

# IMMUNOCORE

## Corporate Presentation

40<sup>th</sup> Annual J.P. Morgan Healthcare Conference  
January 2022

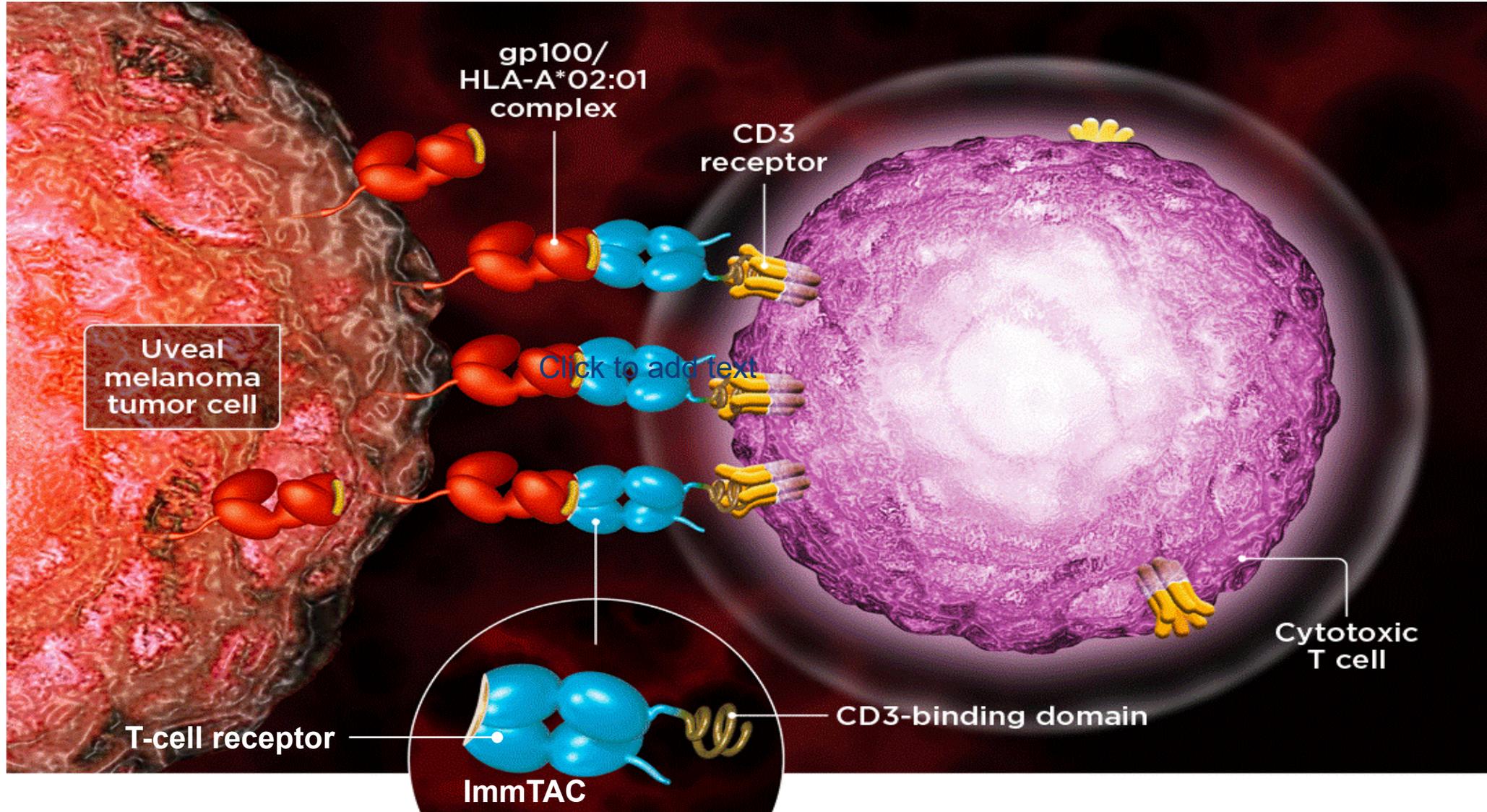


# Forward-Looking Statements

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# Harnessing the immune system to fight disease with targeted, off-the-shelf, bispecific, soluble T cell receptors (TCRs)



## Leader in off-the-shelf bispecific T-cell engagers

First TCR to demonstrate monotherapy overall survival (OS) benefit in solid tumor

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## Clinically-validated platform moving to commercialization in mUM<sup>1</sup>

Potential first FDA approval for a TCR therapeutic

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## Pipeline with potential in multiple indications / therapeutic areas

Oncology (tebentafusp, PRAME, MAGE-A4), infectious and autoimmune diseases

5 clinical stage programs

1. Metastatic uveal melanoma

# Our team

Proven track record with over 25 new medicines for patients



**Bahija Jallal**  
CEO



IMFINZI, FASENRA, LUMOXITI, SELIQ,  
QAIV, SAPHNELO



**Brian Di Donato**  
CFO & Head of Strategy



YERVOY, EMLICITI, LUMOXITI, IMFINZI



**David Berman**  
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YERVOY, EMLICITI, LUMOXITI, IMFINZI



**Mohammed Dar**  
CMO



VOTRIENT, IMFINZI, LUMOXITI



**Andy Hooker**  
VP, CMC & Supply Chain



CIMZIA



**JoAnn Suzich**  
Head of Research



SYNAGIS, FLUMIST, VLP  
technology for HPV vaccines



**Mark Moyer**  
Head of Regulatory



YERVOY, OPDIVO, TAXOTERE, ZOLADEX,  
PLAVIX, JEVTANA, ELOXATIN



**Ralph Torbay**  
Head of Commercial



IMFINZI, TAGRISSO, CALQUENCE, GLEEVEC,  
TASIGNA, ARZERRA, FARYDAK

# Our pipeline

Leading bispecific TCR pipeline with tebentafusp BLA & MAA submissions accepted

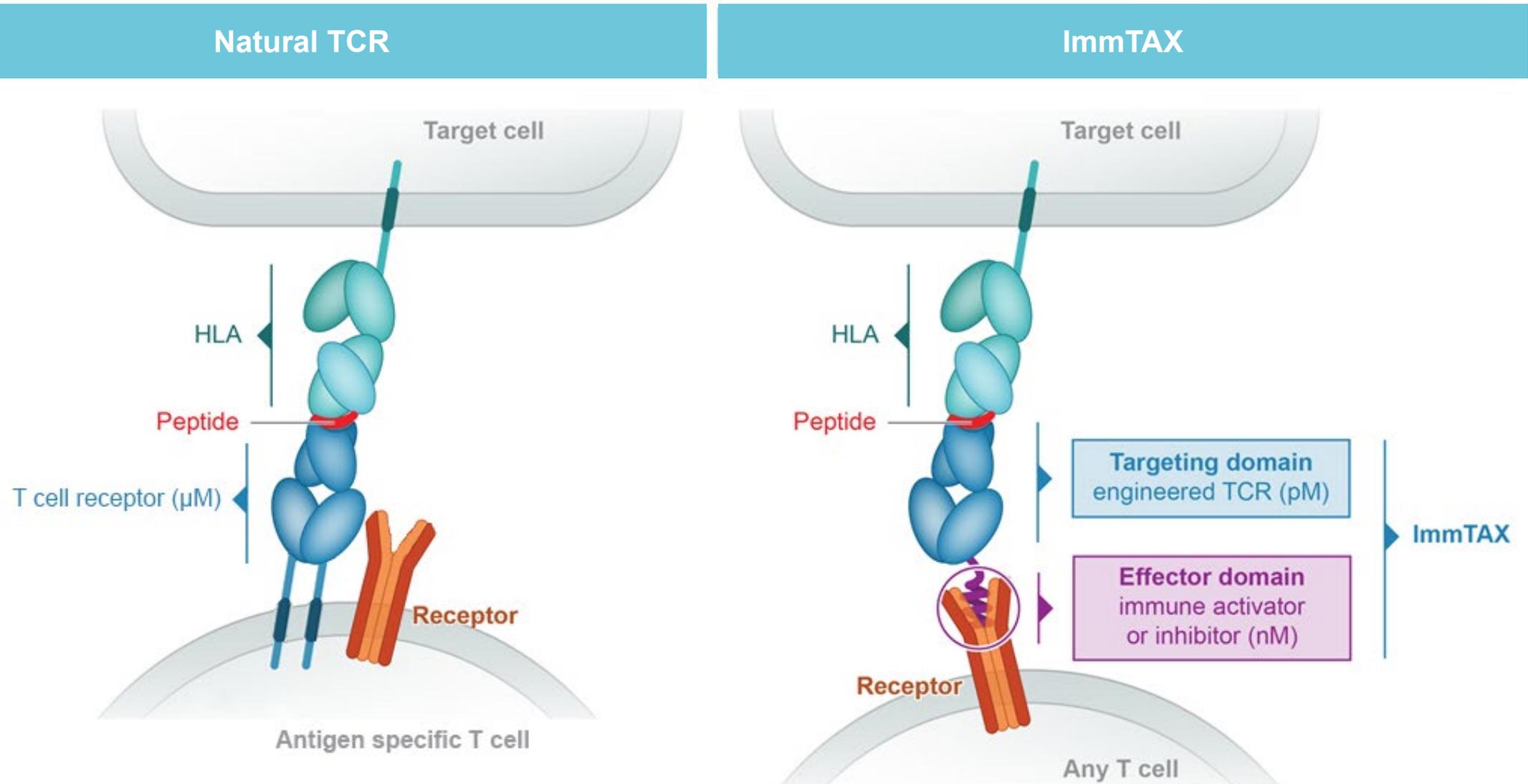
Candidate	Target	Indication	Pre-clinical	Phase 1 / 2	Phase 3	Approved	Anticipated Milestones
<b>Oncology</b>							
Tebentafusp	gp100	Uveal melanoma					<ul style="list-style-type: none"> <li>❖ PDUFA Feb. 2022</li> <li>❖ Commercial launch 1H 2022</li> </ul>
		Cutaneous melanoma					<ul style="list-style-type: none"> <li>❖ Randomized study 4Q 2022</li> </ul>
IMC-F106C	PRAME	NSCLC, breast, endometrial, ovarian, SCLC, melanoma					<ul style="list-style-type: none"> <li>❖ Ph. 1 initial data mid 2022</li> </ul>
IMC-C103C <sup>1</sup>	MAGE-A4	NSCLC, gastric, head & neck, ovarian, synovial sarcoma					<ul style="list-style-type: none"> <li>✓ Initiated ovarian expansion</li> <li>❖ Ph. 1 update 2H 2022</li> </ul>
Candidate #4	Undisclosed	Multiple solid tumors					
Candidate #5	Undisclosed	Colorectal, gastric, pancreatic					
<b>Infectious Diseases</b>							
IMC-I109V	Envelope	Hepatitis B Virus (HBV)					<ul style="list-style-type: none"> <li>❖ Enrolling Ph. 1</li> </ul>
IMC-M113V <sup>2</sup>	Gag	Human Immunodeficiency Virus (HIV)					<ul style="list-style-type: none"> <li>❖ First patient dosing 2Q 2022</li> </ul>

<sup>1</sup> Developed under a co-development/co-promotion collaboration with Genentech. <sup>2</sup> Program is wholly owned, development costs being provided by the Bill & Melinda Gates Foundation (BMGF), Immunocore retain all development and commercialization rights in the developed world.

# Technology Platform

# We pioneered converting membrane-bound T cell receptors

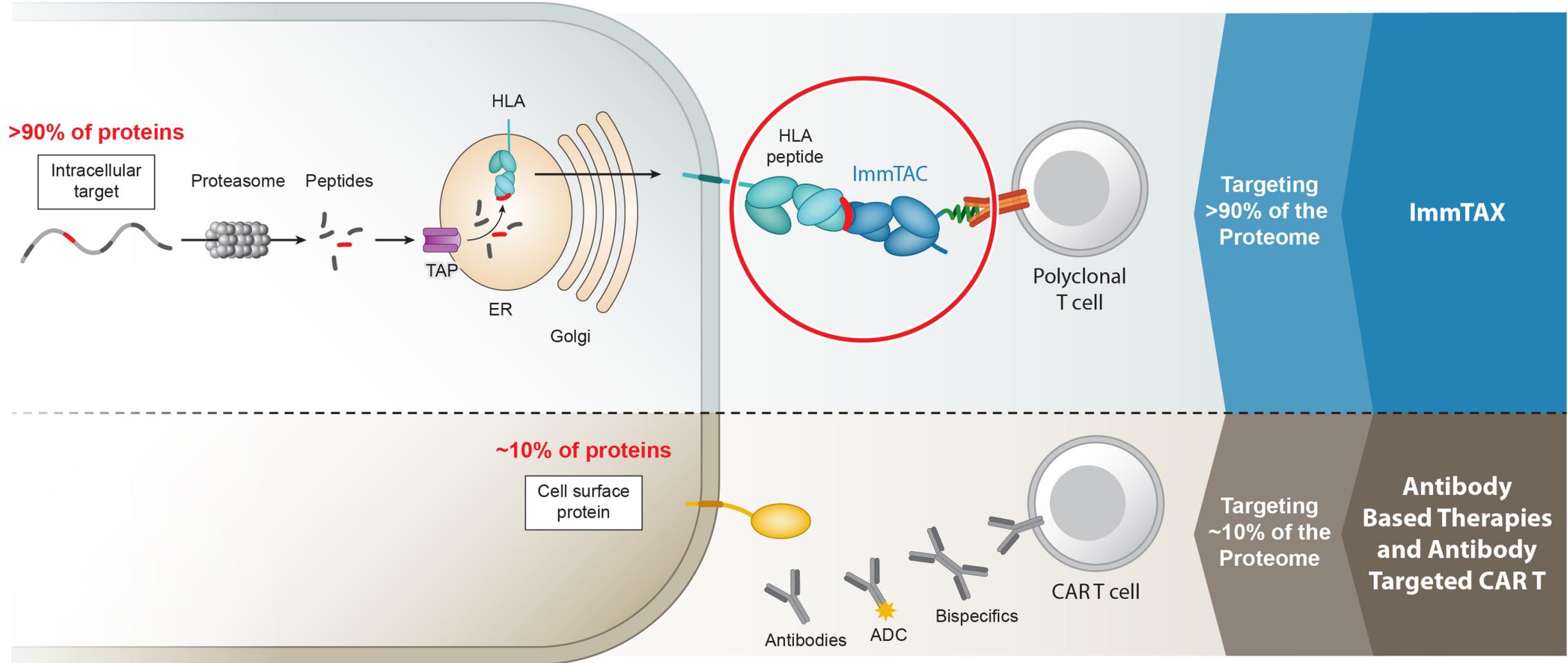
*Into soluble, off-the-shelf, bispecific therapeutics (ImmTAX)*



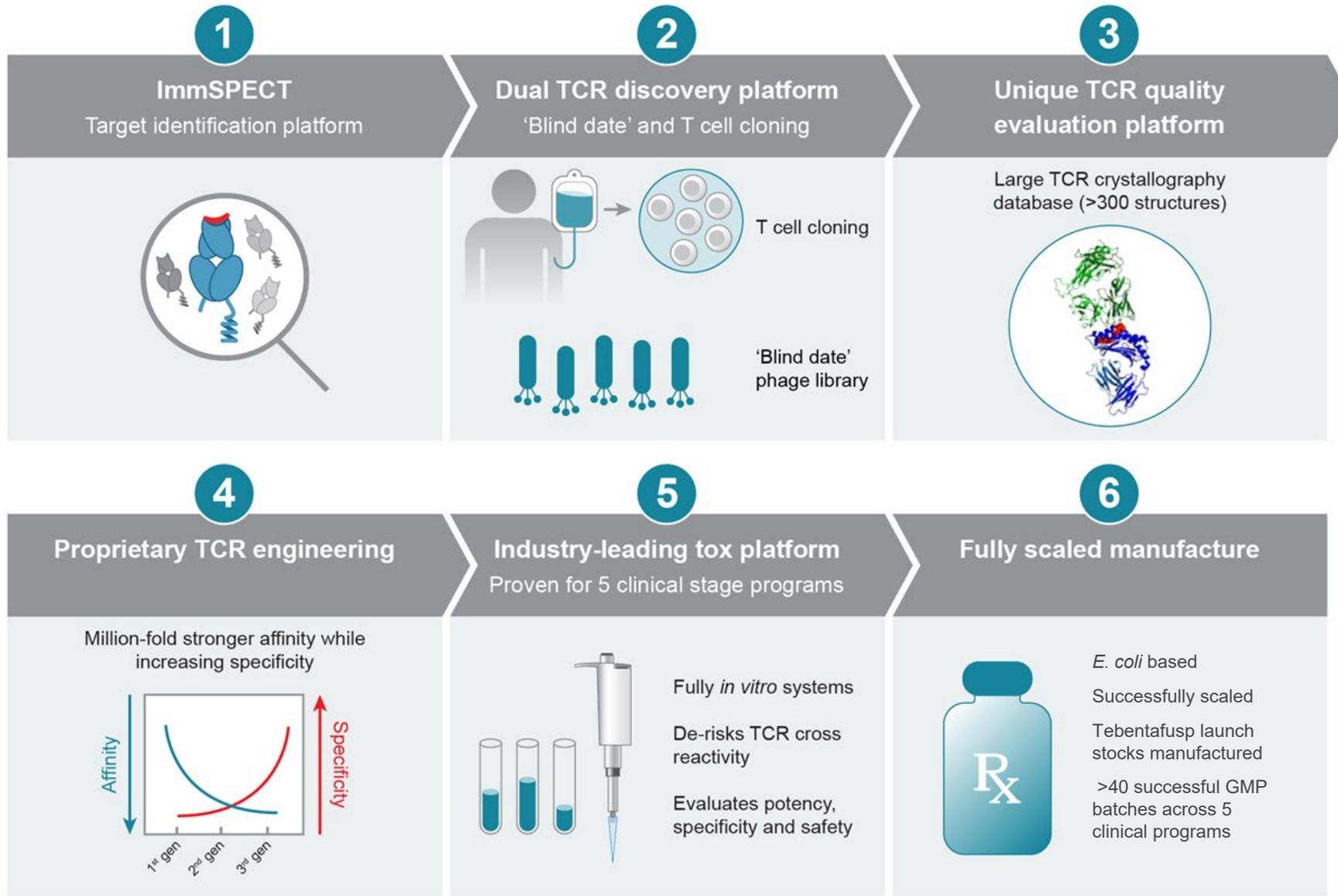
# TCR therapeutics can target nearly the entire human proteome

*Application to oncology, infectious disease and autoimmune*

## Target Cell



# Seamless suite of proprietary technologies spanning target discovery to clinic

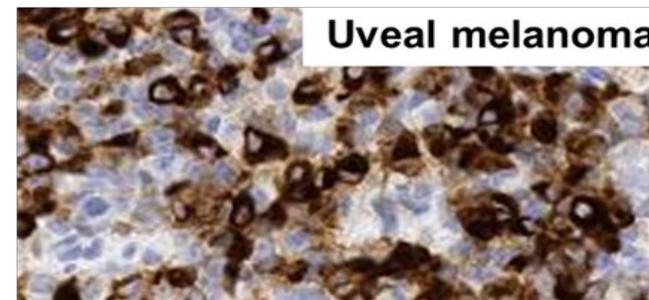
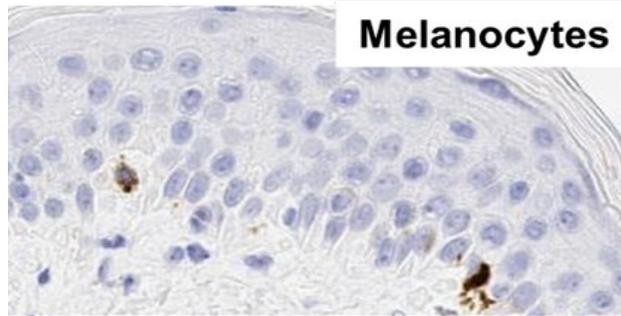


# Tebentafusp in Metastatic Melanoma

# Tebentafusp (Tebe): First-in-class, off-the-shelf, bispecific TCR

Targeting gp100 protein in melanoma

## gp100 protein



## Three melanoma clinical trials



### IMCgp100-01:

Ph 1 in uveal & cutaneous melanoma<sup>1</sup>

**Endpoints: safety and activity**



### IMCgp100-102:

Ph 2 in uveal melanoma<sup>2</sup>

Second or third line in metastatic disease

**Primary endpoint: RECIST ORR**



### IMCgp100-202:

Ph 3 pivotal in uveal melanoma<sup>3</sup>

First line metastatic

**Primary endpoint: Overall Survival**



The NEW ENGLAND  
JOURNAL of MEDICINE

## Metastatic UM:

- HLA-A\*0201+
- No prior systemic therapy in the advanced setting
- No prior liver-directed therapy, except surgery
- Any LDH

378 patients  
Randomized  
2:1

Tebentafusp

## Investigator's Choice (IC):

- Dacarbazine 6%
- Ipilimumab 12%
- Pembrolizumab 82%

Primary endpoint  
Overall Survival (OS)

Data cut-off date: October 13, 2020; data snapshot date: January 22, 2021.

Tebentafusp (n=245)		
Adverse Event (AE), related*	Any grade n (%)	Grade 3/4 n (%) <sup>†</sup>
<b>Any</b>	244 (99.6) <sup>¶</sup>	110 (45) <sup>**</sup>
<b>Cytokine-mediated</b>		
<b>Cytokine release syndrome<sup>‡</sup></b>	217 (89)	2 (1)
<b>Pyrexia</b>	185 (76)	9 (4)
<b>Chills</b>	114 (47)	1 (0.4)
<b>Nausea</b>	105 (43)	2 (1)
<b>Fatigue</b>	101 (41)	7 (3)
<b>Hypotension</b>	93 (38)	8 (3)
<b>Vomiting</b>	64 (26)	1 (0.4)
<b>Headache</b>	53 (22)	1 (0.4)
<b>Skin-related</b>		
<b>Rash<sup>§</sup></b>	203 (83)	45 (18)
<b>Pruritus</b>	169 (69)	11 (5)
<b>Dry skin</b>	72 (29)	0
<b>Erythema</b>	56 (23)	0

IC (n=111)		
AE, related	Any grade n (%)	Grade 3/4 n (%)
<b>Any</b>	91 (82)	19 (17)
<b>Fatigue</b>	29 (26)	1 (1)
<b>Rash</b>	27 (24)	0
<b>Pruritus</b>	23 (21)	0

- Majority AEs in first few weeks
- AEs generally manageable; low related discontinuation rate for tebentafusp (2%) vs. IC (4.5%)
- No tebentafusp-related deaths as assessed by the investigators

\*Table summarizes treatment related AEs that are present at least 20% any grade; <sup>†</sup>Other (2-4%) severe AEs in tebentafusp arm include AST, ALT, lipase, lymphopenia, hyperbilirubinemia, hypophosphatemia, hypertension;

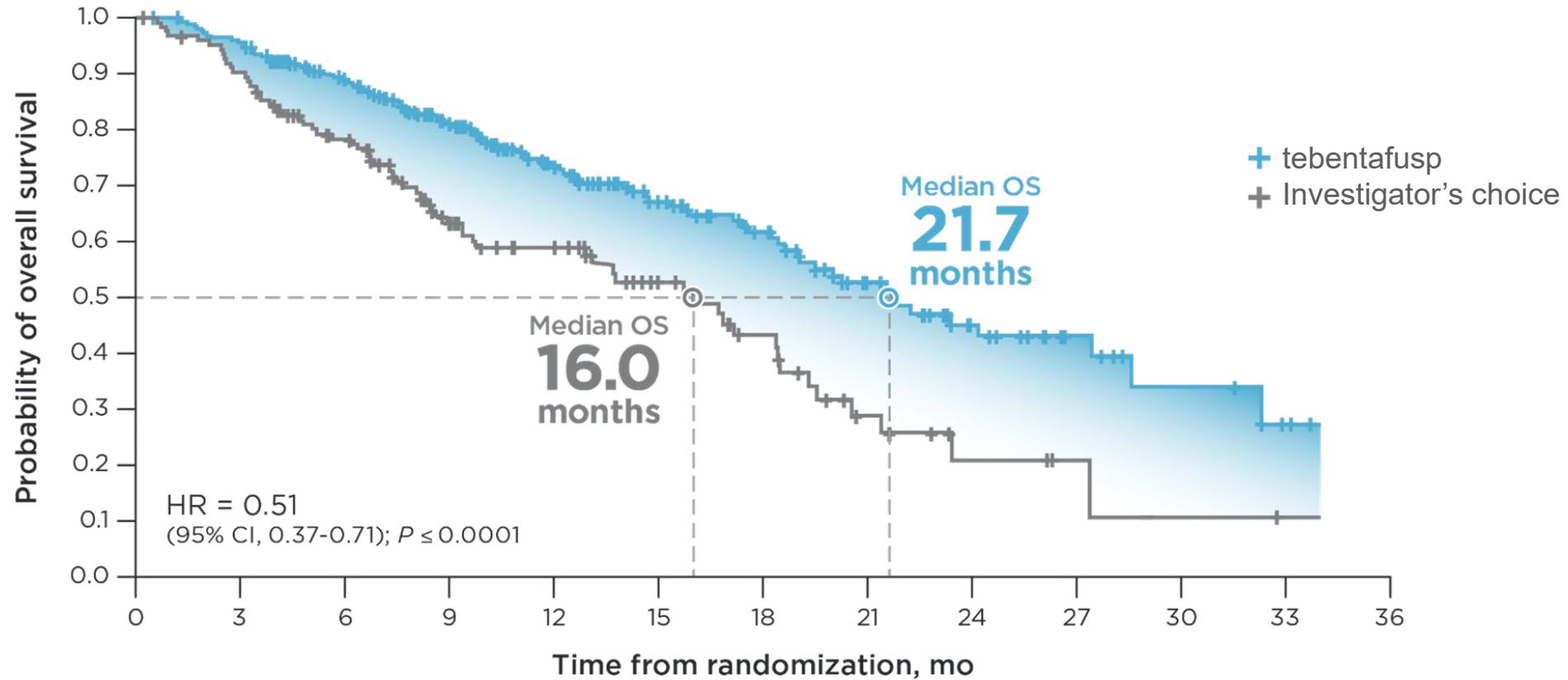
<sup>¶</sup>Includes 1 patient with no related AEs (per Investigator) but with sponsor-adjudicated CRS; <sup>\*\*</sup>Includes 1 patient with related AEs Grade <3 (per Investigator) but with sponsor-adjudicated Grade 3 CRS;

<sup>‡</sup>Cytokine release syndrome was adjudicated by sponsor according to ASTCT Consensus Grading for CRS (Lee et al. 2019); <sup>§</sup>Rash is a composite term for a list of skin toxicities of any grade. AE, adverse event

# Primary Endpoint: Overall Survival (OS) statistically significant

Tebentafusp granted Breakthrough Therapy Designation by FDA

IMCgp100-202 study



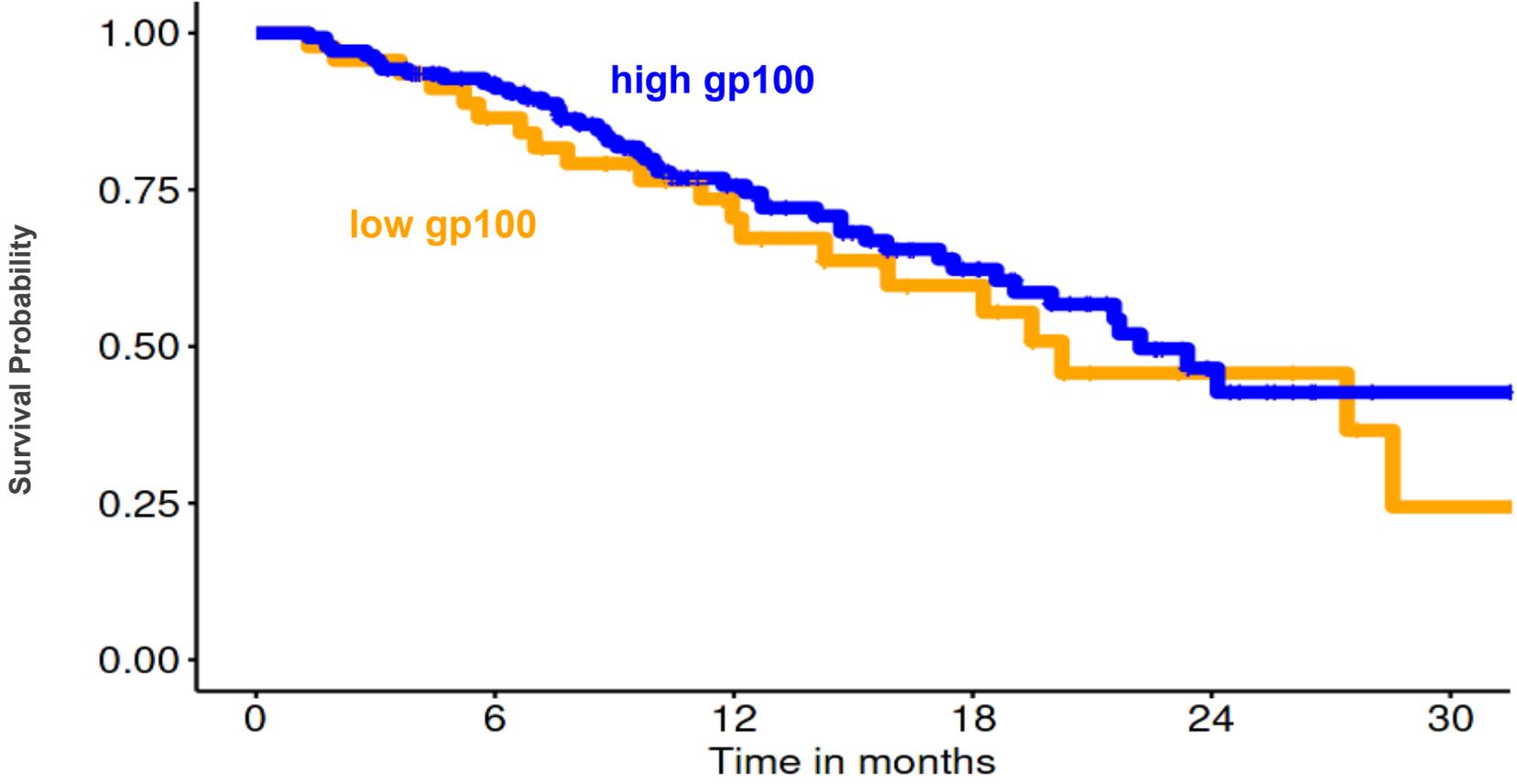
### Number of patients at risk

#### tebentafusp

252 242 221 197 167 132 109 90 71 59 44 33 22 17 9 6 5 0

#### Investigator's choice

126 116 100 86 69 48 43 34 27 20 12 7 4 4 1 1 1 0

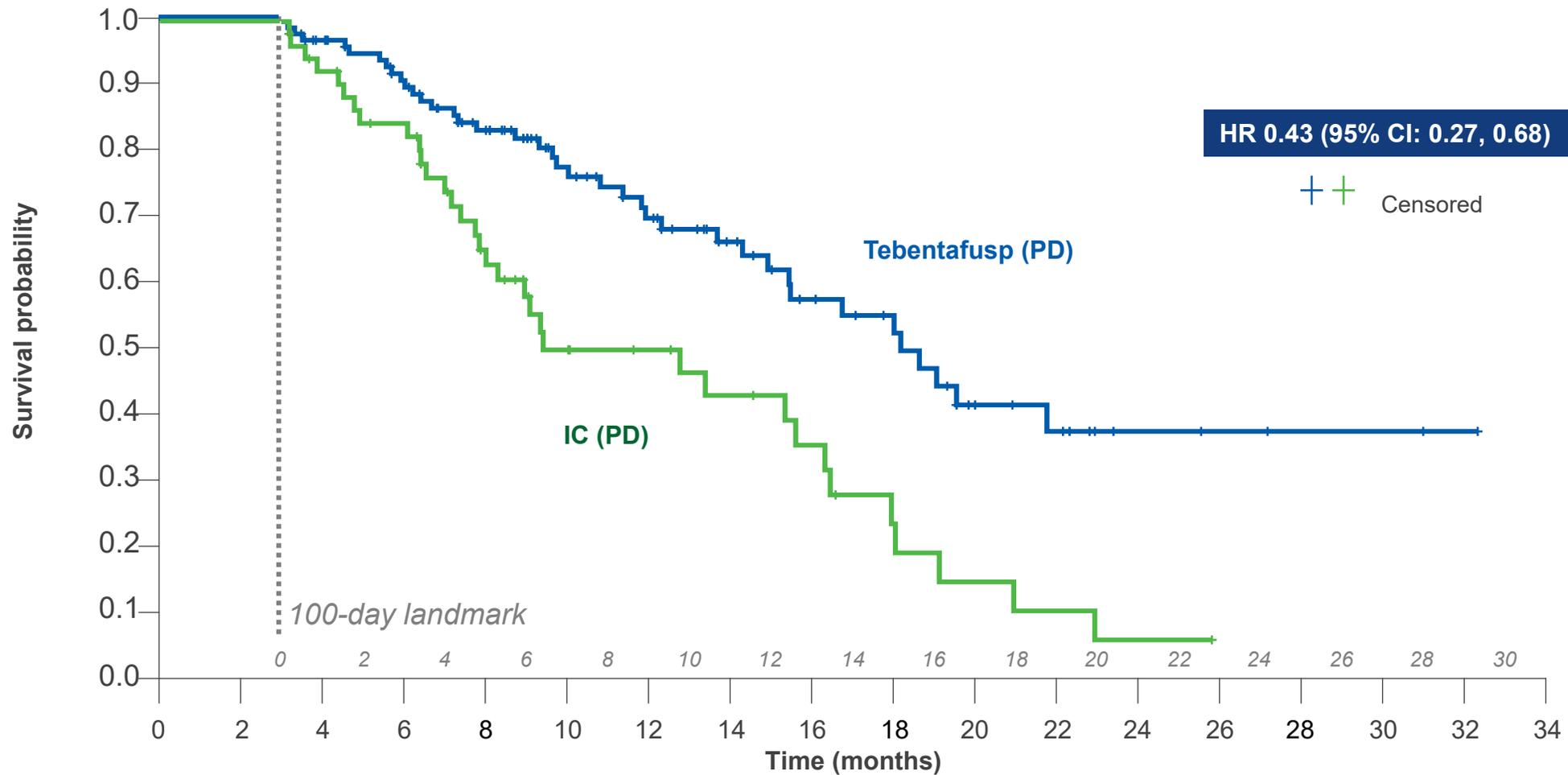


Low gp100 H score < lowest quartile  
High gp100 H score ≥ lowest quartile

# OS benefit in patients with best response of Progressive Disease

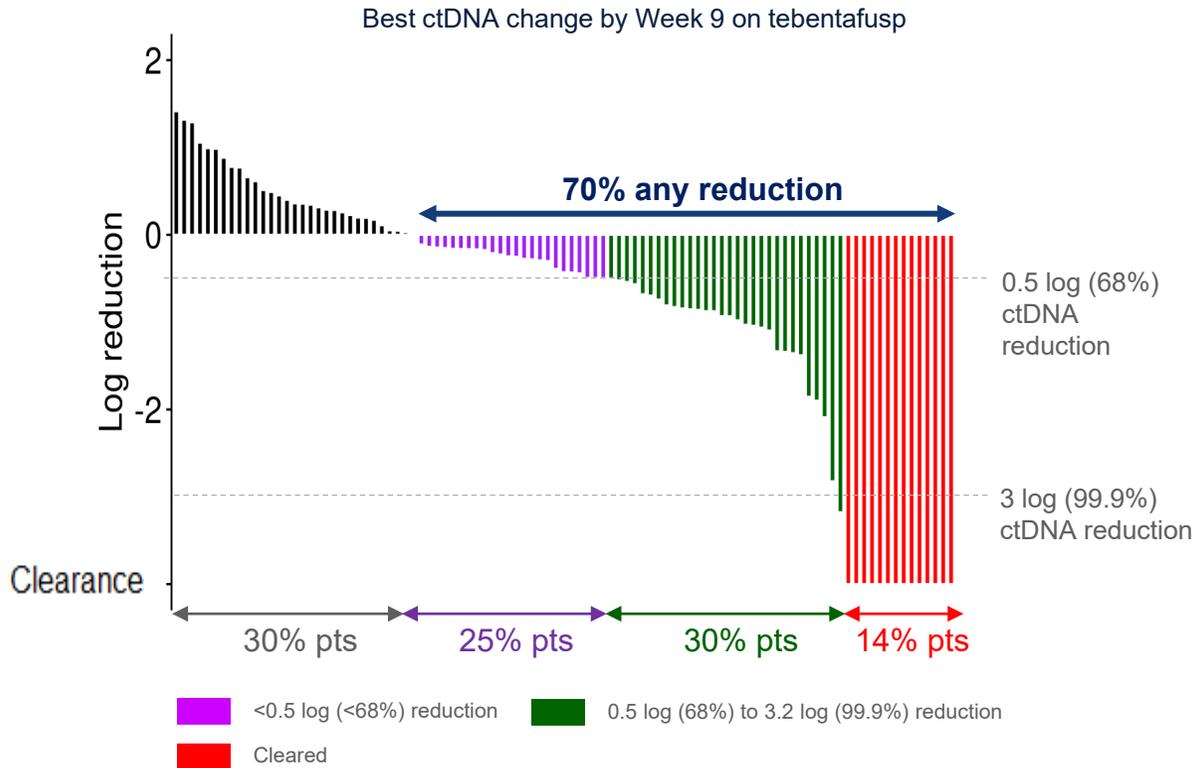
Landmark OS analysis beginning at Day 100

IMCgp100-202 study

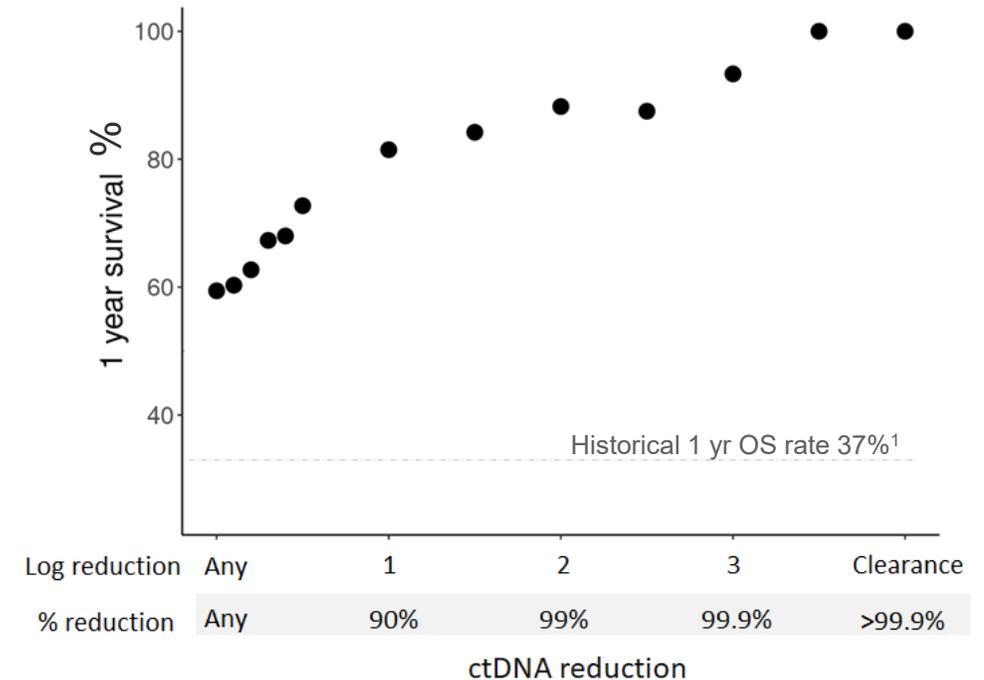


Tebentafusp (PD)	105	92	78	62	47	38	29	22	17	11	6	4	3	2	2	0
IC (PD)	53	42	35	23	16	13	11	6	4	3	2	1	0			

## 70% evaluable patients had any ctDNA reduction

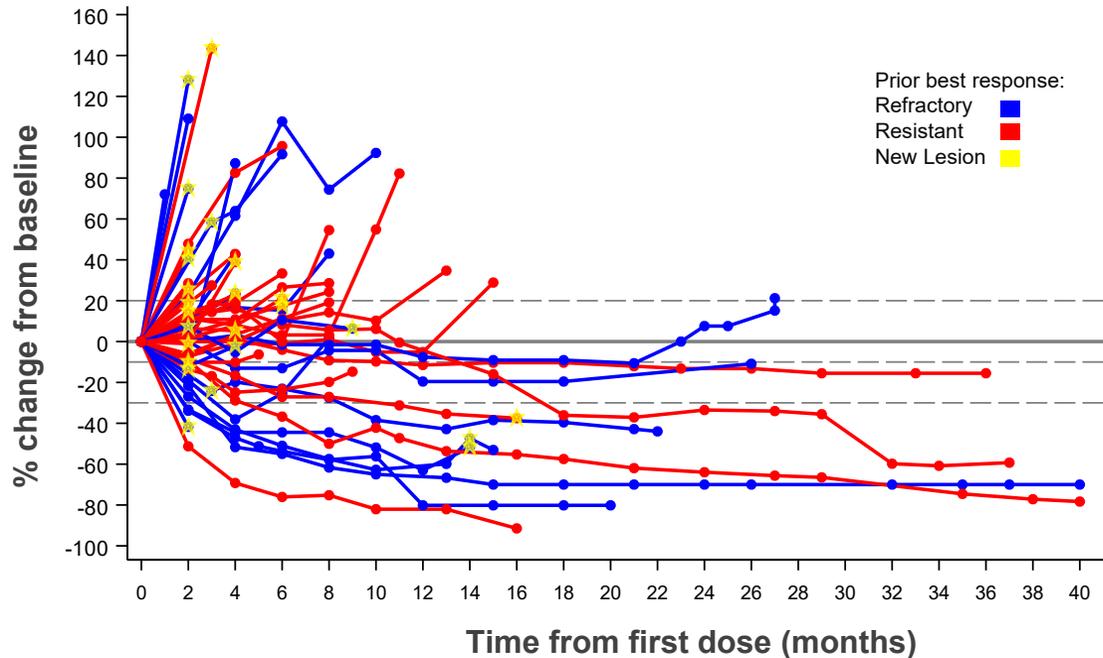


## ctDNA reduction correlates with 1 year OS



ctDNA = circulating tumor DNA

## Durable tumor shrinkage in patients who progressed on prior anti-PD(L)1 tebentafusp + durvalumab\*



\*Study IMCgp100-201: 57 patients in this study received any dose of durvalumab and had a documented best overall response to prior anti-PD(L)1 therapy. Of these 57 patients, 31 received tebentafusp + durvalumab and 26 received tebentafusp + durvalumab + tremelimumab.

Best response to prior anti-PD(L)1: Resistant = best response CR/PR/SD to prior PD(L)1; Refractory = best response of PD to prior anti-PD(L)1

## 1-yr OS

74%, anti-PD(L)1 naïve tebentafusp monotherapy<sup>^</sup>

76%, prior anti-PD(L)1 tebentafusp + durvalumab<sup>†</sup>

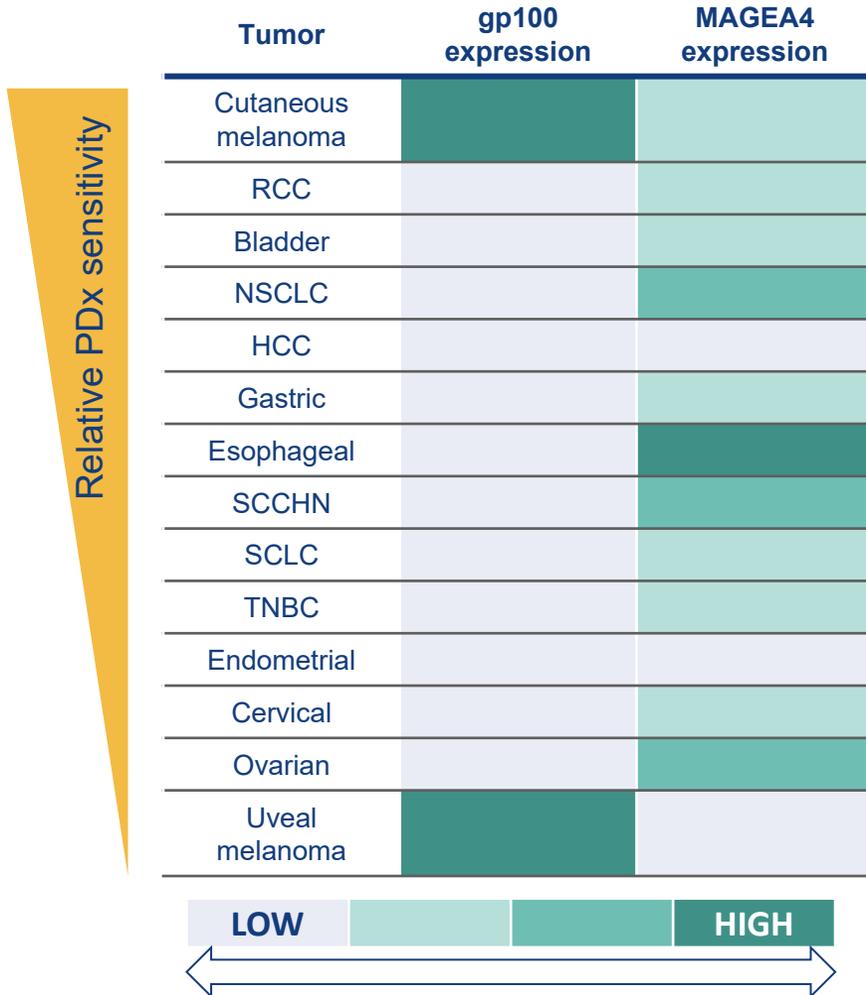
<sup>^</sup> Study IMCgp100-01, n= 49

<sup>†</sup> Study IMCgp100-201, 61 patients received prior anti-PD(L)1 and who received tebentafusp with any dose of durvalumab on this study. Of these 61, 57% patients received tebentafusp + durvalumab and 43% received tebentafusp + durvalumab + tremelimumab.

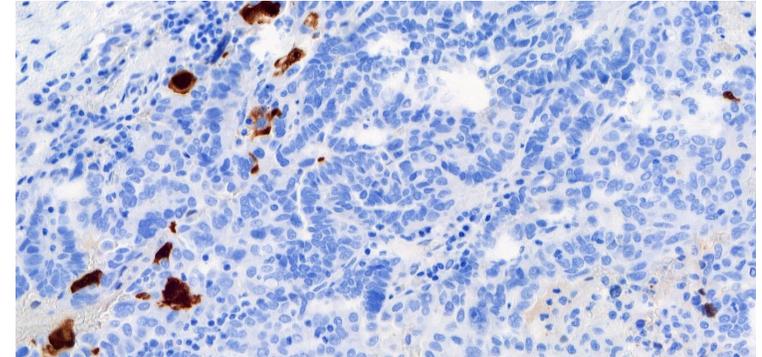
# MAGE-A4 & PRAME

# IMC-C103C targeting MAGE-A4, a cancer testis antigen expressed in multiple solid tumors

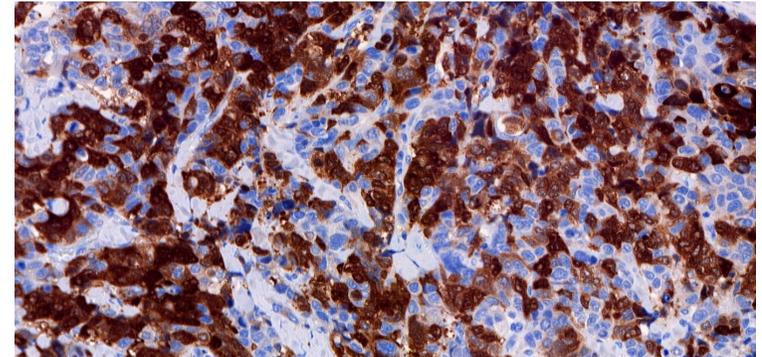
## MAGE-A4 in PDx sensitive and insensitive tumors



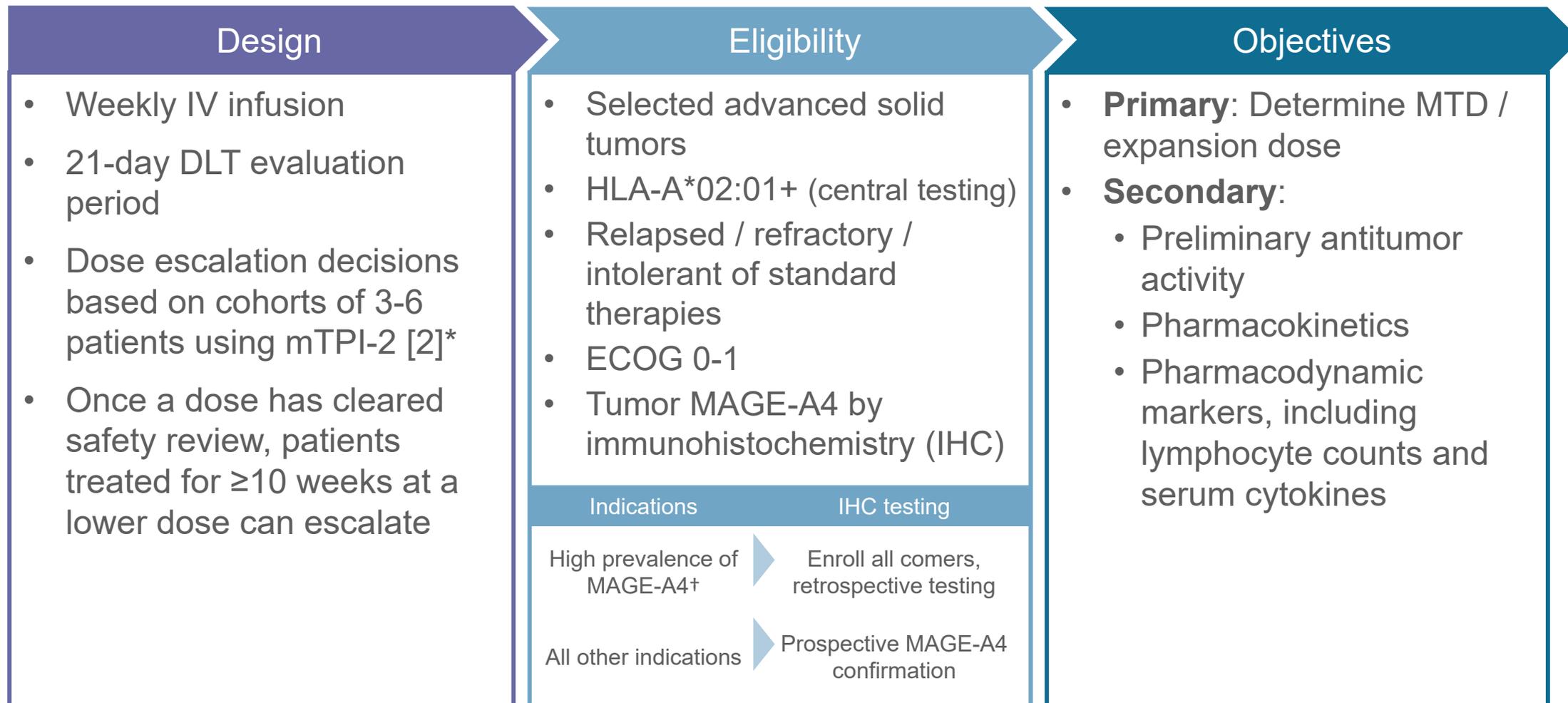
MAGE- A4 low



MAGE- A4 high



# Phase 1: First-in-Human study of IMC-C103C targeting MAGE-A4



\*mTPI-2, modified toxicity probability interval-2 method

<sup>†</sup> e.g., ovarian and synovial

# Dose escalation schema from minimum anticipated biological effect level (MABEL)

	# patients enrolled	Day 1 Dose	Day 8 Dose	Weekly Dose, Starting Day 15	
Fixed Dose	2	0.5 mcg	0.5 mcg	0.5 mcg	Initial dose defined by MABEL No to minimal pharmacodynamic activity
	2	1.5 mcg	1.5 mcg	1.5 mcg	
	3	4.5 mcg	4.5 mcg	4.5 mcg	
	3	15 mcg	15 mcg	15 mcg	Initial pharmacodynamic activity identified
9	15 mcg	45 mcg	45 mcg		
Step-Dose	4	15 mcg	45 mcg	64 mcg	Strong and consistent pharmacodynamic activity
	7*	15 mcg	45 mcg	90 mcg	
	7	15 mcg	45 mcg	140 mcg	
	2	15 mcg	45 mcg	180 mcg	
	5	15 mcg	45 mcg	240 mcg	

\*7 patients assigned to the 90 mcg cohort; however 1/7 discontinued after 15 mcg and never received 45 mcg.

Steroid premedication has been recommended at biologically active doses and, more recently, has been required when the highest dose is given for the first time;

# Safety profile manageable and consistent with mechanism of T cell activation

Preferred Term*	0.5-4.5 mcg (n=7)	15-64 mcg (n=16)	90-240 mcg <sup>§</sup> (n=21)	TOTAL (N=44 <sup>†</sup> )
<b>All Grades</b> (treatment-related events in ≥ 20% of total patients)				
Chills	-	8 (50%)	13 (62%)	21 (48%)
Pyrexia*	2 (29%)	7 (44%)	12 (57%)	21 (48%)
Cytokine release syndrome <sup>‡</sup>	1 (14%)	4 (25%)	11 (52%)	16 (36%)
Headache	1 (14%)	6 (38%)	7 (33%)	14 (32%)
Nausea	1 (14%)	6 (38%)	6 (29%)	13 (30%)
Hypotension*	-	6 (38%)	5 (24%)	11 (25%)
Fatigue	1 (14%)	4 (25%)	5 (24%)	10 (23%)
<b>Grade 3-4</b> (treatment-related events in ≥ 5% of total patients)				
Neutropenia	-	1 (6%)	7 (33%)	8 (18%)
Lymphocyte count decreased	1 (14%)	1 (6%)	2 (10%)	4 (9%)
ALT increased	-	1 (6%)	1 (5%)	2 (5%)
AST increased	-	1 (6%)	1 (5%)	2 (5%)
Headache	-	1 (6%)	1 (5%)	2 (5%)

- **No related AE led to treatment discontinuation**
- **No related AE led to death**

\*Includes events reported as a sign/symptom of CRS

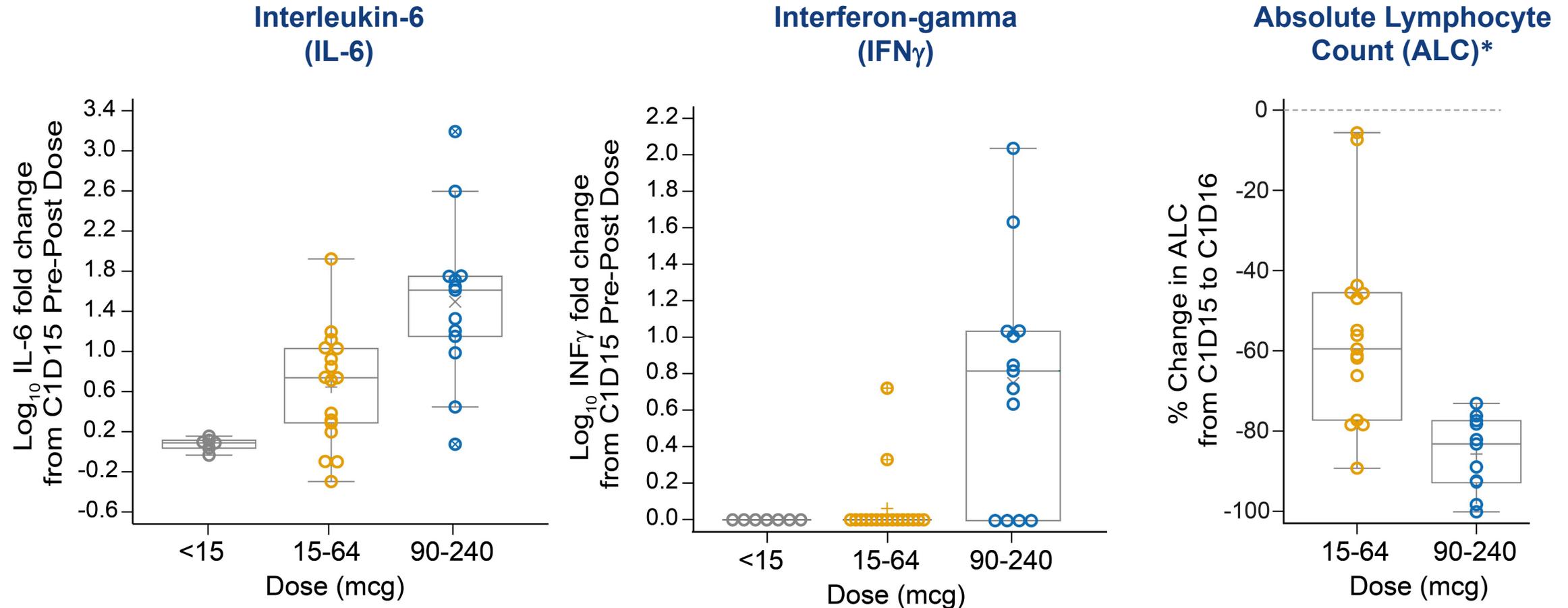
†One patient enrolled at 90 mcg and 9 months after discontinuing study treatment was re-enrolled at 180 mcg

‡Cytokine release syndrome (CRS) was graded by the Investigators using ASTCT criteria (Lee et al. 2019) [3]; all other events were graded using NCI CTCAE v5.0.

§Two DLTs at 240 mcg: Grade 3 AST increased (rapidly resolved; patient continued at 240 mcg until disease progression) and Grade 3 CRS (resolved; patient currently on 140 mcg)

# Consistent and robust evidence of T cell activity at $\geq 90$ mcg IMC-C103C

Assessment after maximal dose (Day 15)



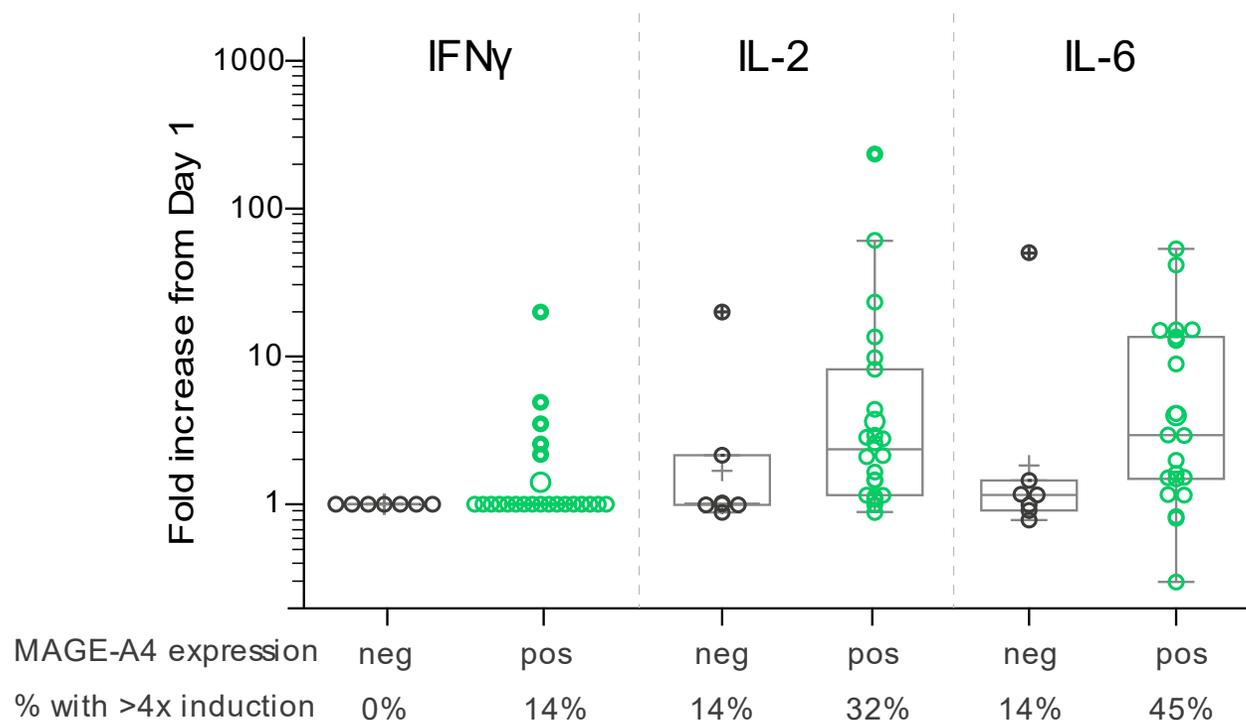
Concentrations < LLOD were set to half LLOD for purposes of deriving fold change  
Fold increase compares pre-dose to maximum post-dose (4hr, 8hr, and 24hr timepoints)  
24 patients evaluable (pre and post-dose cytokine results available for the Day 15 dose)

\* Day 16 ALC was only analyzed following introduction of intra-patient dose escalation; therefore, not collected in first cohorts.

# Cytokine induction primarily in patients with MAGE-A4 positive tumors

Assessment after initial dose, 15 mcg (Day 1)

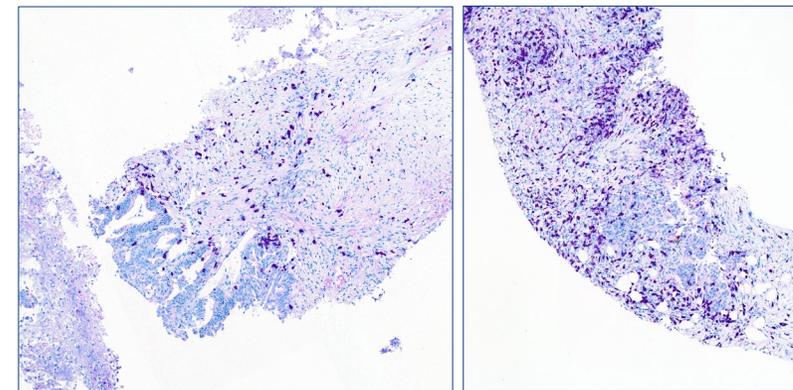
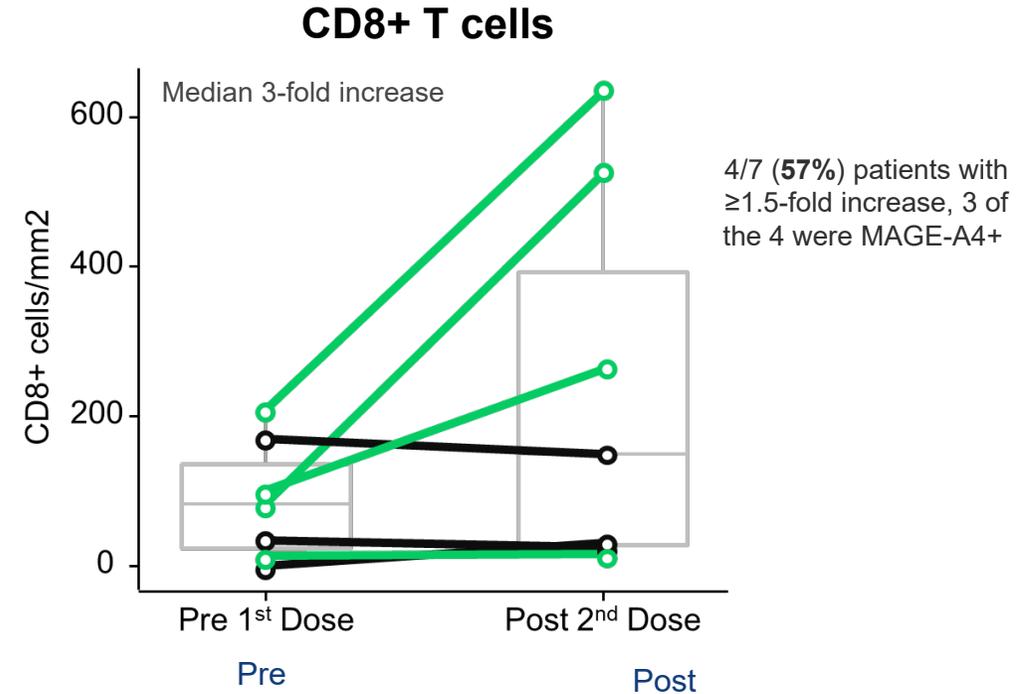
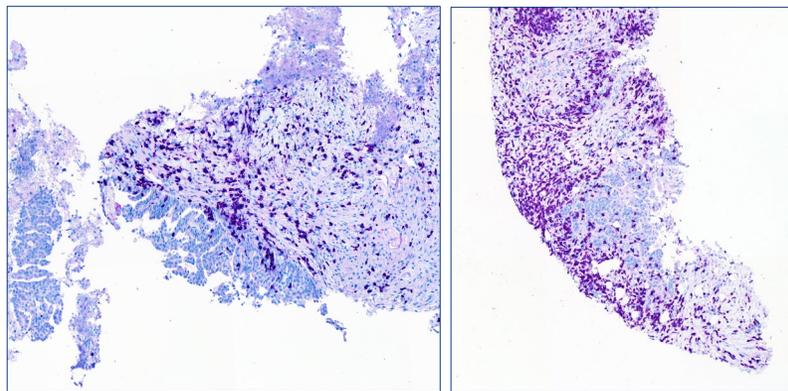
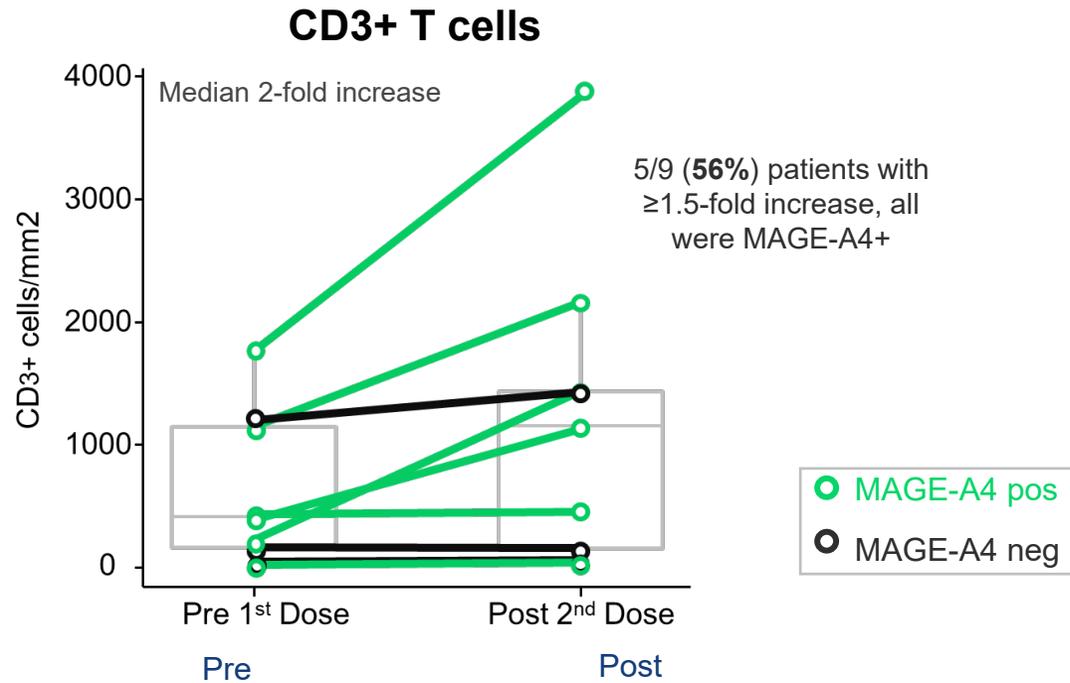
MAGE-A4 positive (H-score > 0)  
MAGE-A4 negative (H-score = 0)



- **IFN $\gamma$  induction** only observed in patients with **MAGE-A4 positive tumors**
- **Median IL-2 and IL-6 induction** higher in patients with **MAGE-A4 positive tumors**

Concentrations < LLOD were set to half LLOD for purposes of deriving fold change  
Fold increase compares pre-dose to maximum post-dose (4hr, 8hr, and 24hr timepoints)  
29 patients evaluable (15 mcg on Day 1, pre and post-dose cytokine results and MAGE-A4 results available)

# Increased T cell infiltration into MAGE-A4 positive tumors



Biopsy after two doses (15 mcg on Day 1 and 45 mcg on Day 8)

# Clinical activity in MAGE-A4 positive ovarian and HNSCC

Majority of evaluable patients had low MAGE-A4 expression

## Efficacy evaluable by MAGE-A4 IHC status all indications at 90-240 mcg

	Negative or NE MAGE-A4	Positive MAGE-A4	H-Score
HNSCC	-	1	285
Esophageal	-	1	175
Urothelial	-	1	3
Ovarian*	7	8	median 35 (range 7-128)
Total	7	11	

\* 17 ovarian patients treated at 90-240 mcg; 15/17 efficacy evaluable and 2/17 (both MAGE-A4 negative) not yet efficacy evaluable. The 15 efficacy evaluable include: MAGE-A4 negative (n=5), not evaluable (NE) by IHC (n=2), and MAGE-A4 positive (n=8)

## Clinical activity in ovarian and HNSCC

Indication	H-Score	Dose	Response	DOR
HNSCC	285	240 mcg	Confirmed PR <sup>^</sup> (ongoing)	2+ mo
Ovarian	19	140 mcg	Overall TL reduction (-44%) but new lesions	
Ovarian	7	140 mcg	Overall TL reduction (-81%) but new lesions	
Ovarian	18	90 mcg	Confirmed PR (ongoing)	4.4+ mo
Ovarian	16	15 mcg	Confirmed PR	8.3 mo

TL, target lesions

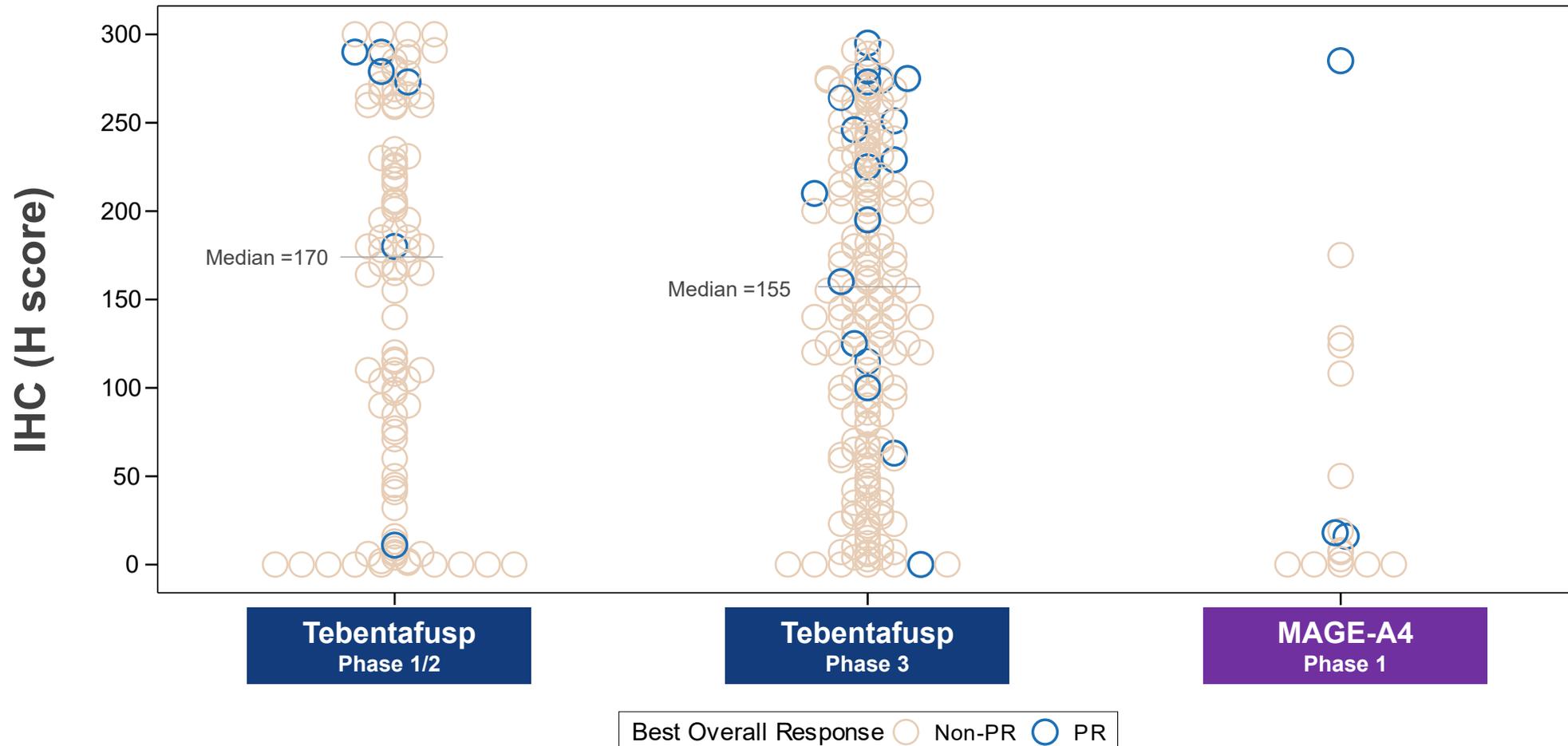
HNSCC, Head and neck squamous cell carcinoma

<sup>^</sup> confirmed after the presentation data cut-off date

- Of 11 MAGE-A4 positive (all indications), **4 had significant tumor shrinkage including 2 confirmed PR**
- Of 2 high MAGE-A4 positive, **1 confirmed PR (HNSCC)**
- Durable partial responses, **includes low and high MAGE-A4 expression**

# Enrichment of tebentafusp RECIST PRs at higher gp100 expression

In Phase 1, most MAGE-A4 patients to date have low or no MAGE-A4 expression



H score: % of tumor cells with 1+, 2+ or 3+ intensity

MAGE-A4 Phase 1 includes 16 efficacy evaluable patients who were evaluable by IHC (90-240 mcg) and single ovarian patient with PR (15 mcg)

# IMC-C103C now demonstrated safety, MoA and clinical activity

## Only clinical off-the-shelf candidate against MAGE-A4

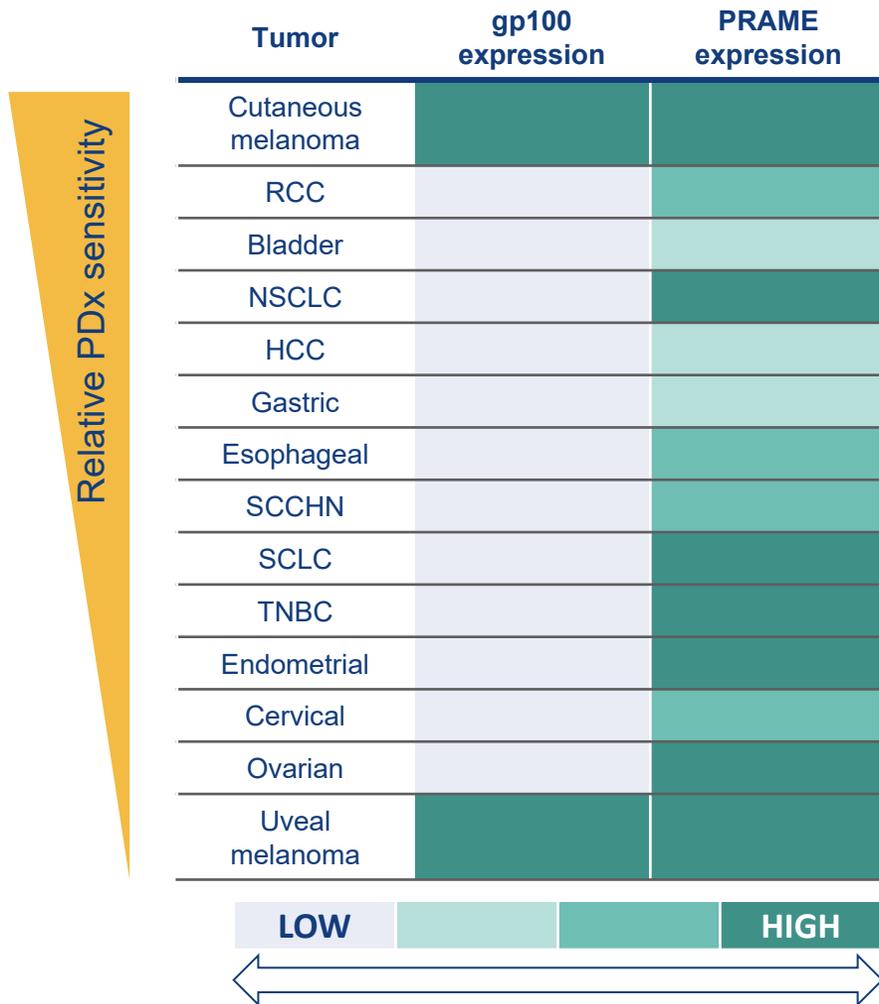
- Manageable safety profile (primarily cytokine-mediated)
- Consistent and robust biomarkers of T cell activation
- Durable PRs in ovarian carcinoma, even with low MAGE-A4 expression, and a confirmed partial response in SCHNN**
- Phase 1 study ongoing in multiple solid tumors
- Initiated first expansion arm in high grade serous ovarian at 140 micrograms**

## Potential for > 75K patients/ yr (G7)

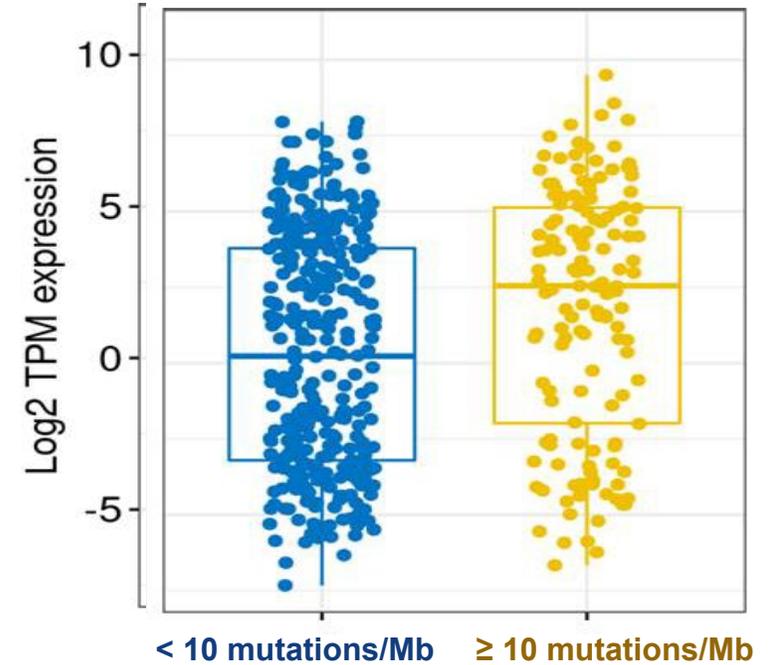
		Annual Metastatic Patients <i>MAGE-A4+ &amp; HLA-A*02:01</i>	
		US	G7
NSCLC	Squamous	8.5k	21k
	Adeno	6.5k	15k
Ovarian		3.5k	8k
SCCHN		3k	8k
Gastric + Esoph Adeno		2k	7.5k
Bladder		2k	5.5k
Esophageal Squamous		1k	5.5k
Select Others		5k	13k

# IMC-F106C targets PRAME, a negative prognostic marker in solid tumors

## Expressed in PDx sensitive and insensitive tumors



## Expressed in low and high TMB tumors (NSCLC)



TMB: tumor mutational burden

# PRAME is largest cancer-testes antigen opportunity

## Ongoing Phase 1 study

- First, and only, off-the-shelf therapeutic against PRAME intracellular protein
- 39 patients enrolled in Phase 1 dose-escalation study\*
- Biomarkers indicate having achieved biologically active doses
- **Initial Phase 1 data expected mid-2022**

## Potential for >150,000 patients/ yr (G7)

		Annual Metastatic Patients <i>PRAME+ &amp; HLA-A*02:01</i>	
		US	G7
NSCLC	Adeno	18.5k	42k
	Squamous	13.5k	32.5k
Ovarian		7.5k	17k
Small Cell Lung Cancer		7.5k	16.5k
Breast	Total	5.5k	14k
	TNBC	2.5k	5.5k
Endometrial		5.5k	11k
Cutaneous Melanoma		5k	10.5k
Select Others		10.5k	33.5k

\*As of December 16, 2021

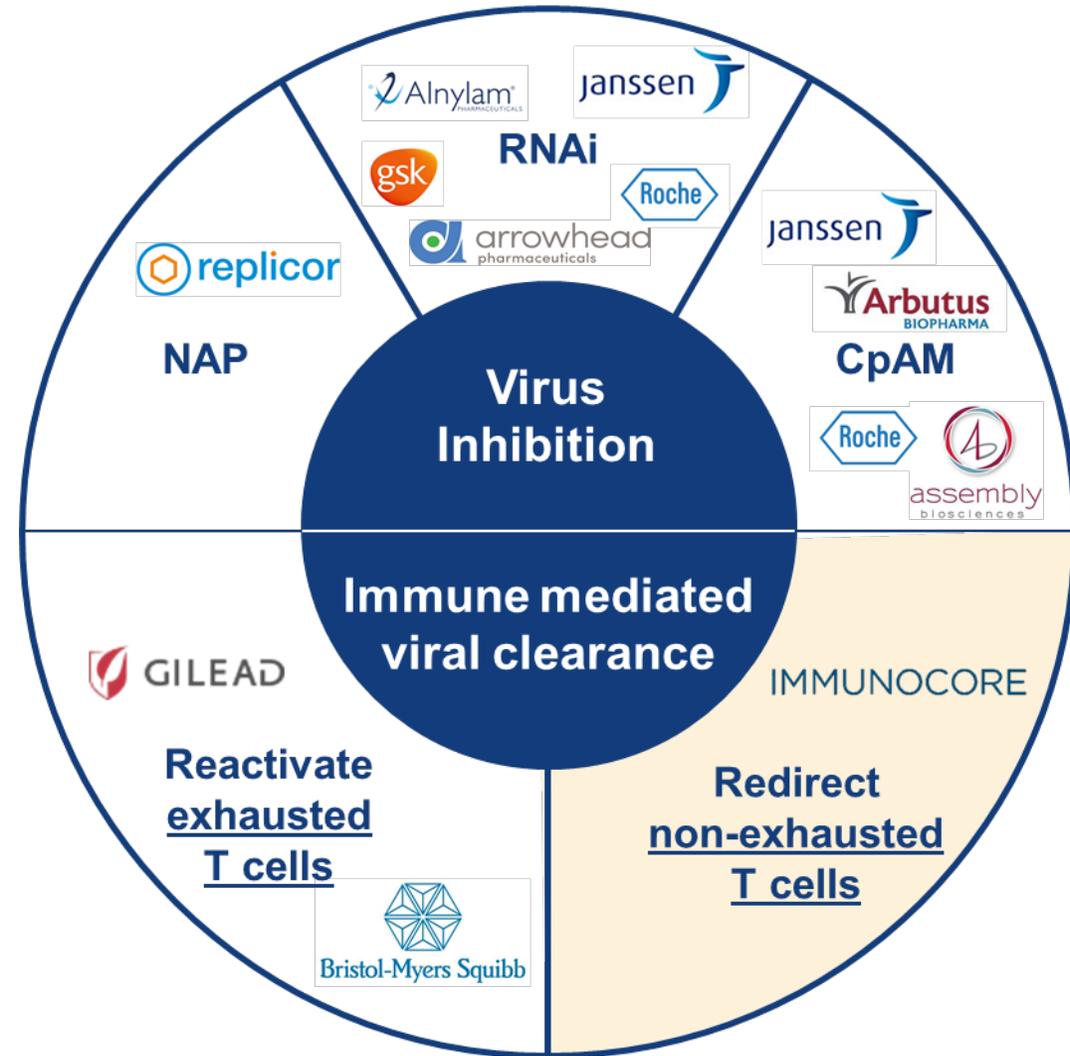
# Validation of ImmTAC platform beyond gp100

	T cell activation	Durable tumor shrinkage	Activity even in low target expression	ctDNA reduction	Overall survival benefit
<b>Tebentafusp</b> gp100	 CLINICAL CANCER RESEARCH	 ESMO IMMUNO-ONCOLOGY VIRTUAL CONGRESS	 	 	 
<b>IMC-C103C</b> MAGE-A4	 ESMO IMMUNO-ONCOLOGY <small>Onsite and Online Congress</small>	 ESMO IMMUNO-ONCOLOGY <small>Onsite and Online Congress</small>	 ESMO IMMUNO-ONCOLOGY <small>Onsite and Online Congress</small>	 <i>To be presented</i>	
<b>IMC-F106C</b> PRAME	 <i>To be presented</i>				

***Immunocore intends to present additional clinical data across all three ImmTAC programs in 2022***

# Potential for functional cure in chronic viral diseases

# Our unique approach for functional cure of chronic HBV



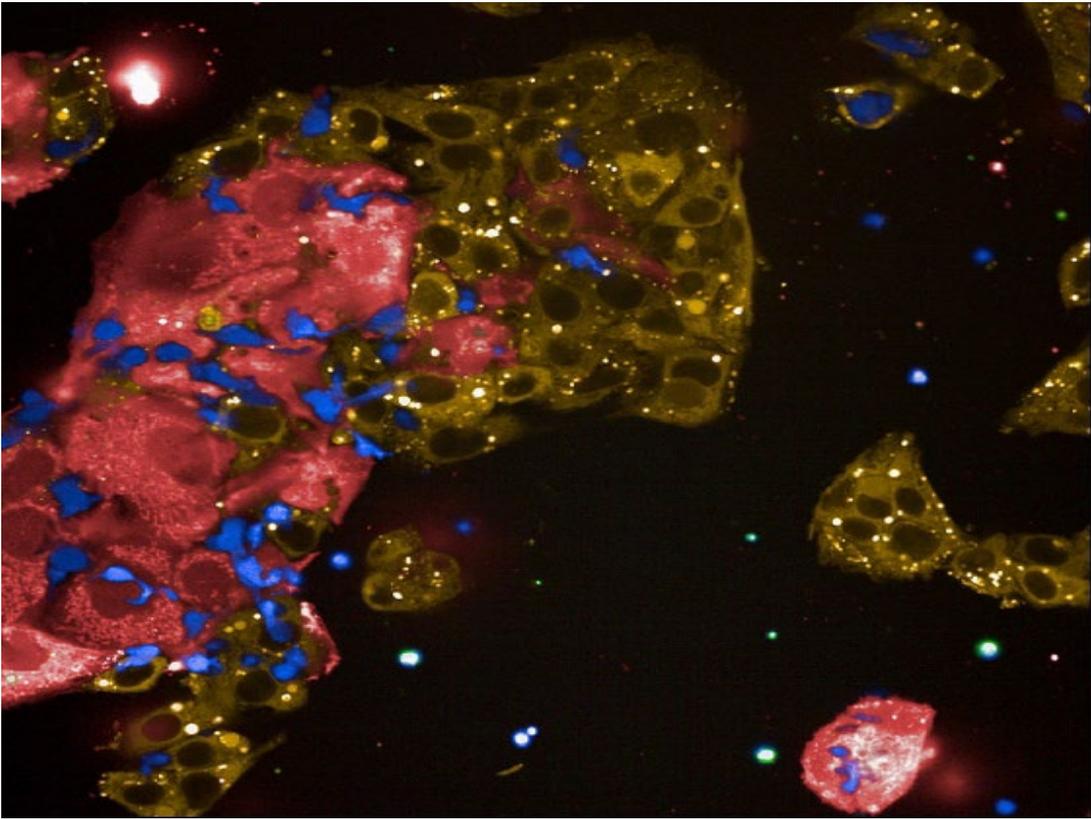
## Key advantages of redirecting non-exhausted T cells

- Same CD3 MoA validated in oncology
- Independent of natural T cell reactivity to Hep B
- Goal is functional cure with finite treatment

## Mass-spectrometry antigen discovery engine for HBV

- Pipeline funnel (e.g., conserved sequences, pHLA presentation/stability, mimetic risk)
- Seven optimal targets identified from envelope, core capsid, and polymerase

# Highly specific killing of cells with integrated HBV DNA

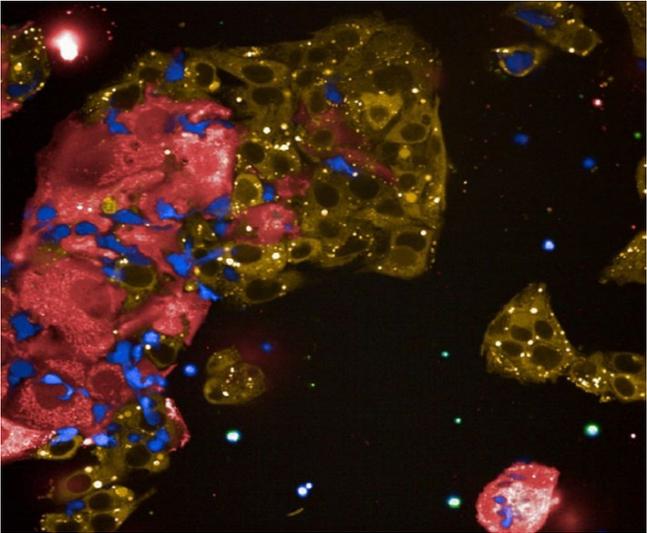


HBV+ cells

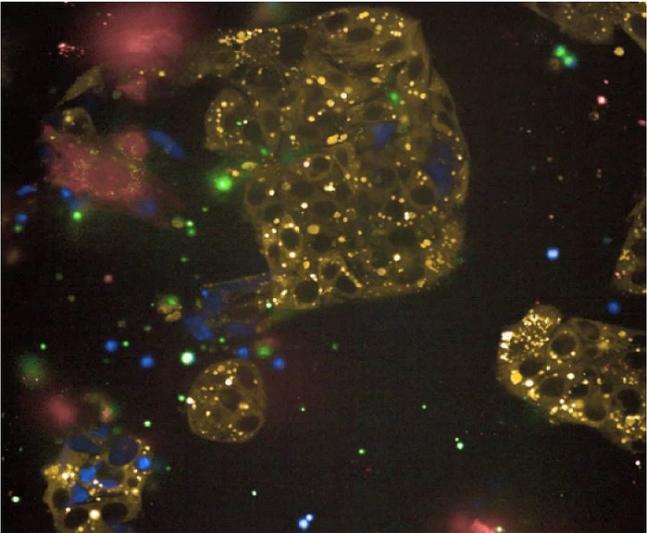
HBV- cells

T cells

Cell death



Co-incubation (start)



HBV+ cell death (end)

# Functional cure program for HIV with goal of eliminating HIV reservoirs



## **Elimination of Latently HIV-infected Cells from Antiretroviral Therapy-suppressed Subjects by Engineered Immune-mobilizing T-cell Receptors**

Hongbing Yang<sup>1</sup>, Sandrine Buisson<sup>2</sup>, Giovanna Bossi<sup>2</sup>, Zoë Wallace<sup>1</sup>, Gemma Hancock<sup>1</sup>, Chun So<sup>1</sup>, Rebecca Ashfield<sup>2</sup>, Annelise Vuidepot<sup>2</sup>, Tara Mahon<sup>2</sup>, Peter Molloy<sup>2</sup>, Joanne Oates<sup>2</sup>, Samantha J Paston<sup>2</sup>, Milos Aleksic<sup>2</sup>, Namir J Hassan<sup>2</sup>, Bent K Jakobsen<sup>2</sup> and Lucy Dorrell<sup>1</sup>

- Same MoA as tebentafusp, but optimized for low target viral peptide presentation
- Bypasses exhausted T cells
- Targets highly conserved & functionally constrained viral epitopes
- Active in ex vivo assays of infected CD4+ T cells from ART-treated HIV patients
- Soluble format access to tissue reservoirs

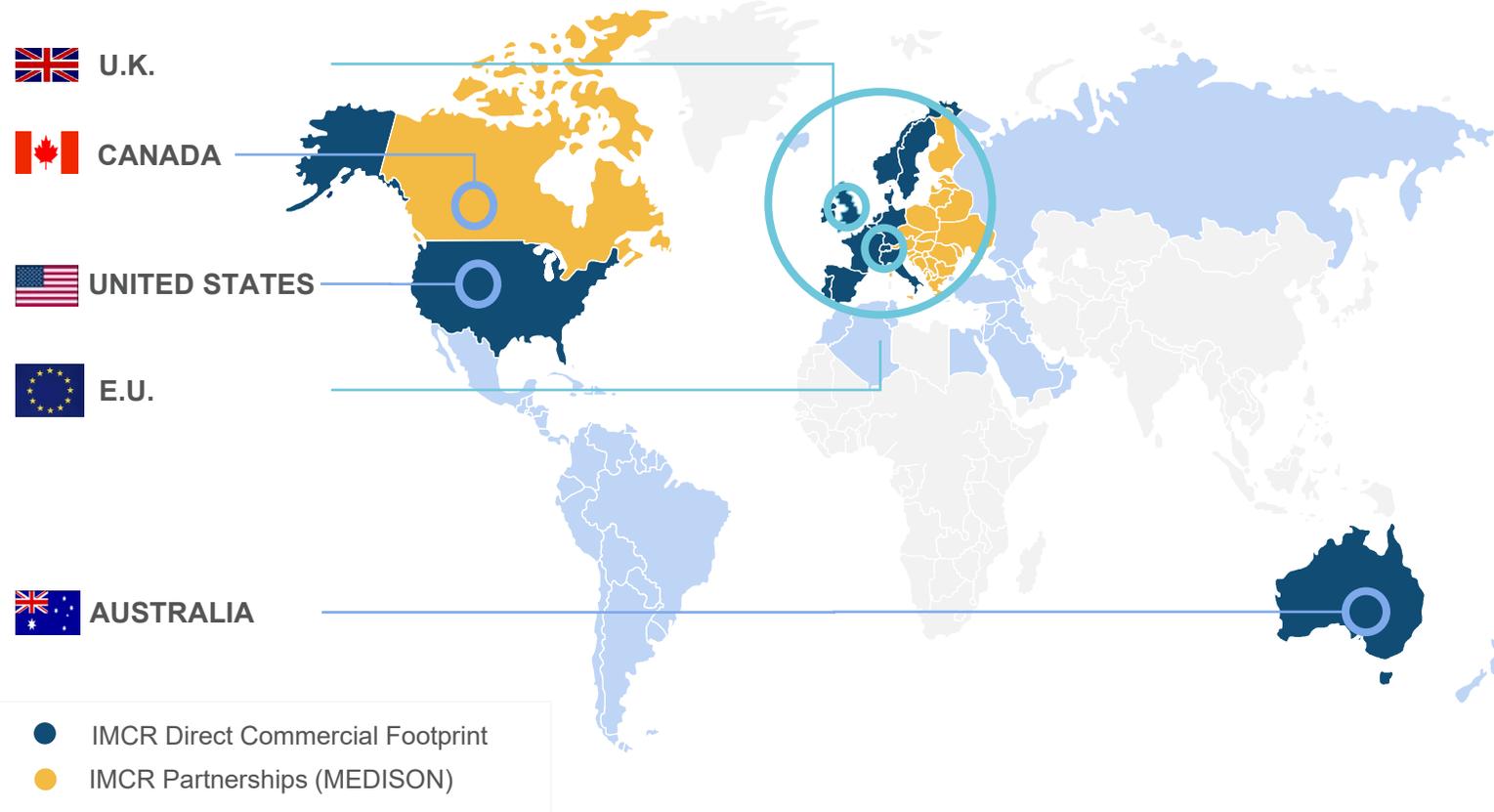
**IMC-M113V CTA accepted in 2021**

# Tebentafusp Launch Readiness & Upcoming Portfolio Milestones

# Our ambition: transform the lives mUM patients around the world

Global regulatory acceptances

## 5 accepted regulatory fillings



**200+** patients on early access program



**US launch-ready**



**Accelerated review in EU**



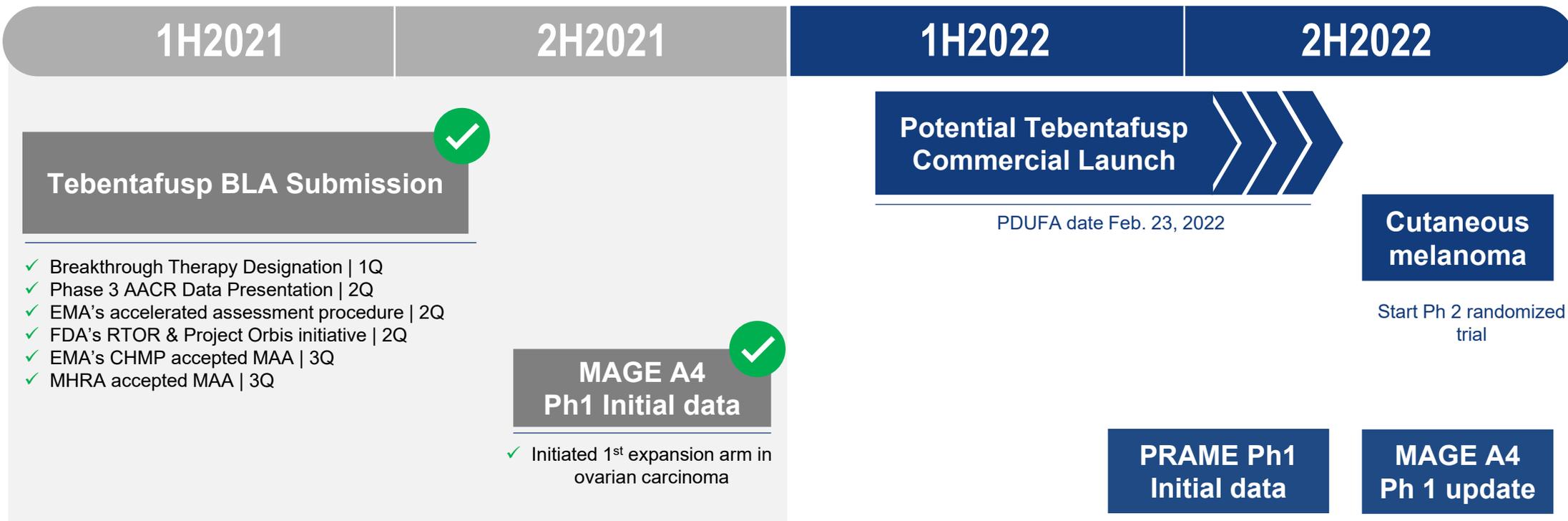
**+22 countries**

in partnership with MEDISON in Canada, Central Eastern Europe, and Israel

**~1,000 patients / year in US and initial priority European markets<sup>1</sup>**

# Portfolio milestones

## Oncology



## Infectious Diseases



**\$320M cash as of YE 2021**

# Immunocore is the most advanced TCR company

- ✓ **First clinically validated TCR platform with survival benefit**
- ✓ **5 clinical-stage programs**
- ✓ **Tebentafusp PDUFA 2/2022, EU & UK MAA submissions accepted**
- ✓ **Multiple value inflection points over the next 12 months**

# IMMUNOCORE

