# UNITED STATES SECURITIES AND EXCHANGE COMMISSION Washington, D.C. 20549

# FORM 6-K

REPORT OF FOREIGN PRIVATE ISSUER PURSUANT TO RULE 13a-16 OR 15d-16 UNDER THE SECURITIES EXCHANGE ACT OF 1934

October 18, 2022	
Commission File Number: 001-3936	3
IMMATICS N.V.	
Paul-Ehrlich-Straße 15 72076 Tübingen, Federal Republic of Ger (Address of principal executive office)	many
Indicate by check mark whether the registrant files or will file annual reports under	er cover of Form 20-F or Form 40-F:
Form 20-F 🔀 For	m 40-F
Indicate by check mark if the registrant is submitting the Form 6-K in paper as permitted by Regulation S-T Rule 101(b)(1): Indicate by check mark if the registrant is submitting the Form 6-K in paper as permitted by Regulation S-T Rule 101(b)(7):	

### INFORMATION CONTAINED IN THIS REPORT ON FORM 6-K

On October 18, 2022, Immatics N.V. (the "Company") made available an updated investor presentation on its website, a copy of which is attached hereto as Exhibit 99.1. The fact that this presentation is being made available and filed herewith is not an admission as to the materiality of any information contained in the presentation. The information contained in the presentation is being provided as of October 18, 2022 and the Company does not undertake any obligation to update the presentation in the future or to update forward-looking statements to reflect subsequent actual results.

### EXHIBIT INDEX

Exhibit No. Description

99.1 Presentation dated October 18, 2022

### SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned, thereunto duly authorized.

IMMATICS N.V. Date: October 18, 2022

By: /s/ Harpreet Singh
Name: Harpreet Singh
Title: Chief Executive Officer

# Immatics Corporate Presentation

October 18, 2022



Delivering the Power of T cells to Cancer Patients

© 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 all-inclusive 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 of IND or CTA filing for pre-clinical stage product candidates, 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", "intend", "estimate", "anticipate", "predict", "potential" or "continue", or the negatives of these terms or variations of them or similar terminology. Such 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, 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 presentation should be regarded as a representation by any person that the forward-looking statements will be achieved. You should not place undue reliance on forward-looking statements, which speak only as of the date they are made. The Company undertakes no duty to update these forward-looking statements.

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.

,

# **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

High rate of confirmed objective responses across multiple solid tumors in early TCR-T clinical development

### Differentiated Platforms

Unique technologies to identify true cancer targets and right TCRs

### Therapeutic Opportunity

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

### Strategic Partnerships

World-leading industry players with synergistic expertise

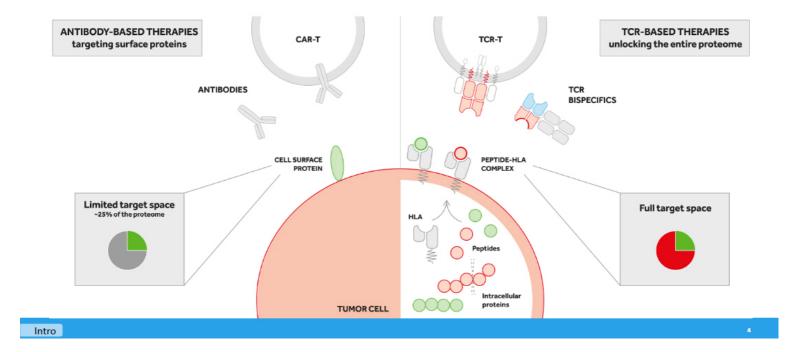
### Solid Cash Runway

To reach multiple value inflections points across our portfolio

Intro

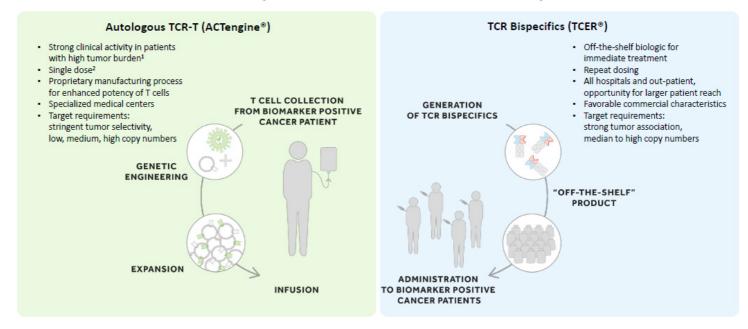


# Our TCR-based Approaches Leverage the Full Target Space beyond the Cancer Cell Surface



# Two Distinct TCR-based Therapeutic Modalities in Clinical Development





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

Intro

<sup>1</sup>Interim data update from the ACTengine<sup>o</sup> IMA203 TCR-T Phase 1 trial with a 50% (6/12) confirmed ORR target dose or above with at least 1 billion infused TCR-T cells across several solid tumor indications, 80% (4/5) confirmed ORR in Phase 1b patients only; <sup>2</sup>Repeat dosing without re-manufacturing possible

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



Modality	Product Candidate	Target		Preclinical	Phase 1a <sup>1</sup>	Phase 1b1	
	IMA203	PRAME	immatics.	+ Cl	heckpoint Inhibitor²		
ACTengine® Autologous ACT	IMA203CD8	PRAME	immatics:				
· ·	IMA201	MAGEA4/8	immatics:				
	IMA204	COL6A3	immatics:				
Autologica ACT	4 programs	Undisclosed	(h Bristol Myers Squibb				
Autologous ACT	2 programs	Undisclosed	GSK				
ACTallo®	IMA30x	Undisclosed	mmatics editas*				
Allogeneic ACT γδ T cells	2 programs	Undisclosed	(*Bristol Myers Squibb"				
	IMA401	MAGEA4/8	(*Bristol Myers Squibb"				
TCER® Bispecifics	IMA402	PRAME	immatics:				
	IMA40x	Undisclosed	immatics:				
Bispecifics	3 programs	Undisclosed	<sup>*Y</sup> Genmab				

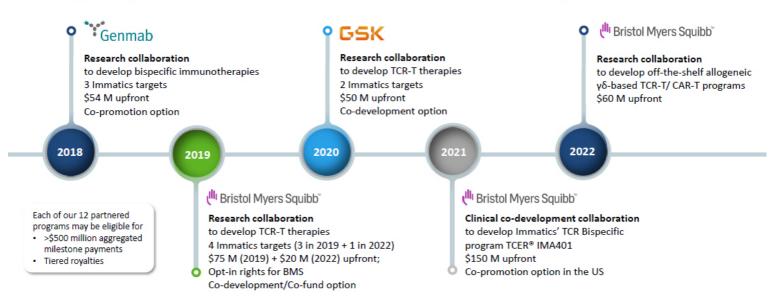
Intro

¹ Phase 1a: Dose escalation, Phase 1b: Dose expansion; ² Opdivoº (nivolumab): programmed death-1 (PD-1) immune checkpoint inhibitor.
¹ Immatics proprietary ACTalloº platform utilizing Editas' CRISPR gene editing technology

# **Strategic Collaborations**



# Synergistic Expertise that Can Foster Transformative Innovations for ACT and Bispecifics



Broadening the clinical framework beyond our pipeline

Intro



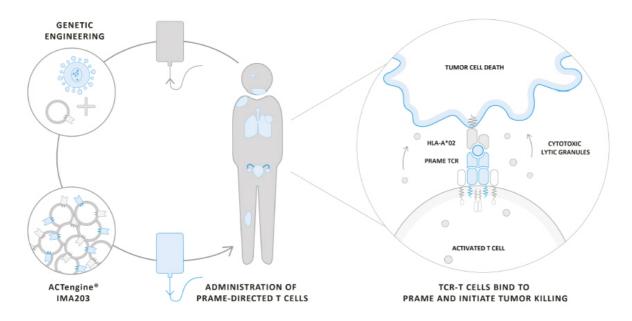


ACTengine® IMA203 – TCR-T Targeting PRAME

# ACTengine® IMA203 Targeting PRAME – Mechanism of Action



Immatics' Leading TCR-T Approach



IMA203

# **Multi-Tumor Target PRAME**

# **Promising Opportunity for TCR-based Therapies**





- HLA-A\*02:01 presented peptide identified by XPRESIDENT® quant. mass spectrometry
- Presented at high target density in tumor tissue (100-1000 copies/cell)
- Homogenously expressed
- Highly cancer-specific, not expressed in normal tissue at relevant levels
- Highly prevalent across many solid cancers
- Potential to reach a large cancer patient population

Uveal Melanoma<sup>2</sup> 90%

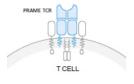
Ovarian Carcinoma 70%

Immatics' clinical trials support high prevalence of PRAME:				
Uterine Carcinoma	90%			
Cut. Melanoma	95%			

:	
_	_

PRAME RNA detection in tumor samples (ISH)

Indication	% PRAME positive patients <sup>1</sup>		
Uterine Carcinoma	100%		
Uterine Carcinosarcoma	100%		
Sarcoma Subtypes	up to 100%		
Cut. Melanoma	95%		
Uveal Melanoma <sup>2</sup>	50%		
Ovarian Carcinoma	80%		
Squamous NSCLC	65%		
TNBC	60%		
Small Cell Lung Cancer	55%		
Kidney Carcinoma	up to 45%		
Cholangiocarcinoma	35%		
Adeno NSCLC	25%		
Breast Carcinoma	25%		
HNSCC	25%		
Esophageal Carcinoma	20%		
HCC	20%		
Bladder Carcinoma	20%		



TUMOR CELL

HLA-A\*02:01

# IMA203 T cell Receptor (TCR):

- Affinity-improved TCR by enhanced TCR chain pairing
- High functional avidity: EC50 ~5 ng/ml
- Off-target toxicity screening against normal tissue peptides selected from our immunopeptidome database to retain specificity

IMA203

# ACTengine® IMA203 - Patient Flow

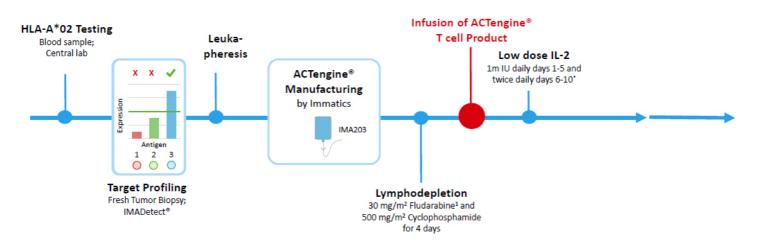


**Screening & Manufacturing Phase** 

**Treatment & Observation Phase** 

Long Term Follow-up

Safety and efficacy monitoring for 12 months



IMA203

\* IL-2 dose reduction from twice daily to daily for the first 5 days and dosing duration from 14 to 10 days introduced prior to treatment of first patients on dose level.

1 Dose reduction of Fludarships (from 40ms/m² to 30ms/m²) was introduced prior to treatment of the first patient on dose level 3.

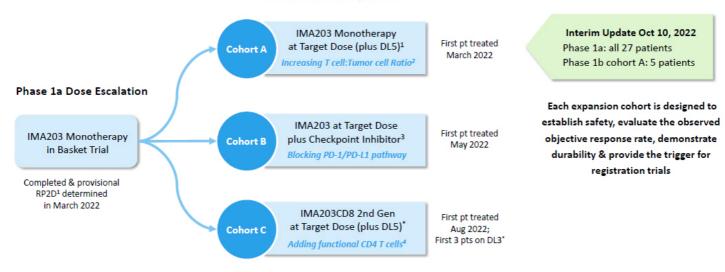
- 1

# IMA203 TCR-T Phase 1 Design



# Three Phase 1b Expansion Cohorts to Establish Durable Objective Responses

### Phase 1b Dose Expansion



IMA203 Treatm

# Moving from Phase 1a to Phase 1b



# Continuous Improvement of Key Aspects that May Influence Clinical Outcome

### Our Focus in Phase 1a

- Safety
- Biological activity
- · Initial signs of clinical activity



### Our Focus in Phase 1b

- Safety
- Durability of response at 6 months and beyond to pave the way for registration trials

### We continue to improve key determinants as we move from Phase 1a into Phase 1b

- 1. Higher T cell dose: Only RP2D or exploratory DL5
- 2. Enhanced cell product: Implementation of manufacturing enhancements (e.g. monocyte depletion, see appendix) focusing on robustness, quality, and speed of product release
- 3. "Real life" patients: Working with more disease area experts to reduce the fraction of very heavily pre-treated patients with extreme disease burden who have exhausted standard of care and have undergone multiple clinical trials

IMA203

# ACTengine® IMA203 - Interim Monotherapy Update



### Phase 1a

# Dose Escalation Data from 27 Patients

- Acceptable & manageable treatment-emergent adverse events (TEAEs)
- DL4 defined as provisional RP2D
- 48% (13/27) <u>initial</u> ORR<sup>1</sup> across all doses and multiple solid cancers
- Limited number of confirmed responses

### Phase 1b Cohort A

# Initial Data from 5 Patients

- Acceptable & manageable TEAEs
- Patients treated at RP2D (DL4) and exploratory DL5
- 80% (4/5) <u>initial</u> ORR<sup>1</sup> in patients with 4 different solid tumors
- 80% (4/5) confirmed ORR<sup>2</sup>: Confirmation of all objective responses after ~3 months; all responses ongoing



### **Key Take Aways**

### **IMA203** Monotherapy

- Favorable tolerability profile
- Confirmed responses in multiple heavily pre-treated solid tumor types (cut. melanoma, uveal melanoma, head and neck cancer, ovarian cancer, synovial sarcoma)
- Positively evolving durability profile for IMA203
  - above 1 bn TCR-T cells (DL4/5)\* in phase 1a and phase 1b: 50% (6/12) confirmed ORR<sup>2</sup>
  - in phase 1b patients only: 80% (4/5) confirmed ORR<sup>2</sup>

Data cut-off = 06-Sept-2022

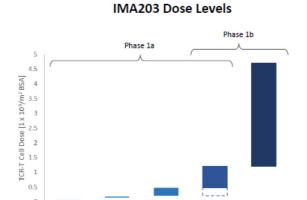
IMA203

ORE: Objective responses rate (partial responses) according to RECIST 1.1 at first scan post infusion ("6 weeks)," confirmed ORE: Confirmed ORE: Confirmed ORE: confirmed operatial responses) according to RECIST 1.1 at first scan post infusion ("12 weeks), 1 patient with 50 at "6 weeks scan with pending "12-week scan considered as non-responder for confirmed ORE; to that transduced visible COBT CoELS, all patients with 50 hazes 1 DUI As and Phazes 1 DUI As and Phazes 1 put Phazes 1 p

# ACTengine® IMA203 Monotherapy – Patient and Product Characteristics



	Phas Dose Esc All pts (DL1-4)	calation	Phase 1b (Cohort A)  Dose Expansion  All pts (DL4/DL5)		
Patients treated	27	7	5		
Prior lines of treatment Mean (min, max)	4.2 (1, 8)	4.6 (1,7)	4.0 (1, 10)		
LDH at baseline >1 x ULN [% of patients]	66.7	85.7	40.0		
Baseline tumor burden Mean target lesion sum of diameter [mm] (min, max)	130.3 (29.0, 219.7)	115.8 (37.0, 197.6)	55.2 (21.0, 102.9)		
<b>Dose</b> Mean transduced viable CD8 T cells infused [x10 <sup>9</sup> ] (min, max)	0.65 (0.08, 2.09)	1.48 (1.07, 2.09)	2.22 (1.30, 4.16)		
Manufacturing Process	Prior ve	ersions <sup>1</sup>	Current version		



32 heavily pre-treated patients, thereof **12 patients at target dose or above**, were infused with IMA203 TCR-T cells targeting PRAME

DL4 was defined as provisional RP2D for Phase 1b, exploration of higher DL5 ongoing

Dose level 2

Data cut-off = 06-Sept-202

Dosel level 5 \*\*

IMA203

\*Except for 1 product for patient at DL3 generated with current manufacturing process; \*DL4: 200m to 1.2bn transduced viable CD8 T cells per m² BSA, all patients in DL4 received cell doses in the upper tier of DL4, above DL3;

\*\*DL5: up to 4.7bn transduced viable CD8 T cells per m² BSA: ULN: Upper limit of normal: BSA: Body surface area: RP2D: Recommended Phase 2 dose: LHD: Lactate dehydrogenase

# IMA203 Tolerability Profile - Most Frequent Adverse Events



Acceptable and Manageable Treatment-emergent Adverse Events (TEAEs)

- · Expected cytopenia (Grade 1-4) associated with lymphodepletion in all patients
- Cytokine release syndrome (CRS): 31 of 32 (97%) patients infused with IMA203 experienced CRS of any Grade
  - 29 patients had Grade 1 or 2 CRS
  - 2 patients had Grade 3 CRS (both in phase 1a); recovered to Grade ≤2 after 3 and 4 days, respectively
- Low-moderate ICANS1: 5 of 32 (16%) patients infused with IMA203 experienced Grade 1 or 2 ICANS (all in phase 1a)
- No dose-dependent increase of CRS and ICANS
- No additional DLT<sup>2</sup>

Data cut-off - 06-Sept-2022

IMA203 One pa

ine patient that started lymphodepletion in Phase 1a died from sepsis of unknown origin and did not receive IMA203 T cells, patient reported earlier and not shown; CRS and ICANS graded by CARTOX criteria (Neelapu et al., 2018); ICANS: Immune effector cell-associated neurotoxicity syndrome; <sup>3</sup> DIT, dose-limiting toxicity, one DIT in phase 1 as 1DL reported on March 17, 2021.

# Frequency of Observed Objective Responses



Improved ORR and Confirmed ORR at Higher Dose and in Phase 1b Cohort A

	Phase 1a		Phase 1a + Phase 1b	Phase 1b only	
	All pts (DL1-4)	DL4 pts only <sup>1</sup>	DL4/DL5 pts only <sup>1</sup>	All pts (DL4/DL5)1	
Patients Treated	27	7	12	5	
ORR (~6 weeks) <sup>2</sup>	48% (13/27)	57% (4/7)	67% (8/12)	80% (4/5)	
cORR (~12 weeks) <sup>3</sup>	19% (5/27)	29% (2/7)	50% (6/12)*	80% (4/5)*	

- Higher ORR and confirmed ORR observed at doses above 1 billion TCR-T cells (DL4, DL5)
- Early trends towards higher ORR and confirmed ORR observed in Phase 1b vs. Phase 1a patients

Data cut-off – 06-Sept-202

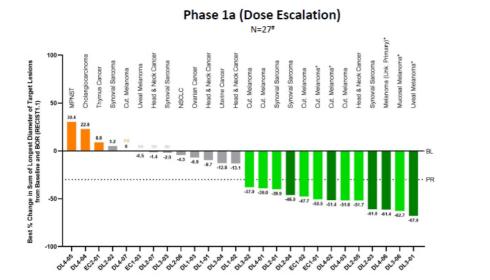
IMA203 respon

\*all patients received > 1.1 to total translucted storage Cust Copies, \*Unit, objective response rate (partial responses) according to RCLS1.1.1 at ITSS scan post initiation (\*) weeks); \*\*Continued usin Copiestive response rate (partial responses) according to According to the Copiestive response rate (confirmed post initiation (\*) \*\*Copiestive response rate (\*) \*\*Copiestive vitin post initiation (\*) \*\*Copiestive response rate (\*) \*\*Copiestive response rate (\*) \*\*Copiestive vitin post initiation (\*) \*\*Copiestive vitin p

# **Best Overall Response**



### IMA203 Continues to Deliver Objective Responses in Major Solid Tumor Types



# Phase 1b (Cohort A) N=1 Tom Baseline and Book (RECISTI.1) Tom Baseline and Book (RECISTI.1) N=1 Tom Baseline and Book (RECISTI.1) Head & Neck Cancer Overran Cancer Ov

Confirmed objective responses across a broad spectrum of different tumor types such as cutaneous melanoma, uveal melanoma, head and neck cancer, ovarian cancer, synovial sarcoma

Data cut-off = 06-Sept-2022

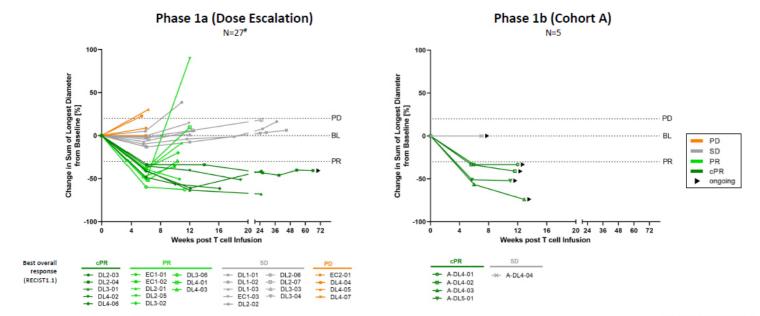
IMA203

\* Maximum change of target lesions and RECIST 1.1 BOR at different timepoints; \*Synovial sarcoma patient (DL3) PD at week 6 not shown as target lesions were not evaluable PD: Progressive disease: SD: Stable disease: PR: Partial response: cPR: Confirmed partial response: BL: Baseline

# **Responses over Time**



# Encouraging Early Signs for Improved Durability at Higher Dose and in Phase 1b Patients



Data cut-off - 06-Sept-2022

\*Synovial sarcoma patient (DL3) PD at week 6 not shown as target lesions were not evaluable; PD: Progressive disease; SD: Stable disease; PR: Partial response; cPR: Confirmed partial response; BL: Baseline 19

### Translational Data Consistent with Clinical Outcomes

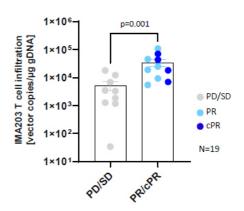


**Supporting Proposed Mechanism of Action for IMA203** 

### High IMA203 T cell engraftment and persistence in peripheral blood

### Persistence over time Peak frequency 1×107 1.5×10°p=0.0003 Vector copies/µg gDNA Vector copies/µg gDNA 1×10 1×104 1×10<sup>e</sup> 1×10<sup>3</sup> 1×10<sup>2</sup> PD/SD 5×10<sup>6</sup> 1×10<sup>1</sup> PR 1×100 cPR N=32 1×10<sup>-1</sup> Phase 1b Cohort A Days post infusion

### IMA203 T cell infiltration into tumor correlates with objective responses1



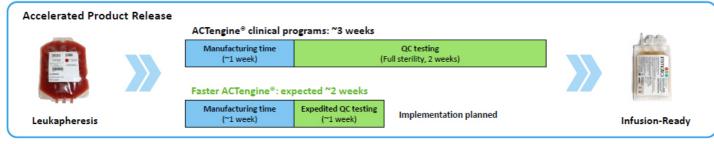
Data cut-off = 06-Sept-2022

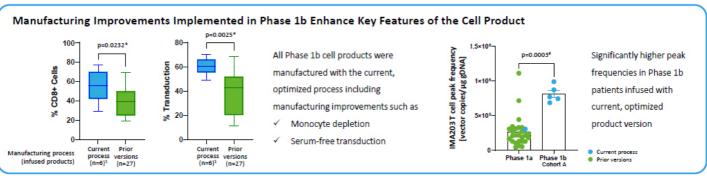
IMA203 Mann-Whitney U test; <sup>1</sup>T cell infiltration for 19 patients (9 non-responder, 10 responder) with 6-week post infusion biopsy available (1 patient with "5-week post infusion biopsy)

# ACTengine® IMA203 Product Manufacturing



Targeting Higher Robustness, Favorable Product Attributes, Faster Turn Around Time





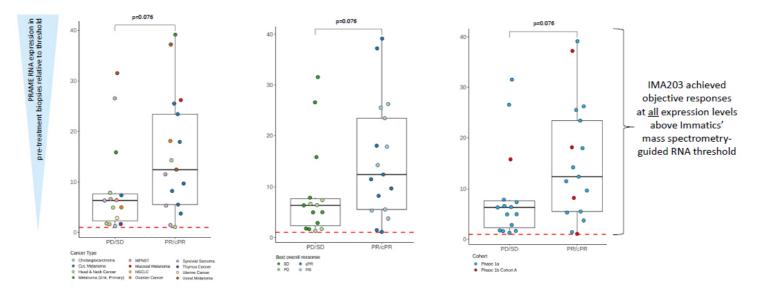
<sup>1</sup>Includes 5 IMA203 products infused into Phase 1b cohort A patients, and 1 product infused into Phase 1a patient at DL3; \* Unpaired t test; \*Mann-Whitney U test, 1 patient in Phase 1a at DL3 received \*0.5 x 10<sup>8</sup> total transduced viable CD8 T cells manufactured with current proces

IMA203

# PRAME Expression in Tumors from Screened Patients (N=32)



Highlighting Tumor Types (left), Type of Best Overall Response (middle) and Study Cohort (right)



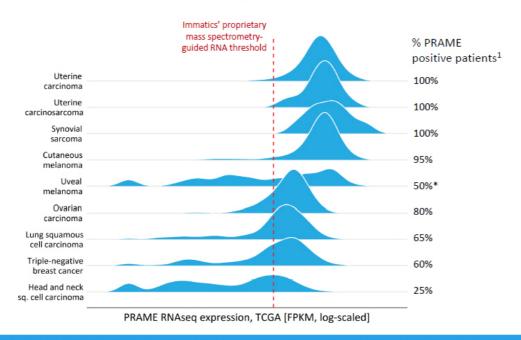
IMA203 has the potential to provide clinical benefit for all PRAME biomarker-positive cancer patients

IMA203 Mann-Whitney U test, p=0.076

# PRAME Expression – RNAseq Data



# Combined with Immatics' Mass Spectrometry-guided RNA Threshold for Prevalence Prediction



IMA203 PRAME

PRAME trape prevalence is souted on TOGA MADescent agreemy with a comparison of the comparison of the

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



~39,000 Patients per Year in the US only

### **Selected Indications**

Initial indications of interest based on PRAME prevalence, patient population size and observed clinical responses Cut. Melanoma
Uveal Melanoma
Ovarian Carcinoma
Uterine Carcinoma
Uterine Carcinosarcoma
Synovial Sarcoma

Synovial Sarcoma Squamous NSCLC Small Cell Lung Cancer Cholangiocarcinoma Adeno NSCLC Breast Carcinoma HNSCC

Incidence	R/R Incidence	PRAME Positive
99,800	7,700	95%
1,500	800	90%
19,900	12,800	80%
62,700	10,700	100%
3,300	1,900	100%
1,000	400	100%
57,000	34,600	65%
31,900	19,400	55%
8,000	7,000	35%
91,200	55,300	25%
290,600	43,800	25% TNBC: 60%
66,500	15,100	25%

Patient Population
Based on R/R Incidence; PRAME and HLA-A*02:01+
2,999
295
4,198
4,387
779
164
9,221
4,375
1,005
5,668
4,490
1,548

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, liver cancer, 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

IMA203

Incidences based on public estimates and Immatics internal model; Relapsed/refractory (R/R) or last-line patient population approximated by annual mortality; Estimated 41% HLA-A\*02.01 positive population in the US; PRAME target prevalence is based on IMADetect and IMA

### IMA203 Monotherapy - Conclusions



ACTengine® IMA203 Targeting PRAME Offers a Unique Opportunity for Solid Cancer Patients

### IMA203 monotherapy Phase 1a and Phase 1b cohort A summary:

- IMA203 continues to be well tolerated with manageable safety profile
- Confirmed responses across a broad spectrum of different solid tumor types in heavily pre-treated patients
- Positively evolving durability profile for patients treated with higher doses and in phase 1b
- Clinical validation of PRAME biomarker threshold and associated prevalences
- We have clinically validated PRAME as one of the largest known T cell targets for solid cancers to date

# IMA203 development strategy:

- Transition to indication-specific development strategy
- Three Phase 1b expansion cohorts ongoing each designed to establish safety, evaluate the observed objective response rate, demonstrate durability & provide the trigger for registration trials

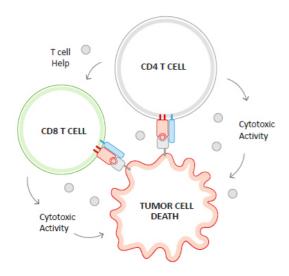
Data highlight the clinical potential of IMA203 TCR-T to achieve meaningful benefit for a large patient population

IMA203

# ACTengine® IMA203CD8 - Next-generation TCR-T



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



- Engagement of CD4 T cells by CD8 co-transduction reported to boost anti-tumor activity in TCR-T trials
- Recent data from leukaemia patients treated with CAR-T suggest a relevant role of engineered CD4 T cells in maintaining durable tumor responses over a long period of time<sup>1</sup>
- Functional superiority of the CD8αβ construct over multiple other CD8 constructs in preclinical experiments
- Proprietary 4-in-1 lentiviral vector to engineer CD4 and CD8 T cells with the PRAME-specific IMA203 TCR and CD8 $\alpha\beta$  construct (IMA203CD8)

IMA203CD8

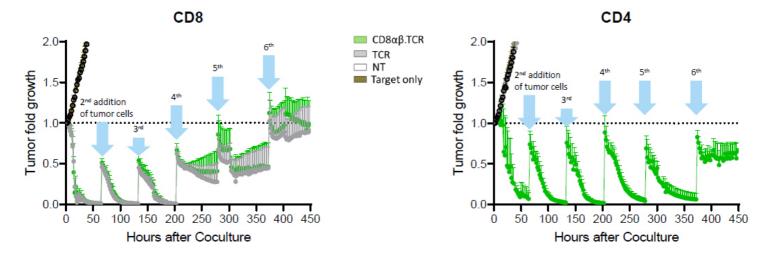
Melaphoret et al. 2022 Natura, Bai et al. 2022 Science Advance

-

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



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



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

IMA203CD8

# **Comprehensive PRAME Strategy**



# To Deliver Meaningful Clinical Benefit to Patients with PRAME-positive Cancers

### Phase 1b Dose Expansion **Upcoming Value Inflection Points** for Our PRAME Programs IMA203 Monotherapy First pt treated at Target Dose (plus DL5)1 Cohort A March 2022 Increasing T cell:Tumor cell Ratio<sup>2</sup> Phase 1a Dose Escalation ACTengine® IMA203: Next data read-outs with meaningful IMA203 at Target Dose IMA203 Monotherapy data across all cohorts in 2023 First pt treated plus Checkpoint Inhibitor<sup>3</sup> Cohort B in Basket Trial May 2022 Blocking PD-1/PD-L1 pathway Completed & provisional RP2D¹ determined IMA203CD8 2nd Gen First pt treated Aug 2022; in March 2022 $\oplus$ at Target Dose (plus DL5)\* **Cohort C** Adding functional CD4 T cells<sup>4</sup> First 3 pts on DL3\* TCER® IMA402: **Entering clinical development** in 2023

22D (target dose) determined at DLA, exploration of higher dose (DLS) ongoing; Demonstrated to be associated with durable response: Locke et al. 2020 Blood Advances; Option® (nivolumab); programmed death-1 (PD-1) immune checkpoint inhibito eatment of n=3 patients ongoing in DL3 prior to patient treatment at Target Dose (DL4), exploration of higher dose (DL5) planned; Demonstrated to be important for long-term remission: Melenhorst et al. 2022 Nature, Bai et al. 2022 Science Advance





ACTengine® IMA201 and IMA204

– TCR-T Targeting MAGEA4/8 and COL6A3

# ACTengine® IMA201 Targeting MAGEA4/8





### TARGE

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

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

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

Identified and validated by XPRESIDENT® quant. mass spectrometry platform

### TCF

High-affinity, specific TCR targeting MAGE4/8

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

Identified and characterized by XCEPTOR® TCR discovery and engineering platform

### CLINICAL DATA

Dose escalation ongoing, target dose level to commence

Too early for assessment of safety or anti-tumor activity

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

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

Status - 02-June-2022

IMA201

<sup>1</sup> Target density: peptide copy number per tumor cell, approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration

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



**Key Features** 

### ARGE

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, nextgeneration TCR engages both, CD8 and CD4 T cells

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

Complete tumor eradication in in vivo mouse models

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

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

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

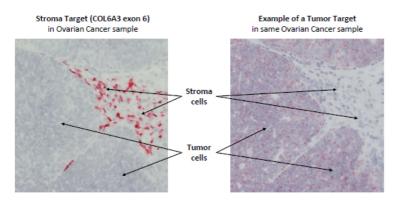
IMA204

<sup>1</sup> Target density: peptide copy number per tumor cell, approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>2</sup> Functional avidity: ECSO half maximal effective concentration and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications with 200 paper and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications with 200 paper and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications with 200 paper and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications with 200 paper and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications and approximate range representing the majority of tumor samples analyzed; <sup>3</sup> Solid paper indications and approximate range representing the rep

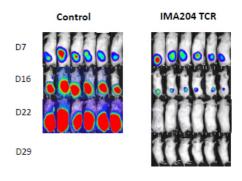
# ACTengine® IMA204 - High Affinity, CD8-independent TCR



# Complete Tumor Eradication in vitro & in vivo1 by Affinity-enhanced IMA204 TCR



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



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

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

IMA204 \* In vivo data in collaboration with Jim Riley, University of Pennsylvania, control: non-transduced T cells. TCR avidity and specificity data not shown, available in IMA204 presentation on Immatics website.

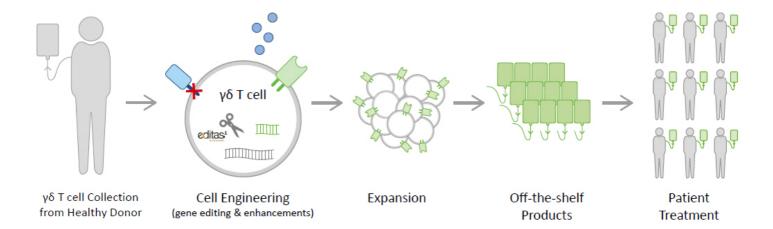




ACTallo® – Our Next-generation Off-the-shelf TCR-T

### ACTallo® – Immatics' Allogeneic Cell Therapy Approach





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

ACTallo® <sup>1</sup> Immatics proprietary ACTallo® platform utilizing Editas' CRISPR gene editing technology

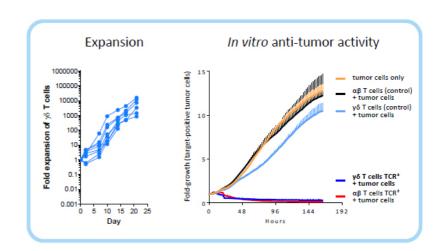
### Why γδ T cells?



### γδ T cells Are Well Suited for an Off-the-shelf Cell Therapy Approach

#### γδ 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 αβ TCR or CAR constructs



ACTallo®

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



Leveraging Complementary Technologies & Capabilities for the Benefit of Cancer Patients

#### **Immatics**

Innovative γδ-derived allogeneic cell therapy platform ACTallo®

State of the art manufacturing & gene editing via collaboration with Editas<sup>1</sup>

Activities for initial 2 BMS programs: Preclinical development

# Off-the-shelf allogeneic TCR-T/CAR-T therapies for patients with solid cancers



- · Initial 2 BMS programs
- Up to 4 additional BMS programs (TCRs developed in the context of the autologous TCR-T collaboration<sup>2</sup> might feed into allogeneic TCR-T programs)
- · Up to 4 Immatics programs

#### **Bristol Myers Squibb**

Complementary next-gen technologies to potentiate anti-tumor activity

> Expertise in oncology drug development and commercialization

Activities for initial 2 BMS programs: Clinical development and commercialization

ACTallo®

<sup>1</sup> Immatics proprietary ACTallo<sup>o</sup> platform utilizing Editas' CRISPR gene editing technology

Exclusive options to 3 Immatics TCR-T targets in 2019, expansion in 2022 by exclusive option to 1 additional Immatics TCR-T targ



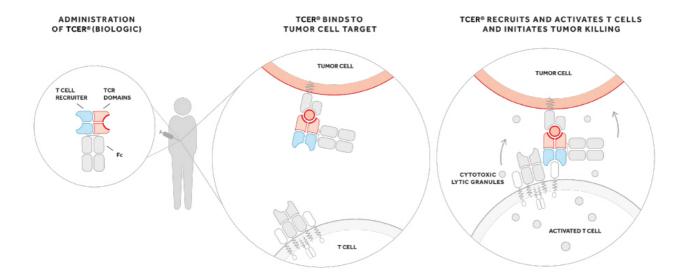


TCER® – TCR Bispecifics

### TCER® – Mechanism of Action



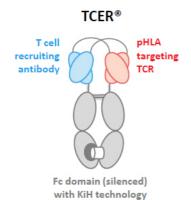
### Immatics' Off-the-Shelf TCR Bispecifics Approach



TCER®

### TCER® - Immatics' Half-Life Extended Bispecifics





#### pHLA targeting TCR

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

#### T cell recruiting antibody

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

#### Next-generation TCER® format

- ✓ Off-the-shelf biologic with antibody-like manufacturability³ 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

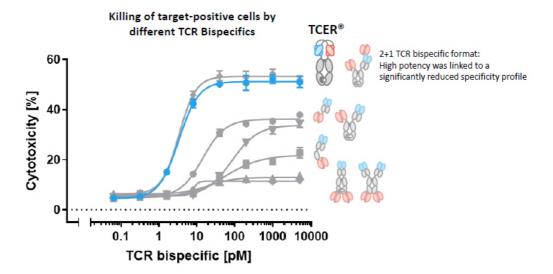
Our TCER® format is designed to maximize efficacy while minimizing toxicities in patients

TCER®

As compared to natural TCR; <sup>2</sup> Based on literature data for other low-affinity recruiters (e.g. Harber et al., 2021, Nature; Trinklein et al., 2019, mAbs); Production in mammalian cells (CHO cells); Based on precinical testing

### 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 specificity¹ 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®

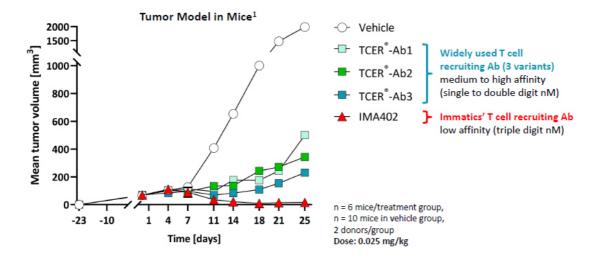
Preclinical data on specificty not show

٠,

### 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®

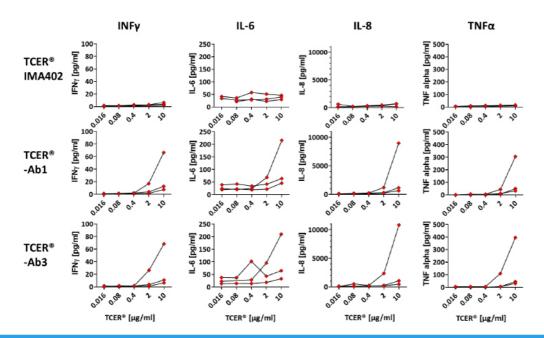
<sup>1</sup> Hs695T xenograft model in NOG mice, tumor volume of group means show

4:

### 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

TCER®

### TCER® Portfolio



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

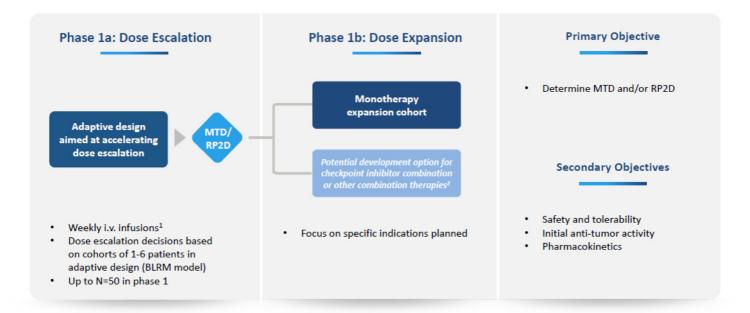
	IMA401	IMA402	IMA40X
	MAGEA4/8	PRAME	Undisclosed targets
Status	Start of Phase 1 trial in May 2022	Submission of CTA/IND application 2Q 2023, Phase 1/2 trial in 2023	TCER® engineering and preclinical testing ongoing
Preclincial Proof-of-concept – Efficacy / Safety	<ul> <li>Complete remission of estab. tumors in xenograft mouse models at low doses</li> <li>Very broad therapeutic window (reactivity tumor compared to normal cells)</li> </ul>		
Half-life	Half-life extended to several days via effector function silenced Fc part		
Clinical Development Strategy	<ul> <li>First-in-human basket trial</li> <li>Adaptive design aiming at fast dose escalation</li> <li>Development strategy includes TCER® as add on to checkpoint inhibitor-based standard of care in early lines of treatment</li> </ul>		

TCER®

A.

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





IMA401

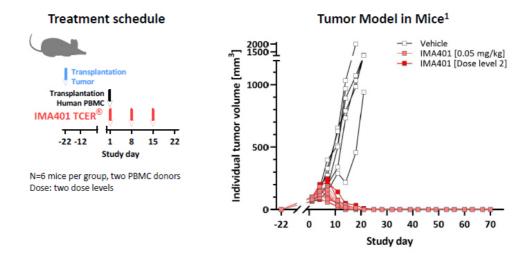
MTD: maximum tolerated dose, RP2D: recommended phase 2 dose; MABEL: minimum anticipated biological effect level; BLRM: Bayesian logistic regression model;

1 Pharmacokinatics data assessed throughout the trial might provide an opportunity to optimize scheduling to a less frequent regimen. 2 Conducted in collaboration with RM

### TCER® IMA401 Targeting MAGEA4/8



**Product Candidate in Clinical Development with Bristol Myers Squibb** 



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

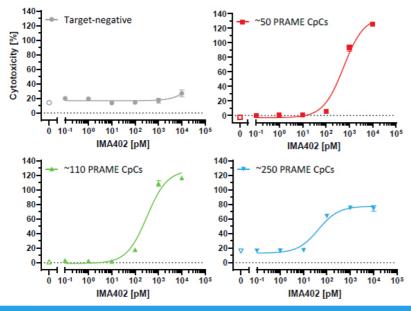
IMA401

<sup>1</sup> Hs695T xenograft model in MHC I/II ko NSG mice, tumor volume of individual mice shown

### TCER® IMA402 Targeting PRAME - Efficacy Assessment in vitro



Tumor Cell Killing at Low Physiological PRAME Peptide Levels

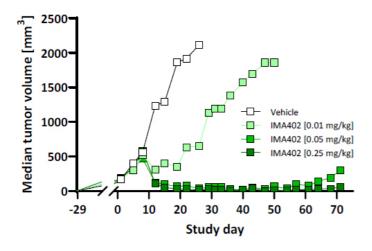


- TCER® IMA402 induces killing of tumor cells with PRAME target copies as low as 50 CpCs
- Physiological PRAME levels detected in majority of cancer tissues from patients are 100 – 1000 CpCs

IMA402

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



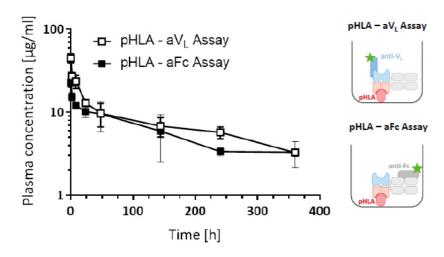


- Dose-dependent efficacy of IMA402 in cell line-derived in vivo mouse model
- Durable shrinkage of large tumors including complete responses over prolonged period
- Sufficiently high drug doses are key to achieving desired anti-tumor effect

IMA402

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





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

IMA402

### TCER® IMA402 Phase 1/2 Clinical Trial to Start in 2023



#### CMC and supply activities on track for clinical trial

- Manufacturing process development completed
- High titer (>3.5 g/L) and good stability allowing liquid formulation



#### **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

#### Phase 1: Dose Escalation

Adaptive design aimed at accelerating dose escalation

- MTD RP2D
- Basket trial in focus indications for accelerated signal finding
- Initially weekly i.v. infusions#
- MABEL-based starting dose
- Dose escalation decisions based on cohorts of 1-6 patients in adaptive design (BLRM model)

#### Phase 2a: Dose Expansion

**Expansion cohort** 

**Expansion cohort** 

**Expansion cohort** 

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

IMA402 MABEL: r



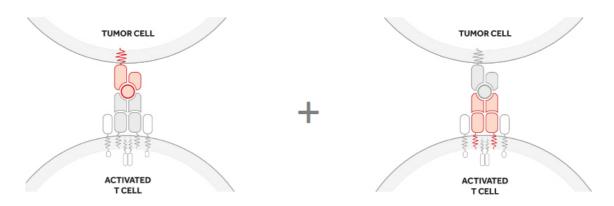


Immatics' Proprietary Target and TCR Discovery Platforms

### **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

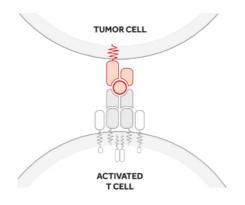
- · recognize the target peptide with high affinity and specificity
- · show selective killing of tumor cells
- are developed to be suitable for two different therapeutic modalities, Cell Therapies and TCR Bispecifics

Technology

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



**Example of MAGEA4/8 Peptide Target** 



Ranking of pHLA targets

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

MAGEA4/8

Immatics

MAGEA4

Commonly used

peptide

MAGEA4 and MAGEA4/8

Peptide (AbsQuant®)

p<0.001<sup>2</sup>

100000-

10000

1000 100 10-

AbsQuant® CpC

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

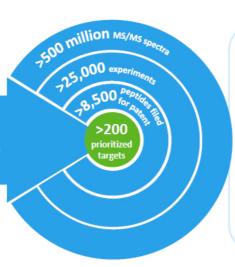
Technology <sup>1</sup>Copy number per tumor cell (CpC) measured on a paired-sample basis by AbsQuant<sup>9</sup>, i.e. comparing MAGEA4 vs. MAGEA4/A8 peptide presentation on same sample, <sup>2</sup>Students paired T test

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



pHLA Database based on primary tissues

>2,500 cancer & normal tissues analyzed by Quantitative, Ultra-Sensitive Mass Spectrometry



#### **200 Prioritized Targets**

#### Grouped in 3 Target Classes:

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

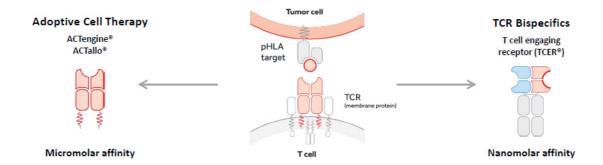
~50% of our prioritized targets are non-HLA-A\*02 restricted, substantially broadening the potential patient reach

Technology 53

### 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 maturate 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 discovery¹ and TCR maturation²

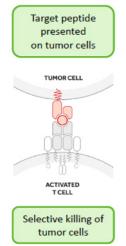
Technology

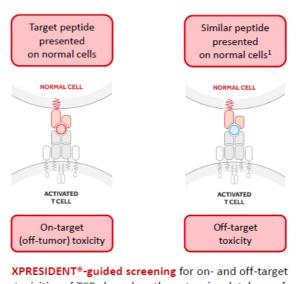
\*XPRESIDENT\*-guided off-target toxicity screening; \*XPRESIDENT\*-guided similar peptide counterselection

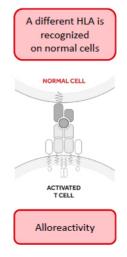
### Optimal Target Selection & TCR Specificity for Minimizing Safety Risks



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







toxicities of TCRs based on the extensive database of peptides presented on normal tissues

Technology

<sup>1</sup>Clinical fatalities have occurred in TCR-T trials using a titin cross-reactive TCR (Cameron et al., Sci Transl Med)





## **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,
Probiodrug)



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



Cedrik Britten
Chief Medical Officer
>14 yrs pharma & biotech
(GSK, BioNTech)



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



Steffen Walter Chief Technology 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)

Corporate

### Strong, Focused and Highly Integrated Trans-Atlantic Organization





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



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



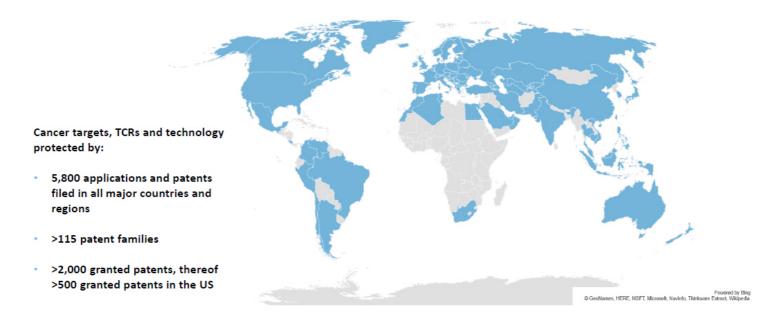
Munich, Germany, ~55 FTEs Various operating functions

Corporate FTE status as of June 2022

#### **Robust IP Portfolio**







Corporate

#### **Near-Term Value Drivers**



#### **Clinical Development of TCR-T and TCR Bispecifics**



#### Advance clinical development of ACTengine® IMA203 candidates

- Multiple IMA203 Ph1b expansion cohorts:
   Monotherapy, checkpoint combination, 2nd-gen approach IMA203CD8
- Next data read-outs with meaningful data across all cohorts in 2023
- · Additional ACTengine® programs in clinical and preclinical development



#### Further clinical development of TCER® candidates

- Ongoing Ph1 trial for IMA401 (MAGEA4/8) (start in May 2022)
- IMA402 Ph 1/2 clinical trial on track to start in 2023; submission of CTA/IND application 2Q 2023
- · Innovative TCER® program(s) IMA40X in preclinical development

Solid cash runway into 2025 to reach multiple value inflections points across our portfolio

Corporate

.

# **Delivering**

the Power of T cells to Cancer Patients



www.immatics.com

Dimmatics. Not for further reproduction or distribution