

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

# Unlocking Immunotherapies for Solid Cancer Patients

Immatics Corporate Presentation, April 2021

# Forward-Looking Statements



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# Unlocking Immunotherapies for Solid Cancer Patients



**Two Transformative Treatment Modalities:  
Adoptive Cell Therapies and TCR Bispecifics**



**Highly Differentiated Technologies to Identify  
True Cancer Targets and the Right TCRs**

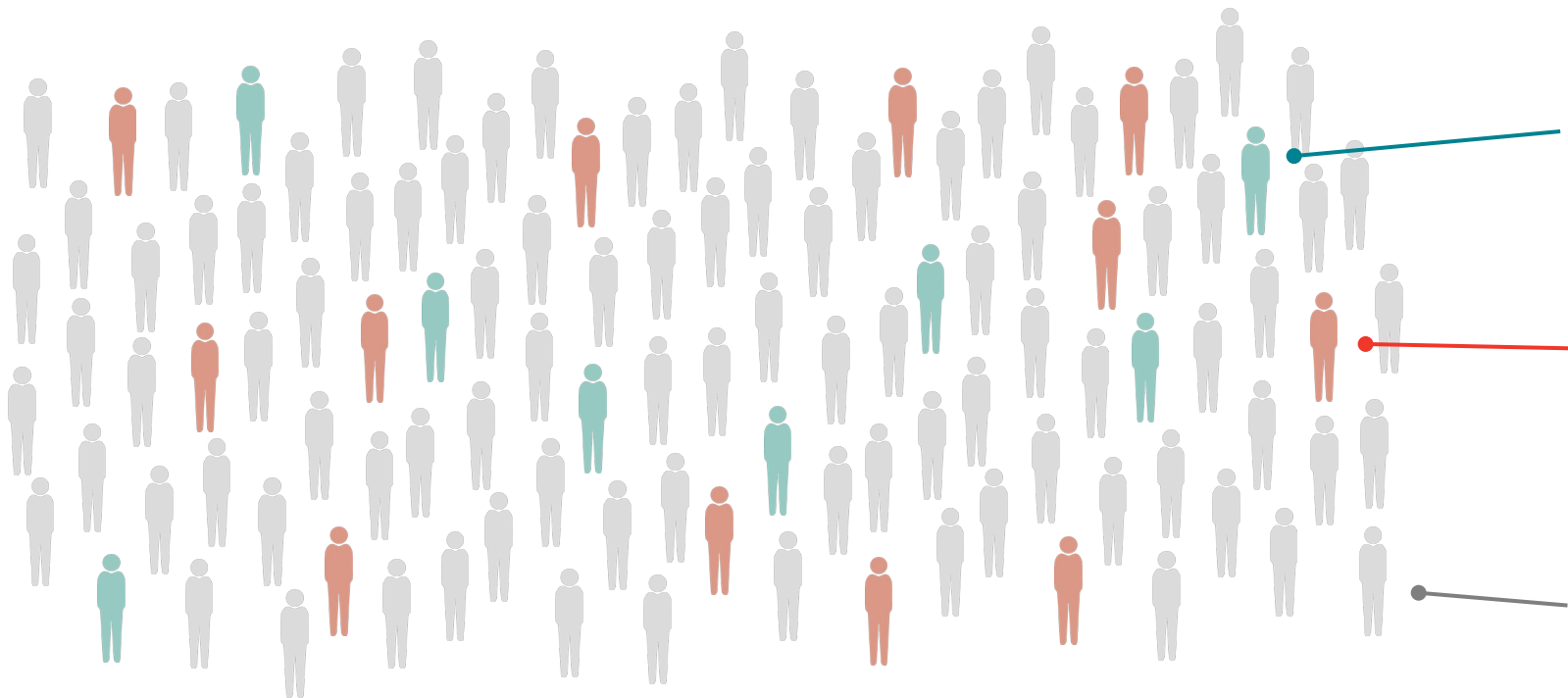


**Strategic Collaborations with World-leading  
Industry Players**

# Limitations of Current Immunotherapies in Solid Cancer Patients

## ... Driven by a Lack of Known Cancer-specific Targets

Most cancer patients do not benefit from current immuno-oncology approaches



**Checkpoint inhibitors**  
mainly effective in tumors with high mutational burden  
**minority of all cancers<sup>1</sup>**

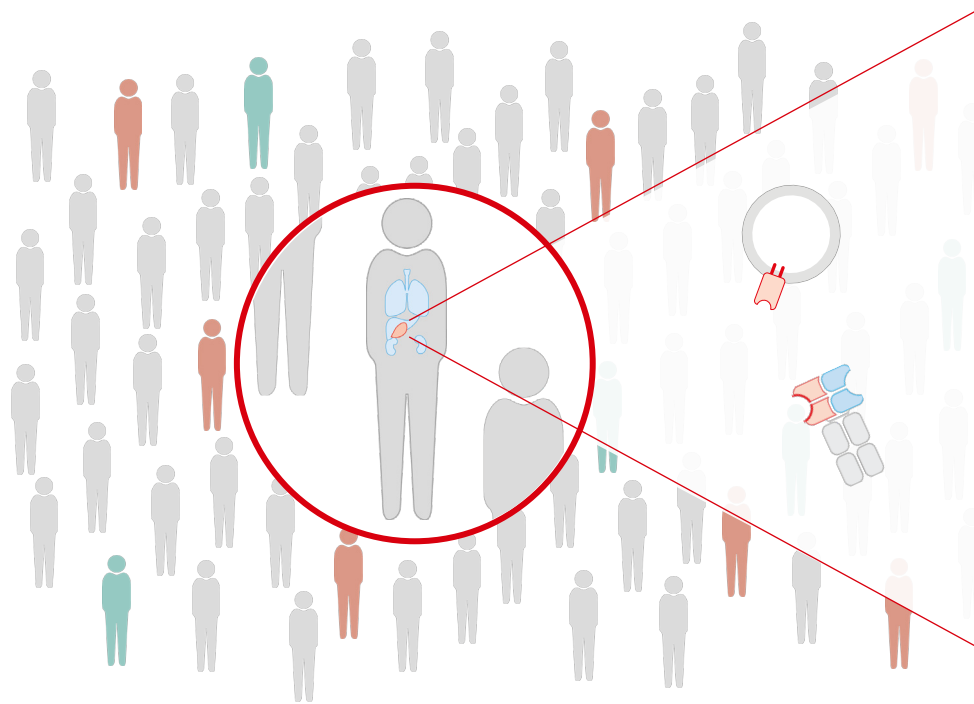
**CAR-T**  
mainly effective in hematological malignancies  
**minority of all cancers<sup>2</sup>**

**Solid tumors**  
limited established treatments & high medical need  
**majority of all cancers**

**We are unlocking immunotherapies for solid cancer patients with high unmet medical need by accessing intracellular cancer targets with TCR-based therapeutics**

# The Immatics Approach to Disrupt Current Tumor Treatment Paradigms

## Based on 5 Defined Principles



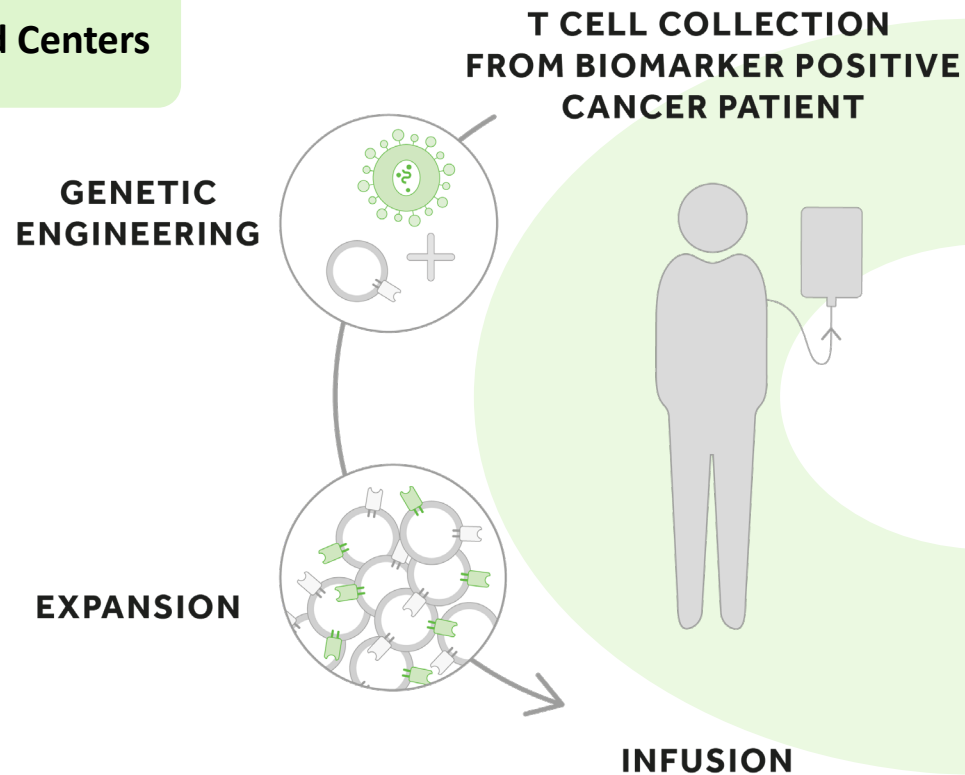
1. True Cancer Targets & Matching Right TCRs
2. Targeted Approach in Two Distinct Modalities: Adoptive Cell Therapy & TCR Bispecifics
3. Optimized Manufacturing to Enhance T cell Persistence & Efficacy
4. Disrupting the Tumor Microenvironment by Targeting Stroma
5. Combating Tumor Heterogeneity & Escape through Multi-Target Approach

# Immatics' Targeted Approach in Two Distinct Modalities

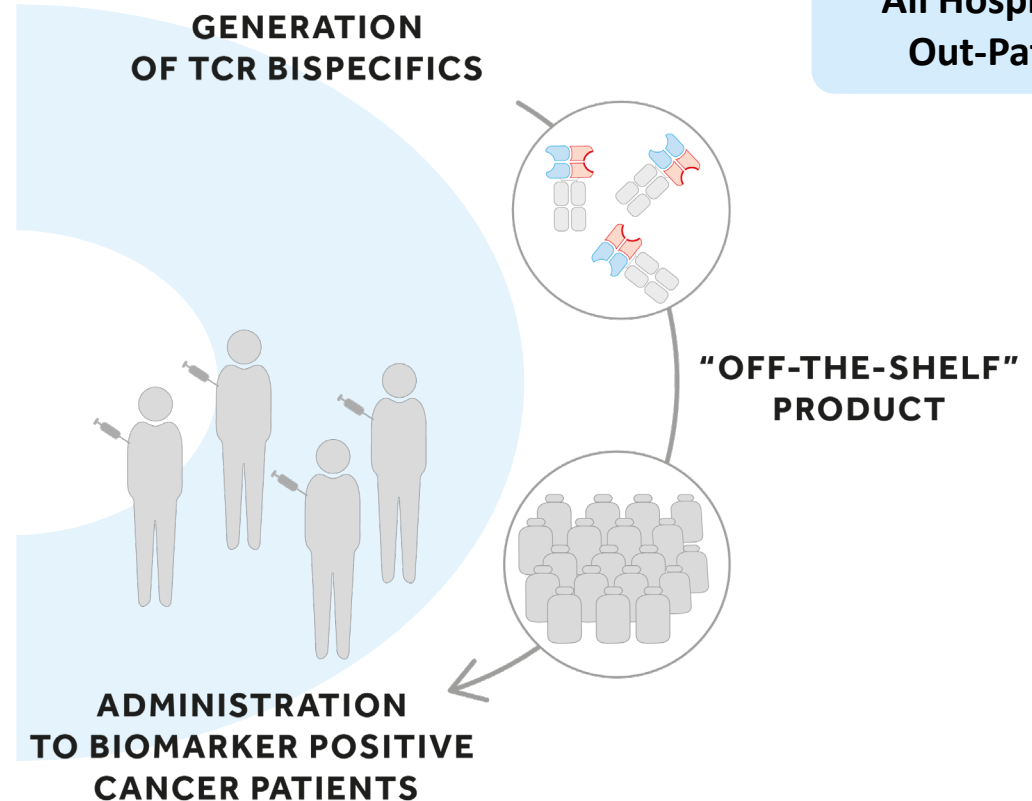
High tumor burden

Specialized Centers

## Adoptive Cell Therapy ACTengine®


















## TCR Bispecifics TCER®



Lower tumor burden<sup>1</sup>

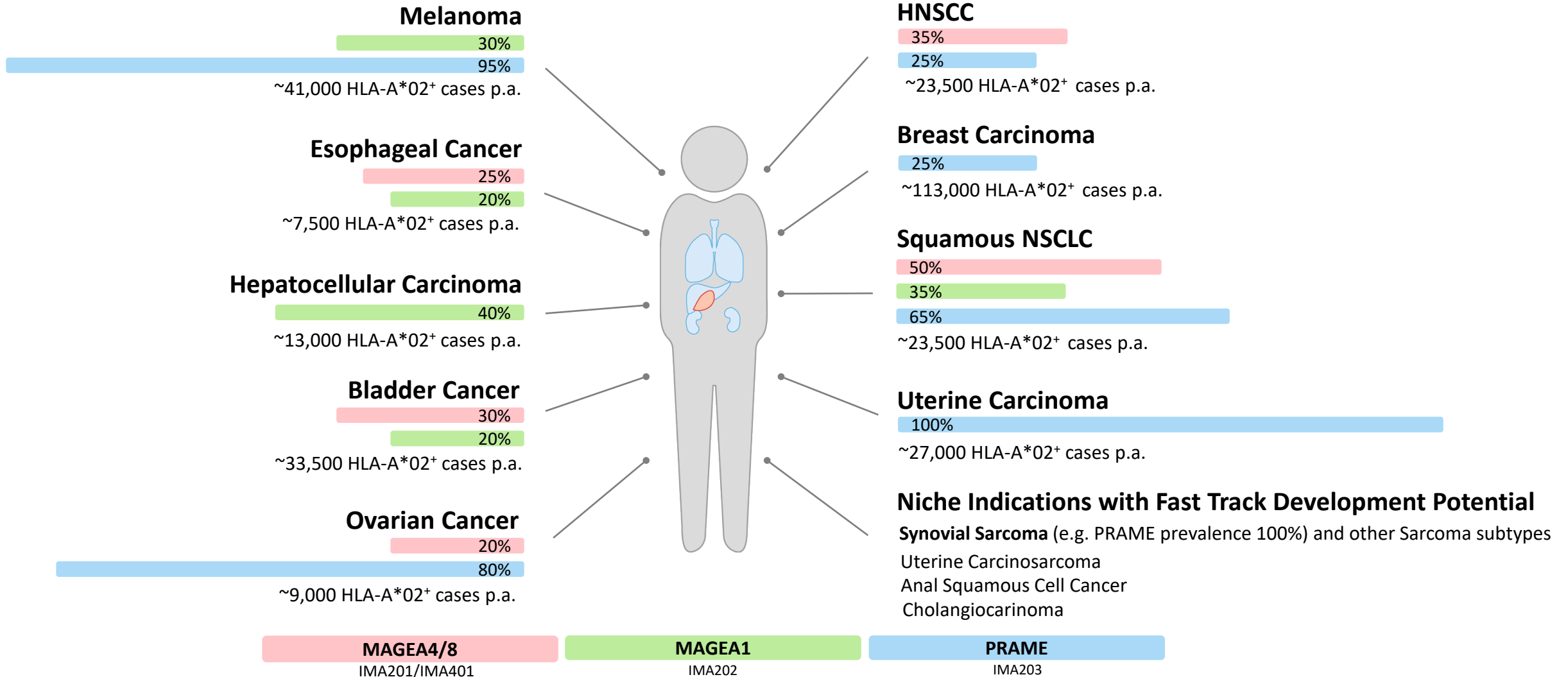
All Hospitals & Out-Patient

# Immatics' Pipeline

Modality	Product Candidate	Status	Preclinical	Phase 1a <sup>1</sup>	Phase 1b <sup>1</sup>	Phase 2	Phase 3
<b>Autologous ACT</b>	ACTengine® IMA201 (MAGEA4/8)	Proprietary					
	ACTengine® IMA202 (MAGEA1)	Proprietary					
	ACTengine® IMA203 (PRAME)	Proprietary					
	ACTengine® IMA204 (COL6A3)	Proprietary					
	ACT programs (Undisclosed)	 Bristol Myers Squibb™					
<b>Allogeneic ACT</b>	ACT programs (Undisclosed)						
	ACTallo® IMA301 (Undisclosed)	Proprietary					
<b>Bispecifics</b>	TCER® IMA401 (MAGEA4/8)	Proprietary					
	TCER® IMA402 (Undisclosed)	Proprietary					
	Bispecific programs (Undisclosed)						
	Bispecific programs (Undisclosed)						

# Significant Addressable Market in Solid Cancers

## High Prevalence of MAGE4/8, MAGEA1 and PRAME in Major Tumor Indications





## Adoptive Cell Therapy

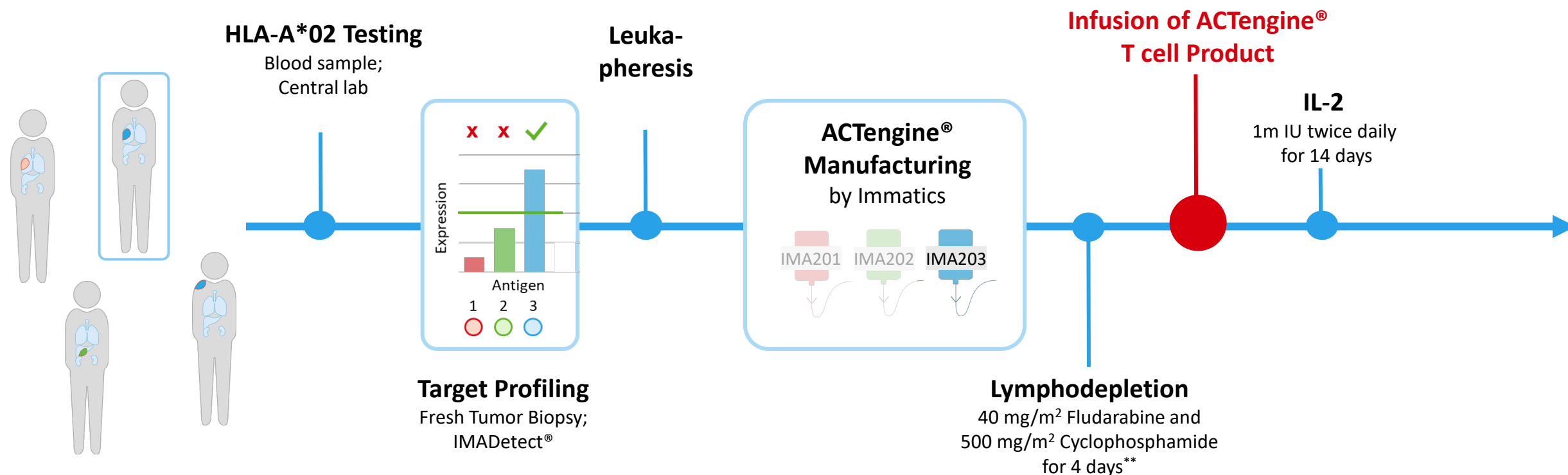
# Key Features of Our Clinical ACTengine® Programs

Differentiated Targets, TCRs and Cellular Manufacturing Designed to Enhance Safety and Activity

	IMA201	IMA202	IMA203
	HLA-A*02-presented peptide derived from		
<b>Peptide Target</b>	<b>MAGEA4/8</b> shown to be naturally and specifically presented on native tumor tissues at differentiated high peptide target density <sup>1</sup> 100-1,000 copies/cell	<b>MAGEA1</b> 50-900 copies/cell	<b>PRAME</b> 100-1,000 copies/cell
<b>T cell Receptor (TCR)</b>	High-affinity specific TCRs with high functional avidity <sup>2</sup>		
	Natural TCR ~10 ng/ml	Natural TCR ~15 ng/ml	Pairing-enhanced TCR ~5 ng/ml
<b>T cell Product</b>	Autologous T cells gene-engineered with lentiviral vector expressing TCR and applying proprietary short-term manufacturing process designed to achieve better T cell engraftment and persistence		
	7-10 days <sup>3</sup>	7-10 days <sup>3</sup>	6-7 days <sup>3</sup>

# ACTengine® Clinical Programs – Clinical Overview & Patient Flow

High Enrollment Efficiency through Combined Screening for Three Targets



14

Patients infused across three TCR-T Programs, as of data cut-off on Feb 16, 2021\*

<1bn

T cells infused per patient at dose levels 1 and 2 – presumed to be sub-therapeutic

# ACTengine® Clinical Programs – Safety Profile

## Treatment-emergent Adverse Events Are Manageable, Transient and Expected for Cell Therapies

### Adverse Events:

- Most frequent adverse events were transient cytopenias associated with lymphodepletion
- Transient CRS<sup>3</sup> (Grade 1-2) in 13/14 infused patients.
- Transient Grade 1 or 2 ICANS in 3/14 infused patients, resolved within 48h in all cases

### Dose-limiting toxicities:

- IMA201 and IMA202: No DLT<sup>5</sup> observed
- IMA203: One transient, Grade 3 atrial fibrillation with onset on day 5 post infusion that resolved within 48h after onset. DLT triggered expansion of dose level 2 from three to six patients

All treatment-emergent adverse events (TEAEs) with grade 1-2 occurring in at least 5 patients (incidence  $\geq 31.3\%$ ) and additionally all events with grade 3-5 regardless of relatedness to study treatment are presented. Data source: clinical and safety database; hematological adverse events were derived from lab values. Grades were determined according to National Cancer Institute Common Terminology Criteria of Adverse Events, version 5.0. Grades for CRS and ICANS were determined according to CARTOX criteria (Neelapu et al, 2018). Patients are counted only once per adverse event and severity classification.

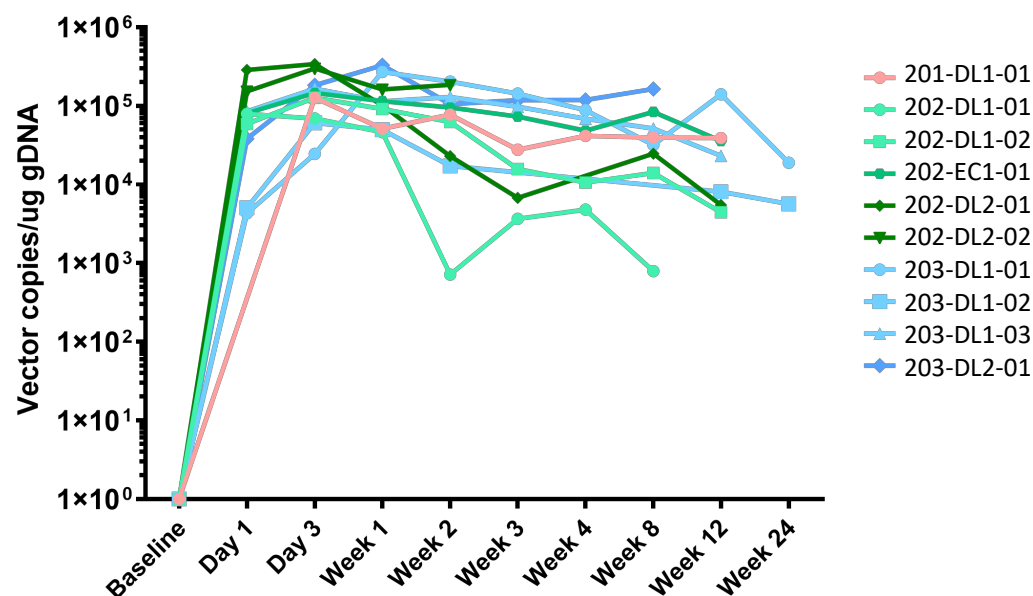
TEAEs by maximum severity (N=16)				
Adverse event	All Grades		$\geq$ Grade 3	
	No.	%	No.	%
<b>Patients with any adverse event</b>	<b>16</b>	<b>100.0</b>	<b>16</b>	<b>100.0</b>
<b>Lymphopenia</b>	16	100.0	16	100.0
<b>Leukopenia</b>	16	100.0	16	100.0
<b>Neutropenia</b>	16	100.0	15	93.8
<b>Anaemia</b>	16	100.0	10	62.5
<b>Thrombocytopenia</b>	15	93.8	6	37.5
Nausea	11	68.8	0	0
Pyrexia	8	50.0	0	0
Vomiting	6	37.5	1	6.3
Fatigue	5	31.3	1	6.3
Hypoxia	5	31.3	1	6.3
Hyponatraemia	5	31.3	0	0
Dyspnoea <sup>1</sup>	3	18.8	1	6.3
Atrial fibrillation	2	12.5	1	6.3
Hypertension	2	12.5	1	6.3
Muscular weakness	2	12.5	1	6.3
Pleural effusion	2	12.5	1	6.3
Tumor pain	2	12.5	1	6.3
Blood alkaline phosphatase increased	1	6.3	1	6.3
Candida infection	1	6.3	1	6.3
Corona virus infection	1	6.3	1	6.3
Febrile neutropenia	1	6.3	1	6.3
Infection	1	6.3	1	6.3
Pneumonia <sup>1</sup>	1	6.3	1	6.3
Sepsis <sup>2</sup>	1	6.3	1	6.3
<b>Adverse Events of Special Interest</b>				
<b>Cytokine release syndrome<sup>3</sup></b>	13	81.3	0	0
<b>ICANS<sup>4</sup></b>	3	18.8	0	0

Data cut-off – February 16, 2021

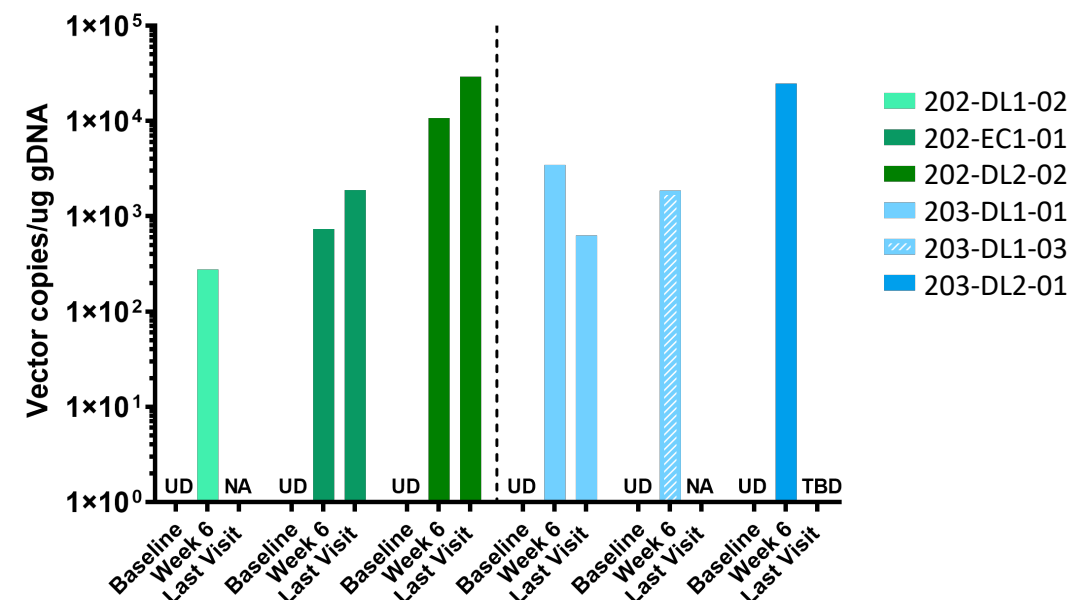
# ACTengine® Clinical Programs – Biological Activity

T cells Robustly Engraft, Persist and Infiltrate into Tumor after Infusion of Low Doses of ACTengine®

**Engraftment & T cell Persistence in the Blood**



**Detection of T cells in the Tumor**



- Robust T cell engraftment and persistence post infusion until the end of the observation period as assessed by qPCR\*
- Engineered T cells are detectable in serial tumor biopsies post T cell infusion in all evaluable patients by qPCR

Data cut-off – February 16, 2021

# ACTengine® Clinical Programs – Best Overall Response (BOR) Assessment



Disease Control in 9 out of 10 Patients at Dose Level 1 and 2 (below 1 Billion Transduced CD8 T cells)

	IMA201	IMA202					IMA203			
Patient	201-DL1-01	202-DL1-01	202-DL1-02	202-EC1-01	202-DL2-01	202-DL2-02	203-DL1-01	203-DL1-02	203-DL1-03	203-DL2-01
Dose level	DL1	DL1	DL1	EC1	DL2	DL2	DL1	DL1	DL1	DL2
Total transduced cells <sup>1</sup>	0.11x10 <sup>9</sup>	0.11x10 <sup>9</sup>	0.09x10 <sup>9</sup>	0.19x10 <sup>9</sup>	0.51x10 <sup>9</sup>	0.65x10 <sup>9</sup>	0.12x10 <sup>9</sup>	0.11x10 <sup>9</sup>	0.08x10 <sup>9</sup>	0.35x10 <sup>9</sup>
Age (gender)	60 (M)	33 (M)	63 (F)	64 (F)	68 (F)	49 (M)	40 (F)	63 (M)	61 (F)	57 (M)
Diagnosis	NSCLC	HNSCC	Squamous Cell Cancer	Melanoma	Squamous Cell Cancer	Melanoma	Head and Neck Cancer	Ovarian Cancer	Synovial Sarcoma	
Prior lines of systemic therapy	4	5	6	4	3	7	6	4	7	2
Prior lines of ICI <sup>2</sup> treatment	1	3	1	2	1	3	2	-	1	-
Disease status at infusion	Patients with recurrent and/or refractory solid tumors									
Best response RECIST1.1	SD	SD	SD	SD	SD	PD	SD	SD	SD	PR <sup>3</sup>

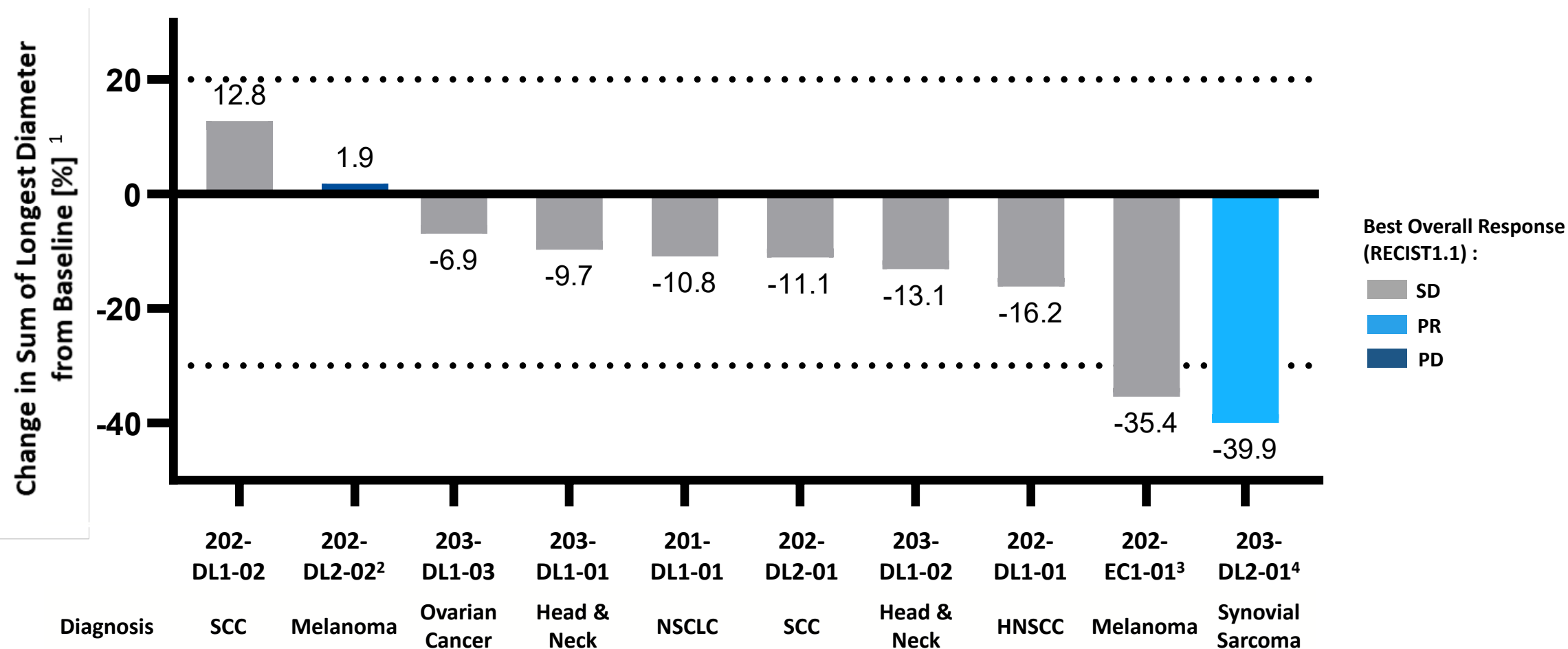
Data cut-off – February 16, 2021

<sup>1</sup> Total infused dose of transduced viable CD8 T cells; <sup>2</sup> Immune checkpoint inhibitor; <sup>3</sup> Unconfirmed as of data cut-off;

DL: Dose level, EC1: Enrichment cohort with intermediate dose level between DL1 and DL2, SD: stable disease, PD, progressive disease, PR: partial response

# ACTengine® Clinical Programs – Change of Sum of Diameters in Target Lesions

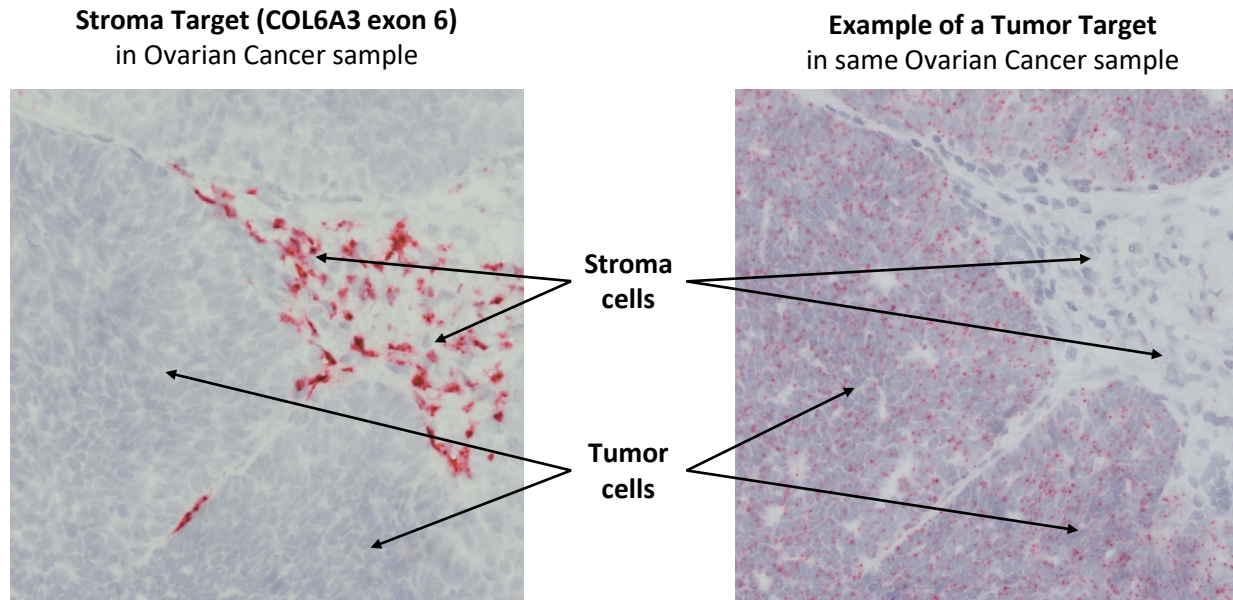
Tumor Shrinkage Observed in 8 of 10 Patients at Low Dose Levels



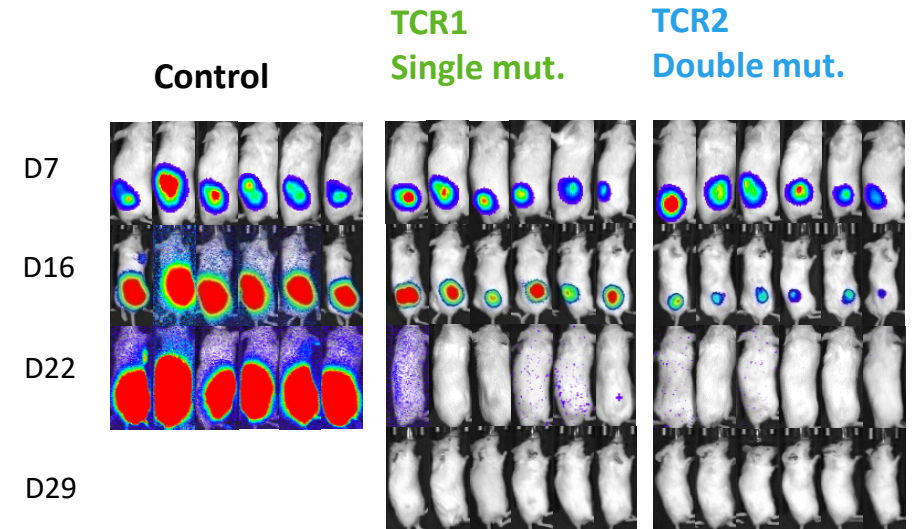
Data cut-off – February 16, 2021

# ACTengine® IMA204 – Targeting Tumor Stroma

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



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



One IMA204 TCR candidate leads to full functionality  
of both CD8 and CD4 T cells

- Final preclinical safety evaluation of two candidate TCRs ongoing
- **IMA204 clinical trial application expected 2021**

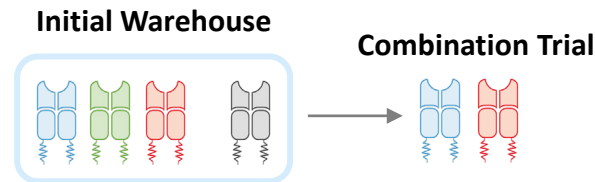
# Combating Tumor Heterogeneity & Escape through Multi-Target Approach

## A Multi-Step Approach towards Highly Personalized Multi-TCR-T Therapy

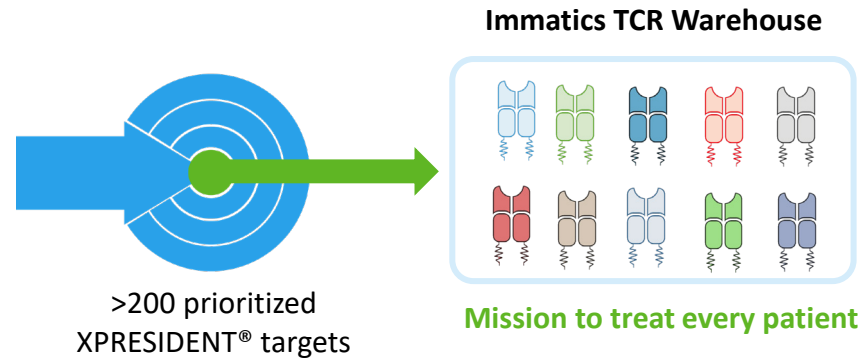


**2 ACTengine® TCR-T**

**3 ACTengine® Multi-TCR-T**



**4 Personalized Multi-TCR-T**



	HLA	Targets	T cells	Status	Objective
<b>1</b>	HLA-A2	Multiple	Endogenous	Completed	Demonstrate feasibility of multi-target concept
<b>2</b>	HLA-A2	Single	Genetically engineered	3 trials ongoing	Deliver significant clinical benefit for patients with certain tumor types
<b>3</b>	HLA-A2	Two	Genetically engineered	Mid-Term Perspective	Expand spectrum of tumor types and increase response durability
<b>4</b>	Multiple	Multiple	Genetically engineered	Long-Term Perspective	Treat every patient regardless of tumor and HLA type

# ACTengine® IMA200 Series – Summary and Future Directions

## First Anti-tumor Activity Consistent with Robust Biological Activity during Early Phases of Dose Escalation

### Key Findings

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Transient and manageable treatment-emergent adverse events as expected for cell therapies



Robust T cell engraftment and persistence post infusion and tumor infiltration in all evaluable patients



Tumor shrinkage observed in 8/10 patients including one unconfirmed partial response



IMA204: Preclinical data: In vivo tumor eradication by targeting the tumor stroma with high-affinity TCRs

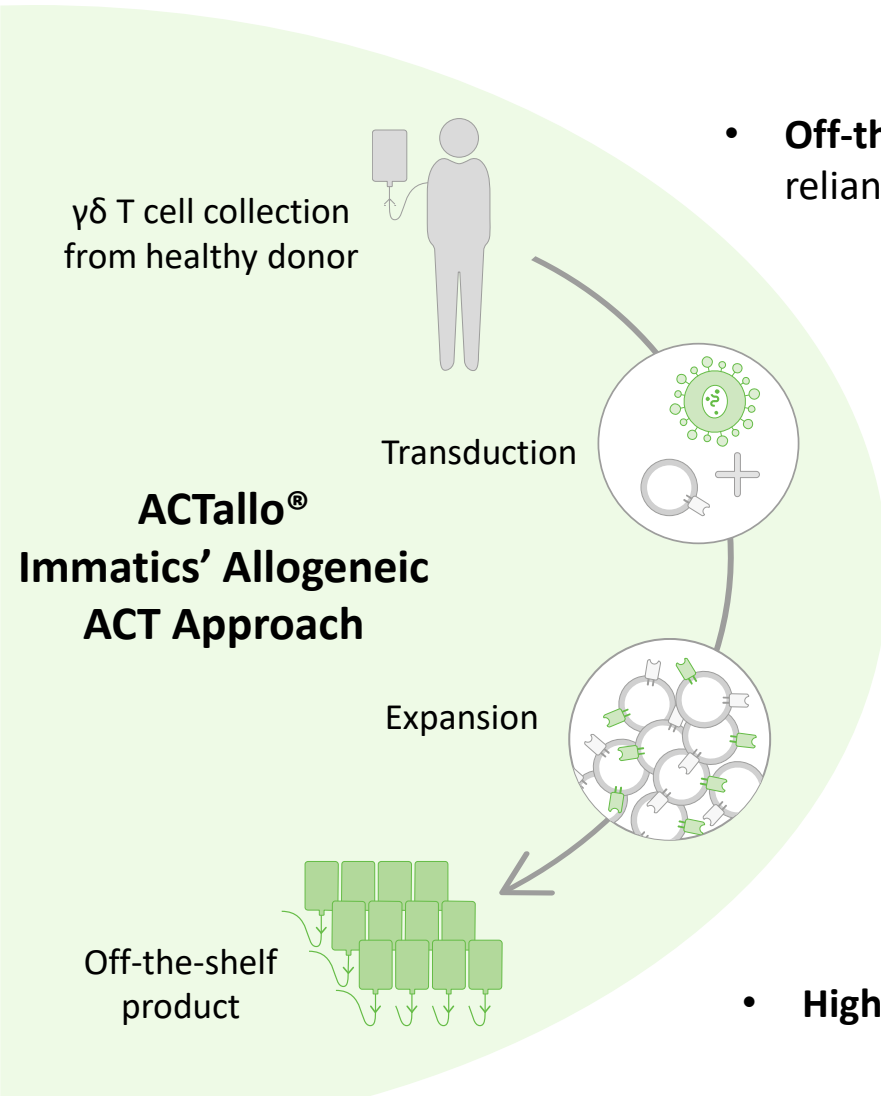
### Next Steps

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- IMA201, IMA202, IMA203 clinical trials
  - Complete Dose Escalation
  - Initiate Dose Expansion and treat patients at target dose
  - Update on patients treated at target dose expected for 2H2021
- IMA204 clinical trial application in 2H2021
- Preparation of first multi-TCR-T study

# ACTallo® IMA301 – Towards Off-the-shelf ACT

## Effective Redirection of $\gamma\delta$ T cells Using $\alpha\beta$ TCR



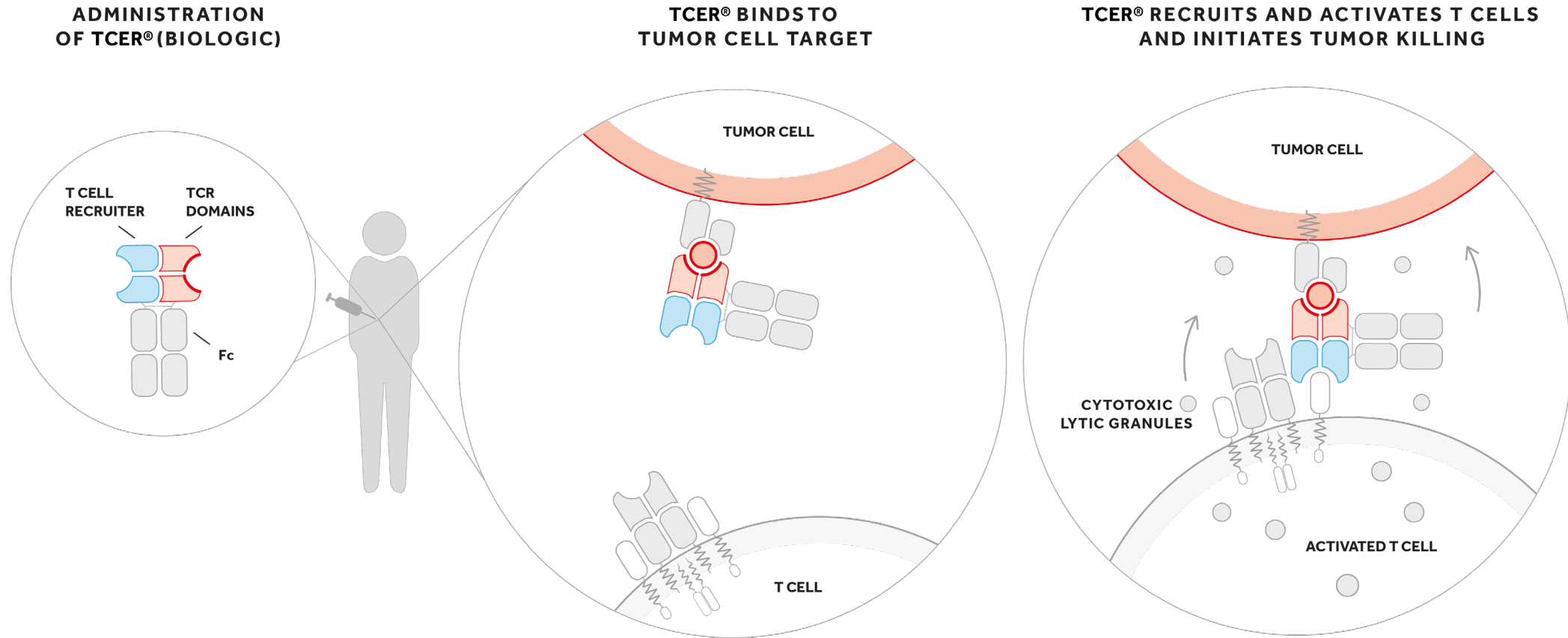
- **Off-the-shelf cell therapy**, applicable without need for personalized manufacturing and not reliant on potentially encumbered immune system of patient
- **$\gamma\delta$  T cells** are abundant, show intrinsic anti-tumor activity, naturally infiltrate solid tumors and do not cause graft-vs-host disease
- **Proprietary manufacturing protocol** delivering robust expansion of  $\gamma\delta$  T cells with the potential for hundreds of doses from one single donor leukapheresis
- **Proprietary single lentiviral vector system** (4-in-1 construct) including TCR and CD8 alpha & beta chains
- **High potency:** TCR transduced  $\gamma\delta$  T cells show similar anti-tumor activity to  $\alpha\beta$  T cells



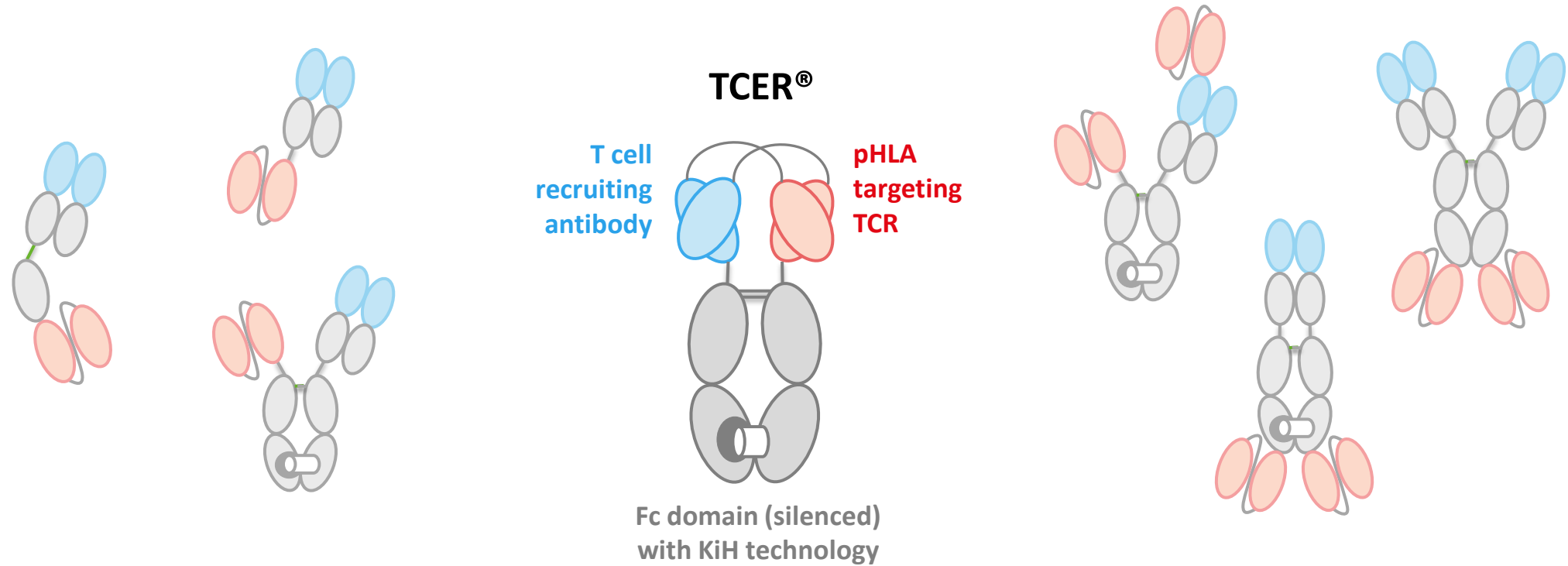
## TCR Bispecifics

# TCER® – Immatics' TCR Bispecifics

## Off-the-shelf Biologics Linking Immune Cells to Tumor Cells



# TCER® – Superior Proprietary TCR Bispecific Format



Potency and stability of proprietary TCER® format is superior to six alternative TCR Bispecific formats<sup>1</sup>

# TCER® – Preclinical POC for IMA401

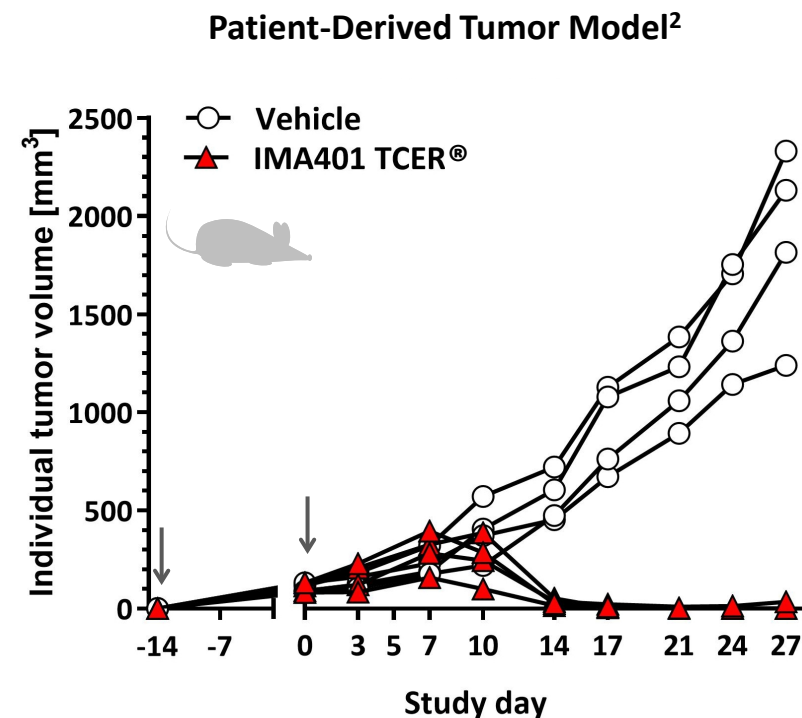
## IMA401 Targeting MAGEA4/8 Results in Tumor Eradication of Established Tumors

### Preclinical Proof-of-Concept Data:

- High **affinity** TCR (2 nM) after >10,000-fold affinity-maturation via yeast display
- High **potency** at low concentrations *in vitro* and *in vivo* in two independent xenograft tumor models (NSCLC and melanoma)<sup>1</sup>
- Distinguished **specificity & broad therapeutic window** ( $\geq 1,000$ -fold concentration difference between tumor vs. healthy cell reactivity)
- Favorable pharmacokinetics with **10-11 days terminal half-life** in mice

### Favorable CMC Characteristics:

- Positive **purity & stability** characteristics with high **production yields** (>2 g/L)



# Advancing TCER® IMA401 Towards Clinical Development

## Recent Achievements and Intended Next Steps for IMA401

### CMC

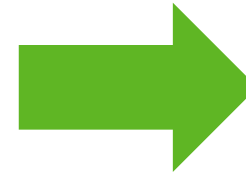
- ✓ Manufacturing process development & pilot run completed
- ✓ Formulation development completed
- Next step: GMP run scheduled for 2Q2021

### Regulatory

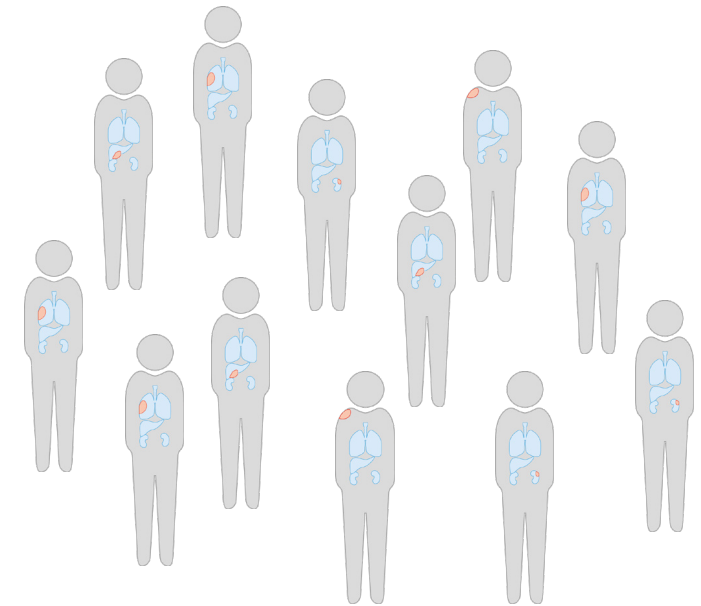
- ✓ Successful scientific advice with German regulatory authority<sup>1</sup>
- Next step: Development on track for clinical trial application **YE 2021**

### Clinical

- Basket trial with adaptive design for dose escalation & expansion cohorts
- Next step: First-in-human clinical trial in preparation



**First-in-human clinical trial  
in patients with MAGEA4/8 positive  
solid tumors**

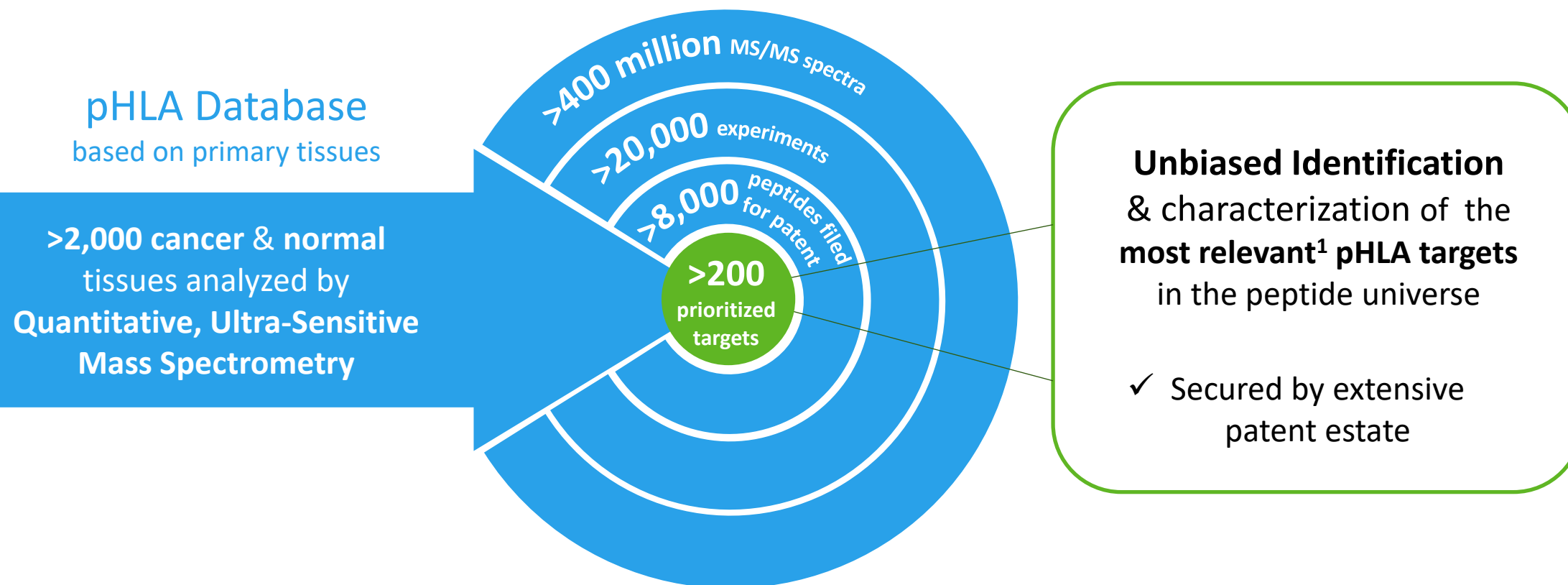




## Discovery Platforms

# XPRESIDENT® – Discovery of True Cancer Targets

Quantitative, Ultra-Sensitive Mass Spectrometry Expertise Developed over Two Decades



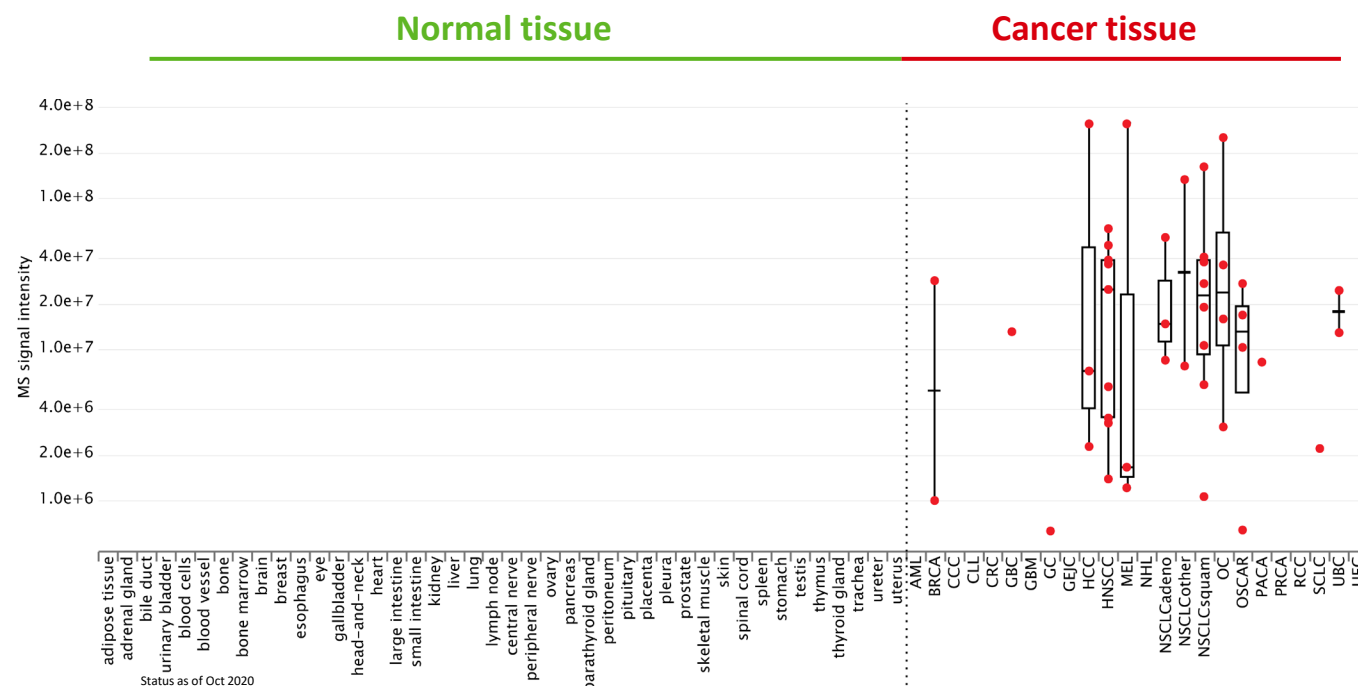
## 200 Prioritized Targets grouped in 3 Target Classes:

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

# MAGEA4/8 Target in IMA201 and IMA401 Programs

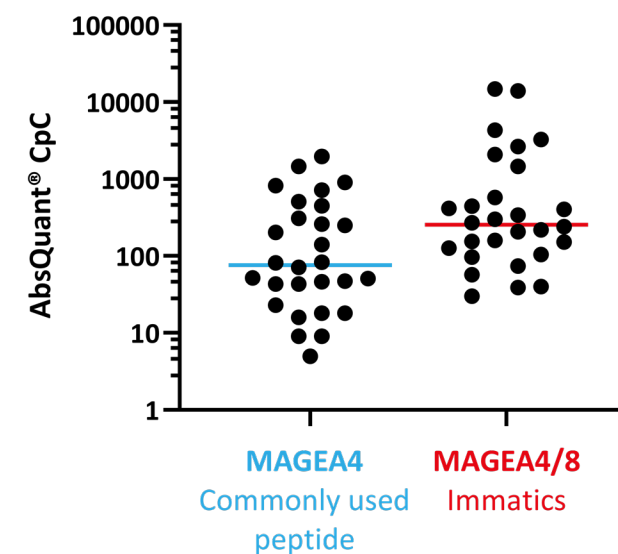
## Unique Target Discovery and Characterization Capabilities

### MAGEA4/8 Peptide (quantitative mass spectrometry detection)



**MAGEA4/8 target peptide is naturally and specifically presented on native tumor tissue vs. various normal tissues**

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



**>5-fold higher target density<sup>1</sup> than a commonly used MAGEA4 target peptide**



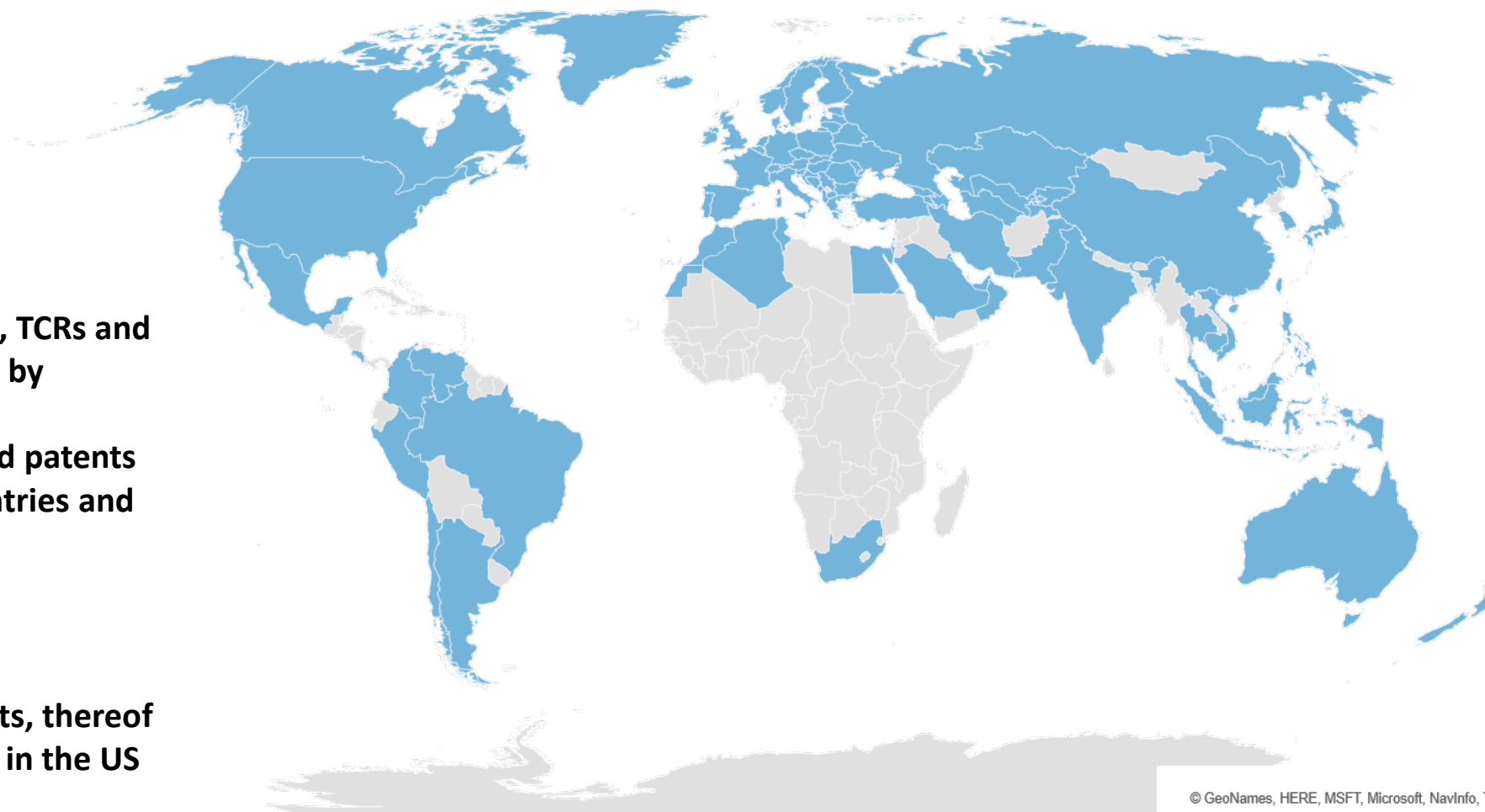


## Corporate Information & Milestones

# Robust IP Portfolio

## Immatics' Patent Estate – Territorial Coverage

- **>8,000 cancer targets, TCRs and technology protected by**
- **3,500 applications and patents filed in all major countries and regions**
- **>100 patent families**
- **>1,550 granted patents, thereof  
>400 granted patents in the US**



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# Strong, Focused and Highly Integrated Trans-Atlantic Organization

## Tübingen, Germany, ~150 FTEs



Senior Leadership, Research and Development (XPRESIDENT®, XCEPTOR®, TCER®), Translational Development, Clinical Operations, Finance, HR, IT, QM

## Houston, Texas , 80 FTEs



Senior Leadership, Research and Development (Adoptive Cell Therapy), CMC, Clinical Operations, Regulatory Affairs, QA/QC, HR, Investor Relations

## Munich, Germany, 20 FTEs



Senior Leadership, Business Development, Intellectual Property, Regulatory Affairs, Communications

# 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  
(Probiodrug, NovImmune, Medigene, InflaRx)



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



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



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



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



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

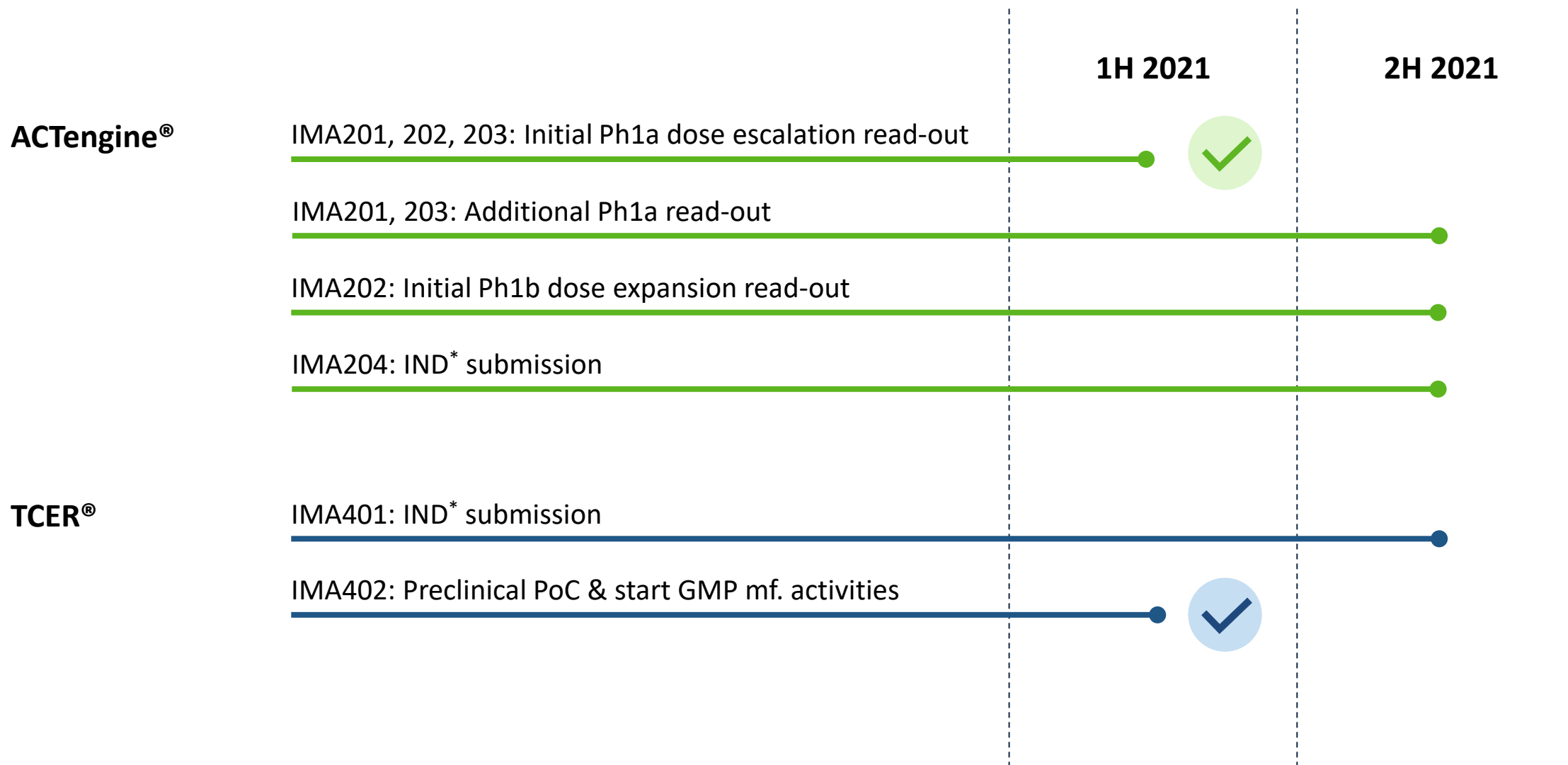


**Edward Sturchio**  
General Counsel  
>15 yrs pharma & biotech experience  
(Schering, Merck, Novartis, Advanced Accelerator Applications, Abeona Therapeutics)




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

# Upcoming R&D Milestones in 2021



# Immatics Key Take-Aways

- Broadly positioned in TCR therapeutics space with two distinct treatment modalities: ACT & TCR Bispecifics
- ACTengine® (TCR-T) IMA200 Clinical Series
  - Proprietary cell manufacturing resulting in younger T cells for better engraftment & persistence
  - First anti-tumor activity observed in three TCR-T trials at early phases of dose escalation – next readout in 2H21
- TCER® - Leading TCR Bispecifics platform with antibody-like stability and half-life
  - Clinical trial application on track in 4Q21 for IMA401 program against high density target
- Differentiated target and TCR discovery platforms secured by a broad patent estate including >200 prioritized targets
- Multiple strategic collaborations with world-leading industry players incl. Amgen, Genmab, BMS and GSK
- Strong cash position of approx. US\$ 285m (as of December 31, 2020) with cash reach into 2023



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

Thank you

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