# **Immatics Corporate Presentation**

January 13, 2025



#### Delivering the Power of T cells to Cancer Patients

 $\ensuremath{\mathbb{C}}$  Immatics. Not for further reproduction or distribution.

#### **Forward-Looking Statement**



This presentation ("Presentation") is provided by Immatics N.V. ("Immatics" or the "Company") for informational purposes only. The information contained herein does not purport to be allinclusive and none of Immatics, any of its affiliates, any of its or their respective control persons, officers, directors, employees or representatives makes any representation or warranty, express or implied, as to the accuracy, completeness or reliability of the information contained in this Presentation.

Forward-Looking Statements. Certain statements in this presentation may be considered forward-looking statements. Forward-looking statements generally relate to future events or the Company's future financial or operating performance. For example, statements concerning timing of data read-outs for product candidates, the timing, outcome and design of clinical trials, the nature of clinical trials (including whether such clinical trials will be registration-enabling), the timing of IND or CTA filing for pre-clinical stage product candidates, the timing of BLA filings for clinical stage product candidates, estimated market opportunities of product candidates, manufacturing timetables, capacity and success rates, the Company's focus on partnerships to advance its strategy, and other metrics are forward-looking statements. In some cases, you can identify forward-looking statements by terminology such as "may", "should", "expect", "plan", "target", "intend", "will", "estimate", "anticipate", "believe", "predict", "potential" or "continue", or the negatives of these terms or variations of them or similar terminology. Such forward-looking statements. These forward-looking statements are subject to risks, uncertainties, and other factors which could cause actual results to differ materially from those expressed or implied by such forward-looking statements. These forward-looking statements are based upon estimates and assumptions that, while considered reasonable by Immatics and its management, are inherently uncertain. New risks and uncertainties may emerge from time to time, and it is not possible to predict all risks and uncertainties. Factors that may cause actual results to differ materially from current expectations include, but are not limited to, various factors beyond management's control including general economic conditions and other risks, uncertainties and factors set forth in the Company's Annual Report on Form 20-F and other filings with the Securities and Exchange Commission (SEC). Nothing in this prese

No Offer or Solicitation. This communication is for informational purposes only and does not constitute, or form a part of, an offer to sell or the solicitation of an offer to sell or an offer to buy or the solicitation of an offer to buy any securities, and there shall be no sale of securities, in any jurisdiction in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such jurisdiction. No offer of securities shall be made except by means of a prospectus meeting the requirements of Section 10 of the Securities Act of 1933, as amended, or in an offering exempt from registration.

Certain information contained in this Presentation relates to or is based on studies, publications, surveys and the Company's own internal estimates and research. In addition, all of the market data included in this presentation involves a number of assumptions and limitations, and there can be no guarantee as to the accuracy or reliability of such assumptions. Finally, while the Company believes its internal research is reliable, such research has not been verified by any independent source. All the scientific and clinical data presented within this presentation are – by definition prior to completion of the clinical trial and a clinical study report – preliminary in nature and subject to further quality checks including customary source data verification.

### **Strategic Priorities in 2025 and Beyond**

Strong Cash Position into 2H 2027 to Deliver on Pipeline

# Commercializing PRAME cell therapy

in 2L cutaneous melanoma

#### RMAT designation<sup>1</sup> by FDA received

Phase 3 SUPRAME trial initiated; Primary endpoint: PFS for full approval

Large addressable patient population: 7,300<sup>\*</sup> 2L patients in US & EU5

Commercial buildout initiated including in-house state-of-the-art TCR-T manufacturing

#### **EXPECTED MILESTONES**

- Interim data read-out: **1Q26**
- Final read-out: 4Q26
- BLA submission: **1Q27**
- Launch: **3Q27**

#### **Expanding the PRAME** commercial opportunity

to earlier lines and additional solid cancer types

IMA203 expansion into **uveal melanoma** through ongoing Phase 1b trial

IMA203CD8 GEN2 in **PRAME+ solid cancers**, starting with gynecologic cancers

IMA402 in 1L cut. melanoma, gynecologic cancers, sqNSCLC, breast cancer & others

#### **EXPECTED MILESTONES**

IMA203 Ph1b data in uveal melanoma: 2025

IMA203CD8 Ph1a data incl. ovarian cancer: 2025

IMA402 Ph1a data in 2L melanoma: 2025

# Leveraging the potential of our proprietary platform

to provide innovative therapeutics and unlock more cancer types

IMA401 in 1L sqNSCLC, HNSCC, bladder cancer & others

Multiplexing of TCR Bispecifics covering multiple targets including PRAME, MAGEA4/8 & other undisclosed targets

Advancement of **mRNA-encoded TCER® molecules** in collaboration with Moderna

#### **EXPECTED MILESTONES**

IMA401 Ph1b data with HNSCC focus: 2025

IMA401 Ph1b data with sqNSCLC focus: 2026



<sup>1</sup> Includes all benefits of Breakthrough Therapy Designation; \* PRAME\*/HLA-A\*02:01\* addressable patient population, source: Clarivate Disease Landscape and Forecast 2025; 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; EU5: France, Germany, Italy, Spain, United Kingdom; PFS: progression-free survival; BLA: Biologics license application; sqNSCLC: squamous non-small-cell lung cancer, HNSCC: head and neck squamous cell carcinoma

## A Transformative Oncology Pipeline Across Modalities and Indications



#### Leveraging the Full Potential of 2 Therapeutic Modalities and 4 Clinical Candidates in Multiple Indications

Target	Product Candidate	Modality	Indication		Preclinical	Phase 1a <sup>1</sup>	Phase 1b <sup>1</sup>	Phase 2	Phase 3
	IMA203	ACTengine®	2L Melanoma	Immatics					
	IMA203	ACTengine®	Uveal melanoma	immatics					
DRAME	IMA203	ACTengine® + mRNA	Undisclosed	immatics moderna					
FICALIVIE		<b>ACTengine</b> <sup>®</sup>	Gynecologic cancers		_				
	(		Other solid cancers						
	IMA402	TCER®	Melanoma, others	immatics					
MAGEA4/8	IMA401	TCER®	HNSCC, sqNSCLC, others	immatics					
	IMA204	ACTengine®	COL6A3+ solid cancers	Immatics					
Other Targets	Undisclosed <sup>2</sup>	TCER®	Undisclosed	moderna					
	Undisclosed	ACTengine®	Undisclosed	ر <mark>ال</mark> ا Bristol Myers Squibb"					
	IMA30x	ACTallo®	Undisclosed	immatics editas <sup>3</sup>					

<sup>1</sup> Phase 1a: Dose escalation, Phase 1b: Dose expansion; <sup>2</sup> mRNA-enabled *in vivo* expressed TCER<sup>®</sup> molecules; <sup>3</sup> Immatics' proprietary ACTallo<sup>®</sup> platform utilizing Editas' CRISPR gene editing technology 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; HNSCC: head and neck squamous cell carcinoma; sqNSCLC: squamous non-small-cell lung cancer

## **Immatics' Clinical Portfolio**

#### 4 Clinically Active TCR Candidates Across 2 Modalities



5

	Cell Therapy			TCR Bispecifics			
	IMA203 (PRAME)	IMA203CD8 (PRAME)		IMA402 (PRAME)	IMA401 (MAGEA4/8)		
<b>Clinical Activity<sup>1</sup></b>	54% (14/26) cORR 12.1 months mDOR 6 months mPFS mOS not reached	41% (14/34) cORR 9.2 months mDOR		Initial clinical signal/first PRs observed and depending on target expression and TCER® dose	29% ORR and 25% cORR in patients with MAGEA4/8 <sup>high</sup> expression at relevant doses 53% DCR 53% tumor shrinkage		
Status	Phase 3 SUPRAME trial has commenced	Dose escalation ongoing		Early dose escalation ongoing	Dose escalation ongoing		
Positioning	Immatics' first TCR therapeutic to access market in 2L cut. melanoma, expansion to uveal melanoma as "add-on"	Enhanced pharmacology provides potential to expand PRAME cell therapy to tumor- agnostic label in PRAME+ solid cancers, starting with gynecologic cancers		Targeting 1L in cut. melanoma, gynecologic cancers, sqNSCLC, breast cancer & others	Targeting 1L sqNSCLC, HNSCC, bladder cancer & others		
Next Data Update	Phase 1b data in uveal melanoma 2025; Phase 3 interim data read- out in Q1 2026	Phase 1a data including ovarian cancer 2025		Phase 1a data to deliver clinical PoC in last-line 2025	Phase 1b data with HNSCC focus 2025		

Expanding the PRAME cell therapy opportunity to earlier lines and additional solid cancer types beyond melanoma Data updates on all clinical assets throughout 2025

<sup>1</sup> Data cut-off dates: IMA203: Aug 23, 2024; IMA203CD8: Sep 30, 2024; IMA402: Nov 6, 2024; IMA401: Jul 23, 2024; 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; cORR: confirmed objective response rate; mDOR: median duration of response; mPFS: median progression-free survival; OS: overall survival; PR: partial response; sqNSCLC: squamous non-small-cell lung cancer; HNSCC: head and neck squamous cell carcinoma; DCR: disease control rate

## **Breadth of PRAME Commercial Opportunity in Solid Cancers**



6

**Based on Positive Data and High Unmet Need** 

Intro



## Leadership in the Development of TCR-based Therapies



**Two Distinct TCR-based Therapeutic Modalities in Clinical Development** 

## **ACTengine®** (Autologous TCR-T)

Intro

## **TCER®** (TCR Bispecifics)



<sup>1</sup> Minimal target product profile (TPP) in monotherapy in 2L settings at recommended phase 2 dose (RP2D), i.e. typically in Ph1b dose expansion, for go-/no-go decision prior entering Ph2 or pivotal trial. Other factors such as mPFS (median progression-free survival) and mOS (median overall survival) in Ph1b vs. Ph1a may also contribute to decision-making. <sup>2</sup> Target prevalence is based on TCGA RNAseq data combined with a proprietary mass spec-guided RNA expression threshold; cORR: confirmed objective **7** response rate; mDOR: median duration of response; g2w: once every two weeks; sqNSCLC: squamous non-small-cell lung cancer, HNSCC: head and neck squamous cell carcinoma





## **ACTengine® IMA203 – TCR-Based Cell Therapy Targeting PRAME**

## The ACTengine® IMA203 Commercial Opportunity in 2L Melanoma



**TCR-Based Cell Therapy Targeting PRAME** 



IMA203 Opportunity

#### 2L Unresectable or Metastatic Cutaneous Melanoma

US	EU5	

~3.7k

~**3.6**k

#### Unresectable or Metastatic Uveal Melanoma



## ACTengine<sup>®</sup> IMA203 TCR-T Monotherapy Targeting PRAME in Melanoma



**Positive Data and High Unmet Need** 





#### SUPRAME Phase 3 trial in 2L melanoma commenced in December 2024

IMA203

<sup>1</sup> Includes all benefits of Breakthrough Therapy Designation; \* PRAME\*/HLA-A\*02:01\* addressable patient population, source: Clarivate Disease Landscape and Forecast 2025; CRS: cytokine release syndrome; ICANS: immune effector cell associated neurotoxicity syndrome; cORR: confirmed objective response rate; mDOR: median duration of response; mPFS: median progression-free survival; OS: overall survival; mFU: median follow-up; 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; EU5: France, Germany, Italy, Spain, United Kingdom

Data cut-off Aug 23, 2024 **10** 

## **ACTengine® IMA203 TCR-T Monotherapy – Patient Flow**





**IMA203** \* 30 mg/m<sup>2</sup> Fludarabine and 500 mg/m<sup>2</sup> Cyclophosphamide for 4 days; \*\* 1m IU daily days 1-5 and twice daily days 6-10, total dose is approx. only 5% of the overall dose for high-dose IL-2 given typically with TIL therapy (Sarnaik et al. 2021 Journal of Clinical Oncology)

## ACTengine<sup>®</sup> IMA203 TCR-T Trial in Melanoma



#### **Heavily Pretreated Patient Population**



	Total Safety Populati	on Melanoma Dose Escalatio Population	on Melanoma Efficacy Population <sup>1</sup>
	All Comers (Phase 1a and Phase 1	Melanoma b) (Phase 1a)	Melanoma (Phase 1b, at RP2D)
Number of patients	TotalN=70MelanomaN=41OtherN=29	TotalN=Cutaneous melanomaN=Uveal melanomaN=Mucosal melanomaN=	I1TotalN=283Cutaneous melanomaN=132Uveal melanomaN=121Melanoma ofN=1unknown primaryMucosal melanomaN=2
Prior lines of systemic treatment (median, min, max)	3 (0, 9)	4 (2, 7)	2 (0, 6)
<b>Thereof CPI</b> (melanoma only) (median, min, max)	2 (0, 4)	2 (1, 4)	1* (0, 4)
LDH at baseline >1 x ULN [% of patients]	64.3	81.8	60.7
<b>Baseline tumor burden</b> Median Target lesion sum of diameter [mm] (min, max)	117.8 (15.0, 309.8)	117.5 (37.0, 211.0)	107.5 (15.0, 309.8)
Liver/brain lesions at baseline [% of patients]	65.7	63.6	82.1
Dose level	DL1-5	EC1/DL3/4	DL4/5
<b>Total infused dose</b> TCR-T cells [x10 <sup>9</sup> ]	2.09 (0.08, 10.2)	0.586 (0.10, 2.09)	4.1 (1.3, 10.2)

IMA203 <sup>1</sup> All infused patients; \*Cutaneous melanoma patients had a median of 2 prior lines of checkpoints, see appendix; RP2D: recommended phase 2 dose; CPI: Checkpoint inhibitors; EC1: 0.06-0.12x10<sup>9</sup> TCR-T cells/m<sup>2</sup> BSA; DL3: 0.2-0.48x10<sup>9</sup> TCR-T cells/m<sup>2</sup> BSA, DL4: 0.2-1.2x10<sup>9</sup> TCR-T cells/m<sup>2</sup> BSA, DL5: 1.201 - 4.7x10<sup>9</sup> TCR-T cells/m<sup>2</sup> BSA

## Most Frequent Adverse Events of IMA203 Across All Dose Levels in Phase 1a/b



N=70 Patients Across All Dose Levels in Phase 1a/b (Total Safety Population)

- Most frequent adverse events were expected cytopenias (Grade 1-4) associated with lymphodepletion in all patients
- Mostly mild to moderate cytokine release syndrome (CRS)
  - 37% (26/70) Grade 1
  - 46% (32/70) Grade 2
  - 11% (8/70) Grade 3\*
- Infrequent ICANS (6% Grade 1, 4% Grade 2, 4% Grade 3)
- No IMA203-related deaths
- Tolerability in the melanoma subset is generally consistent with the full IMA203 monotherapy tolerability profile

Favorable tolerability profile for IMA203 monotherapy at recommended Phase 2 dose (1x10<sup>9</sup> to 10x10<sup>9</sup> TCR-T cells) supporting potential outpatient administration

## Tolerability Profile of IMA203 Across All Dose Levels in Phase 1a/b



All ≥Grade 3 Adverse Events (N=70<sup>1</sup>)

Adverse event	≥ Gra	ade 3
(System organ class, Preferred term)	No.	%
Patients with any adverse event	70	100.0
Adverse Events of Special Interest	9	12.9
Cytokine release syndrome	8	11.4
ICANS <sup>2</sup>	3	4.3
Blood and lymphatic system disorders	70	100.0
Neutropenia	62	88.6
Lymphopenia	39	55.7
Leukopenia	38	54.3
Anaemia	36	51.4
Thrombocytopenia	24	34.3
Febrile neutropenia	2	2.9
Cytopenia	1	1.4
Leukocytosis	1	1.4
Infections and infestations	10	14.3
Urinary tract infection	2	2.9
Appendicitis	1	1.4
COVID-19	1	1.4
Cytomegalovirus infection reactivation	1	1.4
Enterococcal infection	1	1.4
Human herpesvirus 6 encephalitis	1	1.4
Infection	1	1.4
Orchitis	1	1.4
Sepsis <sup>2,3</sup>	1	1.4
Septic shock <sup>2</sup>	1	1.4
Investigations	10	14.3
Alanine aminotransferase increased	6	8.6
Aspartate aminotransferase increased	5	7.1
Blood creatinine increased	2	2.9
Blood alkaline phosphatase increased	1	1.4
Blood bilirubin increased	1	1.4
Blood fibrinogen decreased	1	1.4
Lymphocyte count increased	1	1.4
Respiratory, thoracic and mediastinal disorders	10	14.3
Нурохіа	4	5.7
Pleural effusion	2	2.9
Bronchial obstruction	1	1.4
Dyspnoea	1	1.4
Epistaxis	1	1.4
Laryngeal Inflammation	1	1.4
Respiratory failure	1	1.4

TEAEs by maximum severity for all patients in Phase 1a and Phase 1b (N=70<sup>1</sup>)

Adverse event	≥Gra	ade 3
(System organ class, Preferred term)	No.	%
table continued		
Metabolism and nutrition disorders	7	10.0
Hypokalaemia	3	4.3
Hyponatraemia	3	4.3
Hypophosphataemia	2	2.9
Dehydration	1	1.4
Failure to thrive	1	1.4
Vascular disorders	7	10.0
Hypertension	6	8.6
Hypotension	1	1.4
Renal and urinary disorders	6	8.6
Acute kidney injury	4	5.7
Nephritis	1	1.4
Proteinuria	1	1.4
Gastrointestinal disorders	5	7.1
Abdominal pain	3	4.3
Diarrhoea	1	1.4
Ileus	1	1.4
Vomiting	1	1.4
General disorders and administration site conditions	4	5.7
Faligue	1	1.4
Pyrevia	1	1.4
Swelling face	1	1.4
Skin and subcutaneous tissue disorders	4	5.7
Rash maculo-papular	3	4.3
Eczema	1	1.4
Cardiac disorders	3	4.3
Atrial fibrillation <sup>4</sup>	3	4.3
Eye disorders	2	2.9
Periorbital oedema	1	1.4
Ulcerative keratitis	1	1.4
Injury, poisoning and procedural complications	2	2.9
Humerus fracture	1	1.4
Infusion related reaction	1	1.4
Musculoskeletal and connective tissue disorders	2	2.9
Back pain	1	1.4
Muscle spasms	1	1.4

Adverse event	≥ Grade 3			
(System organ class, Preferred term)	No.	%		
table continued				
Nervous system disorders	2	2.9		
Headache	1	1.4		
Posterior reversible encephalopathy syndrome	1	1.4		
Endocrine disorders	1	1.4		
Inappropriate antidiuretic hormone secretion	1	1.4		
Hepatobiliary disorders	1	1.4		
Cholangitis	1	1.4		
Immune system disorders	1	1.4		
Haemophagocytic lymphohistiocytosis	1	1.4		
Reproductive system and breast disorders	1	1.4		
Vaginal haemorrhage	1	1.4		

All treatment-emergent adverse events (TEAEs) with  $\geq$  Grade 3 regardless of relatedness to study treatment. 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 Cytokine release syndrome and ICANS were determined according to CARTOX criteria (Neelapu et al., 2019). Patients are counted only once per adverse event and severity classification. Based on interim data extracted from open clinical database (23-Aug-2024); <sup>1</sup> Two patients with disease progression after first IMA203 infusion received exploratory second IMA203 infusion. They had these  $\geq$  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> Fatal adverse events were not considered related to any study drug; <sup>3</sup> Patient died from sepsis of unknown origin and did not receive IMA203 TCR-T cells; <sup>4</sup> DLT: Dose limiting toxicity in phase 1a at DL2 reported on March 17, 2021 ICANS: Immune effector cell-associated neurotoxicity syndrome

## **Clinical Anti-Tumor Activity of IMA203 Monotherapy in Melanoma**



**Objective Responses in Heavily Pretreated Patients in Phase 1b (N=28<sup>#</sup>)** 

IMA203



ding for two melanoma patients at data-cut; \* Maximum change of target lesions and RECIST1.1 response at different timepoints. \*\* Tumor shrinkage of target lesions; <sup>1</sup>Patient is off study at data cut-off; <sup>2</sup>Patient out of study due to PD (external assessming 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 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 PD at any prior timepoint, patients with ongoing lation; 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

Data cut-off Aug 23, 2024 15

## **Duration of IMA203 Monotherapy Responses in Melanoma**



#### Durable Responses 2+ Years after Treatment in Heavily Pretreated Patients in Phase 1b (N=28<sup>#</sup>)



"First tumor assessment post infusion pending for two melanoma patients at data-cut; \* Tumor shrinkage of target lesions; <sup>1</sup>Patient out of study due to PD (external assessment) <sup>2</sup>Patient is off study at data cut-off; Initial ORR: Objective response rate according to RECIST 1.1 at any post infusion scans or patients with PD at any prior timepoint, patients with ongoing unconfirmed ORR cloulation; Duration of response (DOR) in confirmed ores is defined as time from first documented response until disease progression/death. Patients with ongoing response will be consored at data-cut-off. Blockage: SD: Stable Disease: PR: Partial Resonse: PR: Confirmed Partial Resonse: DCR: Disease control rate mEL's median DOR is analyzed by using the Kaplan-Meier method; Overall survival (OS) and progression-free survival (PFS) confirmed at data-cut: Bl: Raseline PD: Progressive Disease: PR: Partial Resonse: PR: Confirmed Partial Resonse: DCR: Disease control rate mEL's median follow-un

IMA203

Data cut-off Aug 23, 2024 16

## Significant Shift in PFS and OS Between Dose Escalation & Dose Expansion



mPFS of 6 Months and mOS Not Reached in Melanoma Efficacy Population



**Overall Survival** 

- Significant shift in mPFS and mOS between melanoma patients treated during the dose escalation and dose expansion phase
- mPFS in dose escalation is comparable to reported data in 2L+ cut. melanoma population<sup>\*</sup>
- mOS in dose escalation is shorter than reported mOS for 2L+ cut. melanoma population<sup>\*</sup> ٠
- All patients in the dose escalation group died, and 20/28 patients are alive in dose expansion .

Overall survival (OS) and progression-free survival (PFS) censored at data-cut; \* These data are derived from different clinical trials at different points in time with differences in trial design and IMA203 patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

## IMA203 Phase 1b in Melanoma: Overview of Studies



#### **PFS and OS Data in Melanoma Cohorts**

Drug Product	Phase	Ν	Melanoma patient population	Prior lines of therapies	<b>mPFS</b> (months)	<b>mOS</b> (months)
IMA203 in Melanoma	1b (Dose Expansion)	28	46% cutaneous 43% uveal 11% other	4% n=0, 18% n=1, 32% n=2, 29% n=3:, 4% n=4, 11% n=5, 4% n=6 86% received prior CPI (median of 1 prior line of CPI in overall population, median of 2 prior lines of CPI in cut. melanoma) Median of 2 prior lines, median of 2 prior lines in cut. melanoma	6.0	not reached
IMA203 in Melanoma	1a (Dose Escalation)	11	73% cutaneous 18% uveal 9% other	0% n=1, 27% n=2, 73% n>2 prior lines 100% received prior CPI (median of 2 prior lines of CPI, median of 2.5 prior lines of CPI in cut. melanoma) Median of 4 prior lines, median of 4.5 prior lines in cut. melanoma	2.6	6.3
IMA201/202/203 combined in Melanoma	1a (Dose Escalation)	19	63% cutaneous 11% uveal 26% other	0% n=1, 16% n=2, 84% n>2 prior lines 100% received prior CPI (median 3 prior lines of CPI) Median of 4 prior lines, median of 4.5 prior lines in cut. melanoma	2.5	5.3
Lifileucel (C-144-01, Cohort 2+4) <sup>1</sup>	2	153	54% cutaneous 0% uveal 45% other	median of 3 prior lines (min/max: 1/9) 100% received prior CPI	4.1	13.9
Tilsotolimod + Ipilimumab (ILLUMINATE-301) <sup>2</sup>	3	238	85% cutaneous 0% uveal 15% other	57% n=1, 27% n=2, 12% n>2 prior lines 99% received prior CPI	2.9	11.6
Nivolumab + Relatlimab (RELATIVITY-020, D1 Cohort) <sup>3</sup>	1/2	354	68% cutaneous 0% uveal 32% other	46% n=1, 35% n=2, 19% n≥3 prior lines 99% received prior CPI	2.1	14.7

These data are derived from different clinical trials at different points in time with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

# Enhanced mPFS of >1 Year in Melanoma Patients with Deep Responses N=26<sup>#</sup>



	Ν	mPFS
Dose Escalation IMA203	11	2.6 months
Dose Expansion IMA203 <50% tumor size reduction (including tumor size increase)	14*	5.7 months
Dose Expansion IMA203 >50% tumor size reduction	12	13.4 months

- 46% (12/26) patients have a deep response (≥50% tumor reduction)
- This subgroup of patients has highly medically meaningful mPFS of more than 1 year
- Patients with <50% tumor reduction (including tumor size increase) still observe a more than 2x longer mPFS as compared to patients treated in dose escalation with suboptimal doses

Immatic

## **SUPRAME: Registration-Enabling Randomized Phase 3 Trial**



#### Trial Design Following Recent Type D Meeting with FDA and SA Meeting with PEI<sup>1</sup>





IMA203

<sup>1</sup>Scientific Advice Meeting with Paul-Ehrlich-Institute, the German regulatory authority; <sup>2</sup> FDA requires demonstration of "no overall survival detriment" as endpoint; 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; mPFS: median progression-free survival, ORR: objective response rate; DOR: Duration of response; BLA: Biologics license application

## **Cell Therapy Manufacturing Facility**



#### To Support IMA203 BLA and Commercialization

- ~100,000 sq ft state-of-the-art research & GMP manufacturing facility
- Modular design for efficient and cost-effective scalability
   total of 8 manufacturing suites, plus further expansion space
- Capacity sufficient to serve early-stage and registration-directed clinical trials as well as planned commercial supply
- In-house manufacturing and QC allows full control of process, product and costs
- Located in the Houston Metropolitan Area, Texas, offering economic labor and operating costs and talent pool highly qualified in cell therapy manufacturing & QC







ACTengine<sup>®</sup> IMA203CD8 Expansion of the PRAME Commercial Opportunity Beyond Melanoma

# **Expansion of PRAME Commercial Opportunity Beyond Melanoma**



Second Generation ACTengine<sup>®</sup> IMA203CD8 Leveraging CD8 and CD4 T Cells



(0)

## IMA203CD8 Opportunity

## **2L Solid Tumors**

	US	EU5
Ovarian	2k	2k
Uterine	2k	2k
sqNSCLC	7k	10k
HNSCC	<b>2</b> k	2k
Breast	5k	8k
Others	16k	18k

- Co-transduction of CD8αβ alongside PRAME TCR adds functional CD4<sup>+</sup> T cells designed to boost cytotoxicity
- Proof of concept from preclinical experiments<sup>1</sup> and CD19 CAR T cell studies in leukemia<sup>2</sup>
- First clinical data with IMA203CD8 in Phase 1a dose escalation indicates potential for deeper responses and targeting both high and medium-level PRAME indications

All patient numbers refer to PRAME<sup>+</sup>/HLA-A<sup>\*</sup>02:01<sup>+</sup> patients in the US and EU5 in 2025; Source: Clarivate Disease Landscape and Forecast; EU5: France, Germany, Italy, Spain, United Kingdom <sup>1</sup> Bajwa et al. 2021 Journal for **IMA203CD8** Immunotherapy of Cancer; <sup>2</sup> Melenhorst et al. 2022 Nature, Bai et al. 2022 Science Advances; 2L: patients with unresectable or metastatic melanoma who have received at least 1 prior therapy; sqNSCLC: squamous non-small-cell lung cancer. HNSCC: head and neck squamous cell carcinoma

## **ACTengine® IMA203CD8 TCR-T Monotherapy Targeting PRAME**



#### **Summary: Clinical Data & Next Steps**



Dose escalation ongoing to investigate full clinical potential in hard-to-treat solid tumors outside of melanoma



Data cut-off Sep 30, 2024 24



## **Tolerability of IMA203CD8 Monotherapy**

All ≥Grade 3 Adverse Events (N=44)

#### TEAEs by maximum severity for all patients (N=44)

Adverse event		Grade 3	Adverse event	≥ Grade 3	
(System organ class, preferred term)	No.	%	(System organ class, preferred term)	No.	%
Patients with any adverse event	44	100.0	table continued		
Adverse events of special interest	7	15.9	Immune system disorders	4	9.1
Cytokine release syndrome <sup>1</sup>	6	13.6	Haemophagocytic lymphohistiocytosis <sup>2</sup>	4	9.1
Immune effector cell-associated neurotoxicity syndrome	1	2.3	Infections and infestations	4	9.1
Blood and lymphatic system disorders	44	100.0	Pneumonia	2	4 5
Neutropenia	40	90.9	Infection	1	2.3
Anaemia	25	56.8	Sensis <sup>3</sup>	1	2.3
Lymphopenia	25	56.8	Systemic candida	- 1	2.3
Thrombocytopenia	15	34.1	Gastrointestinal disorders	3	6.8
Leukopenia	11	25.0	Diarrhoea	2	4 5
Febrile neutropenia	2	4.5	Abdominal nain	1	23
Investigations	9	20.5	Skin and subcutaneous tissue disorders	3	6.8
Alanine aminotransferase increased	5	11.4	Pach	3	4.5
Aspartate aminotransferase increased	5	11.4		2	4.5
Blood creatinine increased	2	4.5	Alopecia	1	2.3
Blood alkaline phosphatase increased	1	2.3	Rash maculo-papular	1	2.3
Blood bilirubin increased	1	2.3	Vascular disorders	3	6.8
Gamma-glutamyltransferase increased	1	2.3	Hypertension	3	6.8
Metabolism and nutrition disorders	6	13.6	Nervous system disorders	2	4.5
Hypophosphataemia	2	4.5	Neurotoxicity <sup>2</sup>	1	2.3
Acidosis	1	2.3	Syncope	1	2.3
Decreased appetite	1	2.3	Renal and urinary disorders	2	4.5
Hyperglycaemia	1	2.3	Acute kidney injury	1	2.3
Hypermagnesaemia	1	2.3	Urinary tract obstruction	1	2.3
Hypoalbuminaemia	1	2.3	Henatobiliary disorders	1	2.3
General disorders and administration site conditions	5	11.4	Henatic function abnormal	1	2.3
Fatigue	5	11.4	Reproductive system and breast disorders	1	2.3
Oedema peripheral	1	2.3	Reproductive system and breast disorders	1	2.3
Musculoskeletal and connective tissue disorders	5	11.4		T	2.3
Bone pain	3	6.8			
Myalgia	2	4.5			
Back pain	2	45			

#### • Overall manageable tolerability profile

- Expected cytopenia
- Mostly mild to moderate CRS:
  - 36% (16/44) Grade 1
  - 48% (21/44) Grade 2
  - 11% (5/44) Grade 3
  - 2% (1/44) Grade 4
- DLTs in 2 patients at DL4b as previously reported by the Company:
  - Patient DL4b-01: high *in vivo* T cell expansion, Grade
     4 neurotoxicity, Grade 4 CRS, Grade 3 HLH
  - Patient DL4b-04: Grade 3 CRS defined by Grade 3 ALT resolved to Grade 2 within 10 days; no need for vasopressors or ventilation
- No IMA203CD8-related patient death<sup>3</sup>
- Consecutive modification I/E criteria + IL2 scheme
- **Dose escalation ongoing** based upon manageable tolerability in patients at DL4a

All treatment-emergent adverse events (TEAEs) with  $\geq$  Grade 3 regardless of relatedness to study treatment that occurred in at least 1 patient are presented; <sup>1</sup> DLT: Dose limiting toxicity in patient DL4b-04. <sup>2</sup> DLTs in patient DL4b-01; CRS: cytokine release syndrome, HLH: hematophagocytic lymphohistiocytosis

2.3

IMA203CD8

Arthralgia

#### Clinical Anti-Tumor Activity of IMA203CD8 Monotherapy (N=41) Ongoing Dose Escalation



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 (mFU) is analyzed by using the reverse Kaplan-Meier method; PD: Progressive Disease; SD: Stable Disease; PR: Partial Response; cPR: Confirmed Partial Response; CR: complete response; BL: Baseline; BOR: Best Overall Response; DCR: disease control rate; NSCLC: non-small-cell lung cancer

**IMA203CD8** 



cORR 41% (14/34)

 median DOR
 9.2 months

 (min, max)
 2.0+, 23.5+

 mFU
 13.1 months

10/17 responses ongoing including 3 confirmed responses at 1+ year

Deep responses with ≥50% tumor size reduction in 11/17 responders incl. 2 patients with complete response of target lesions

41% (17/41)

84% (32/38)

85% (34/40)

ORR

Tumor shrinkage<sup>3</sup>

DCR<sup>4</sup> (at week 6)



## **Duration of IMA203CD8 Monotherapy Responses (N=41)**

#### **Ongoing Dose Escalation**

IMA203CD8



## **Opportunity of IMA203CD8 in Medium-Level PRAME Expressing Indications**





IMA203CD8 offers similar responses at 1.5 x 10<sup>9</sup> total infused dose as IMA203 at 3x higher dose. With higher doses currently being explored,

IMA203CD8 may offer an enhanced opportunity to treat cancers with both high and medium-level PRAME expression including ovarian cancer, uterine cancer, sqNSCLC,

triple-neg. breast cancer and others. Next clinical data update including focus on ovarian cancer in **2025**.

IMA203CD8

\* Patients treated at RP2D during Ph1b with evaluable post baseline assessments at data-cut off IMA203: Aug 23, 2024; BOR: best of response; PD: progressive disease; SD: stable disease; (c)PR: (confirmed) partial response; sqNSCLC: squamous non-small-cell lung cancer





## **TCER® IMA402 - Off-the-Shelf TCR Bispecific Targeting PRAME**



## **Expansion of the PRAME Commercial Opportunity to 1L Tumors**

**Off-the-Shelf Biologic Approach to Target First Line Setting** 



Repeat dosing

IMA402

• Patient reach also into community setting

Others

25k

**32k** 



**Dose escalation** with higher DLs **ongoing** to leverage PRAME potential in advanced stage indications

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

![](_page_31_Picture_1.jpeg)

- ECOG status 0-1
- Received or not eligible for all available indicated standard of care treatments

![](_page_31_Figure_4.jpeg)

#### Total safety population (N=33)

- MABEL-based starting dose
- Dose escalation based on cohorts of 1-6 patients using adaptive design (BLRM model)
- Weekly infusions<sup>2</sup> with potential to explore less frequent dosing based on PK data

- MTD not yet determined
- Dose escalation ongoing at DL9

Immoti

![](_page_32_Picture_0.jpeg)

## **IMA402** Demonstrates Favorable Tolerability in N=33 Patients

#### **Most Frequent Related AEs were Lymphopenia and CRS**

Treatment-related AEs <sup>1</sup> , n [%]	All Grades	≥ Grade 3
Lymphopenia	17 [52]	10 [30]
Cytokine release syndrome	16 [48]	1 [3]
Arthralgia	9 [27]	0
Fatigue	9 [27]	0
Pruritus	7 [21]	0
Rash	7 [21]	0
Aspartate aminotransferase increased	6 [18]	2 [6]
Alanine aminotransferase increased	5 [15]	1 [3]
Pyrexia	5 [15]	0
Anaemia	4 [12]	2 [6]
Vomiting	4 [12]	0
C-reactive protein increased	3 [9]	0
Headache	3 [9]	0
Rash maculo-popular	3 [9]	0
Neutropenia	2 [6]	2 [6]
Stomatitis	2 [6]	1 [3]
Blood creatinine increased	1 [3]	1 [3]
Electrocardiogram abnormal	1 [3]	1 [3]
Gamma-glutamyltransferase increased	1 [3]	1 [3]
Hypertension	1 [3]	1 [3]
Immune-mediated arthritis	1 [3]	1 [3]
Tumor lysis syndrome	1 [3]	1 [3]
Tumor pain	1 [3]	1 [3]

TEAEs, n [%]	All Grades	≥ Grade 3
Any	33 [100]	17 [52]
Treatment-related	32 [97]	15 [45]

- Data here includes patients up to DL8
- Favorable tolerability profile
- Most frequent/relevant related AEs were
  - transient lymphopenia,
  - mostly mild to moderate CRS (42% Grade 1, 3% Grade 2, 0% Grade 3, 3% Grade 4), majority at first dose
  - one DLT: Grade 4 CRS (fully resolved)
- No IMA402-related Grade 5 events
- As of Jan 10, dose escalation remains ongoing at DL9 (5 mg)
- MTD not reached

#### Early Signs of Clinical Activity Associated with PRAME Expression and IMA402 Dose

![](_page_33_Picture_1.jpeg)

![](_page_33_Figure_2.jpeg)

- Melanoma patient with confirmed partial response ongoing at 3 months (DL7, see next slide)
- Melanoma patient with -27.5% tumor shrinkage at first scan (DL8)
- Uveal melanoma patient with -25.0% tumor shrinkage deepening over time (started at DL4 and currently at DL7, see next slide)
- Ovarian cancer patient with -13% tumor shrinkage ongoing at 3 months (started at DL6 and currently at DL7)
- Next data update(s) throughout 2025 with initial focus on cut. melanoma

## **Exemplary Patient Cases Suggesting Dose-Dependent Tumor Response**

![](_page_34_Picture_1.jpeg)

Patients with Disease Control (RECIST1.1) at Relevant Doses (DL7+)

![](_page_34_Figure_3.jpeg)

#### **Patient Characteristics & Outcomes**

52-year-old female with cutaneous melanoma

Lesions in lung, lymph nodes, gall bladder, fat tissue, pancreas

1 prior line of therapy and maintenance with anti-PD-1

Patient received DL7 from start (after step-up dosing)

Ongoing cPR at 3 months post treatment start with -40.2% reduction of target lesion size

![](_page_34_Figure_10.jpeg)

#### **Patient Characteristics & Outcomes**

46-year-old female with uveal melanoma

#### Lesions in liver

3 prior lines of therapy with anti-PD1 and tebentatafusp

Patient received DL4 and went up to DL7 through intra-patient dose escalation

Ongoing SD at 8+ months post-treatment start with -25% reduction of target lesion size

![](_page_35_Picture_0.jpeg)

![](_page_35_Picture_1.jpeg)

## **TCER® IMA401 - Off-the-Shelf TCR Bispecific Targeting MAGEA4/8**

# IMA401 All patient numbers refer to MAGEA4/8<sup>+</sup>/HLA-A<sup>\*</sup>02:01<sup>+</sup> patients in the US and EU5 in 2025; Source: Clarivate Disease Landscape and Forecast; EU5: France, Germany, Italy, Spain, United Kingdom; sqNSCLC: squamous non-small-cell lung cancer, HNSCC: head and neck squamous cell carcinoma

# The IMA401 Commercial Opportunity in Solid Cancers

**TCER® IMA401 Targeting MAGEA4/8** 

![](_page_36_Figure_3.jpeg)

![](_page_36_Picture_4.jpeg)

![](_page_37_Figure_0.jpeg)

AE: Adverse Event; CRS: Cytokine Release Syndrome; (c)ORR: (confirmed) objective response rate; PR: Partial Response; DCR: disease control rate; CPI: checkpoint inhibitors; q4w: once every four weeks; HNSCC: Head and neck squamous cell carcinoma; sqNSCLC: squamous non-small-cell lung cancer

![](_page_38_Picture_0.jpeg)

## Trial Design – IMA401-101 Phase 1a Dose Escalation

#### First-in-Human Basket Trial Targeting the MAGEA4/8 Peptide in Solid Tumors

![](_page_38_Figure_3.jpeg)

IMA401

#### Total safety population (N=35)

• MABEL-based starting dose

- Dose escalation based on cohorts of 1-6 patients using adaptive design (BLRM model)
- Four initial q1w step dosings<sup>1</sup> up to target dose, q2w after reaching target dose<sup>2</sup>
- MTD not yet determined
- Dose escalation ongoing to optimize dosing intervals and schedule

#### Objectives

#### Primary:

Determine MTD and/or RP2D

#### Secondary:

- Tolerability
- Pharmacokinetics
- Initial anti-tumor activity

#### Key Eligibility Criteria

- Recurrent and/or refractory solid tumors
- HLA-A\*02:01 positive
- MAGEA4/8-positive as confirmed by mRNA-based assay<sup>3</sup>
- ECOG status 0-2
- Received or not eligible for all available indicated standard of care treatments

# immatics

## IMA401 Demonstrates Manageable Tolerability in N=35 Patients

#### Most Frequent Related AEs were Lymphopenia, CRS and Neutropenia

Treatment-related AEs <sup>1</sup> , n [%]	All Grades	≥ Grade 3
Lymphopenia	12 [34]	11 [31]
Cytokine release syndrome	11 [31]	0
Neutropenia	8 [23]	5 [14]
Facial pain	6 [17]	2 [6]
Anaemia	5 [14]	4 [11]
Thrombocytopenia	5 [14]	2 [6]
Headache	5 [14]	1 [3]
Hypertension	4 [11]	2 [6]
Leukopenia	4 [11]	2 [6]
Fatigue	4 [11]	0
Nausea	3 [9]	0
Нурохіа	2 [6]	1 [3]
Aspartate aminotransferase increased	1 [3]	1[3]
Febrile neutropenia	1 [3]	1[3]
Pneumonia	1 [3]	1[3]
Sinus tachycardia	1 [3]	1[3]

IMA401

TEAEs, n [%]	All Grades	≥ Grade 3
Any	32 [91]	26 [74]
Treatment-related	28 [80]	19 [54]

- Overall manageable tolerability profile
- Most frequent/relevant related AEs were
  - transient lymphopenia
  - mild to moderate CRS (23% Grade 1, 9% Grade 2, no
     Grade ≥ 3), majority at first dose
  - neutropenia<sup>2</sup> occurred mostly at initial target dose and fully resolved in all cases except one (see below)
  - one possibly related death (pneumonia in the context of lung tumor progression and concurrent neutropenia) as previously reported<sup>3</sup>
- MTD not reached based on the BLRM

## **Objective Responses are Associated with Target Expression**

![](_page_40_Picture_1.jpeg)

**Exploratory Analysis in Patients with MAGEA4/8<sup>high</sup> Expression at Relevant IMA401 Doses (DL6-7; N=17)** 

![](_page_40_Figure_3.jpeg)

## IMA401 Demonstrates Initial Anti-Tumor Activity in Multiple Tumor Types

![](_page_41_Picture_1.jpeg)

#### Exploratory Analysis in Patients with MAGEA4/8<sup>high</sup> Expression at Relevant IMA401 Doses (DL6-7; N=17\*)

![](_page_41_Figure_3.jpeg)

Cancer Indications: Cut.: Cutaneous; HNSCC: Head & Neck Squamous Cell Carcinoma; LCNEC: Large Cell Neuroendocrine Carcinoma; Muc.: Mucosal; NET CUP: Neurodendocrine Tumor, Cancer of Unknown Primary; SCLC: Small Cell Lung Cancer; sqNSCLC: Squamous Non-small Cell Lung Cancer; TNBC: Triple Negative Breast Cancer.

IMA401

\* Patients in this analysis are part of the efficacy analysis set with at least one post-treatment tumor assessment and had received IMA401 infusions at ≥1 mg and showed MAGEA4/8 target expression higher than the MAGEA4/8 qPCR threshold (n=17); 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; two patients not included in tumor shrinkage calculation or shown in the figures as they had clinical progression and post-treatment tumor assessment is not available; PR: Partial Response; cPR: Confirmed Partial Response; SD: Stable Disease

Data cut-off Jul 23, 2024

42

![](_page_42_Picture_0.jpeg)

![](_page_42_Picture_1.jpeg)

**The Immatics Opportunity** 

## The Immatics Opportunity

#### **Delivering the Power of T Cells to Cancer Patients**

![](_page_43_Picture_2.jpeg)

![](_page_43_Picture_3.jpeg)

#### Leveraging 2 TCR modalities to target solid cancers

TCR-T cell therapy (ACTengine<sup>®</sup>) and TCR Bispecifics (TCER<sup>®</sup>) directed against pHLA targets to address late-stage and early-stage solid cancers

#### Delivering on the promise of cell therapy

IMA203 with compelling clinical activity in 2L melanoma IMA203 SUPRAME Phase 3 trial has commenced IMA203CD8 data support investigation beyond melanoma

![](_page_43_Picture_8.jpeg)

#### ✓ = ✓ = ✓ = ✓ = ✓ =

#### Achieving robust cell therapy product manufacturing

Manufacturing process optimized for product efficacy Manufacturing facility for clinical-stage and planned commercial supply

#### **Progressing to commercial stage**

Buildout of commercial organization has commenced IMA203 received an RMAT<sup>1</sup> designation from the FDA

![](_page_43_Picture_14.jpeg)

![](_page_43_Picture_15.jpeg)

# Delivering off-the-shelf Bispecifics to broaden the solid cancer opportunity

Initial clinical data for TCER<sup>®</sup> IMA402 (PRAME) and IMA401 (MAGEA4/8) support exploring indication expansion and earlier treatment lines

#### Solid financial position and focus on clinical-stage assets

![](_page_43_Picture_19.jpeg)

![](_page_43_Picture_20.jpeg)

![](_page_44_Picture_0.jpeg)

# the Power of T cells to Cancer Patients

![](_page_44_Picture_2.jpeg)

www.immatics.com

![](_page_44_Picture_4.jpeg)

© Immatics. Not for further reproduction or distribution.

![](_page_44_Picture_6.jpeg)