carcinoma	18%



PRAME is one of the most promising and most prevalent, clinically validated solid tumor targets known to date

**Leverage the full potential of targeting PRAME by continued evaluation of the best suited therapeutic modality (ACTengine® vs. TCER® or both) for each cancer type**

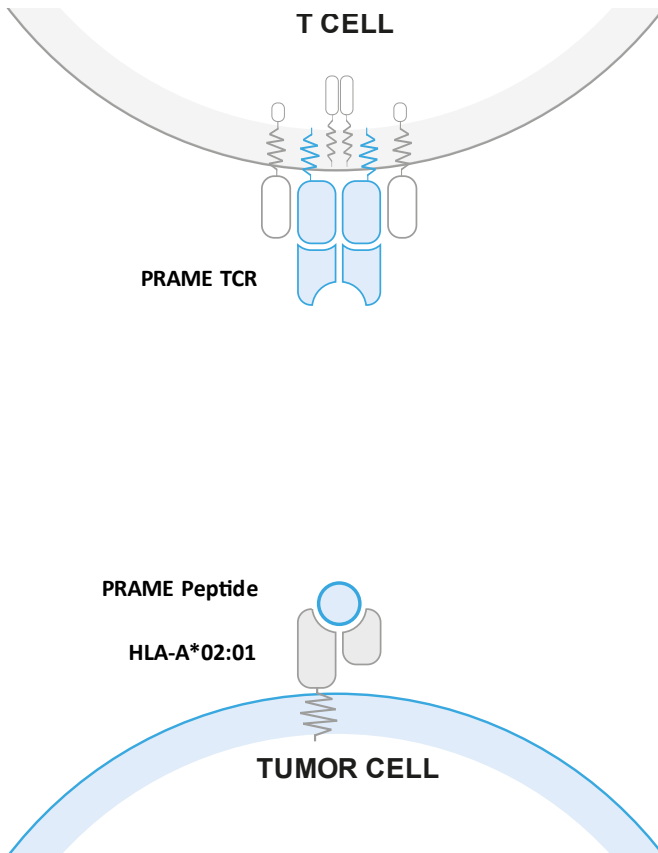




## ACTengine® IMA203 – TCR-T Targeting PRAME

# The Multi-Cancer Opportunity of PRAME

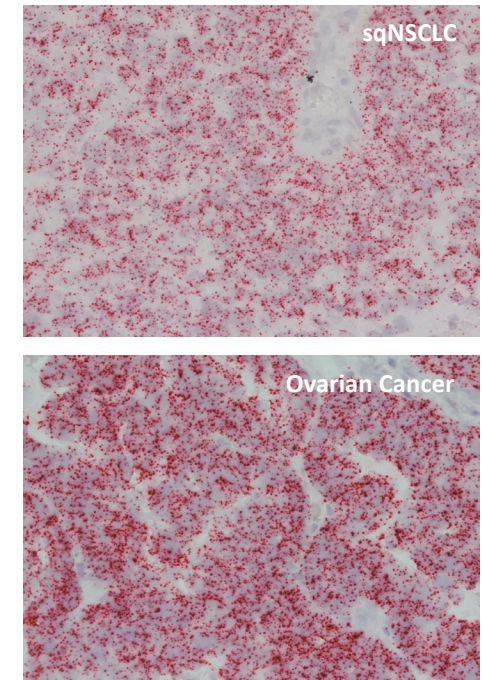
One of the Most Promising Solid Tumor Targets for TCR-based Therapies Known To Date



**PRAME fulfills all properties of an ideal target for TCR-based therapies**

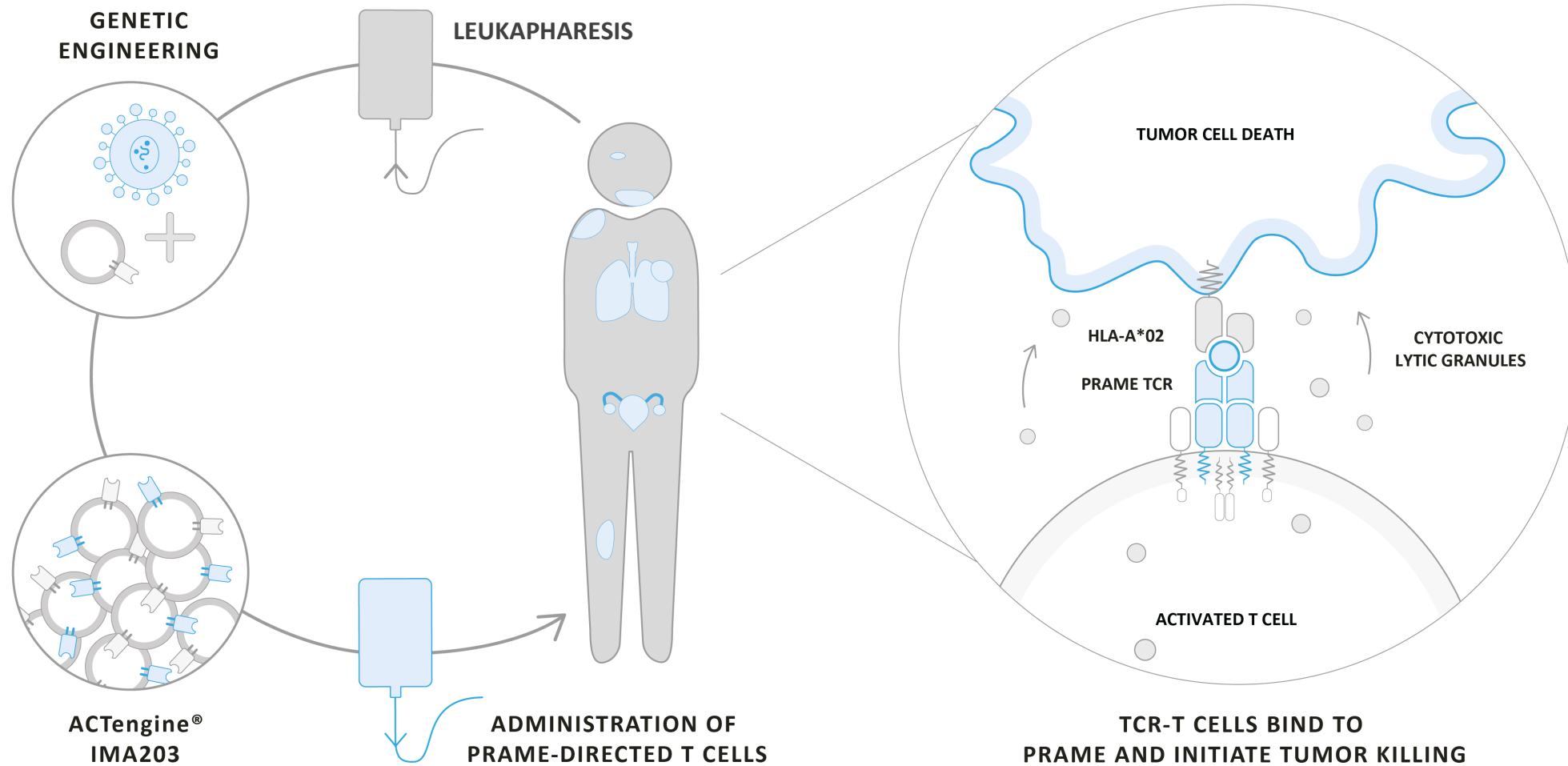
- ✓ High prevalence
- ✓ High target density
- ✓ Homogeneous expression
- ✓ “Clean” expression profile
- ✓ Clinical proof-of-concept

PRAME RNA detection in tumor samples (ISH)



# ACTengine® IMA203 Targeting PRAME – Mechanism of Action

## Immatics' Leading TCR-T Approach



# ACTengine® IMA203/IMA203CD8 TCR-T Monotherapy – Patient Flow

## Screening & Manufacturing Phase

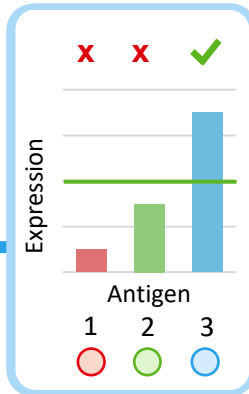
## Treatment & Observation Phase

## Long Term Follow-up

Safety and efficacy monitoring for 12 months

### HLA-A\*02 Testing

Blood sample;  
Central lab



### Target Profiling

IMADetect® mRNA assay using  
Immatics' MS-guided threshold;  
Biopsy or archived tissue

#### Patient screening data from Immatics' clinical trials:

Cut. Melanoma	95% (58/61)
Uveal Melanoma	91% (30/33)
Uterine Carcinoma	89% (8/9)
Ovarian Carcinoma	82% (23/28)

### Leukapheresis

### Manufacturing by Immatics

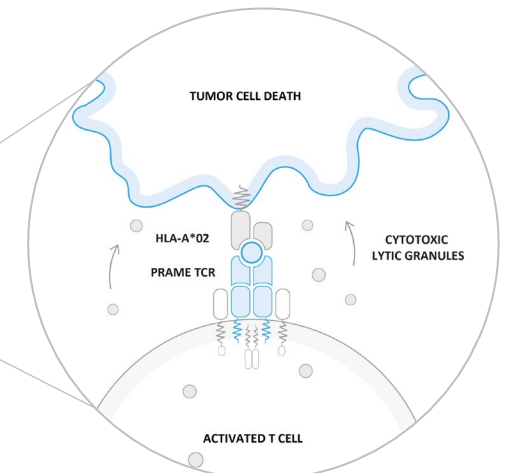
#### Short process time of 14 days

7-day manufacturing process  
applying CD8/CD4 T cell selection  
7-day QC release testing

### Lymphodepletion\*

### Low dose IL-2\*\*

### Infusion of ACTengine® IMA203 TCR-T Product



# IMA203 GEN1 – Melanoma as First Indication for Pivotal Development

Patient Numbers*	ALL	Melanoma	Ovarian Cancer	Synovial Sarcoma	H&N Cancer	Others
Phase 1a RP2D	7	5	0	0	0	2
Cohort A RP2D	18	8	4	3	1	2

Patient characteristics	All comers Cohort A	Melanoma pts Ph1a & Cohort A at RP2D	Ovarian cancer pts Ph1a & Cohort A at RP2D
<b>Efficacy population*</b>	<b>18</b>	<b>13</b>	<b>4</b>
<b>Prior lines of treatment</b>	<b>3</b>	<b>4</b>	<b>4.5</b>
Median (min, max)	(0, 10)	(0, 7)	(3, 10)
<b>LDH at baseline</b>	<b>50.0</b>	<b>53.9</b>	<b>100.0</b>
>1 x ULN [% of patients]			
<b>Baseline tumor burden</b>	<b>58.9</b>	<b>52.0</b>	<b>108.8</b>
Target lesion sum of diameter [mm] (median, min, max)	(21.0, 207.3)	(21.0, 178.7)	(50.6, 207.3)

All 8 cut. melanoma patients were CPI-refractory and 5 of 8 were BRAF-inhibitor pretreated

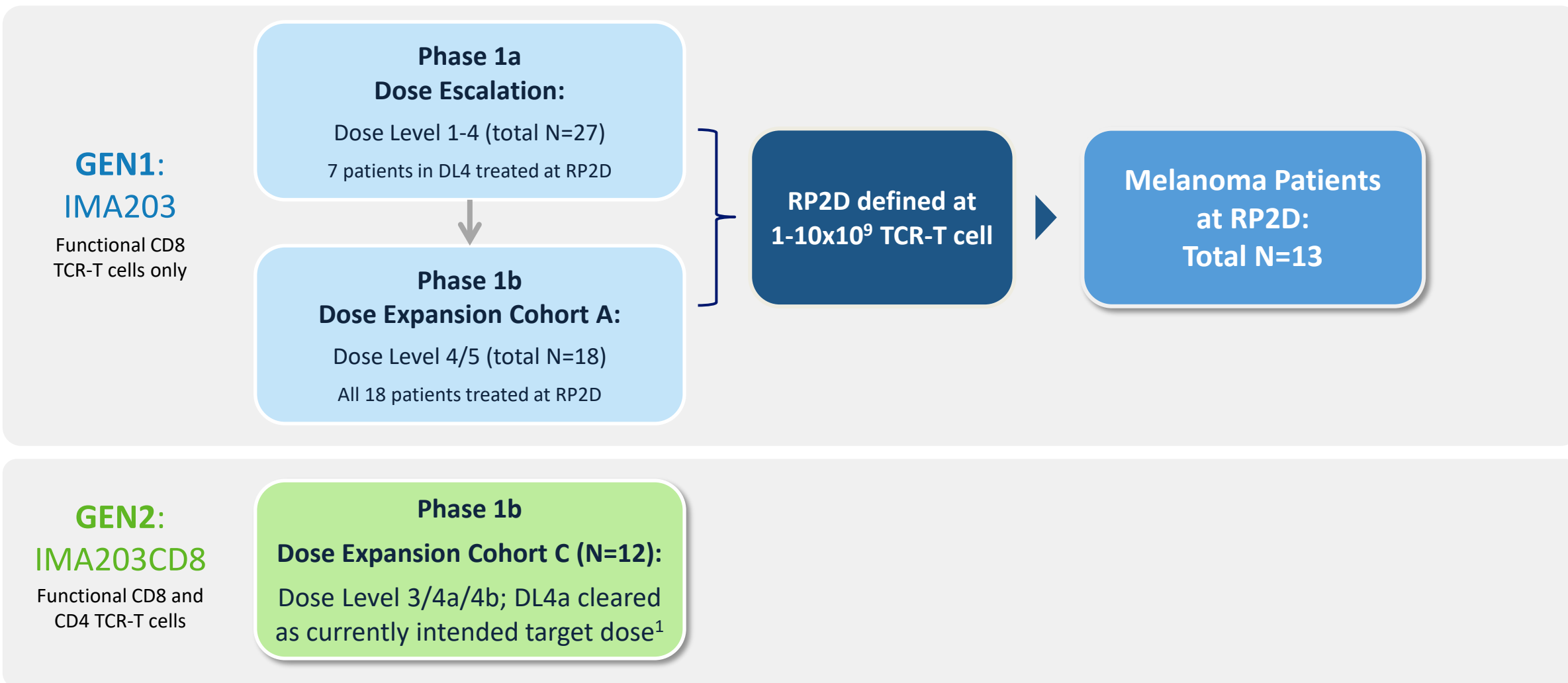
All ovarian cancer patients were platinum-resistant

- Sub-group analysis per tumor type at target dose includes data from Phase 1a plus Cohort A at RP2D
- Melanoma patient number (N=13) and characteristics allow such sub-group analysis for initial assessment of anti-tumor activity
- For other tumor types, appropriate patient numbers and characteristics have not yet been achieved



# ACTengine® IMA203 / IMA203CD8 TCR-T Trial in Advanced Solid Tumors

## Overview



Phase 1a and Cohort A data set in appendix

# Overview of Patient Characteristics and Responses

## Heavily Pretreated Patient Population across Clinical Trial Cohorts

	IMA203 GEN1		IMA203CD8 GEN2
	All Comers (N=45)	Melanoma Subgroup (N=13 of 45)	All Comers (N=12)
	Phase 1a	Cohort A	Cohort C
Efficacy population*	N=27 Thereof N=7 at RP2D	N=18 at RP2D	N=13 at RP2D
Prior lines of systemic treatment (median, min, max)	4 (1, 8)	3 (0, 10)	4 (0, 7)
LDH at baseline >1 x ULN [% of patients]	66.7	50.0	53.8
Baseline tumor burden Median target lesion sum of diameter [mm] (min, max)	133.0 (29, 219.7)	58.9 (21, 207.3)	52.0 (21.0, 178.7)
Dose level	DL1-4	DL4/5	DL4/5
ORR	48% (13/27)	50% (9/18)	62% (8/13)
cORR	19% (5/27)	47% (8/17)	50% (6/12)
mDOR [months]	4.4 (2.4, 23.0)	Not reached	Not reached
mFU [months]	Not defined#	10.8	14.4
			4.8

\* Patients with at least one available tumor response assessment post infusion; # All patients were PD at data cut-off; Initial ORR: Objective response rate according to RECIST 1.1 at any post infusion scan; Confirmed ORR (cORR): Confirmed objective response rate according to RECIST 1.1 for patients with at least two available post infusion scans or patients with progressive disease (PD) at any prior timepoint, patients with ongoing unconfirmed PR not included in cORR calculation; Duration of response (DOR) in confirmed responders is defined as time from first documented response until disease progression/death. Patients with ongoing response will be censored at date of data cut-off. Median DOR is analyzed by using the Kaplan-Meier method; Median Follow-up is analyzed by using the reverse Kaplan-Meier method; DOR: Duration of Response; FU: Follow-up

# ACTengine® IMA203 TCR-T Interim Update

Delivering a Meaningful Benefit to Patients with an Unmet Medical Need



## IMA203 GEN1 Monotherapy

Phase 1a & Cohort A – Focus on Melanoma at RP2D

## IMA203CD8 GEN2 Monotherapy

Cohort C – First Data Set on 2<sup>nd</sup> Generation

## Summary & Next Development Steps

# IMA203 GEN1 in All Melanoma Patients at RP2D – Most Frequent Adverse Events

N=16 Patients in Safety Population<sup>1</sup>



- **Expected cytopenia (Grade 1-4)** associated with lymphodepletion in all patients
- **Mostly mild to moderate cytokine release syndrome (CRS)**
  - 63% (10/16) with Grade 1 CRS
  - 31% (5/16) with Grade 2 CRS
  - 6% (1/16) with Grade 3 CRS (Phase 1a patient; recovered to Grade 2 after 3 days, no need for vasopressors and/or ventilation)
  - No dose-dependent increase of CRS
- **One non-serious, mild (Grade 1) ICANS<sup>2</sup> in DL5**
- **No dose-limiting toxicity**
- **No IMA203-related deaths**
- full IMA203 GEN1 monotherapy safety profile (generally consistent with safety in melanoma subset), see next slide

**IMA203 GEN1 monotherapy continues to be well tolerated  
at total doses between 1-10x10<sup>9</sup> TCR-T cells (RP2D)**

# IMA203 GEN1 across All Dose Levels – Tolerability Data

## Phase 1a Dose Escalation and Cohort A – All ≥Grade 3 Adverse Events (N=49)

TEAEs by maximum severity for all patients in Phase 1a dose escalation and Cohort A dose expansion (N=49)<sup>1</sup>

Adverse event (System organ class, Preferred term)	≥ Grade 3		Adverse event (System organ class, Preferred term)	≥ Grade 3	
	No.	%		No.	%
<b>Patients with any adverse event</b>	<b>49</b>	<b>100.0</b>	<b>table continued...</b>		
<b>Adverse Events of Special Interest</b>	<b>2</b>	<b>4.1</b>	<b>General disorders and administration site conditions</b>	<b>4</b>	<b>8.2</b>
Cytokine release syndrome	2	4.1	Condition aggravated <sup>4</sup>	1	2.0
ICANS <sup>2</sup>	0	0.0	Fatigue	1	2.0
<b>Blood and lymphatic system disorders</b>	<b>48</b>	<b>98.0</b>	Pyrexia	1	2.0
Neutropenia	36	73.5	Swelling face	1	2.0
Lymphopenia	27	55.1	<b>Metabolism and nutrition disorders</b>	<b>4</b>	<b>8.2</b>
Leukopenia	26	53.1	Hypokalaemia	3	6.1
Anaemia	24	49.0	Failure to thrive	1	2.0
Thrombocytopenia	17	34.7	Hypophosphataemia	1	2.0
Cytopenia	1	2.0	<b>Gastrointestinal disorders</b>	<b>2</b>	<b>4.1</b>
Leukocytosis	1	2.0	Abdominal pain	1	2.0
Lymphocytosis	1	2.0	Diarrhoea	1	2.0
<b>Investigations</b>	<b>9</b>	<b>18.4</b>	Vomiting	1	2.0
Neutrophil count decreased	4	8.2	<b>Injury, poisoning and procedural complications</b>	<b>2</b>	<b>4.1</b>
Alanine aminotransferase increased	2	4.1	Humerus fracture	1	2.0
Aspartate aminotransferase increased	2	4.1	Infusion related reaction	1	2.0
White blood cell count decreased	2	4.1	<b>Renal and urinary disorders</b>	<b>2</b>	<b>4.1</b>
Blood alkaline phosphatase increased	1	2.0	Acute kidney injury	1	2.0
Blood creatinine increased	1	2.0	Proteinuria	1	2.0
Blood fibrinogen decreased	1	2.0	<b>Skin and subcutaneous tissue disorders</b>	<b>2</b>	<b>4.1</b>
<b>Infections and infestations</b>	<b>7</b>	<b>14.3</b>	Rash maculo-papular	2	4.1
Appendicitis	1	2.0	<b>Cardiac disorders</b>	<b>1</b>	<b>2.0</b>
COVID-19	1	2.0	Atrial fibrillation <sup>3</sup>	1	2.0
Enterococcal infection	1	2.0	<b>Endocrine disorders</b>	<b>1</b>	<b>2.0</b>
Infection	1	2.0	Inappropriate antidiuretic hormone secretion	1	2.0
Orchitis	1	2.0	<b>Eye disorders</b>	<b>1</b>	<b>2.0</b>
Sepsis <sup>4,5</sup>	1	2.0	Ulcerative keratitis	1	2.0
Septic shock <sup>4</sup>	1	2.0	<b>Hepatobiliary disorders</b>	<b>1</b>	<b>2.0</b>
Urinary tract infection	1	2.0	Cholangitis	1	2.0
<b>Respiratory, thoracic and mediastinal disorders</b>	<b>6</b>	<b>12.2</b>	<b>Immune system disorders</b>	<b>1</b>	<b>2.0</b>
Hypoxia	3	6.1	Contrast media allergy	1	2.0
Bronchial obstruction	1	2.0	<b>Musculoskeletal and connective tissue disorders</b>	<b>1</b>	<b>2.0</b>
Laryngeal inflammation	1	2.0	Muscle spasms	1	2.0
Pleural effusion	1	2.0	<b>Nervous system disorders</b>	<b>1</b>	<b>2.0</b>
Respiratory failure	1	2.0	Headache	1	2.0
<b>Vascular disorders</b>	<b>6</b>	<b>12.2</b>	<b>Reproductive system and breast disorders</b>	<b>1</b>	<b>2.0</b>
Hypertension	4	8.2	Vaginal haemorrhage	1	2.0
Hypotension	2	4.1			

- Well tolerated at doses as high as ~10x10<sup>9</sup> TCR-T cells
- No AE ≥Grade 3 was observed with a frequency ≥10% when excluding expected cytopenia associated with lymphodepletion
- No IMA203-related Grade 5 Adverse Events

All treatment-emergent adverse events (TEAEs) with ≥ Grade 3 regardless of relatedness to study treatment that occurred in at least 1 patient (except for ICANS, where only Grade 1-2 occurred; listed for completeness due to being an adverse event of special interest) are presented. 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, version 5.0. Grades for CRS and ICANS were determined according to CARTOX criteria (Neelapu et al., 2018). Patients are counted only once per adverse event and severity classification. Based on interim data extracted from open clinical database (30-Sep-2023); <sup>1</sup> Two patients with disease progression after first IMA203 infusion received exploratory second IMA203 infusion. They had these ≥ Grade 3 TEAEs only after second infusion, which are included in the table: First patient: Abdominal pain, Cytokine release syndrome, Diarrhoea, Hypokalaemia, Proteinuria; Second patient: Humerus fracture, Muscle spasms, Neutropenia, Thrombocytopenia; <sup>2</sup> ICANS: Immune effector cell-associated neurotoxicity syndrome; <sup>3</sup> DLT: Dose limiting toxicity in phase 1a at DL2 reported on March 17, 2021; <sup>4</sup> Fatal Adverse events were not considered related to any study drug; <sup>5</sup> Patient died from sepsis of unknown origin and did not receive IMA203 TCR-T cells.

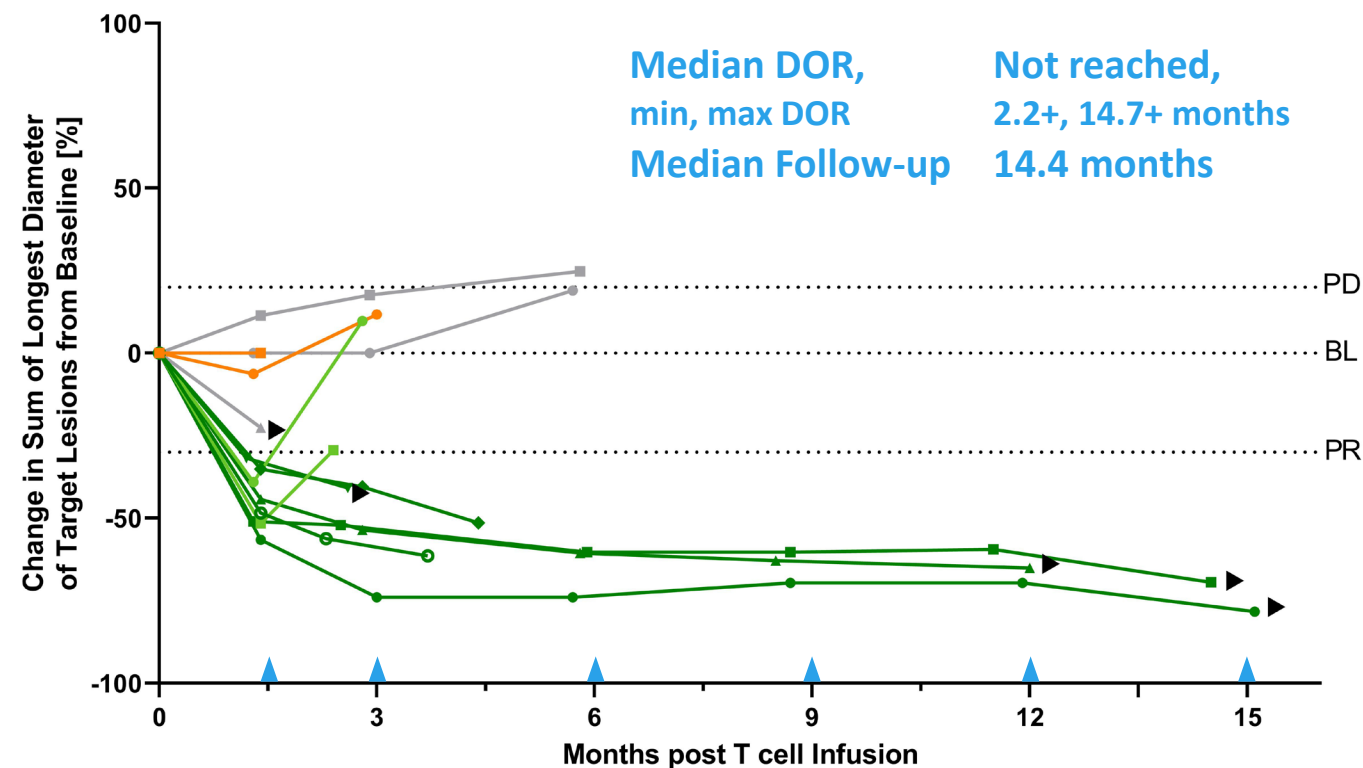
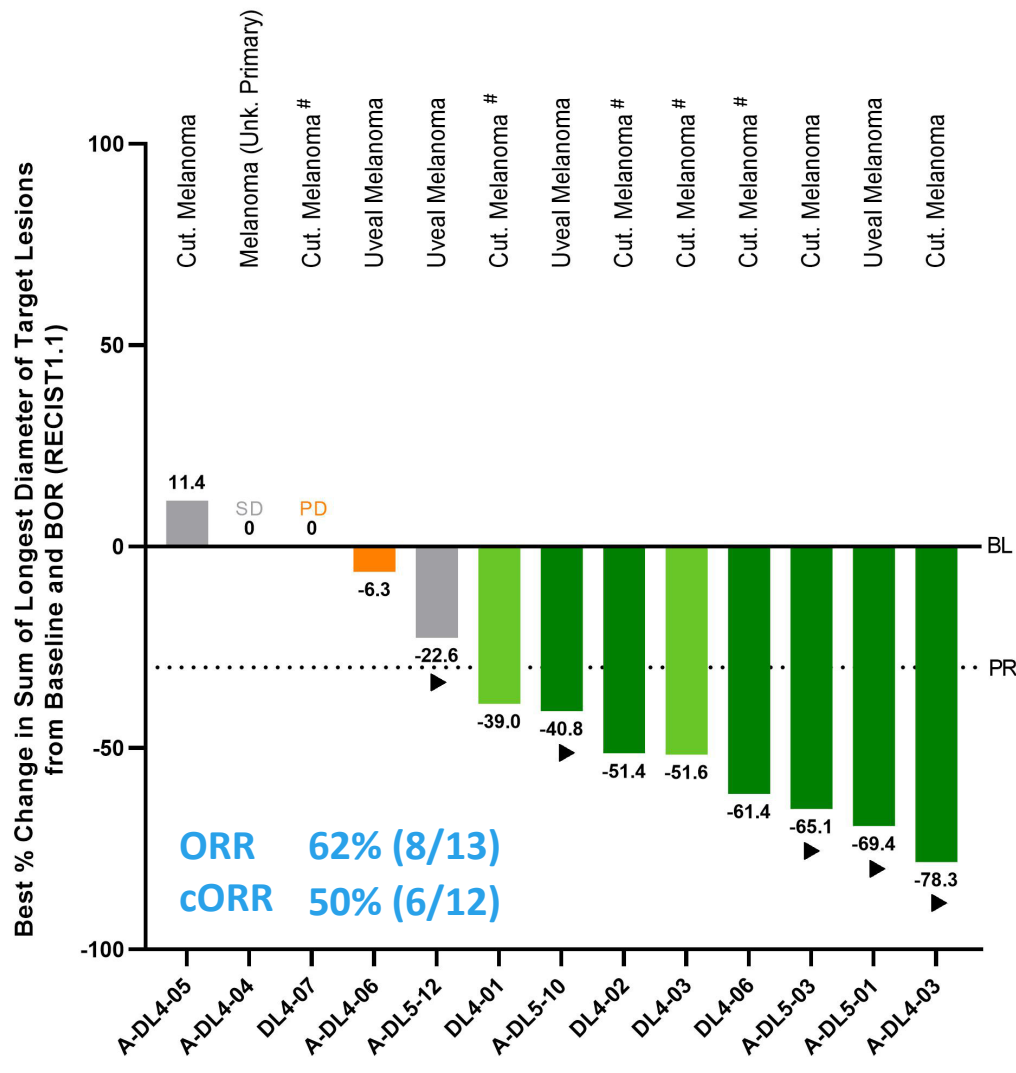


# IMA203 GEN1 in All Melanoma Patients at RP2D (N=13) – BOR and Response over Time

## Durable Responses 15+ Months after Treatment



Immatics®



# IMA203 GEN1 in Melanoma Targeted to Enter Registration-Enabling Randomized Phase 2/3 Trial in 2L+ Melanoma in 2024

## Clinically and Commercially Attractive Features of IMA203

≥95% of cutaneous melanoma patients are PRAME-positive
Well tolerated Mostly mild to moderate CRS, infrequent & mild ICANS
Promising anti-tumor activity (cORR, mDOR)
Leukapheresis as source for cell product, no surgery required
Short manufacturing time of 7 days plus 7 days of QC release testing
Low dose IL-2 post IMA203 infusion with better tolerability profile than high dose IL-2

## High Medical Need in Cutaneous and Uveal Melanoma

	Cutaneous Melanoma	Uveal Melanoma
Patient Population	<b>2L+</b> CPI-refractory, BRAF/MEK inhibitor-refractory if BRAF mutation+	<b>2L+</b> Kimmtrak-refractory, CPI/chemotherapy-refractory
IMA203 Opportunity	<b>~3,000</b> HLA-A*02:01 and PRAME-positive cutaneous melanoma patients annually in the US <sup>1</sup>	<b>~300</b> HLA-A*02:01 and PRAME-positive uveal melanoma patients annually in the US <sup>2</sup>

# ACTengine® IMA203 TCR-T Interim Update

Delivering a Meaningful Benefit to Patients with an Unmet Medical Need

## IMA203 GEN1 Monotherapy

Phase 1a & Cohort A – Focus on Melanoma at RP2D

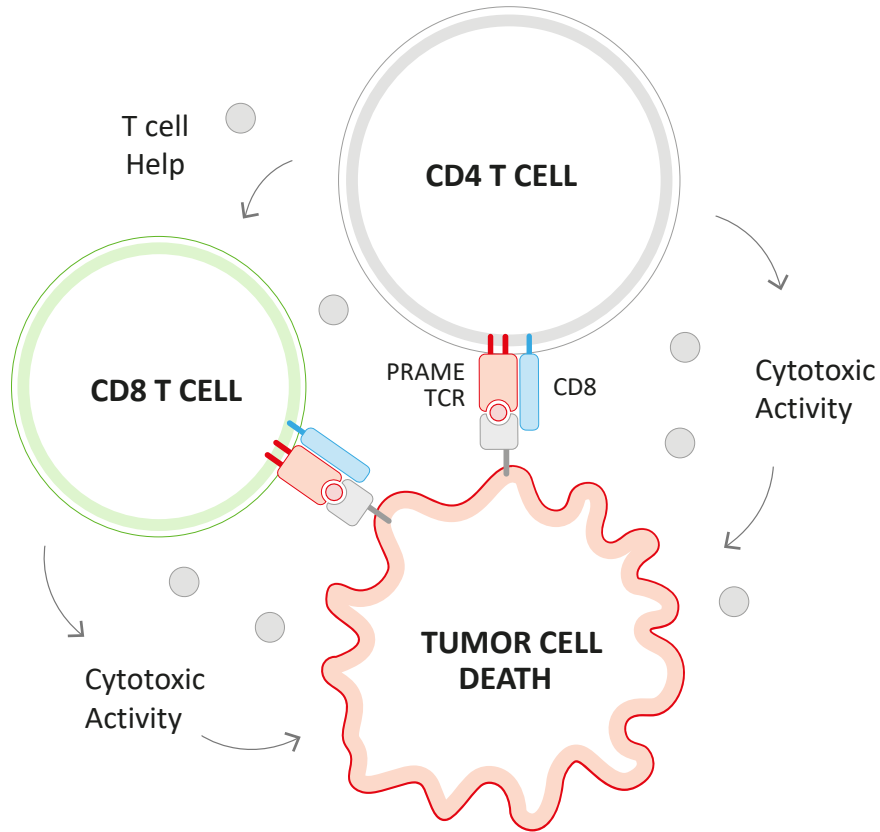
## IMA203CD8 GEN2 Monotherapy

Cohort C – First Data Set on 2<sup>nd</sup> Generation

## Summary & Next Development Steps

# IMA203CD8 GEN2 – IMA203 TCR-T Monotherapy Leveraging CD8 and CD4 cells

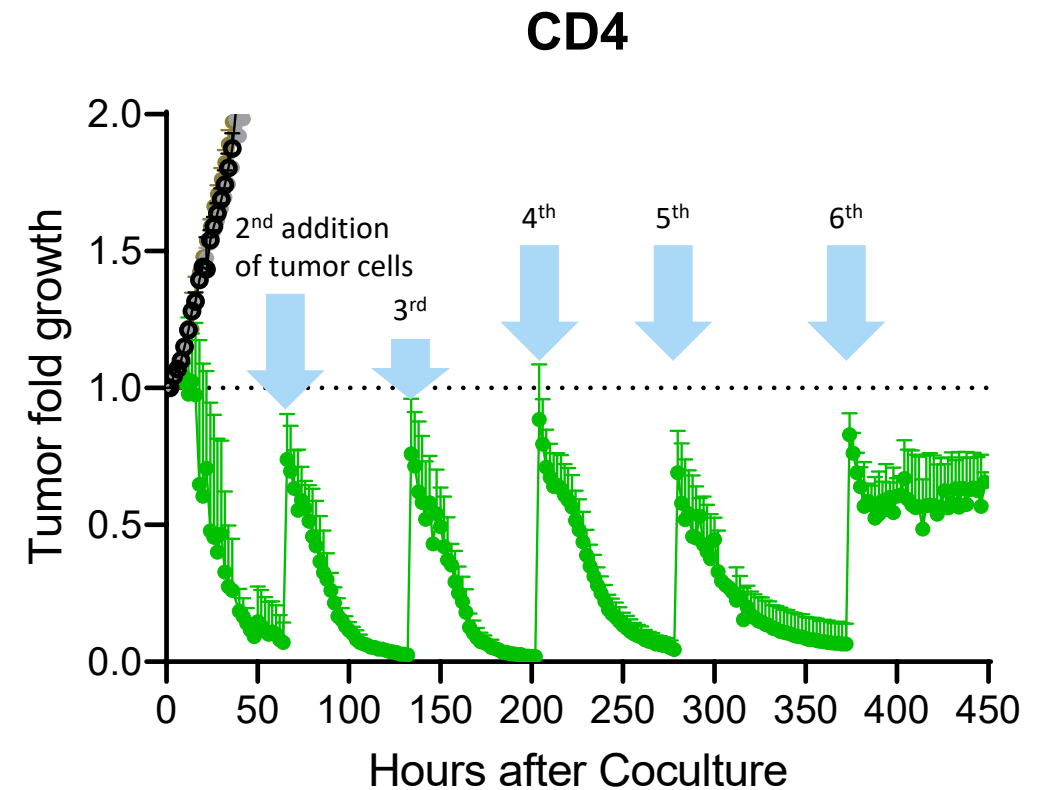
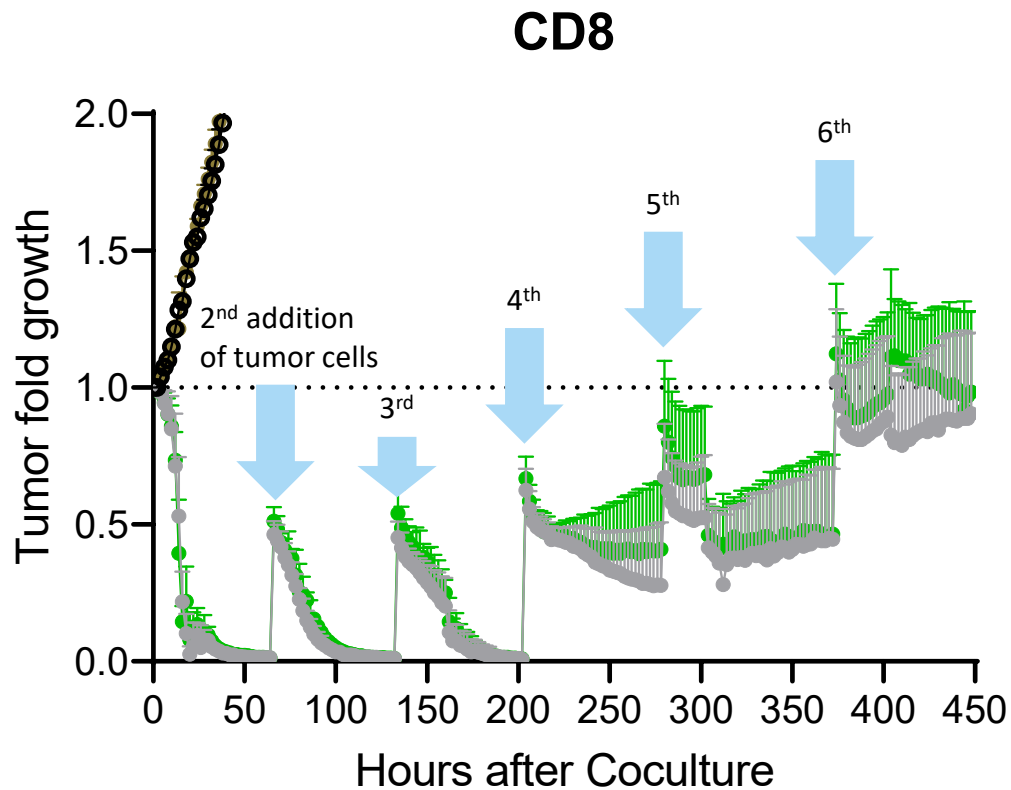
Differentiated Pharmacology Compared to 1<sup>st</sup>-Generation TCR-only Approaches



- IMA203CD8 GEN2 designed to broaden the clinical potential of IMA203 TCR-T monotherapy by adding functional CD4 T cells via co-transduction of CD8 $\alpha\beta$  alongside PRAME TCR
- Activated CD4 T cells aid activity of other immune cells by releasing cytokines and acquire cytotoxic functions
- Functional CD4 T cells mediate longer anti-tumor activity than CD8 T cells and potentiate the anti-tumor activity of the cell product in preclinical studies<sup>1</sup>
- Data from CD19 CAR-T-treated leukaemia patients suggest a relevant role of engineered CD4 T cells in long-term durability<sup>2</sup>

# IMA203CD8 GEN2 – Preclinical Assessment of Anti-Tumor Efficacy

Functional CD4 T cells Mediate Longer Anti-Tumor Activity than CD8 T cells *in vitro*





# IMA203CD8 GEN2 in Cohort C (N=12) – Most Frequent Adverse Events

## Manageable Tolerability in 12 Patients Treated with IMA203CD8 at 3 Escalating Dose Levels<sup>1</sup>

- **Expected cytopenia (Grade 1-4)** associated with lymphodepletion in all patients
- Cytokine release syndrome (CRS) in 92% (11/12) of patients:  
Trend towards **more severe CRS at higher doses, in all cases well manageable**
  - 67% (8/12) with Grade 1 or 2 CRS (4 in DL3, 3 in DL4a, 1 in DL4b)
  - 17% (2/12) with Grade 3 CRS (2 in DL4b; patient C-DL4b-04, see also description below)
  - 8% (1/12) with Grade 4 CRS (1 in DL4b, patient C-DL4b-01, see also description below)
- **One patient with neurotoxicity (see below), no ICANS<sup>2</sup> or neurotoxicity reported for the other patients**
- **Dose-limiting toxicities (DLTs) at Dose Level 4b** were observed in 2 of 4 patients
  - 1) In patient C-DL4b-01 treated with highest possible dose at DL4b, high biological activity (*in vivo* T cell expansion) observed; patient developed Grade 4 neurotoxicity and Grade 4 CRS on day 6 after infusion, combined with Grade 3 Hemophagocytic Lymphohistiocytosis (HLH)
  - 2) Patient C-DL4b-04 treated at DL4b developed Grade 3 CRS with transient Grade 3 liver enzyme (ALT) increase that resolved to Grade 2 within 10 days; no need for vasopressors or ventilation at any time
- **No high-grade CRS, no neurotoxicity and no DLTs were reported for 4 patients treated at DL3 and 4 patients treated at DL4a**
- **No IMA203CD8-related deaths<sup>3</sup>**
- **Expanded DL4a dose cohort ongoing**

## IMA203CD8 GEN2 monotherapy shows a manageable tolerability profile

# Tolerability Data – Cohort C IMA203CD8 GEN2

## All ≥Grade 3 Adverse Events (N=12)

TEAEs by maximum severity for all patients in Cohort C (N=12)

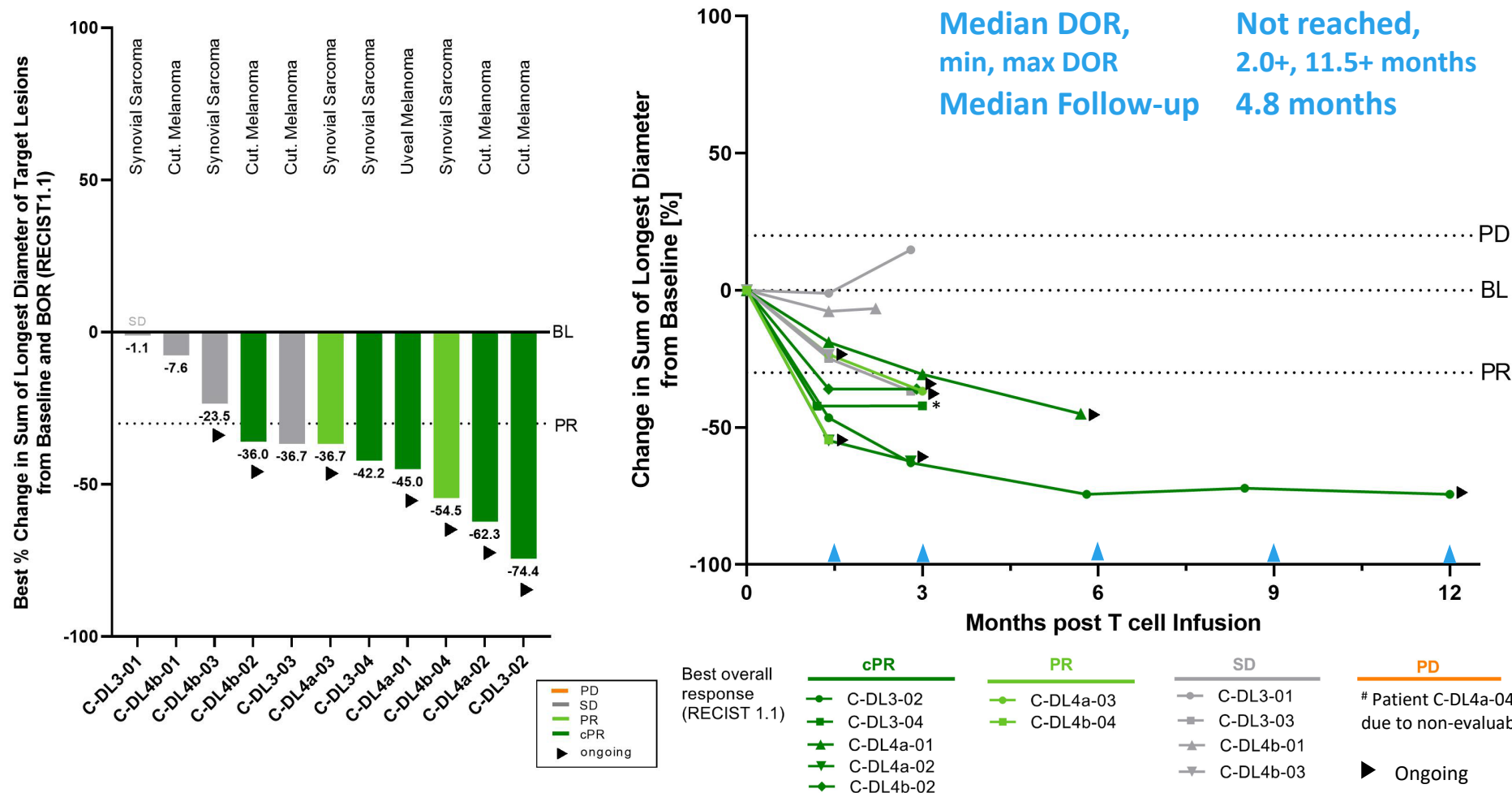
Adverse event (System organ class, preferred term)	≥ Grade 3	
	No.	%
<b>Patients with any adverse event</b>	<b>12</b>	<b>100.0</b>
<b>Adverse events of special interest</b>	<b>3</b>	<b>25.0</b>
Cytokine release syndrome <sup>1</sup>	3	25.0
Immune effector cell-associated neurotoxicity syndrome	0	0.0
<b>Blood and lymphatic system disorders</b>	<b>11</b>	<b>91.7</b>
Neutropenia	9	75.0
Anaemia	8	66.7
Lymphopenia	8	66.7
Thrombocytopenia	4	33.3
Leukopenia	2	16.7
<b>Investigations</b>	<b>4</b>	<b>33.3</b>
Aspartate aminotransferase increased	2	16.7
Neutrophil count decreased	2	16.7
Alanine aminotransferase increased	1	8.3
Blood alkaline phosphatase increased	1	8.3
Blood bilirubin increased	1	8.3
Gamma-glutamyltransferase increased	1	8.3
<b>Metabolism and nutrition disorders</b>	<b>2</b>	<b>16.7</b>
Hypermagnesaemia	1	8.3
Hypoalbuminaemia	1	8.3
Hypophosphataemia	1	8.3
<b>Nervous system disorders</b>	<b>2</b>	<b>16.7</b>
Neurotoxicity <sup>2</sup>	1	8.3
Syncope	1	8.3
<b>Immune system disorders</b>	<b>1</b>	<b>8.3</b>
Haemophagocytic lymphohistiocytosis <sup>2</sup>	1	8.3
<b>Infections and infestations</b>	<b>1</b>	<b>8.3</b>
Infection	1	8.3

- Manageable tolerability
- Most frequent ≥Grade 3 AEs were expected cytopenia associated with lymphodepletion
- No IMA203CD8-related Grade 5 Adverse Events<sup>1</sup>
- Dose escalation ongoing

All treatment-emergent adverse events (TEAEs) with ≥ Grade 3 regardless of relatedness to study treatment that occurred in at least 1 patient (except for ICANS, where no event was documented; listed for completeness due to being an adverse event of special interest) are presented. 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, version 5.0. Grades for CRS and ICANS were determined according to CARTOX criteria (Neelapu et al., 2018). Patients are counted only once per adverse event and severity classification. Based on interim data extracted from open clinical database (30-Sep-2023); <sup>1</sup> DLT: Dose limiting toxicity in patient DL4b-04. <sup>2</sup> DLTs in patient DL4b-01;

# IMA203CD8 GEN2 in Cohort C (N=12<sup>#</sup>) – BOR and Response over Time

Deepening of Response from SD to PR in 2 Patients, 6 Responses Ongoing



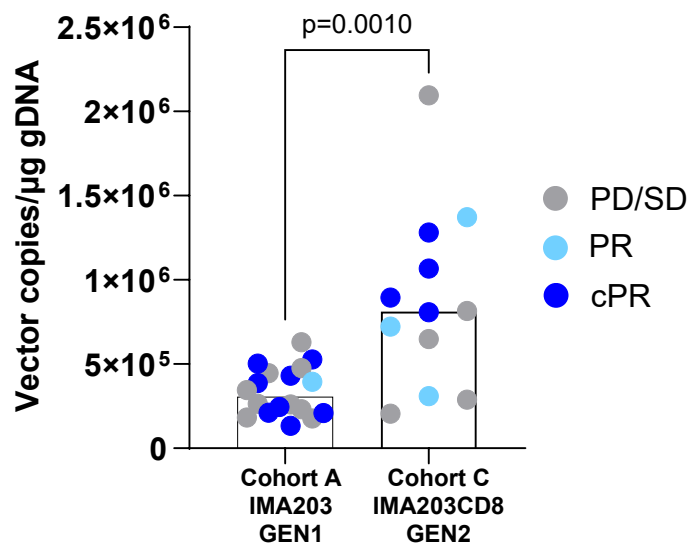
**ORR 58% (7/12)**  
**cORR 56% (5/9)**

- 6 out of 7 responses ongoing
- 11/12 patients show tumor shrinkage
- Deepening of response from SD to PR in two patients (C-DL4a-01, C-DL4a-03)
- Ongoing durable response 12+ months after infusion

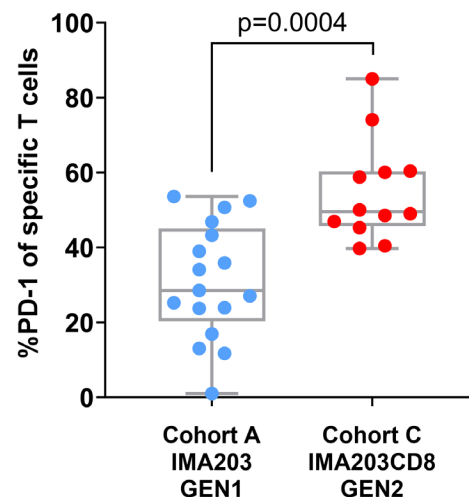
# IMA203CD8 GEN2: Translational Data Shows Enhanced Pharmacology

## Cohort A IMA203 GEN1 (All Patients at RP2D) vs Cohort C IMA203CD8 GEN2

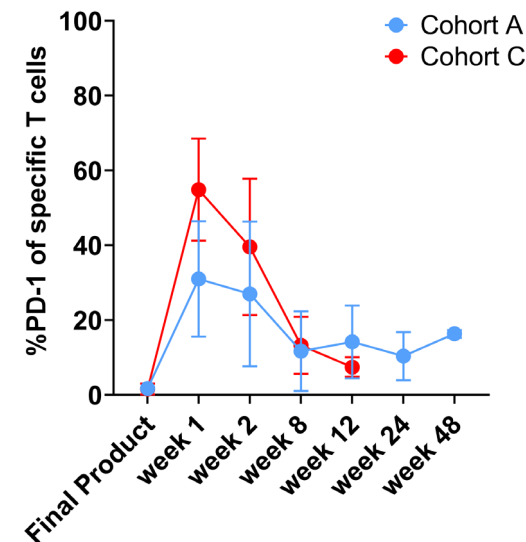
Higher peak expansion ( $C_{max}$ )  
of IMA203CD8 T cells  
when normalized to infused dose



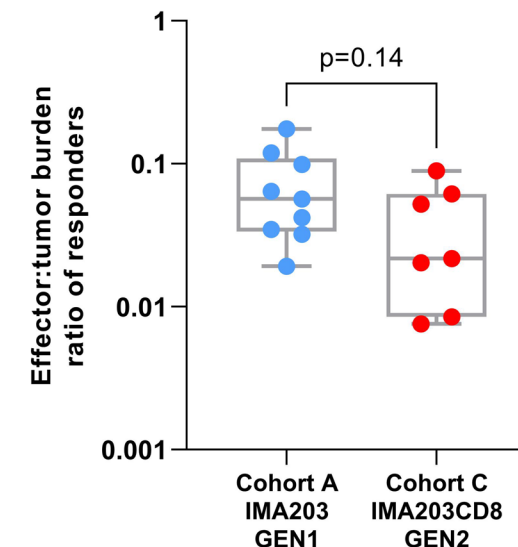
Higher activation levels in  
IMA203CD8 T cells at week 1...



...without exhaustion  
over time



Trend towards responses at  
lower cell dose and higher  
tumor burden with IMA203CD8



Initial translational data indicates higher biological and clinical activity of IMA203CD8 GEN2

# ACTengine® IMA203 TCR-T Interim Update

Delivering a Meaningful Benefit to Patients with an Unmet Medical Need



## IMA203 GEN1 Monotherapy

Phase 1a & Cohort A – Focus on Melanoma at RP2D

## IMA203CD8 GEN2 Monotherapy

Cohort C – First Data Set on 2<sup>nd</sup> Generation

## Summary & Next Development Steps

# ACTengine® IMA203 / IMA203CD8 TCR-T Monotherapy Targeting PRAME

## Summary of GEN1 and GEN2 Clinical Data and Planned Next Steps

### IMA203 GEN1 Monotherapy in Melanoma at RP2D

- Well tolerated, mostly mild to moderate CRS, infrequent & mild ICANS
- **50% (6/12) cORR, mDOR not reached at mFU of 14.4 months**
- **Durability with ongoing responses at 15+ months in some patients**
- RP2D defined at  $1-10 \times 10^9$  total TCR-T cells
- FDA RMAT designation received in multiple PRAME expressing cancers including cutaneous and uveal melanoma

#### Next Step

Ongoing alignment with FDA on patient population, trial design, CMC targeting registration-enabling randomized Phase 2/3 trial in 2L+ melanoma

### IMA203CD8 GEN2 Monotherapy

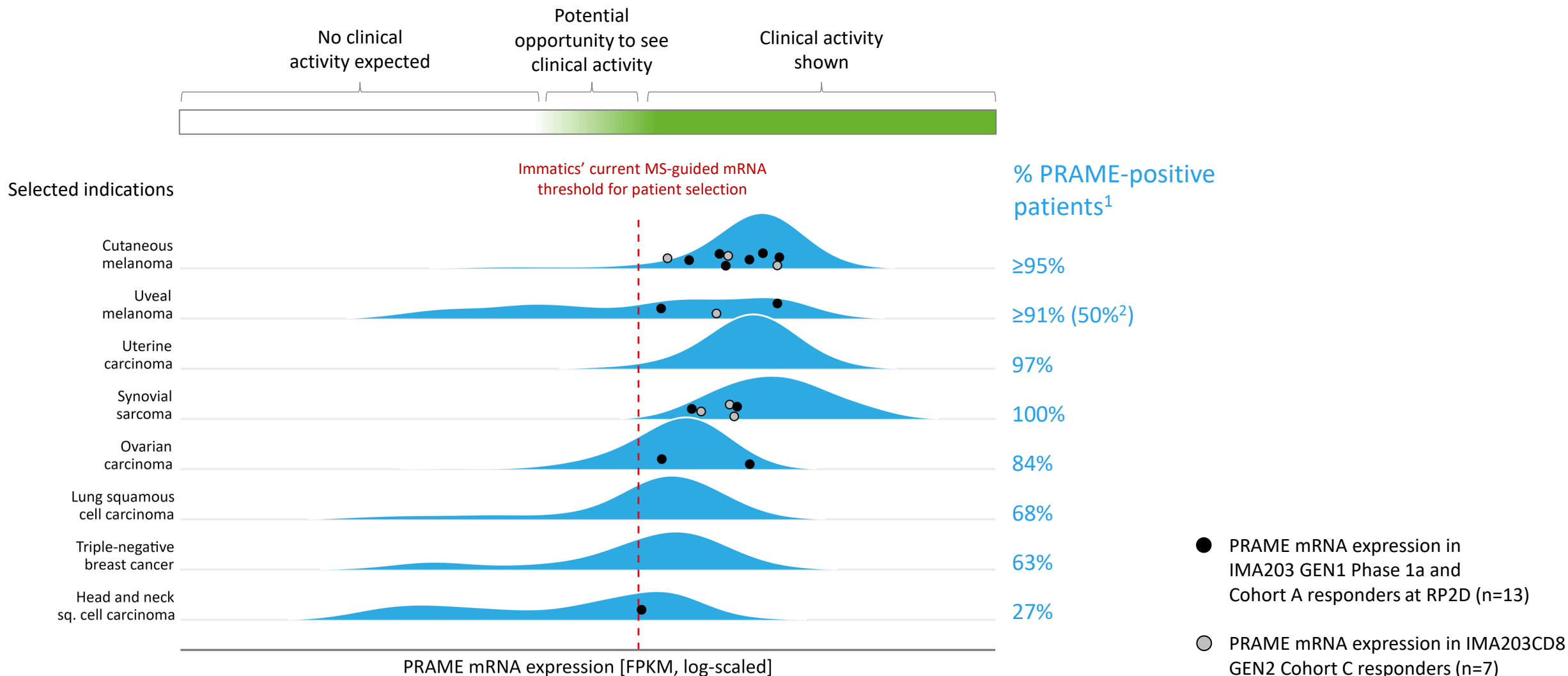
- Enhanced primary and secondary pharmacology when compared to GEN1
- Manageable tolerability (2 DLTs at DL4b, dose escalation ongoing)
- Initial clinical activity observed with differentiated response pattern
  - 56% (5/9) cORR
  - 6 out of 7 responses ongoing at data cut-off, durable response at 12+ months
  - SD converting to PR over time (N=2)
  - Enhanced biological efficacy with PRs at lower T cell:tumor cell ratio compared to IMA203 GEN1

#### Next Step

Clinical footprint expansion outside of melanoma in addition to treating melanoma patients

# Potential of IMA203 in Additional Solid Cancer Indications

Based on PRAME Expression in IMA203 GEN1 and IMA203CD8 GEN2 Responders





# ACTengine® IMA203 / IMA203CD8 TCR-T Monotherapy Targeting PRAME

Leveraging the Full Breath of PRAME in Three Steps

## Development Strategy

### Step 1 2024

IMA203 GEN1 in cutaneous melanoma (potentially bundled with uveal melanoma) as first tumor type targeted to enter registration-enabling trial

### Step 2 2024

Signal finding in ovarian cancer and uterine cancer in dedicated dose expansion cohorts with IMA203CD8 GEN2

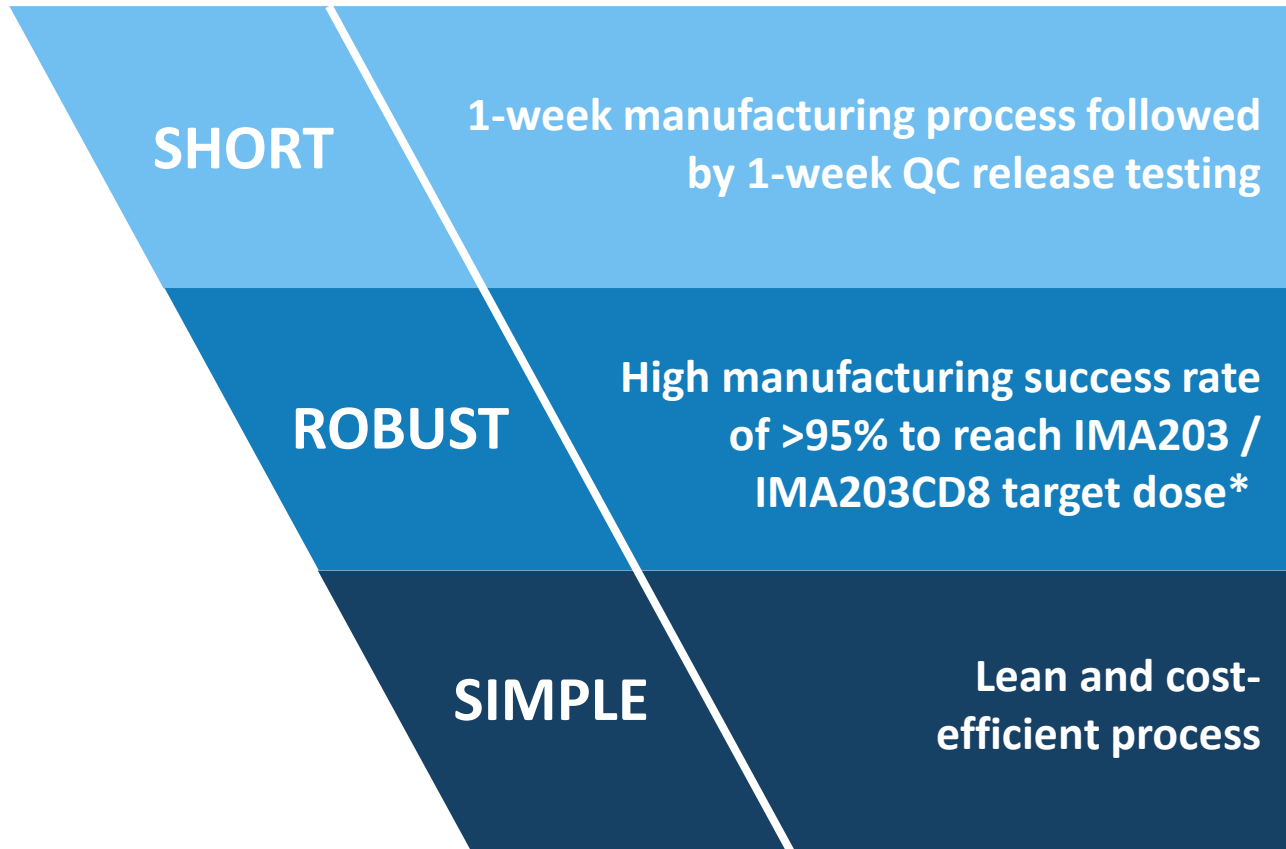
### Step 3

Pursue tumor-agnostic label in PRAME+ solid cancers to leverage full breadth of PRAME - including NSCLC, triple-negative breast cancer and others

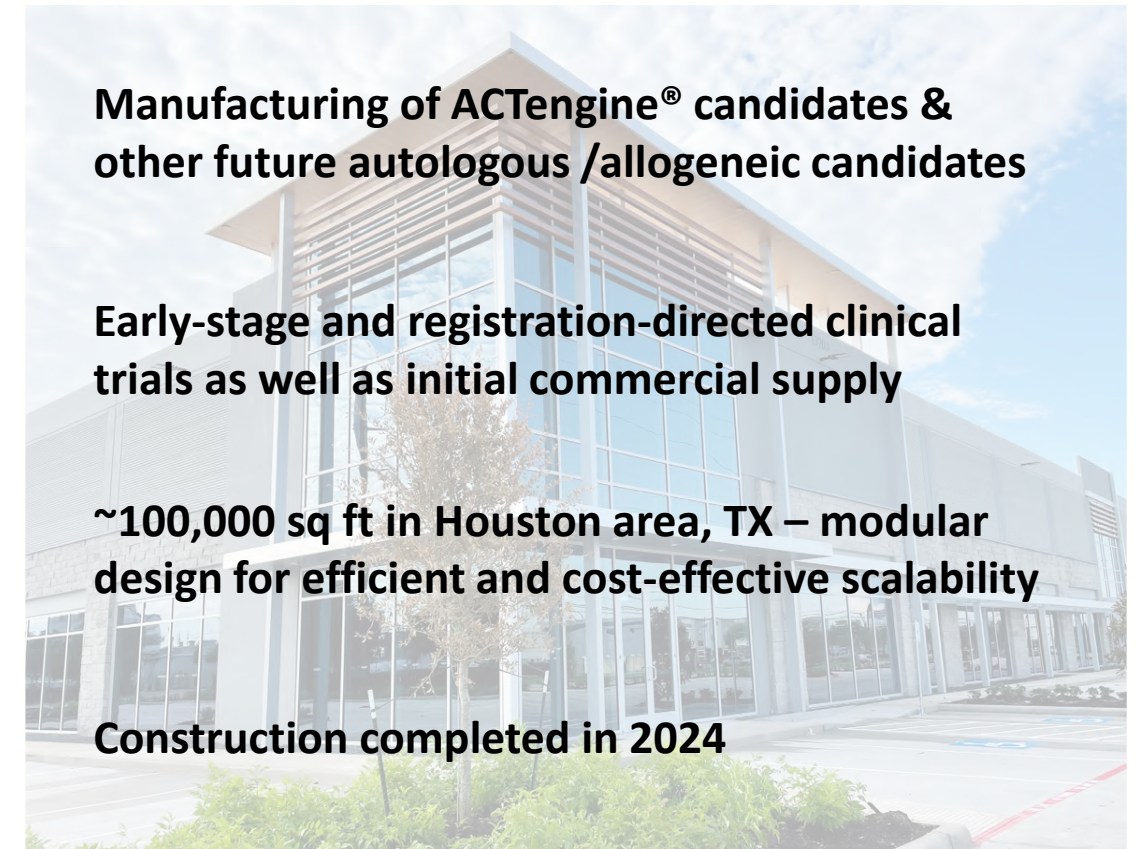
# ACTengine® IMA203 TCR-T Product Manufacturing

## Enhancing Manufacturing Process and Capabilities

### Proprietary Manufacturing Process



### State-of-the-art Research & GMP Manufacturing Facility



# IMA203 TCR-T Has the Potential to Reach a Large Patient Population

~39,000 Patients per Year in the US only

## Selected Indications

	<u>Incidence</u>	<u>R/R Incidence</u>	<u>PRAME Positive</u>
Cut. Melanoma	99,800	7,700	95%
Uveal Melanoma	1,500	800	91%
Ovarian Carcinoma	19,900	12,800	84%
Uterine Carcinoma	62,700	10,700	97%
Uterine Carcinosarcoma	3,300	1,900	100%
Squamous NSCLC	57,000	34,600	68%
Small Cell Lung Cancer	31,900	19,400	45%
Adeno NSCLC	91,200	55,300	25%
HNSCC	66,500	15,100	27%
Breast Carcinoma	290,600	43,800	26% TNBC: 63%
Synovial Sarcoma	1,000	400	100%
Cholangiocarcinoma	8,000	7,000	33%

## Patient Population

Based on R/R Incidence;  
PRAME and HLA-A\*02:01+

2,999  
298  
4,408  
4,255  
779  
9,646  
3,579  
5,668  
1,672  
4,669  
164  
947

**TOTAL ~39,000**  
**annually in the US**

## Multiple opportunities to broaden patient reach and patient benefit:

- Expand beyond US population
- Expand into other indications such as kidney, esophageal, bladder, other liver cancers, other sarcoma subtypes through indication-specific or indication-agonistic label expansion
- Move into earlier lines of therapy (R/R Incidence → Incidence)
- Inclusion of patients with lower PRAME-threshold



## ACTengine® IMA204 – TCR-T Targeting COL6A3 Exon 6

# 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 – 76%  
Breast Carcinoma – 77%  
Stomach Carcinoma – 67%  
Sarcoma – 63%  
Colorectal Carcinoma – 60%  
Esophageal Carcinoma – 60%  
Squamous NSCLC – 55%  
Adeno NSCLC – 57%  
HNSCC – 56%  
Uterine Carcinosarcoma – 50%  
Mesothelioma – 44%  
Cholangiocarcinoma – 36%  
Melanoma – 35%  
Bladder Carcinoma – 34%  
Ovarian Carcinoma – 31%

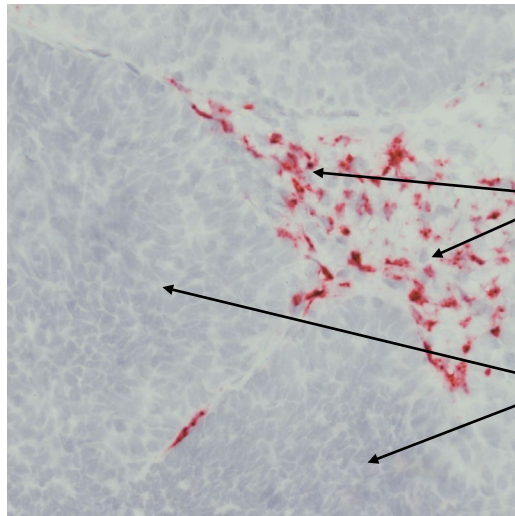
IMA204 provides a promising therapeutic opportunity for a broad patient population as monotherapy or in combination with TCR-T cells directed against tumor targets



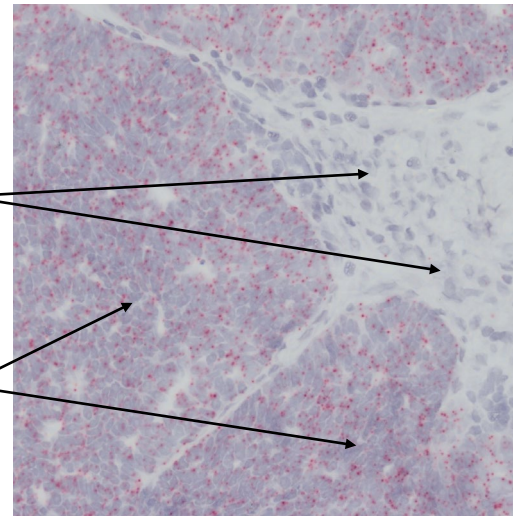
# ACTengine® IMA204 – High Affinity, CD8-independent TCR

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

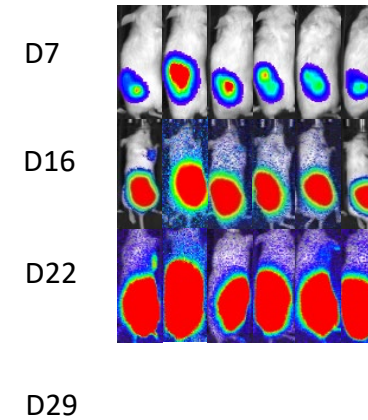


Stroma  
cells

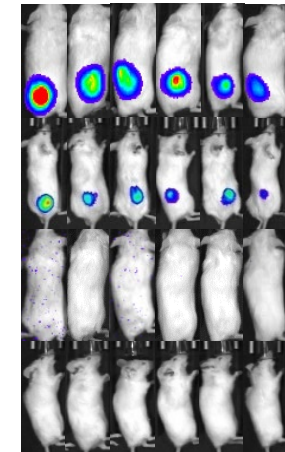
Tumor  
cells

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

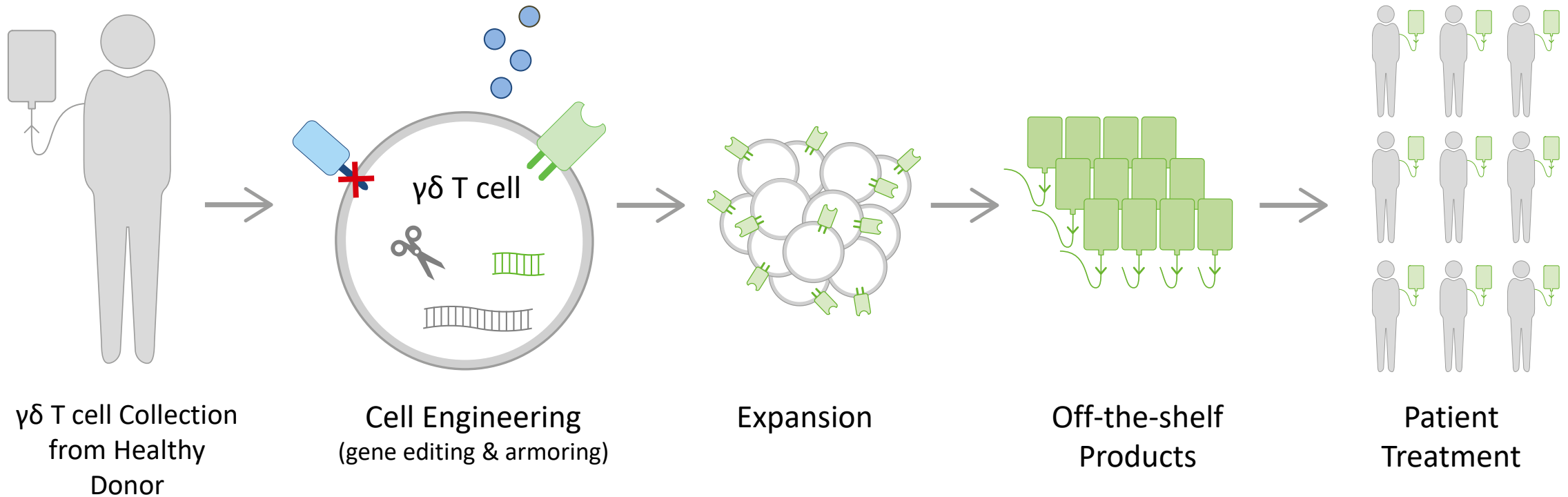
**Affinity matured CD8-independent, next-generation TCR engages both CD4 and CD8 T cells without the need of CD8 co-transduction**





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

# ACTallo® – Immatics' Allogeneic Cell Therapy Approach



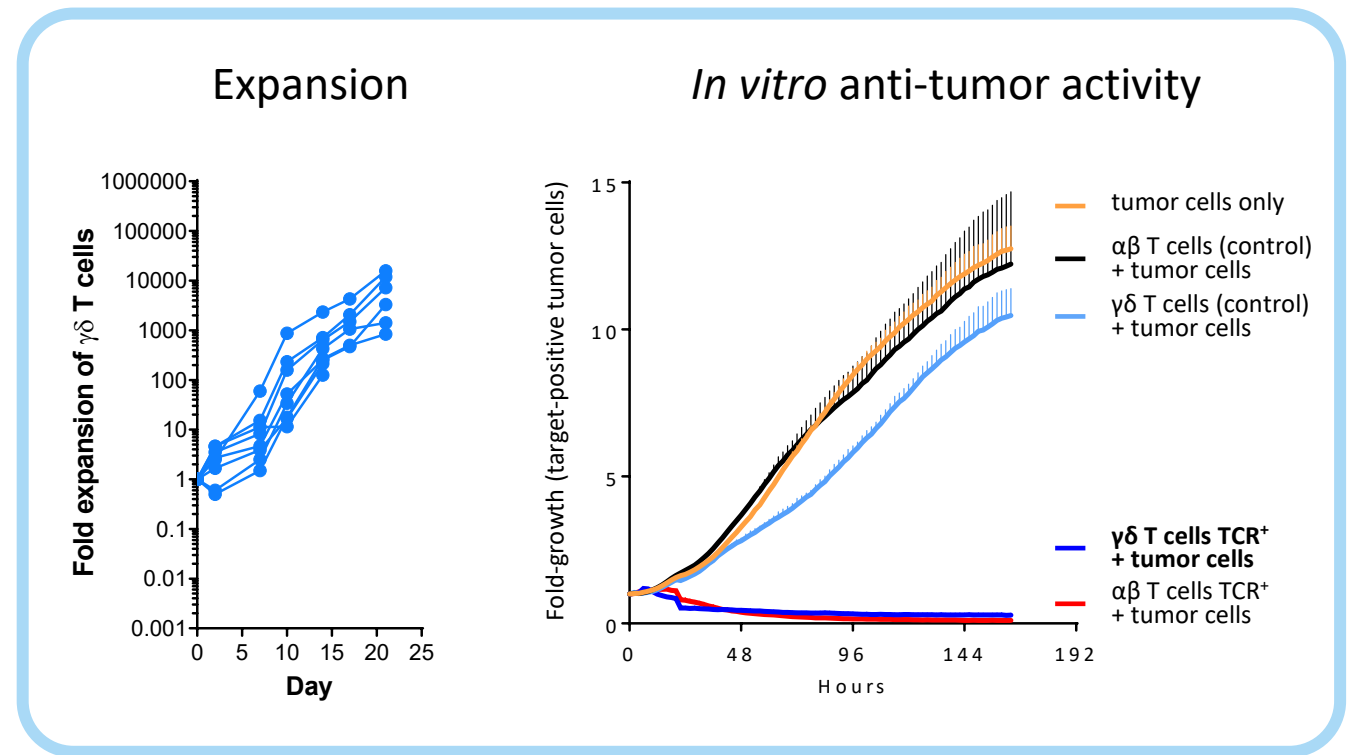
- **Off-the-shelf cell therapy**, no need for personalized manufacturing → reduced logistics and time to application
- **Potential for hundreds of doses** from one single donor leukapheresis → lower cost of goods
- **Use of healthy donor material** provides standardized quality and quantity of starting material
- Strategic collaborations combining Immatics' proprietary ACTallo® platform with Bristol Myers Squibb's next-gen technologies and Editas Medicine's CRISPR gene editing technology to develop next-gen allogeneic  $\gamma\delta$  TCR-T/CAR-T programs

# 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

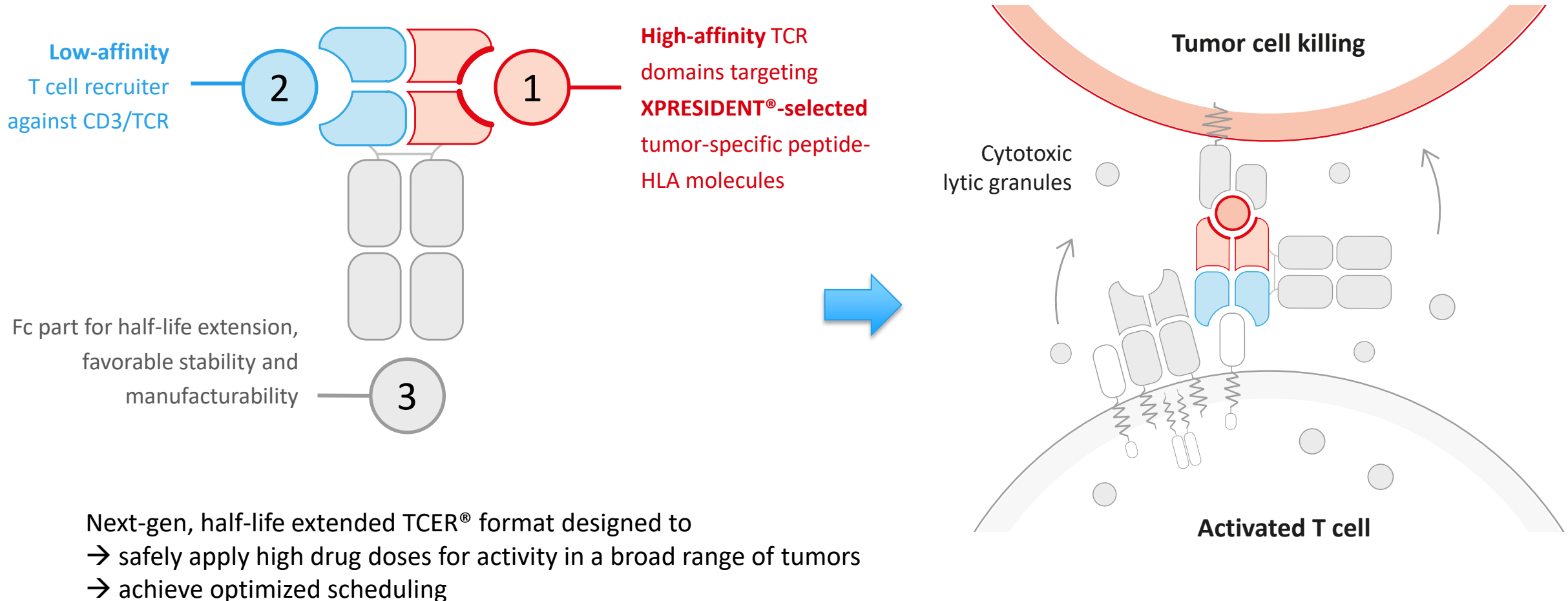




## TCER® – TCR Bispecifics

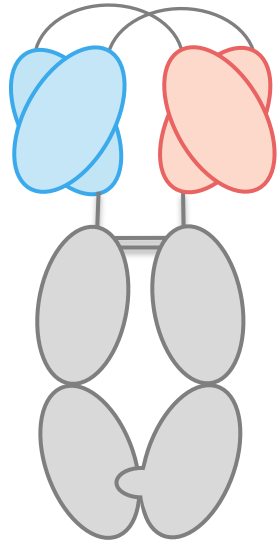
# TCER® – Immatics' Next-generation, Half-Life Extended Bispecifics

## Proprietary TCER® Format Consisting of Three Distinct Elements





# TCER® – Immatics' Next-generation, Half-Life Extended Bispecifics



1

## pHLA targeting TCR

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

2

## T cell recruiting antibody

- ✓ **Low-affinity** (triple digit nM) 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

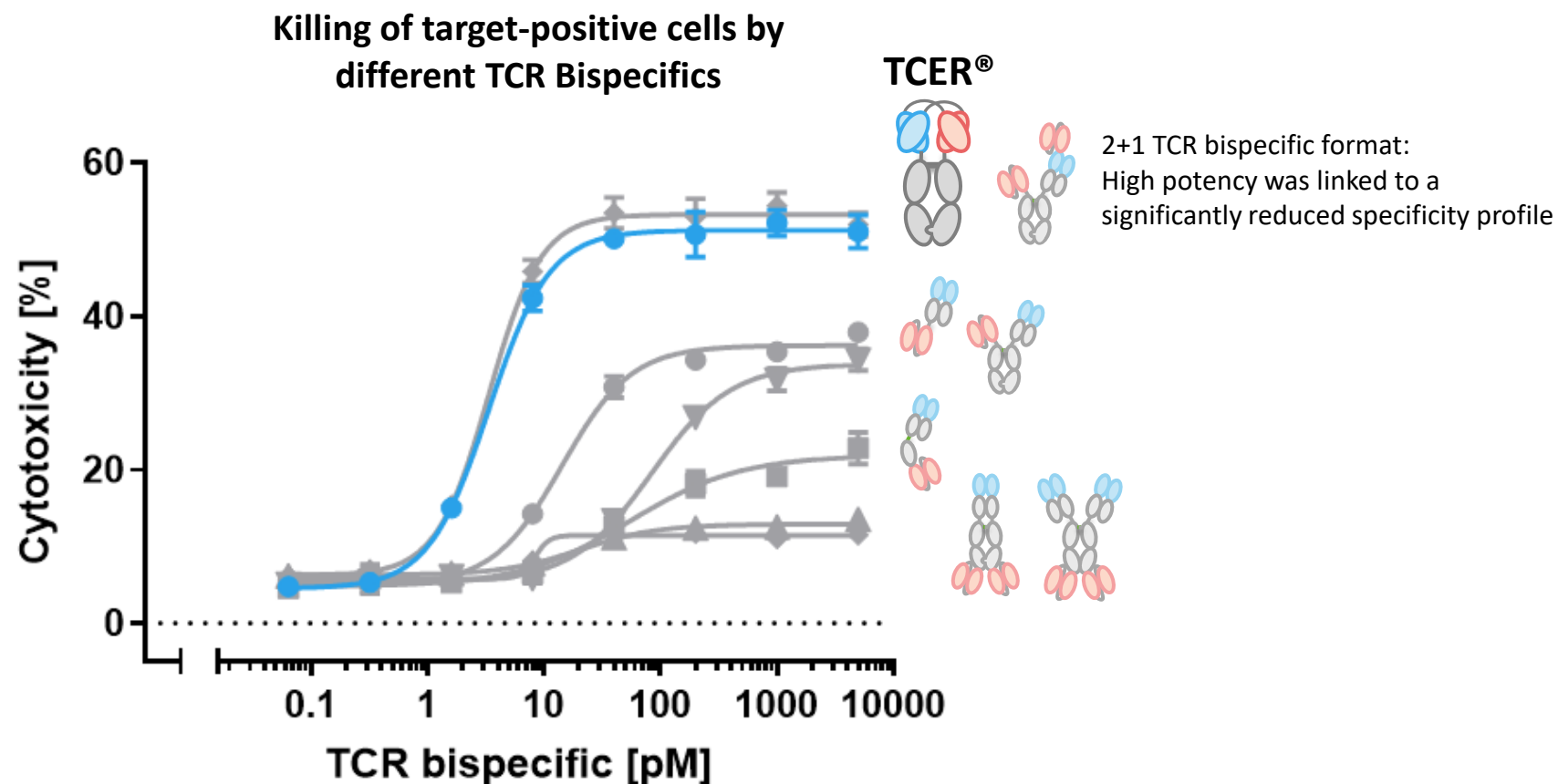
3

## 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®



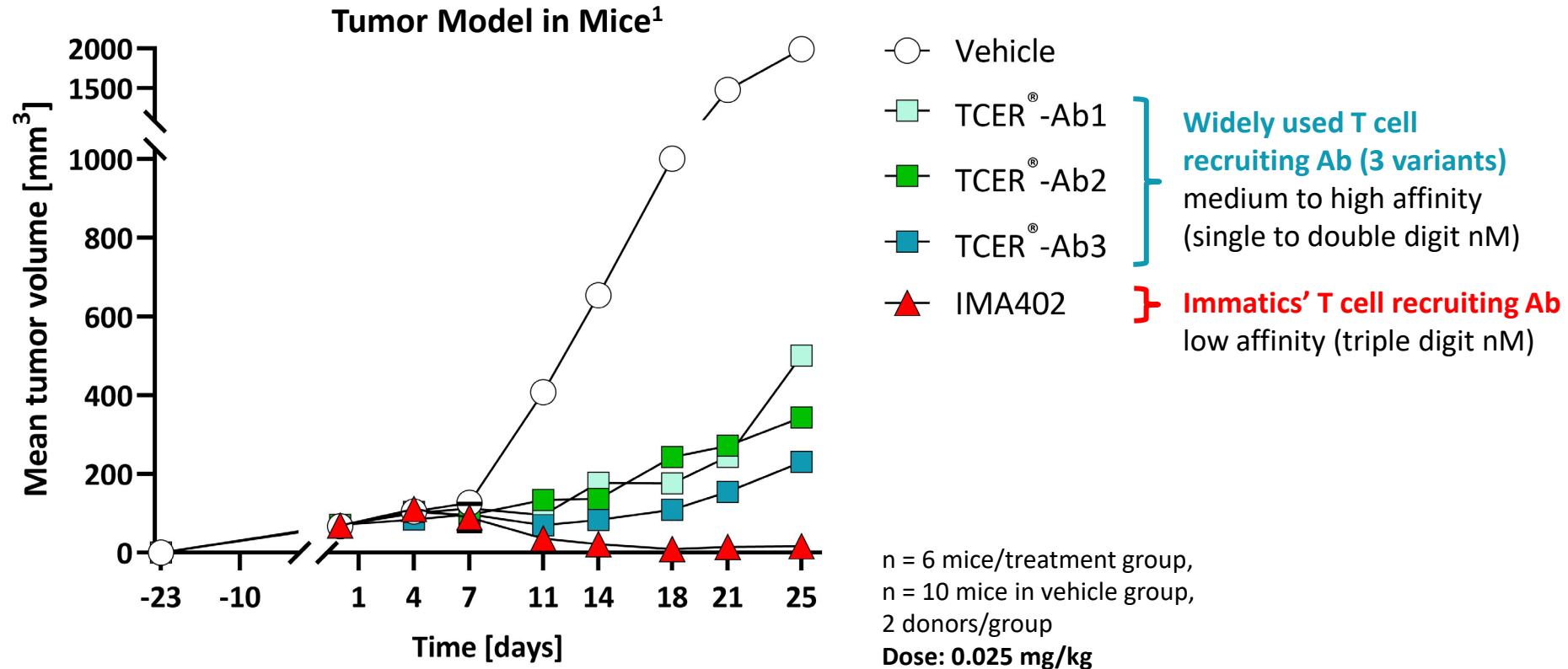
- Seven different TCR Bispecific formats were evaluated with a pHLA targeting TCR and the identical T cell recruiting antibody
- TCER® format had higher combination of potency and specificity<sup>1</sup> than six alternative TCR Bispecific format designs evaluated

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



# TCER® Format Is Designed for Optimized Efficacy and Safety

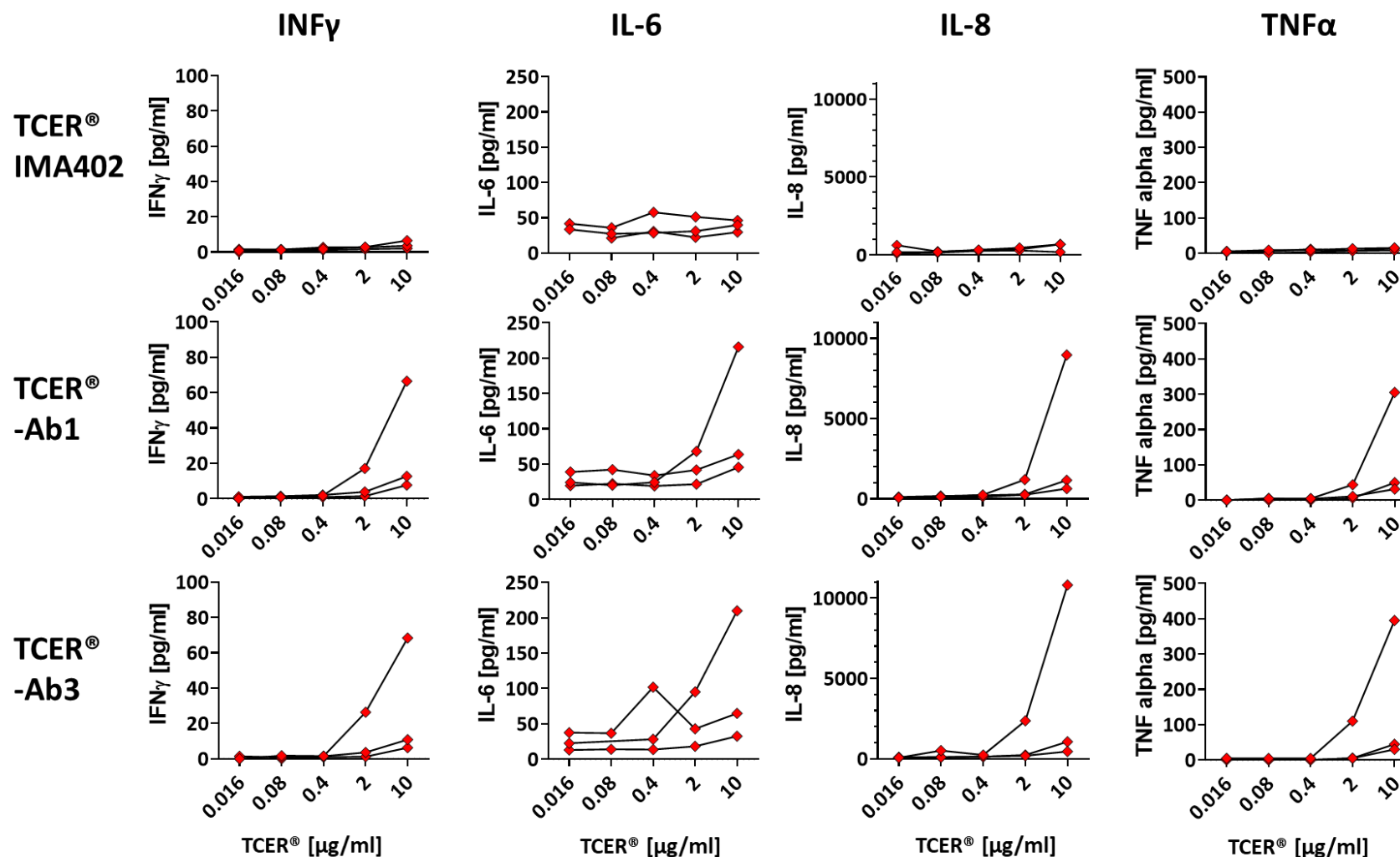
## Superior Tumor Control Using a Novel, Low-Affinity Recruiter



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® Format Is Designed for Optimized Efficacy and Safety

Reduced Target-Unrelated Recruiter-Mediated Cytokine Release using a Low-Affinity Recruiter



Whole blood cytokine release assay  
N=3 HLA-A\*02-positive donors  
N=16 cytokines tested,  
4 exemplary cytokines shown

# Our TCER® Portfolio

## Broad Pipeline of Next-Gen Half-Life Extended TCR Bispecifics

CLINICAL

**IMA401**

Bristol Myers Squibb

- MAGEA4/8 peptide presented by HLA-A\*02:01
- Dose escalation ongoing, first clinical data expected 2H 2024

**IMA402**

- PRAME peptide presented by HLA-A\*02:01
- Start of clinical trial in Aug 2023, first clinical data expected 2H 2024

PRECLINICAL

**IMA40x**

Several innovative programs

- Undisclosed peptides presented by HLA-A\*02:01 and other HLA-types
- TCER® engineering and preclinical testing ongoing

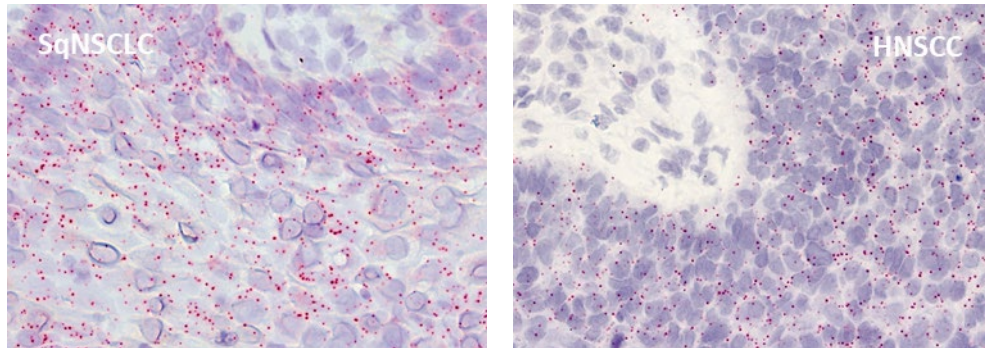
Potential for addressing different indications and large patient populations with novel, off-the-shelf TCR Bispecifics

The current collaboration with Moderna includes the development of mRNA-enabled *in vivo* expressed TCER® molecules

# TCER® IMA401 Targeting MAGEA4/8

Homogeneous Expression, Broad Prevalence and High Copy Number Target

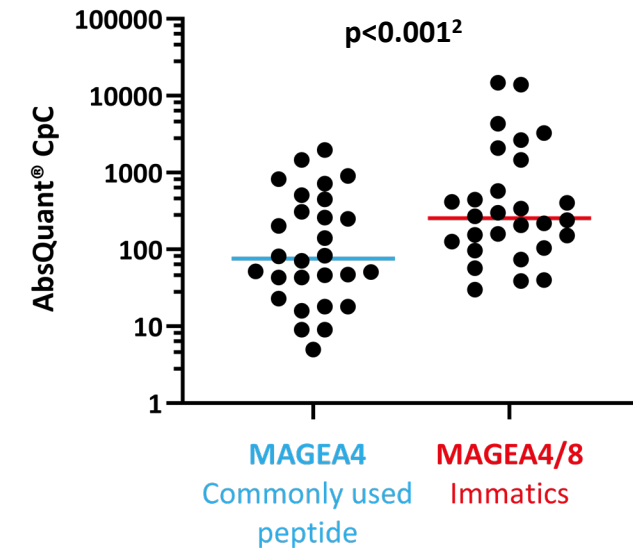
## MAGEA4 RNA detection in tumor samples (ISH)



## MAGEA4/8 target prevalence in selected cancer indications

Indications	Target prevalence [%]
Squamous non-small cell lung carcinoma	52%
Head and neck squamous cell carcinoma	36%
Bladder carcinoma	29%
Uterine carcinosarcoma	29%
Esophageal carcinoma	23%
Ovarian carcinoma	23%
Melanoma	18%
<i>plus several further indications</i>	

## MAGEA4 and MAGEA4/8 Peptide (AbsQuant®)



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

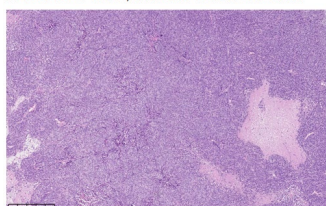
# TCER® IMA401 (MAGEA4/8) – Assessment of Anti-Tumor Activity *in vitro*

## Patient-Derived Tumor Model

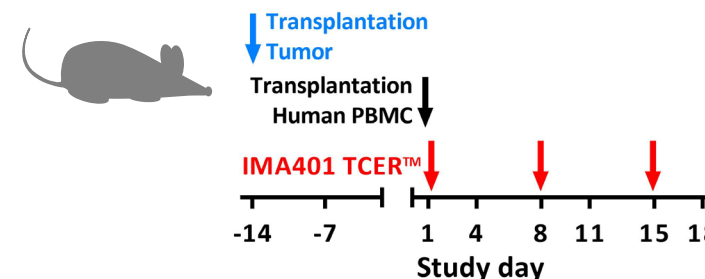
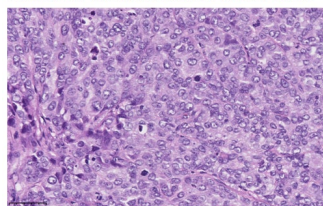
### NSCLC adenocarcinoma:

- Male, Caucasian, age 58, no therapy prior to surgery
- Site of origin: lung, differentiation poor
- Date of surgery: 1987, Freiburg Medical Center
- Volume doubling time: 7.3 day
- Histology:
  - Stroma content, 4%
  - Vascularization, high
  - Grading, undifferentiated

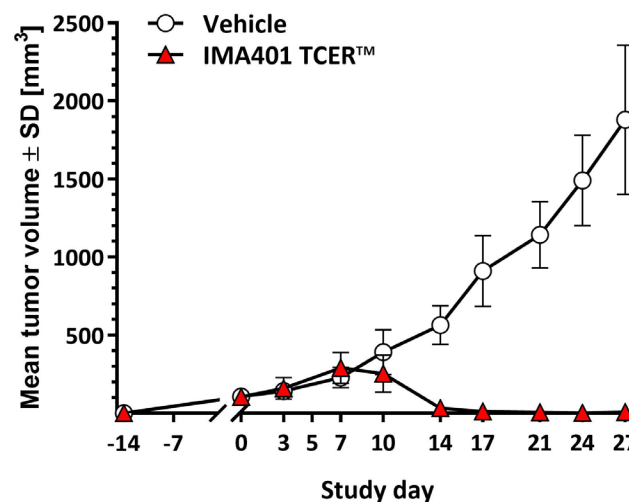
PASSAGE: 12N2, MAGNIFICATION: 5.0X



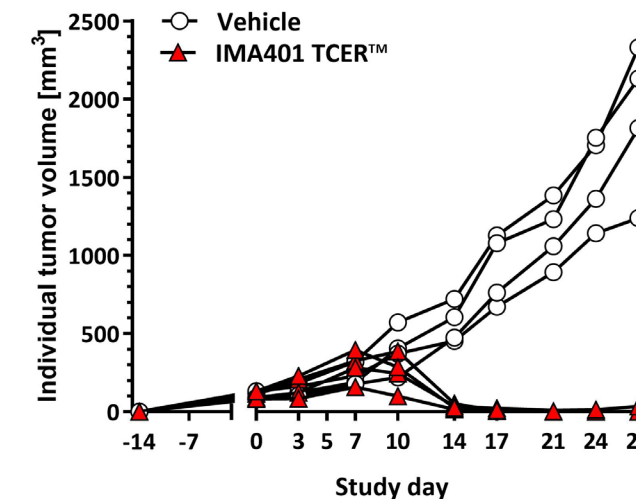
PASSAGE: 12N2, MAGNIFICATION: 40.0X



Group averages



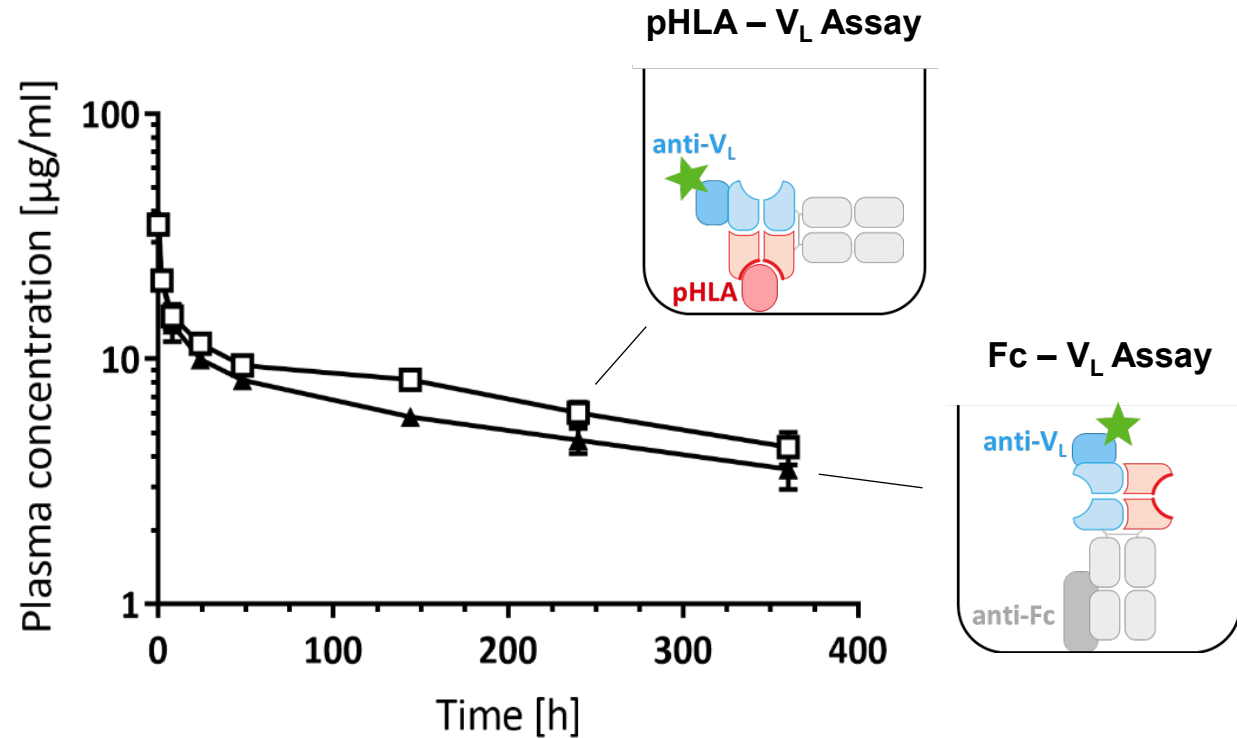
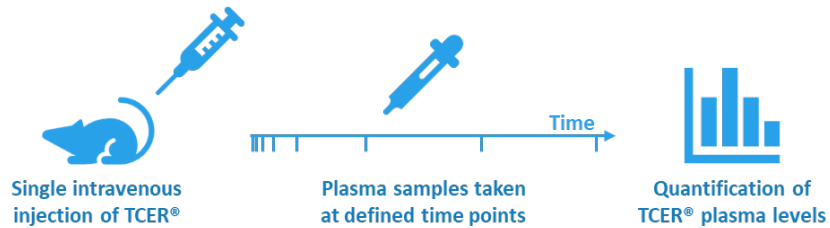
Individual mice  
Two PBMC donors



- TCER® IMA401 shows **high anti-tumor activity** in **Patient-derived xenograft model** of non-small cell lung adenocarcinoma
- **Remission observed in all mice (3 out of 4 mice with complete remission)**

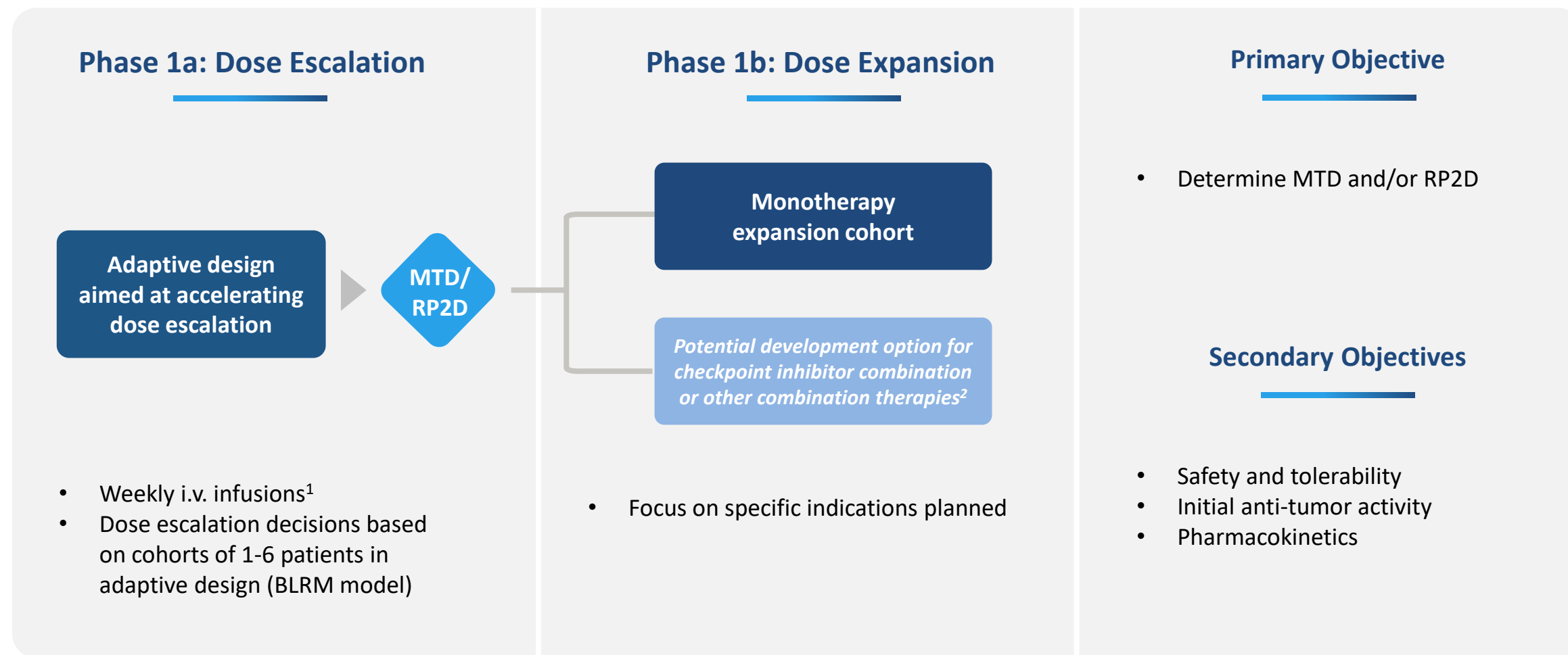
# TCER® IMA401 (MAGEA4/8) – Pharmacokinetics

## PK Analysis in NOG Mice



- Two different PK assays established to ensure functional integrity of protein domains
- **Terminal half-life in mice: 10-11 days**

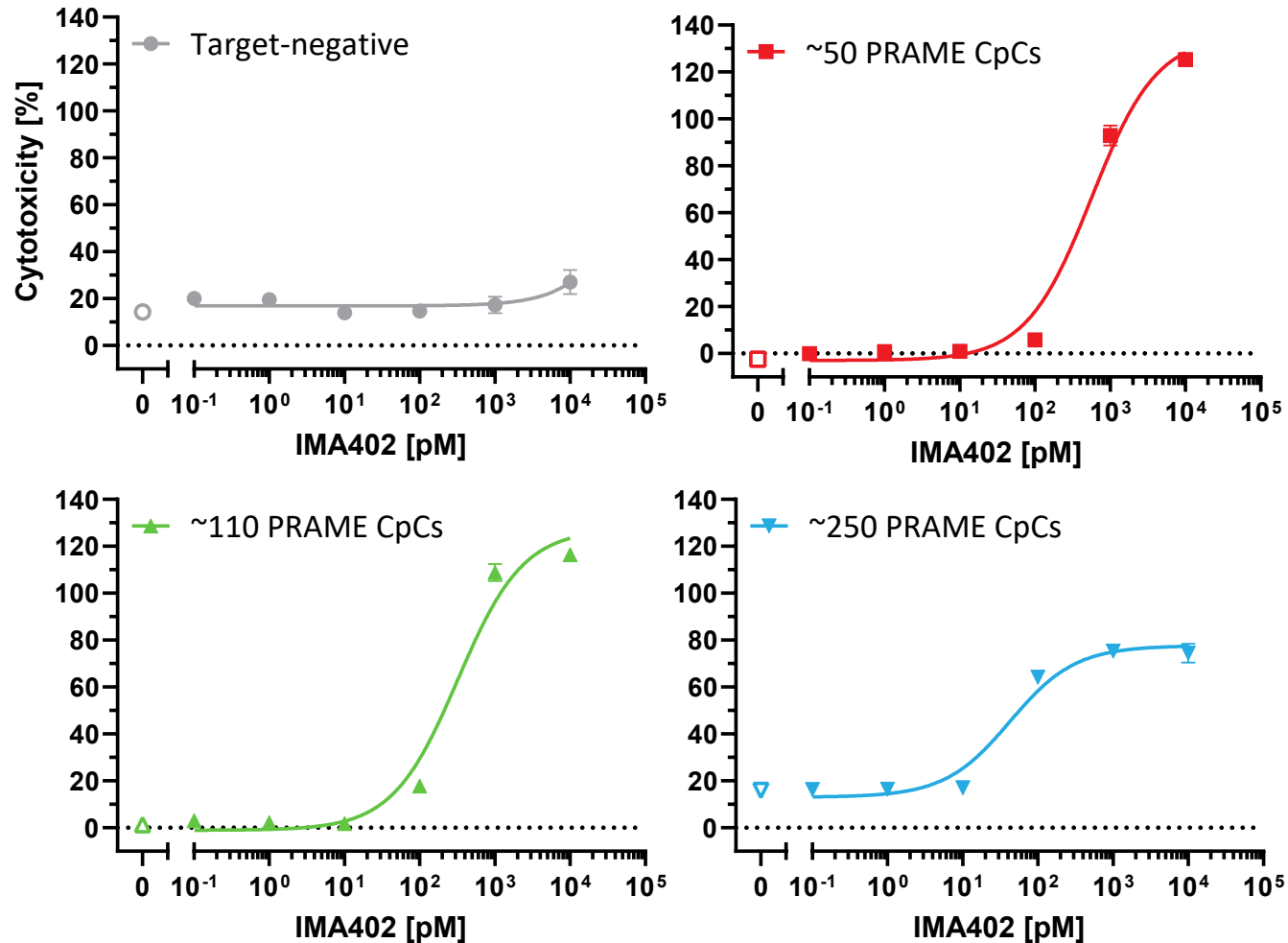
# Phase 1 Clinical Trial to Evaluate TCER<sup>®</sup> IMA401 Targeting MAGEA4/8





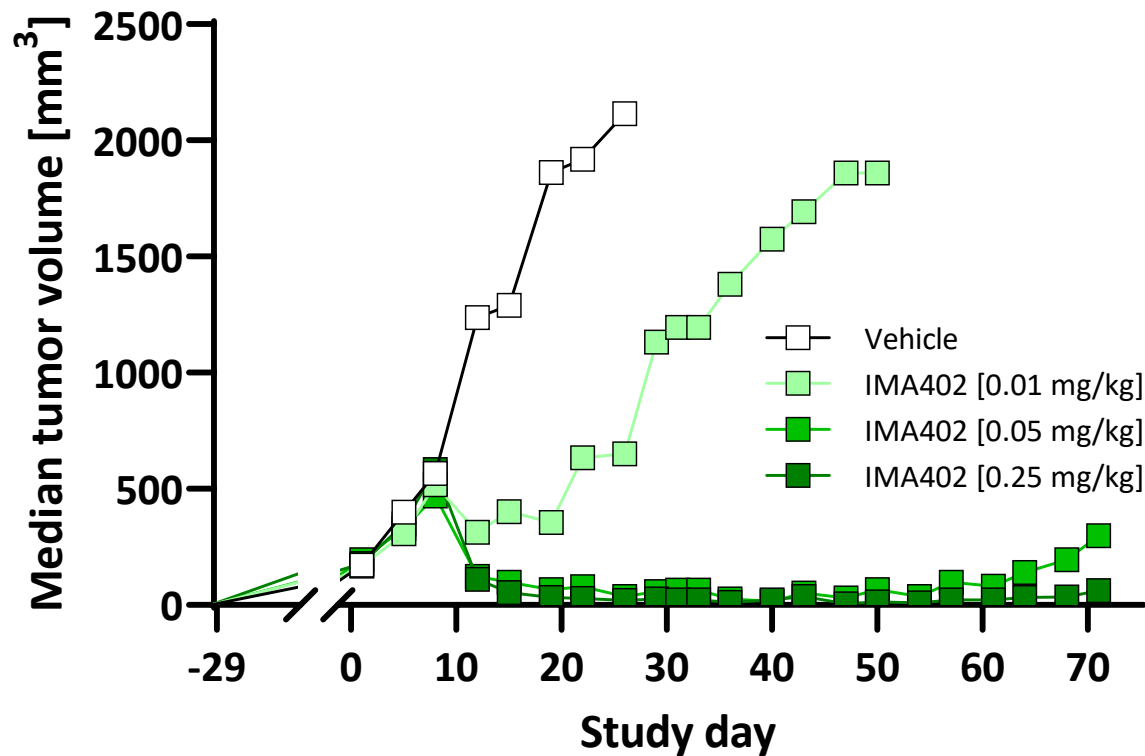
# TCER® IMA402 Targeting PRAME – Efficacy Assessment *in vitro*

## Tumor Cell Killing at Low Physiological PRAME Peptide Levels



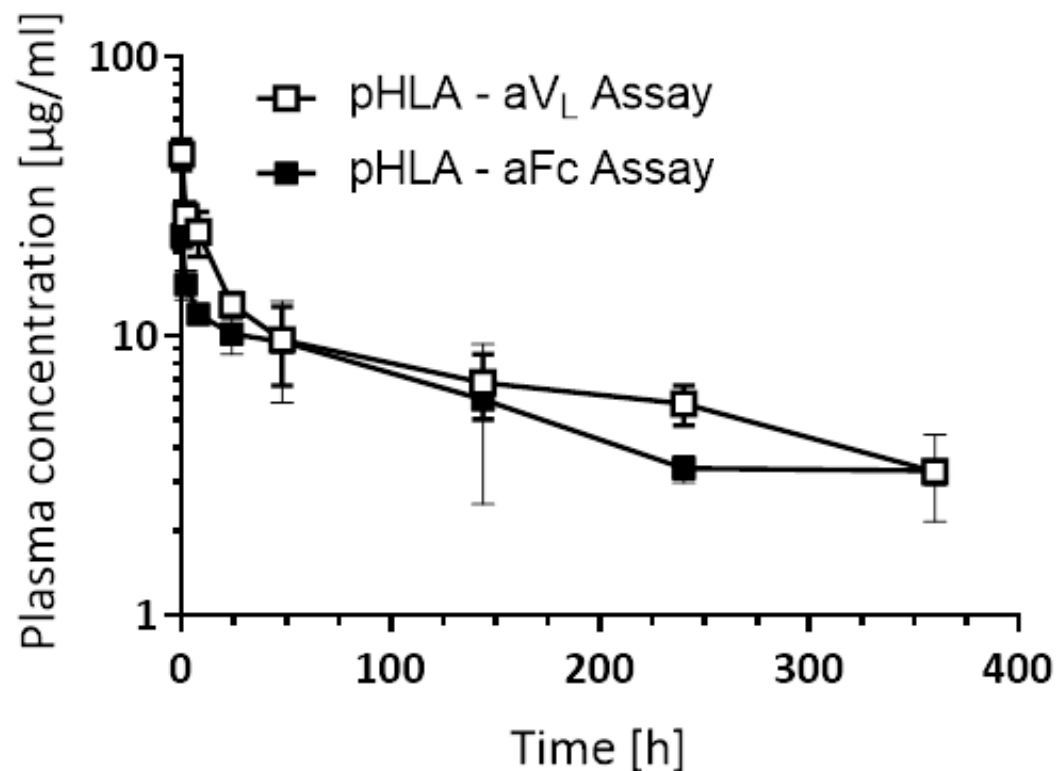
- TCER® IMA402 induces killing of tumor cells with PRAME target copies as low as 50 CpCs
- Physiological PRAME levels detected in majority of cancer tissues from patients are 100 – 1000 CpCs
- Preclinical activity profile enables targeting of a broad variety of tumor indications, such as lung cancer, breast cancer, ovarian cancer, uterine cancer, melanoma and others

# TCER® IMA402 Achieves Durable Tumor Control of Large Tumors *in vivo*

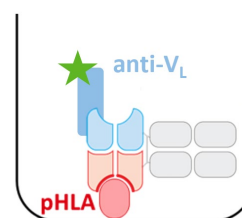


- Dose-dependent efficacy of IMA402 in cell line-derived *in vivo* mouse model
- Durable shrinkage of large tumors including complete responses over prolonged period
- Sufficiently high drug doses are key to achieving desired anti-tumor effect

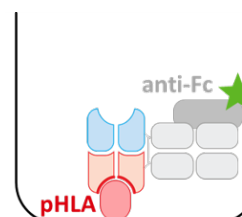
# Half-life Extended Format of IMA402 Confers Terminal Half-life of >1 Week



pHLA – aV<sub>L</sub> Assay



pHLA – aFc Assay



- IMA402 shows a terminal serum half-life of  $\approx 8$  days in mice
- IMA402 will be initially dosed weekly in the clinical trial
- Dosing frequency may be adapted based on clinical data

# Phase 1/2 Clinical Trial to Evaluate TCER® IMA402 Targeting PRAME

First Clinical Data Planned in 2H 2024

## Trial Overview

Phase 1/2 clinical trial to evaluate safety, tolerability and anti-tumor activity of IMA402

- HLA-A\*02:01-positive patients with PRAME-expressing recurrent and/or refractory solid tumors
- Initially weekly i.v. infusions
- Potential for early adjustment of treatment interval based on PK data of half-life extended TCER® format

## Phase 1: Dose Escalation

Adaptive design aimed at accelerating dose escalation

MTD/  
RP2D

- Basket trial in focus indications to accelerate signal finding
- Ovarian cancer, lung cancer, uterine cancer, melanoma, others

## Phase 2a: Dose Expansion

Expansion cohort

Expansion cohort

Expansion cohort

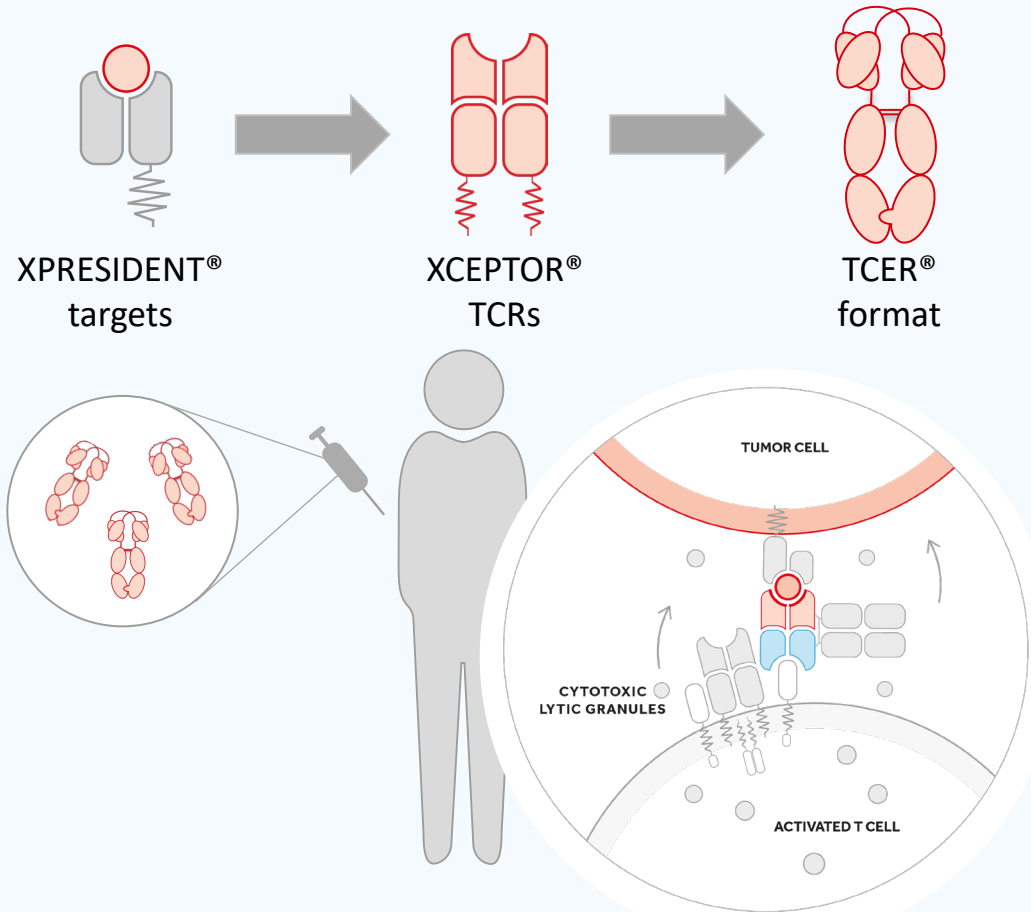
- Specific indications plus ongoing basket
- Combination therapies
- Optional dose/application optimization

# In Vivo Expressed TCER<sup>®</sup> Molecules Targeting Cancer-specific pHLA Targets

Combining Immatics' Target and TCR Platforms with Moderna's mRNA Technology

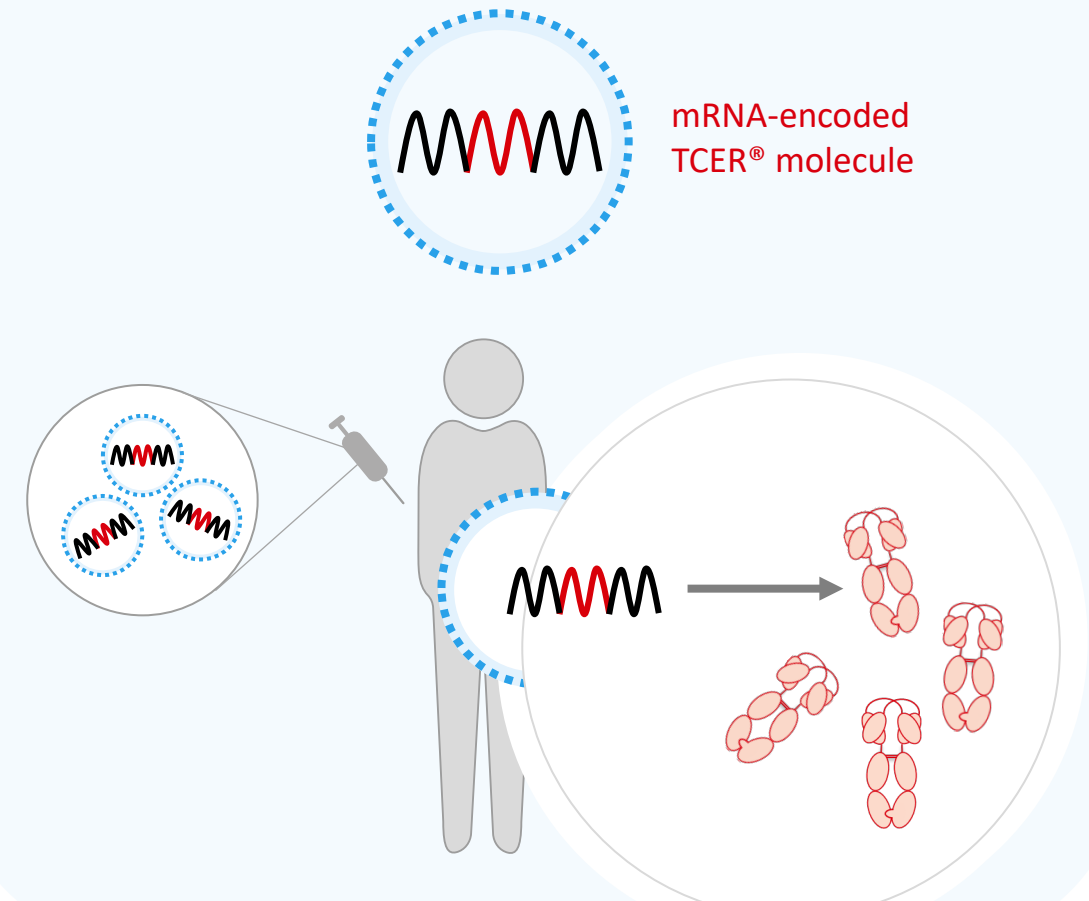
## Immatics

Proprietary cancer targets & TCR Bispecifics format



## Moderna

Delivery of TCER<sup>®</sup> biologics through mRNA

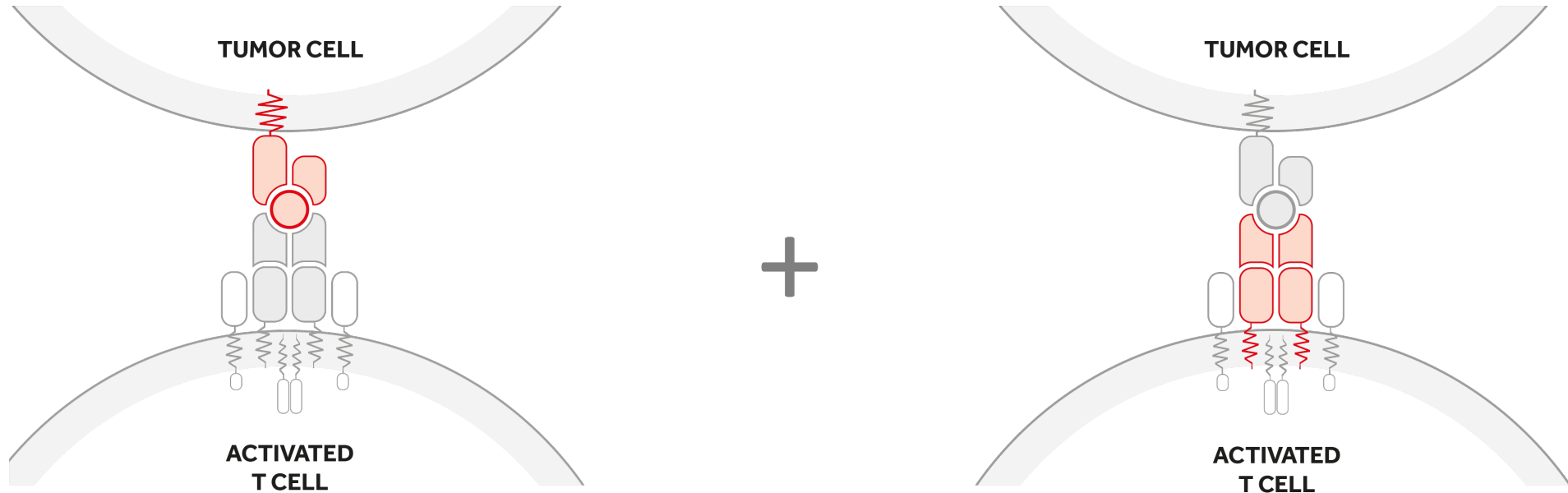




## Immatics' Proprietary Target and TCR Discovery Platforms

# 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

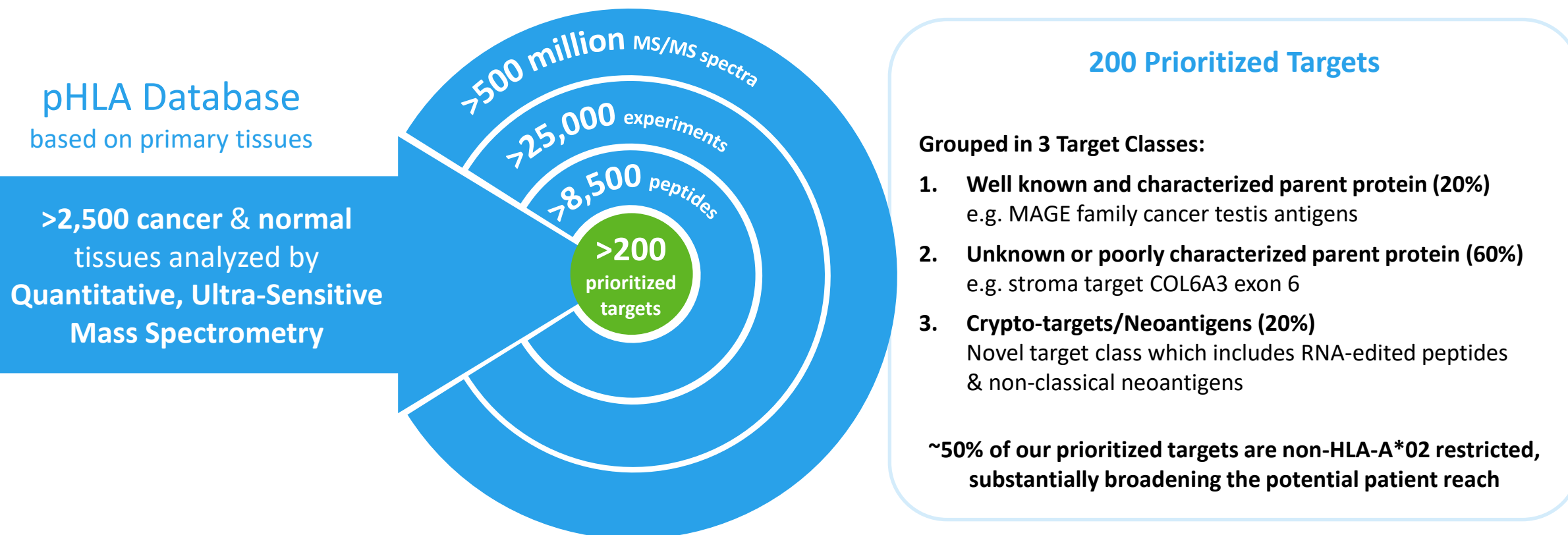
## Right TCRs via XCEPTOR® technology platform

- 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



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

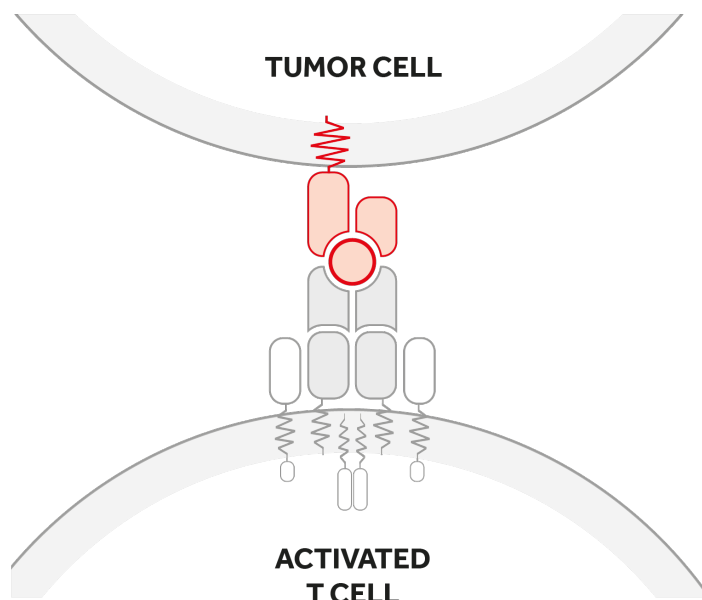
XPRESIDENT® Target Platform



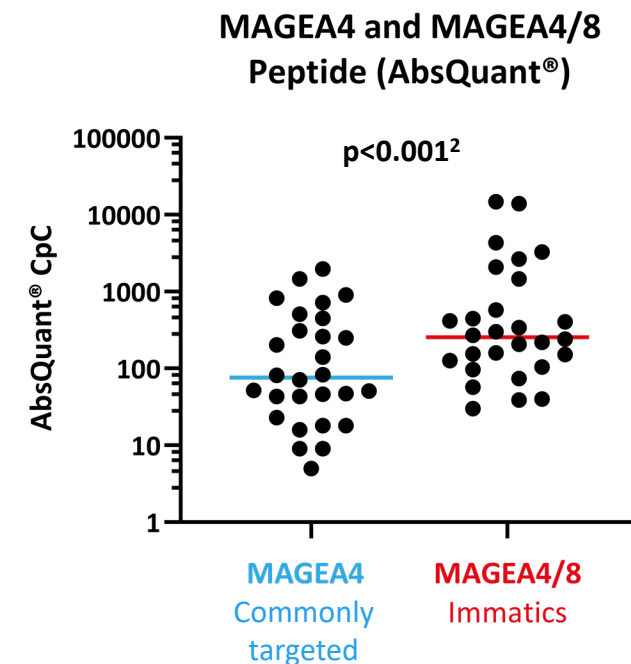
This large data set is leveraged by our bioinformatics & AI-platform XCUBE™ – „AI is where the data is®“

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

## Example of MAGEA4/8 Peptide Target



Ranking of  
pHLA targets

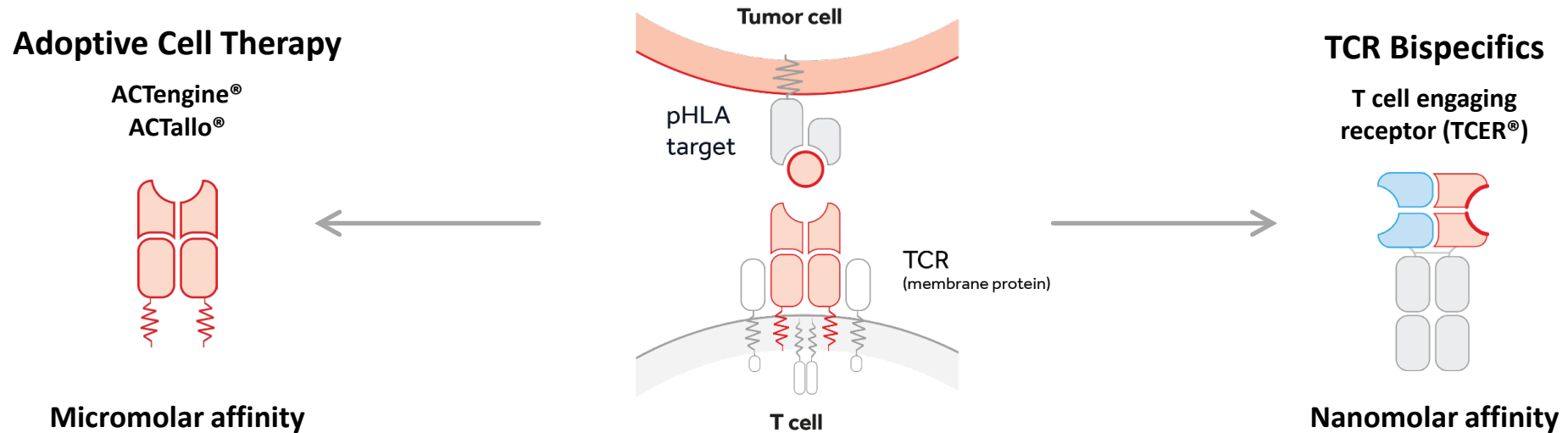


XPRESIDENT® quantitative information on target density<sup>1</sup> between peptides originating from the same source protein

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

# Development of the Right TCR – XCEPTOR® Technology

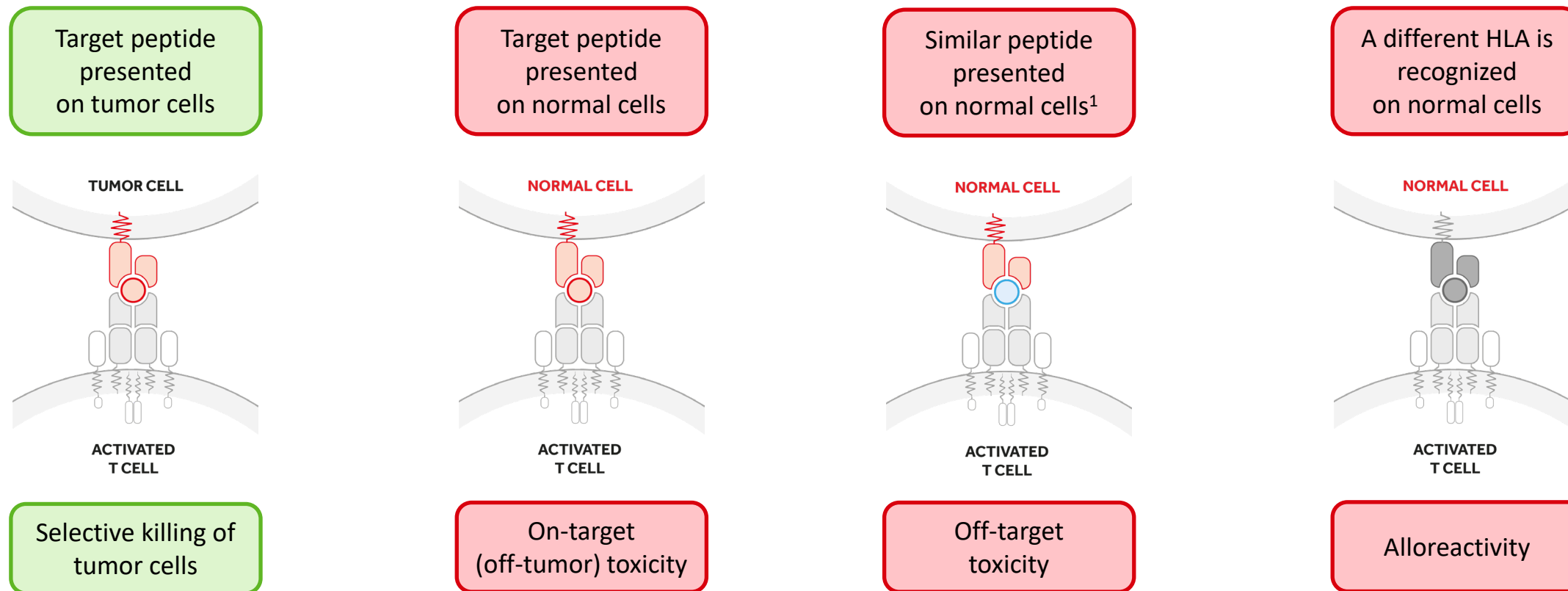
## TCR Discovery and Engineering for ACT and TCR Bispecifics



- 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> (empowered by our bioinformatics & AI-platform XCUBE™)

# Optimal Target Selection & TCR Specificity for Minimizing Safety Risks

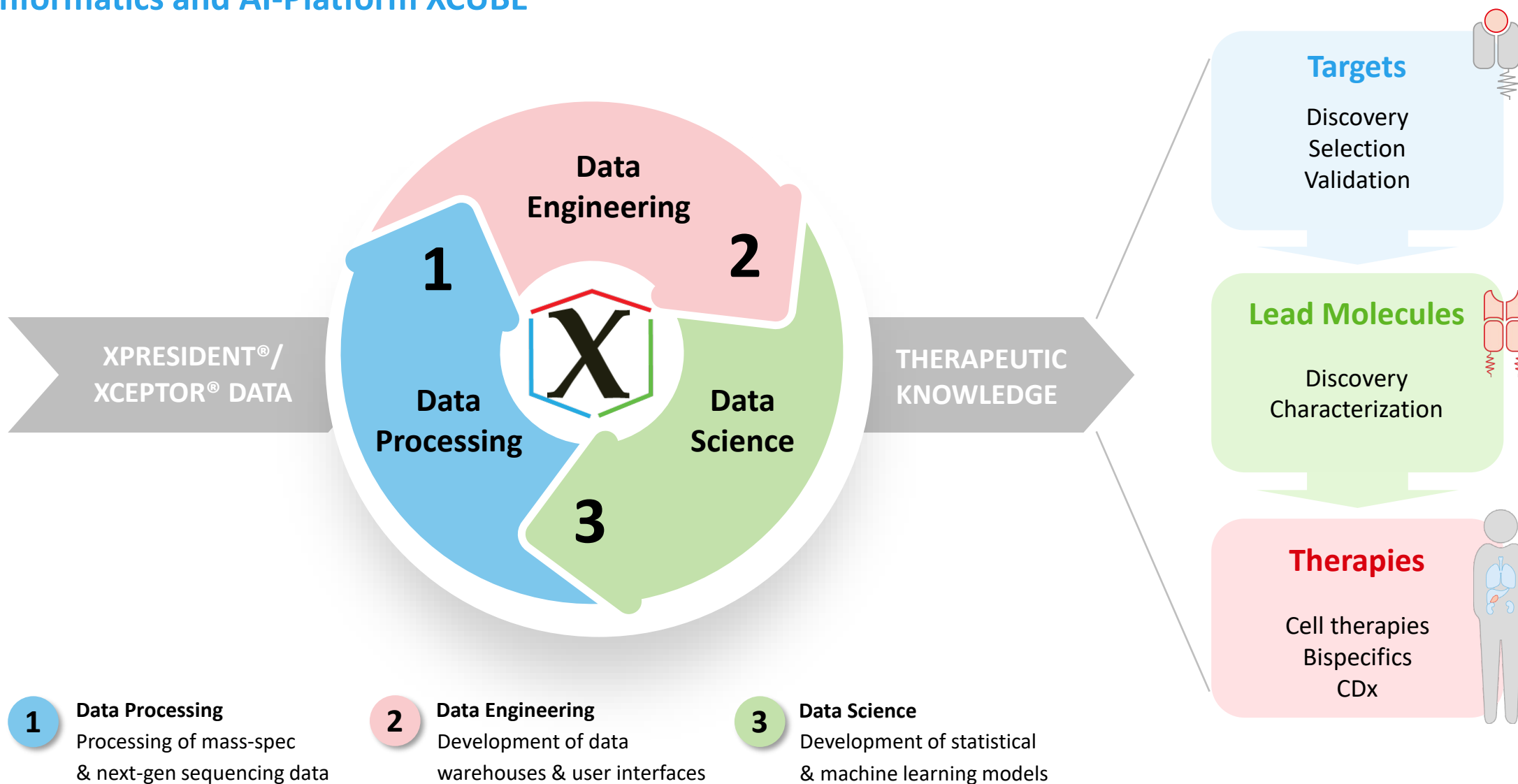
Unique Interplay between Technology Platforms Allows Early De-risking for Clinical Development



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

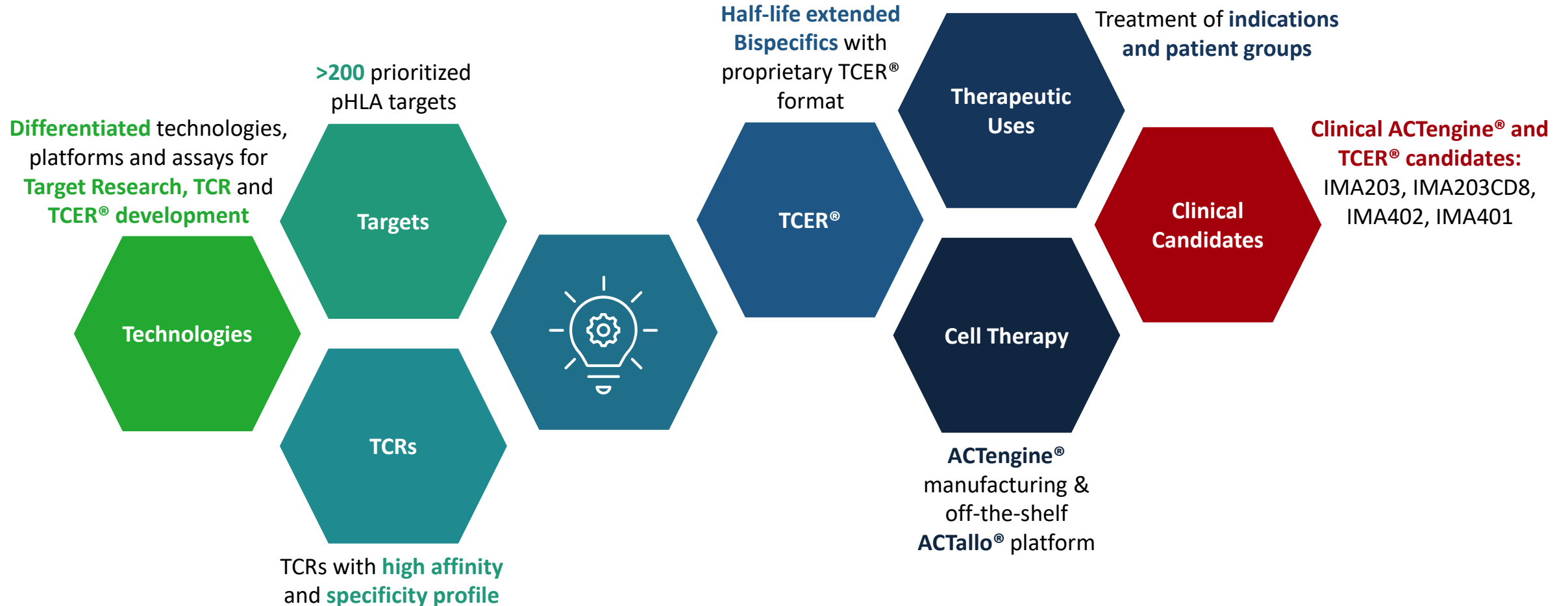
# “AI Is Where the Data Is®”

Bioinformatics and AI-Platform XCUBE™



# Immatics' Robust Intellectual Property Portfolio

## Protection Strategy of Key Assets in Major Markets and Beyond





## 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  
(InflaRx, Medigene, NovImmune, Probiobug)



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



**Cedrik Britten**  
**Chief Medical Officer**  
>15 yrs pharma & biotech experience  
(GSK, BioNTech)



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



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



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



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



**Jordan Silverstein**  
**Head of Strategy**  
>10 yrs biotech experience  
(InflaRx, AAA)

# Strong, Focused and Highly Integrated Trans-Atlantic Organization



# Delivering

the Power of T cells  
to Cancer Patients

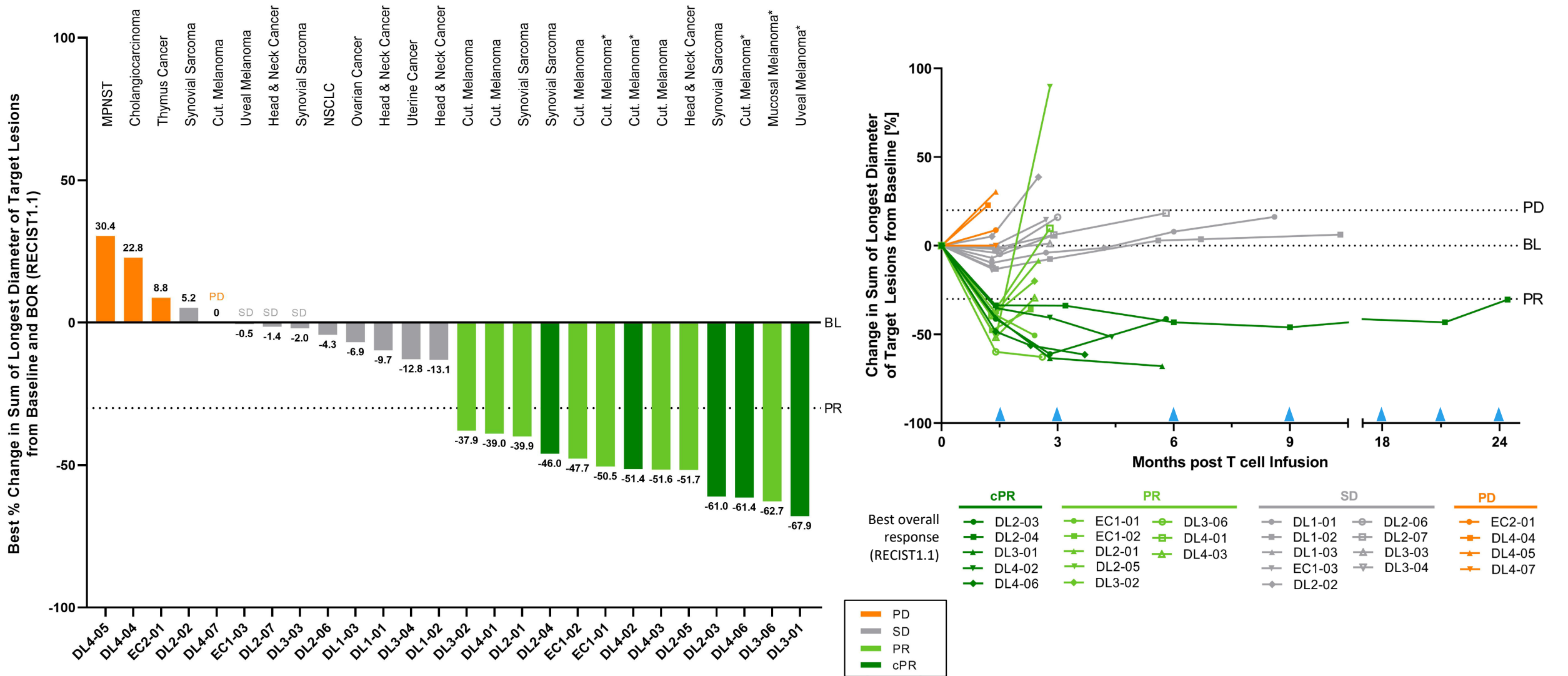
## Appendix

[www.immatics.com](http://www.immatics.com)





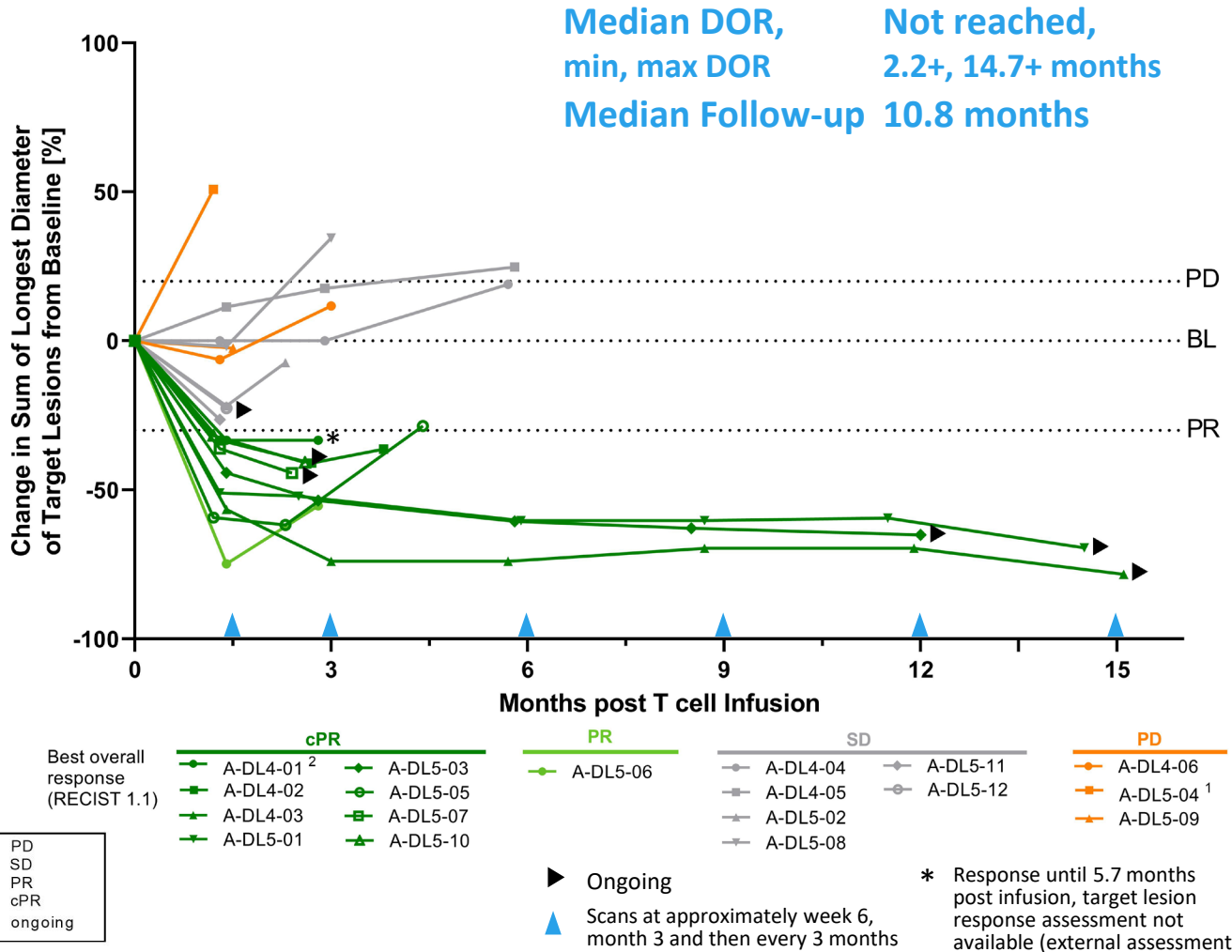
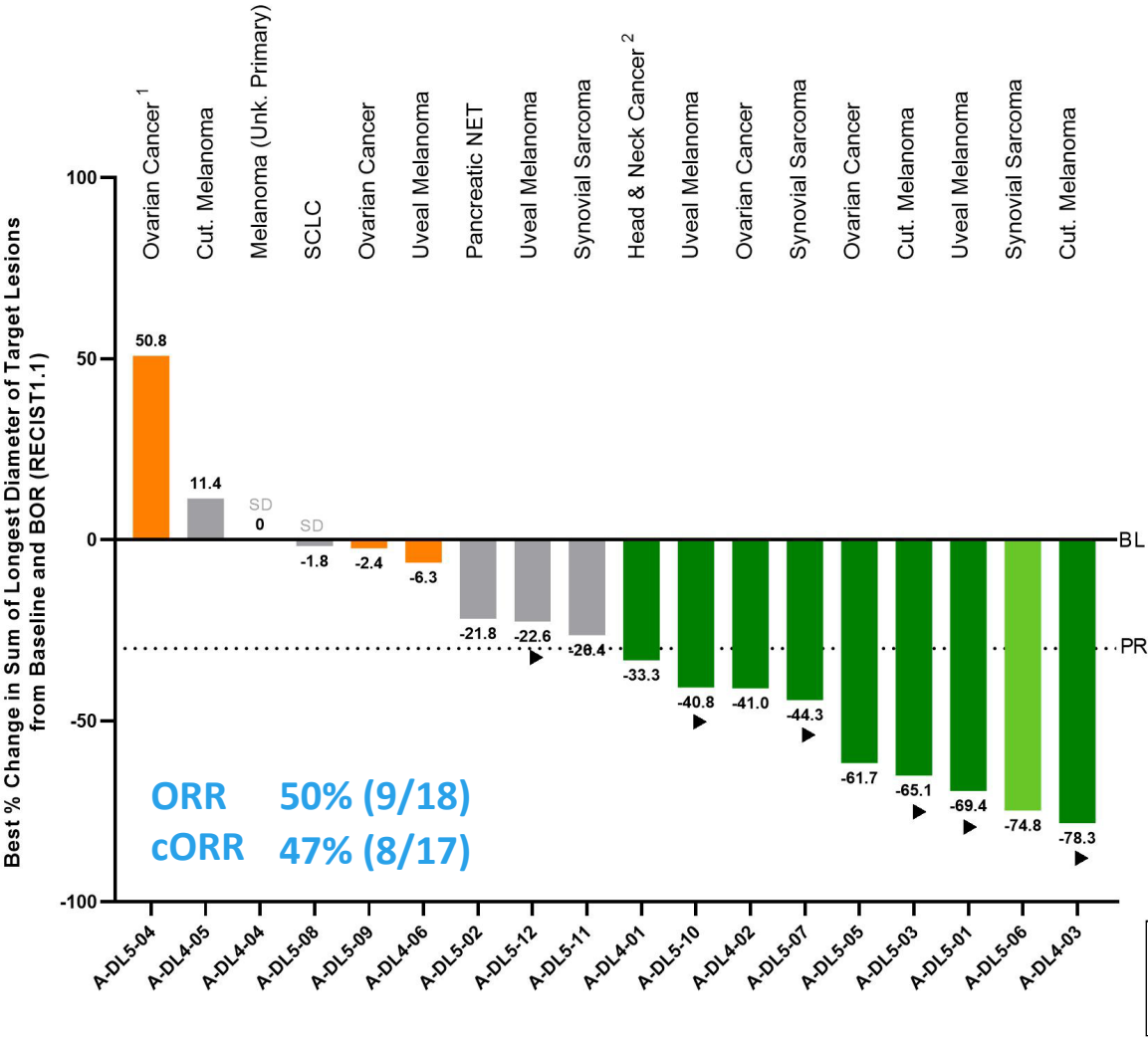
# IMA203 GEN1 in Phase 1a Dose Escalation (N=27#) – BOR and Response over Time





# IMA203 GEN1 in Cohort A (N=18) – BOR and Response over Time

## Objective Responses across Multiple Solid Cancer Types



<sup>1</sup> Patient received one dose nivolumab erroneously; <sup>2</sup> Progressive disease at month 6 due to unequivocal progression of non-target lesions, target lesions not evaluable due to external assessment; Initial ORR: Objective response rate according to RECIST 1.1 at any post infusion scan; Confirmed ORR (cORR): Confirmed objective response rate according to RECIST 1.1 for patients with at least two available post infusion scans or patients with progressive disease (PD) at any prior timepoint, patients with ongoing unconfirmed PR not included in cORR calculation; Duration of response (DOR) in confirmed responders is defined as time from first documented response until disease progression/death. Patients with ongoing unconfirmed PR not included in cORR calculation; Median Follow-up is analyzed by using the reverse Kaplan-Meier method; PD: Progressive Disease; SD: Stable Disease; PR: Partial Response; cPR: Confirmed Partial Response; BL: Baseline; BOR: Best Overall Response; DOR: Duration of Response

Data cut-off Sep 30, 2023 70

# IMA203 GEN1 in Cohort A – Most Frequent Adverse Events

N=21 Patients in Safety Population<sup>1</sup>

- **Expected cytopenia (Grade 1-4)** associated with lymphodepletion in all patients
- **Mild-moderate cytokine release syndrome (CRS) in 90% (19/21) of patients**
  - 43% (9/21) with Grade 1 CRS
  - 48% (10/21) with Grade 2 CRS
  - No dose-dependent increase of CRS
- **One non-serious, mild (Grade 1) ICANS<sup>2</sup> in DL5**
- **No dose-limiting toxicity**
- **No IMA203-related deaths**

**IMA203 GEN1 monotherapy continues to be well tolerated at total doses between 1-10x10<sup>9</sup> TCR-T cells (RP2D)**

<sup>1</sup> Three cutaneous melanoma patients treated with IMA203, and pending post infusion scan included in safety population, but not efficacy population;

<sup>2</sup> ICANS: Immune Effector Cell-Associated Neurotoxicity Syndrome; CRS and ICANS graded by CARTOX criteria (Neelapu *et al.*, 2018)

# IMA203 GEN1 at RP2D – Tolerability Data

## Phase 1a DL4 and Cohort A – All ≥Grade 3 Adverse Events (N=28)

TEAEs by maximum severity for all patients in Ph1a dose escalation DL4 and Ph1b Cohort A dose expansion (RP2D, N=28) <sup>1</sup>					
Adverse event (System organ class, Preferred term)	≥ Grade 3		Adverse event (System organ class, Preferred term)	≥ Grade 3	
	No.	%		No.	%
Patients with any adverse event	28	100.0	table continued...		
Adverse Events of Special Interest	1	3.6	General disorders and administration site conditions	1	3.6
Cytokine release syndrome	1	3.6	Pyrexia	1	3.6
ICANS <sup>2</sup>	0	0.0	Hepatobiliary disorders	1	3.6
Blood and lymphatic system disorders	27	96.4	Cholangitis	1	3.6
Neutropenia	18	64.3	Injury, poisoning and procedural complications	1	3.6
Anaemia	14	50.0	Humerus fracture	1	3.6
Leukopenia	13	46.4	Musculoskeletal and connective tissue disorders	1	3.6
Lymphopenia	11	39.3	Muscle spasms	1	3.6
Thrombocytopenia	9	32.1	Nervous system disorders	1	3.6
Leukocytosis	1	3.6	Headache	1	3.6
Lymphocytosis	1	3.6	Skin and subcutaneous tissue disorders	1	3.6
Investigations	7	25.0	Rash maculo-papular	1	3.6
Neutrophil count decreased	4	14.3			
Alanine aminotransferase increased	2	7.1			
Aspartate aminotransferase increased	2	7.1			
White blood cell count decreased	2	7.1			
Blood alkaline phosphatase increased	1	3.6			
Infections and infestations	3	10.7			
Infection	1	3.6			
Septic shock <sup>3</sup>	1	3.6			
Urinary tract infection	1	3.6			
Respiratory, thoracic and mediastinal disorders	3	10.7			
Hypoxia	2	7.1			
Laryngeal inflammation	1	3.6			
Vascular disorders	3	10.7			
Hypotension	2	7.1			
Hypertension	1	3.6			
Metabolism and nutrition disorders	2	7.1			
Failure to thrive	1	3.6			
Hypokalaemia	1	3.6			
Hypophosphataemia	1	3.6			
Eye disorders	1	3.6			
Ulcerative keratitis	1	3.6			

All treatment-emergent adverse events (TEAEs) with ≥ Grade 3 regardless of relatedness to study treatment that occurred in at least 1 patient (except for ICANS, where only Grade 1-2 occurred; listed for completeness due to being an adverse event of special interest) are presented. 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, version 5.0. Grades for CRS and ICANS were determined according to CARTOX criteria (Neelapu et al., 2018). Patients are counted only once per adverse event and severity classification. Based on interim data extracted from open clinical database (30-Sep-2023); <sup>1</sup> One patient in Phase 1a DL4 with disease progression after first IMA203 infusion received exploratory second IMA203 infusion and had these ≥ Grade 3 TEAEs only after second infusion, which are included in the table: Humerus fracture, Muscle spasms, Neutropenia, Thrombocytopenia; <sup>2</sup> ICANS: Immune effector cell-associated neurotoxicity syndrome; <sup>3</sup> Fatal Adverse events were not considered related to any study drug

- IMA203 was well tolerated at doses as high as ~10x10<sup>9</sup> TCR-T cells
- Most frequent ≥Grade 3 AEs were expected cytopenia associated with lymphodepletion
- No IMA203-related Grade 5 AEs



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