#### 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

January 9, 2023

Commission File Number: 001-39363

# **IMMATICS N.V.**

Paul-Ehrlich-Straße 15 72076 Tübingen, Federal Republic of Germany (Address of principal executive office)

Indicate by check mark whether the registrant files or will file annual reports under cover of Form 20-F or Form 40-F:

Form 20-F	$\times$
	-

Form 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): 🗆

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

On January 9, 2023, 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 January 9, 2023 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.Description99.1Presentation dated January 9, 2023

#### 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.

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

Date: January 9, 2023

# Immatics Corporate Presentation

January 09, 2023



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### **Building a Leading TCR Therapeutics Company**





Two Clinical-Stage Modalities

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



Clinical PoC for Cell Therapy

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

# Our Near-Term Focus – Clinical Development of Our Lead Assets from Our Autologous TCR-T (ACTengine®) and TCR Bispecifics (TCER®) Pipeline







# 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>®</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 1	a1
	ACTengine <sup>®</sup> IMA203	PRAME	immatics	+ (	Checkpoint Inhib	oito
Autologica ACT	ACTengine <sup>®</sup> IMA203CD8	PRAME	immatics			
Autologous ACT	ACTengine <sup>®</sup> IMA201	MAGEA4/8	immatics			
	ACTengine <sup>®</sup> IMA204	COL6A3	immatics			
	Multiple programs	Undisclosed	🐴 Bristol Myers Squibb'			
Allogeneic ACT	ACTallo® IMA30x	Undisclosed	immátics editas*			
γδ T cells	Multiple programs	Undisclosed	💾 Bristol Myers Squibb'			
	TCER® IMA401	MAGEA4/8	( <sup>4)</sup> Bristol Myers Squibb'			
Discosifies	TCER® IMA402	PRAME	immatics			
Bispecifics	TCER <sup>®</sup> IMA403	Undisclosed	immátics			
	Multiple programs	Undisclosed	Genmab			

Intro

<sup>1</sup> Phase 1a: Dose escalation, Phase 1b: Dose expansion; <sup>2</sup> Opdivo<sup>9</sup> (nivolumab): programmed death-1 (PD-1) immune checkpoint inhibitor; <sup>1</sup> Immatics proprietary ACTallo<sup>9</sup> platform utilizing Editas' CRISPR gene editing technology

7





ACTengine<sup>®</sup> IMA203 – TCR-T Targeting PRAME

# ACTengine® IMA203 Targeting PRAME – Mechanism of Action

Immatics' Leading TCR-T Approach



immatics

### Multi-Tumor Target PRAME

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



#### PRAME Peptide Target

- 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

# PRAME TCR

TUMOR CELL

HLA-A\*02:01

PRAME Peptide

#### 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

#### PRAME RNA detection in tumor samples (ISH)



Patient screening data from Immatics' clinical trials support

high prevalence of PRAME: Uterine Carcinoma 90% Cut. Melanoma 95% Uveal Melanoma<sup>2</sup> 90% Ovarian Carcinoma 70%

	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%



# ACTengine<sup>®</sup> IMA203 – Patient Flow



### IMA203 TCR-T Phase 1 Design

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



IMA203 <sup>1</sup>RP2D (target dose) determined at DL4, exploration of higher dose (DL5) ongoing, <sup>2</sup>Demonstrated to be associated with durable response. Locke *et al.* 2020 Blood Advances, <sup>3</sup>Opdivo<sup>®</sup> (nivolumab): programmed death-1 (PD-1) immune checkpoint inhibitor; Treatment of n=3 patients at DL3 completed, enrollment at Target Dose DL4 ongoing, exploration of higher dose (DL5) planned; <sup>4</sup>Demonstrated to be important for long-term remission: Melenhorst *et al.* 2022 Nature, Bai *et al.* 2022 Science Advances



# 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

### ACTengine® IMA203 – Interim Monotherapy Update





10RR: Objective response rate (partial responses) according to RECIST 1.1 at first scan post infusion ("6 weeks); <sup>2</sup> confirmed Objective response rate (confirmed partial responses) according to RECIST 1.1 at second scan post infusion ("12 weeks); <sup>1</sup> 1 patient with SD at "6-week scan with pending "12-week scan considered as non-responder for confirmed ORR: 'Total transduced viable CDB T cells, all patients in Phase 1a DL4 and Phase 1b DL4/DL5; RP02: Recommended Phase 2 dose; DL: Dose level

### ACTengine® IMA203 Monotherapy – Patient and Product Characteristics



	Phas Dose Es	e 1a calation	Phase 1b (Cohort A) Dose Expansion
	All pts (DL1-4)	DL4 pts only	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)
Dose 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	ersions1	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

Data cut-off - 06-Sept-2022

1Except for 1 product for patient at DL3 generated with current manufacturing process; <sup>1</sup>DL4: 200m to 1.2bn transduced viable CD8 T cells per m<sup>3</sup> 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 m<sup>3</sup> BSA; ULN: Upper limit of normal; BSA: Body surface area; RP2D: Recommended Phase 2 dose; LHD: Lactate dehydrogenase



- 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 ICANS<sup>1</sup>: 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 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); 16

# **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) <sup>1</sup>	
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) <sup>*</sup>	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-2022

 1All patients received >1 x 10<sup>9</sup> total transduced viable CD8 T cells; <sup>2</sup> ORR: Objective response rate (partial responses) according to RECIST 1.1 at first scan post infusion (~5 weeks); <sup>3</sup> Confirmed ORR (cORR): Confirmed objective response rate (confirmed objective response rate (confirmed partial responses) according to RECIST 1.1 at second scan post infusion (~12 weeks); \*1 patient with SD at ~ 6-week scan with pending ~12-week scan considered as non-responder for cORR.
 17

### **Best Overall Response**

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



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

MA203	* Maximum change of target lesions and RECIST 1.1 BOR at different timepoints; <sup>4</sup> 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	18

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#### **Responses over Time**

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



IMA203 \*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



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



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#### PRAME Expression – RNAseq Data

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



IMA203 <sup>1</sup>PRAME target prevalence is based on TCGA RNAseq data combined with a proprietary mass spectrometry-guided RNA expression threshold; \* PRAME target prevalence in useal melanoma based on IMADetect\* qPCR testing of screening biopsies from clinical trial patients (n=21) demonstrates substantial higher prevalence of 90%, TCGA, early & late-stage primary tumor samples, Immatics clinical trials: late-stage/metastatic tumor samples, Role of PRAME in metastasis of useal melanoma: Field et al. 2016 Clinical Cancer Research

### 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

immatics

Data cut-off - 06-Sept-2022

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

~39,000 Patients per Year in the US only

Sel	ected Indications	Incidence	<u>R/R Incidence</u>	PRAME Positive	Patient Population Based on R/R Incidence; PRAME and HLA-A*02:01+
total indications of interest	Cut. Melanoma	99,800	7,700	95%	2,999
Initial Indications of Interest	Uveal Melanoma	1,500	800	90%	295
based on PRAIVE prevalence,	Ovarian Carcinoma	19,900	12,800	80%	4,198
patient population size and	Uterine Carcinoma	62,700	10,700	100%	4,387
observed clinical responses	Uterine Carcinosarcoma	3,300	1,900	100%	779
	Synovial Sarcoma	1,000	400	100%	164
Squamous NSCLC Small Cell Lung Cancer		57,000	34,600	65%	9,221
		31,900	19,400	55%	4,375
	Cholangiocarcinoma	8,000	7,000	35%	1,005
	Adeno NSCLC	91,200	55,300	25%	5,668
Breast Carcinoma		290,600	43,800	25% TNBC: 60%	4,490
	HNSCC	66,500	15,100	25%	1,548

<u>TOTAL ~39,000</u> 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  $\rightarrow$  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-4\*02:01 positive population in the US; PRAME target prevalence is based on IMADetect\* QPCR testing of screening biopsies from clinical trial patients (n=21) 24



#### 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



#### 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 CD8αβ construct (IMA203CD8)

IMA203CD8 <sup>1</sup> Melenhorst *et al.* 2022 Nature, Bai *et al.* 2022 Science Advances

### ACTengine<sup>®</sup> 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

27



### **Comprehensive PRAME Strategy**

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



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ACTengine<sup>®</sup> IMA201 and IMA204 – TCR-T Targeting MAGEA4/8 and COL6A3

### ACTengine<sup>®</sup> IMA201 Targeting MAGEA4/8

**Key Features** 



Status – 02-June-2022

30

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: EC50 half maximal effective concentration; <sup>3</sup>Solid cancer indications with 20% or more target expression, Target prevalence for selected cancer indications based on mRNA expression (TCGA and Immatics inhouse data)

# ACTengine<sup>®</sup> IMA204 First-in-Class TCR-T Targeting Tumor Stroma Key Features

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TARGET	TCR	PRECLINICAL DATA	PATIENT POPULATION <sup>3</sup>
HLA-A*02-presented peptide derived from <b>COL6A3 exon 6</b> Naturally and specifically presented on tumors at high target density <sup>1</sup> : <b>100-700 copies/cell</b> Novel <b>tumor stroma target</b> identified and validated by XPRESIDENT <sup>®</sup> quant. mass spectrometry platform	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	CD8-independent, next- generation TCR engages both, CD8 and CD4 T cells <i>In vitro</i> anti-tumor activity against target-positive cell lines in CD8 and CD4 T cells Complete tumor eradication in <i>in vivo</i> mouse models	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: EC50 half maximal effective concentration; <sup>3</sup>Solid cancer indications with 20% or more target expression, Target prevalence for selected cancer indications based on mRNA expression (TCGA and Immatics inhouse data)



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

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



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

IMA204 <sup>1</sup> 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. 32





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

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

# immatics



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

ACTallo<sup>®</sup>

34

# 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









# TCER<sup>®</sup> – TCR Bispecifics

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



**Proprietary TCER® Format Consisting of Three Distinct Elements** 



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





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

\* As compared to natural TCR; \* 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<sup>®</sup> format had higher combination of potency and specificity<sup>1</sup> than six alternative TCR Bispecific format designs evaluated Flexible Plug-and-play platform: TCER<sup>®</sup> format successfully validated for different TCRs & different T cell recruiting antibodies

TCER<sup>®</sup> \* Preclinical data on specificty not shown

39



### TCER<sup>®</sup> 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<sup>®</sup> molecules designed with higher-affinity variants of a widely used recruiter

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

### TCER<sup>®</sup> Format Is Designed for Optimized Efficacy and Safety

immatics

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



# Our TCER<sup>®</sup> Portfolio

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

	IMA401 M Bristol Myers Squitbri	<ul> <li>MAGEA4/8 peptide presented by HLA-A*02:01</li> <li>Start of clinical trial in May 2022, dose escalation ongoing</li> </ul>	
	IMA402	<ul> <li>PRAME peptide presented by HLA-A*02:01</li> <li>Start of clinical trial planned for 2023</li> </ul>	Potential for addressing
	IMA403	<ul> <li>Undisclosed peptide presented by HLA-A*02:01</li> <li>Preclinical PoC studies ongoing</li> </ul>	large patient populations with novel, off-the-shelf TCR Bispecifics
	IMA40x Several innovative programs	<ul> <li>Undisclosed peptides presented by HLA-A*02:01 and other HLA-types</li> <li>TCER<sup>®</sup> engineering and preclinical testing ongoing</li> </ul>	
<b>TCER</b> <sup>®</sup>			42

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



Tumor Cell Killing at Low Physiological PRAME Peptide Levels



IMA402 CpC: Target peptide copy numbers per tumor cell

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





- 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











- 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



46

### Advancing TCER® IMA402 Towards Clinical Development

**Recent and Upcoming Activities** 

#### **Recent activities**

- Completion of IND-enabling data package
- ✓ Manufacturing of GMP batch completed with high titer (>3.5 g/L) and high yield
- ✓ Scientific advice with regulatory authorities

#### **Upcoming activities**

- CTA/IND submission planned for 2Q 2023
- Start of patient treatment planned in 2023

IMA402 TCER® Ph1/2 clinical trial in patients with solid tumors



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





IMA402 MABEL: minimum anticipated biological effect level

#### Accelerated Development of TCER<sup>®</sup> IMA402

TCER<sup>®</sup> IMA402 Phase 1/2 Clinical Trial Design





IMA402 MABEL: minimum anticipated biological effect level; DLT period: Evaluation period for potential dose limiting toxicities (DLT) in a patient





Immatics' Proprietary Target and TCR Discovery Platforms

#### **True Cancer Targets & Matching Right TCRs**





#### 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

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





Technology



# Immatics' Unique Capability – Identification of the most Relevant Target Example of MAGEA4/8 Peptide Target



#### Development of the Right TCR - XCEPTOR<sup>®</sup> 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<sup>®</sup> and XCEPTOR<sup>®</sup> during TCR discovery<sup>1</sup> and TCR maturation<sup>2</sup>

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

Immatics

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



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



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

54

# **Robust IP Portfolio**

Immatics' Patent Estate – Territorial Coverage





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regions





# **Corporate Information & Milestones**

### Experienced Global Leadership Team Across Europe and the US





Corporate

# Strong, Focused and Highly Integrated Trans-Atlantic Organization



Corporate FTE status as of June 2

# Delivering

the Power of T cells to Cancer Patients







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58