

**IMMUNOCORE** 

# Science to Transform Lives

NOVEMBER 2022

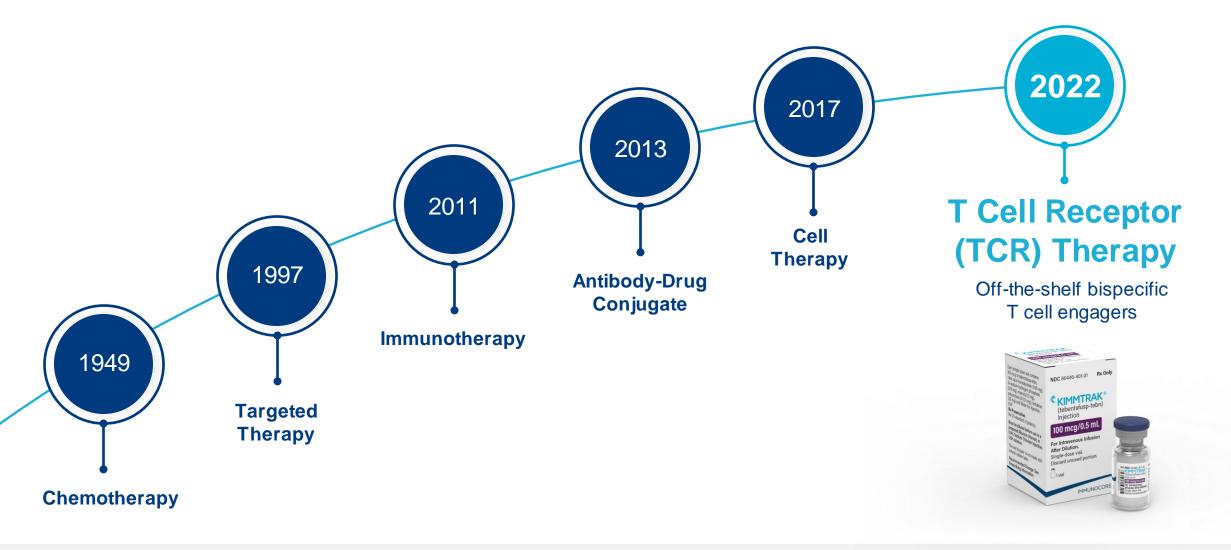
# **Forward-looking statement**

This presentation contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. Words such as "may," "can," "will," "believe," "expect," "plan," "anticipate" and similar expressions (as well as other words or expressions referencing future events or circumstances) are intended to identify forward-looking statements. All statements, other than statements of historical facts, included in this presentation are forward-looking statements. These statements include, but are not limited to, statements regarding the marketing, therapeutic potential, and expected clinical benefits of Immunocore's product candidates, including extended overall survival benefit and reduction in circulating tumor DNA; statements regarding the durability, efficacy and toleration of Immunocore's product candidates; expectations regarding the commercialization of KIMMTRAK including potential growth opportunities and trends and increasing access to KIMMTRAK as the product launch continues in further quarters; expectations regarding the value proposition of KIMMTRAK in metastatic uveal melanoma (mUM); expectations regarding the potential market size and opportunity for Immunocore's other product candidates, including statements with respect to potential patient population: expectations regarding future milestones and value inflection points; future development plans of tebentafusp and Immunocore's other product candidates; the design, progress, timing, scope and results of Immunocore's clinical trials, including the timing for patient enrollment, the initiation of a Phase 2/3 clinical trial for previously treated, advanced melanoma patients and for providing a Phase 1 update for MAGE-A4 and the validation of the ImmTAC platform. These forward-looking statements are based on management's current expectations and beliefs and are subject to a number of risks, uncertainties and important factors that may cause actual events or results to differ materially and adversely from those expressed or implied by any forwardlooking statements, many of which are beyond Immunocore's control. These include, without limitation, risks and uncertainties related to the impact of worsening macroeconomic conditions and the ongoing and evolving COVID-19 pandemic, the war in Ukraine or global geopolitical tension on Immunocore's business, strategy and anticipated milestones, including Immunocore's ability to conduct ongoing and planned clinical trials; Immunocore's ability to obtain and maintain regulatory approval of its product candidates; Immunocore's ability to obtain clinical supply of current or future product candidates or commercial supply of KIMMTRAK or any future approved products, including as a result of supply chain disruptions; Immunocore's ability and plans to launch, market and sell KIMMTRAK or any future approved products, to continue to establish and expand a commercial infrastructure; Immunocore's ability to successfully expand the approved indications for KIMMTRAK, or obtain marketing approval for KIMMTRAK in additional geographies in the future; the delay of any current or planned clinical trials, whether due to the COVID pandemic, patient enrollment delays or otherwise; unexpected safety or efficacy data observed during preclinical studies or clinical trials and Immunocore's ability to successfully demonstrate the safety and efficacy of its product candidates and gain approval of its product candidates on a timely basis, if at all; competition with respect to market opportunities; actions of regulatory agencies, which may affect the initiation, timing and progress of clinical trials or future regulatory approval; Immunocore's ability to obtain, maintain and enforce intellectual property protection for KIMMTRAK or any product candidates it is developing; Immunocore's need for and ability to obtain additional funding on favorable terms or at all, including as a result of worsening macroeconomic conditions such as rising inflation and interest rates, volatility in the capital markets and related market uncertainty; and the success of Immunocore's current and future collaborations, partnerships or licensing arrangements. These and other risks and uncertainties are described in greater detail in the section titled "Risk Factors" in Immunocore's filings with the Securities and Exchange Commission, including Immunocore's most recent Annual Report on Form 20-F, as supplemented by its most recent filings that Immunocore has made or may make with the SEC in the future. Such risks may be amplified by the COVID-19 pandemic and its potential impact on Immunocore's business and the overall global economy. Any forwardlooking statements represent Immunocore's views only as of the date of this presentation and should not be relied upon as representing its views as of any subsequent date. Immunocore does not assume any obligation to update any forward-looking statements, except as may be required by law.

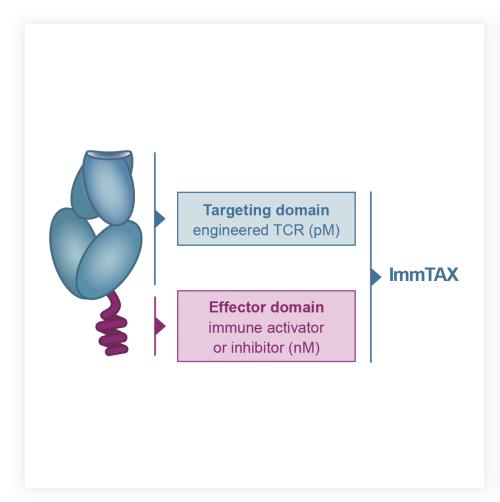
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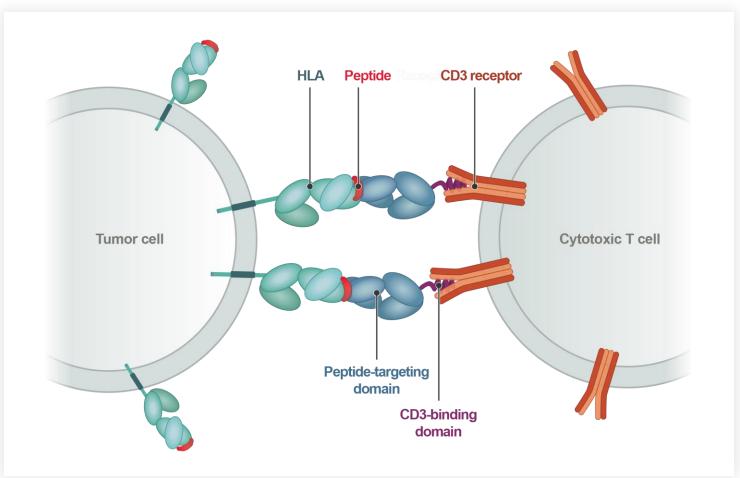
KIMMTRAK™ is a trademark owned or licensed to Immunocore.

# We are defining a new frontier of cancer treatment



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





#### Our team



**Bahija Jallal** 

CEO





**Brian Di Donato** 

CFO & Head of Strategy





**David Berman** 

Head of R&D





**Mohammed Dar** 

CMO





**Andy Hooker** 

VP, CMC & Supply Chain







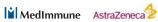


SYNAGIS, FLUMIST, VLP technology for HPV vaccines



**JoAnn Suzich** 

Head of Research





**Mark Moyer** 

Head of Regulatory



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



**Ralph Torbay** 

Head of Commercial





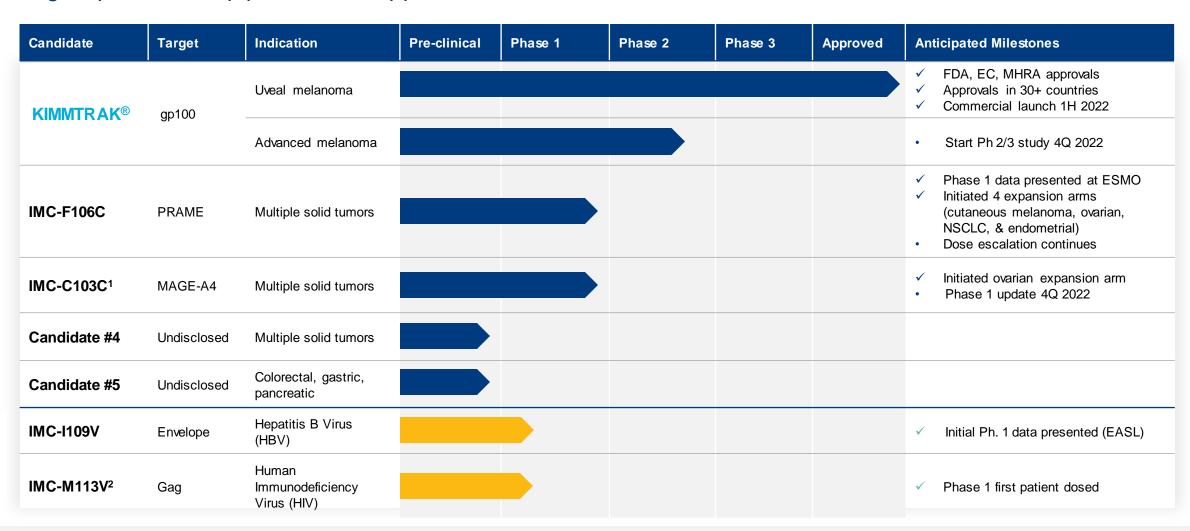
IMFINZI, TAGRISSO, CALQUENCE, GLEEVEC, TASIGNA, ARZERRA, FARYDAK

Regulatory approval of KIMMTRAK® in unresectable or metastatic uveal melanoma (mUM) in 30+ countries

### Our pipeline

ONCOLOGY

#### Leading bispecific TCR pipeline; FDA approval for KIMMTRAK®

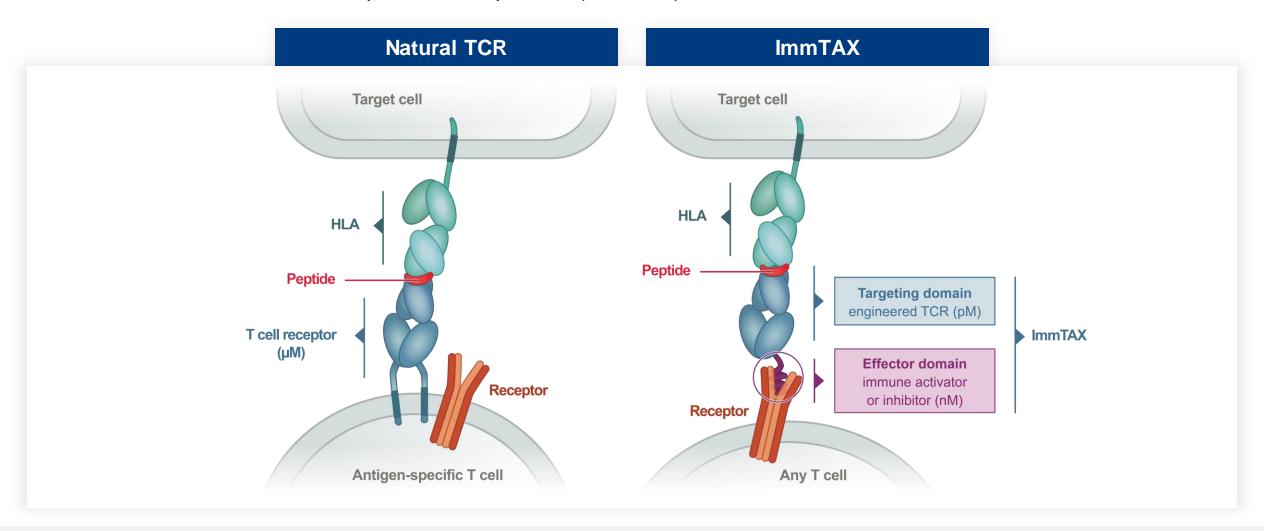


<sup>1.</sup> Developed under a co-development/co-promotion collaboration with Genentech; 2. Program is w holly owned, development costs being provided by the Bill & Melinda Gates Foundation (BMGF), Immunocore retains all development and commercialization rights in the developed w orld.



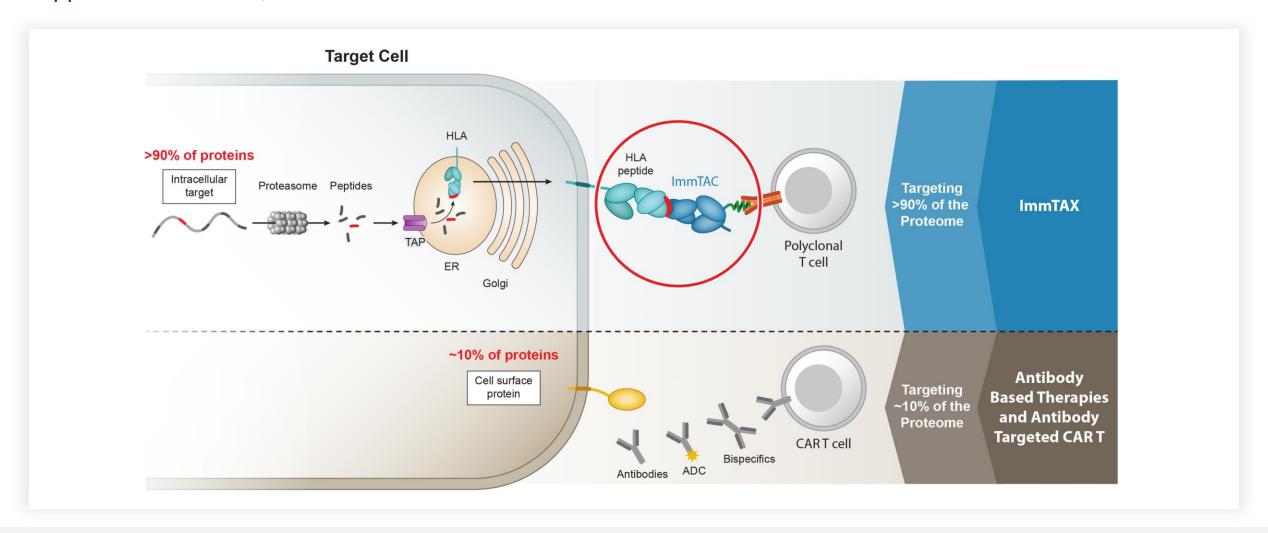
# We pioneered converting membrane-bound T cell receptors...

...into soluble, off-the-shelf, bispecific therapeutics (ImmTAX)



# TCR therapeutics can target nearly the entire human proteome

Application to cancer, autoimmune and infectious diseases





### Metastatic Uveal Melanoma (mUM): is an ultra-rare and aggressive tumor

Targeting gp100 protein in melanoma



**Originates from** melanocytes within the uveal tract of the eye

Median age at diagnosis is 62 years<sup>1</sup>

may develop metastatic disease; 50% liver primary site of metastasis<sup>1</sup>



HLA-02 mUM ~1,000 pts per year in the US/EU<sup>2</sup>



Until KIMMTRAK, no approved treatment<sup>3</sup>

Historic median survival with metastatic disease<sup>2</sup>

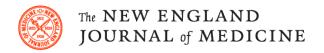
~12 months

# Primary endpoint: Overall Survival (OS) statistically significant

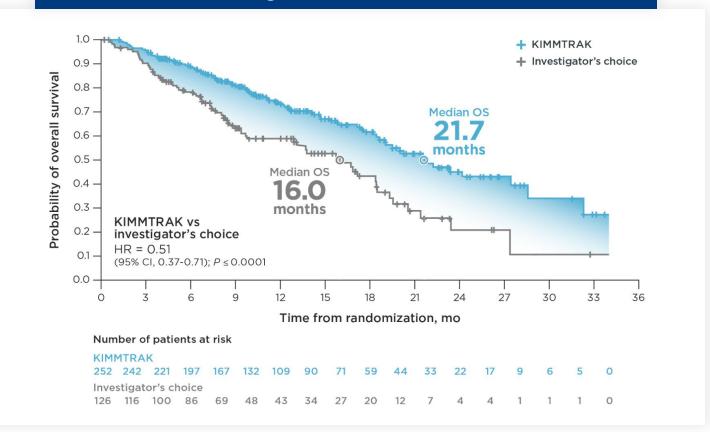
KIMMTRAK®: First-in-class, off-the-shelf, bispecific TCR

# KIMMTRAK was proven to extend median OS by 6 months

- > 21.7 months median OS
- > 0.51 hazard ratio



# Overall Survival benefit in patients treated with KIMMTRAK or investigator's choice in first-line



# Safety profile of KIMMTRAK was predictable and manageable

#### **Key KIMMTRAK findings**

- Adverse Reactions (ARs) consistent with mechanism of action
- Majority of ARs in first 3 weeks
- Low discontinuation rate: KIMMTRAK 2% vs. Investigator's Choice of 4.5%
- No treatment related deaths

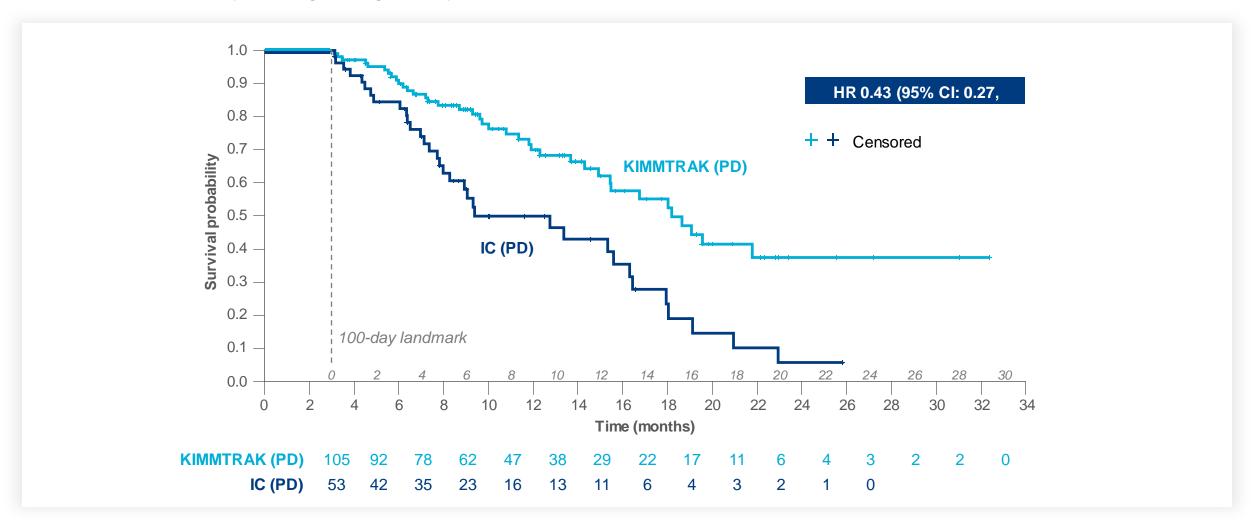
378 previously untreated mUM patients randomized 2:1 KIMMTRAK vs. Investigator's Choice (pembrolizumab 82%, ipilimumab 13%, dacarbazine 6%)

	KIMMTRAK (n = 245)*		
Adverse Reactions (AR)	Any Grade,%	Grade 3 or 4, %	
Any	244 (99.6)	110 (45)	
Cytokine release syndrome <sup>a</sup>	89	0.8	
Rash⁵	83	18.4	
Pyrexia	76	3.7	
Pruritus	69	4.5	
Fatigue <sup>b</sup>	64	5.7	
Nausea	49	2	
Chills	48	0.4	
Hypo-/hyperpigmentation <sup>b</sup>	47	0.4	
Abdominal pain <sup>b</sup>	45	2.9	
Edema <sup>b</sup>	45	0	

<sup>\*</sup> KIMMTRAK. US Package insert. Immunocore Ltd.; 2021. Adverse reactions listed are those with any grade >45%; a. Represents algorithmic identification of CRS cases based on ASTCT grading criteria (Lee et al. 2019). b. Represents a composite of multiple related terms.

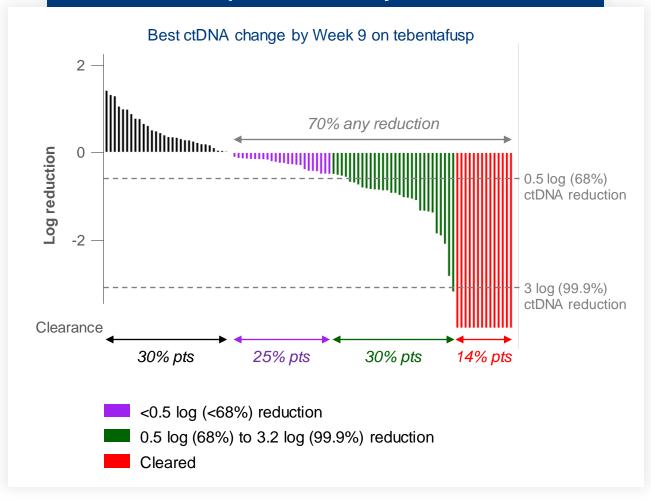
# OS benefit in patients with best response of Progressive Disease

Landmark OS analysis beginning at Day 100

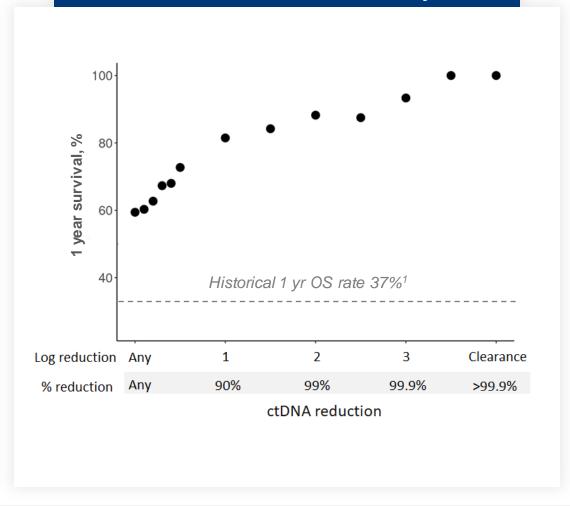


# Circulating tumor (ctDNA) better surrogate of OS than RECIST

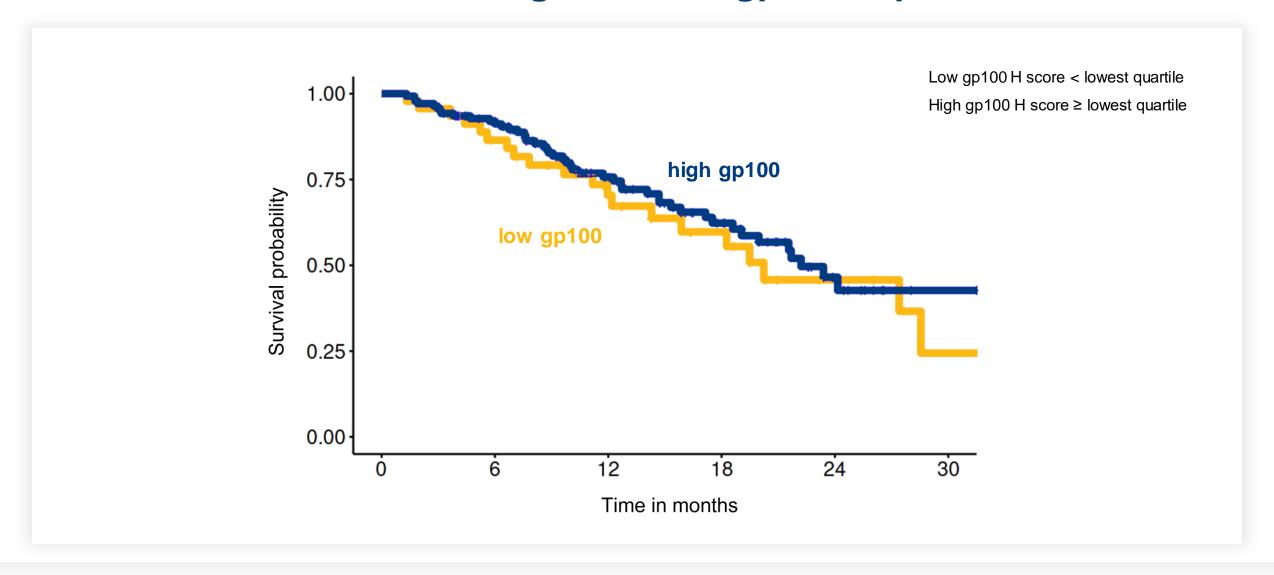
#### 70% evaluable patients had any ctDNA reduction<sup>1</sup>



#### ctDNA reduction correlates with 1 year OS

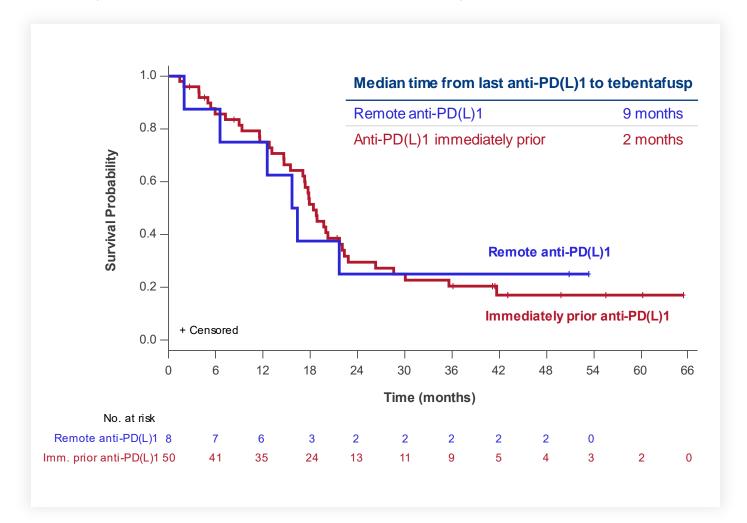


# KIMMTRAK OS benefit in high and low gp100 expression



### Tebentafusp active in cutaneous melanoma

OS by whether prior anti-PD(L)1 therapy was remote or most recent therapy

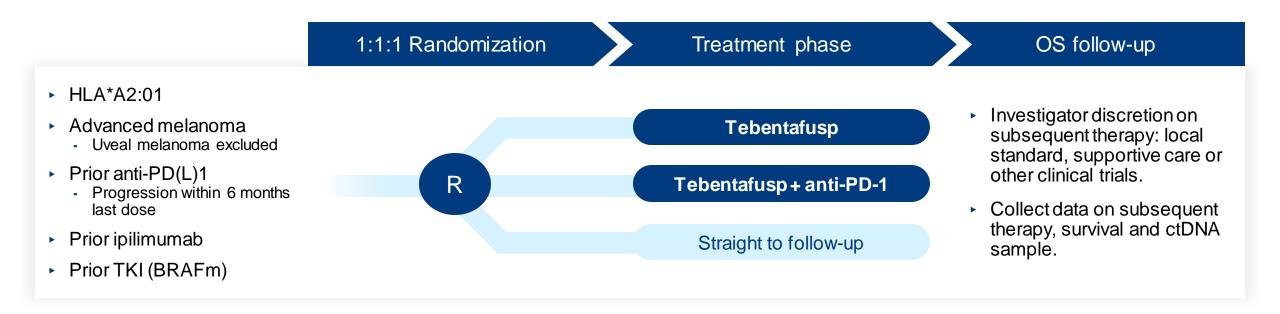


Time from prior anti-PD(L)1	1-yr OS	2-yr OS
Remote	75%	22%
Immediately prior	<b>75%</b>	23%
Benchmark	55%	N/A

Time since last dose of prior anti-PD(L)1 does not impact OS

# Phase 2/3 trial for previously treated, advanced melanoma patients

Randomization to 'real world' treatment as a control arm | Initiation of trial expected Q4 2022

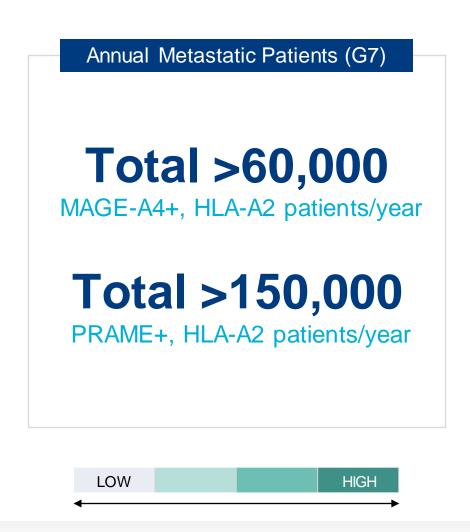


Phase	Primary endpoint	Per arm size	
2	ctDNA and OS	40	Optionality to review Phase 2 data to inform changes to Phase 3, including line of prior therapy, dropping an Arm and optimize powering of study
3	OS	170	



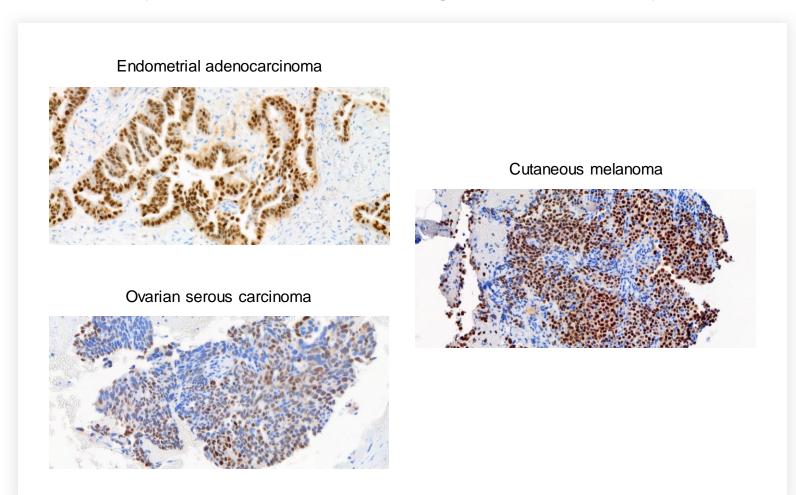
# PRAME & MAGE-A4 expressed in multiple solid tumor types

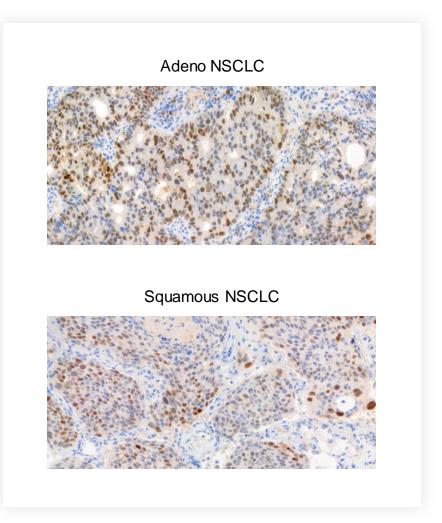
**gp100 PRAME** MAGE-A4 **Tumor** expression expression expression Cutaneous melanoma Relative Checkpoint Inhibitor sensitivity **RCC** Bladder **NSCLC** HCC Gastric Esophageal **SCCHN SCLC TNBC** Endometrial Cervical Ovarian Uveal melanoma



# IMC-F106C: ImmTAC targeting HLA-A2-presented peptide from PRAME

Most broadly expressed cancer-testis antigen in several tumor types but with minimal normal tissue expression





# IMC-F106C-101 PRAME Phase 1 study design

#### **Tumor assessment every 9 weeks**

Screening Treatment Follow-up

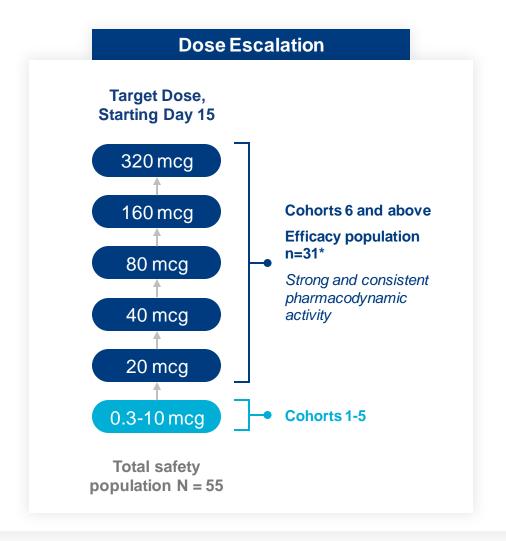
Weekly IV infusion with intra-patient dose escalation (over 3 weeks)

#### Key eligibility criteria

- HLA-A\*02:01 (central testing)
- Select advanced solid tumors
- Tumor PRAME by immunohistochemistry
  - High PRAME prevalence: enroll all comers; test retrospectively
  - All other indications: prospective confirmation of PRAME

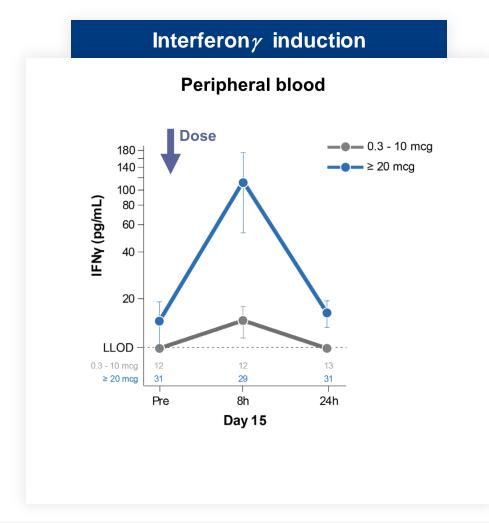
#### Key objectives

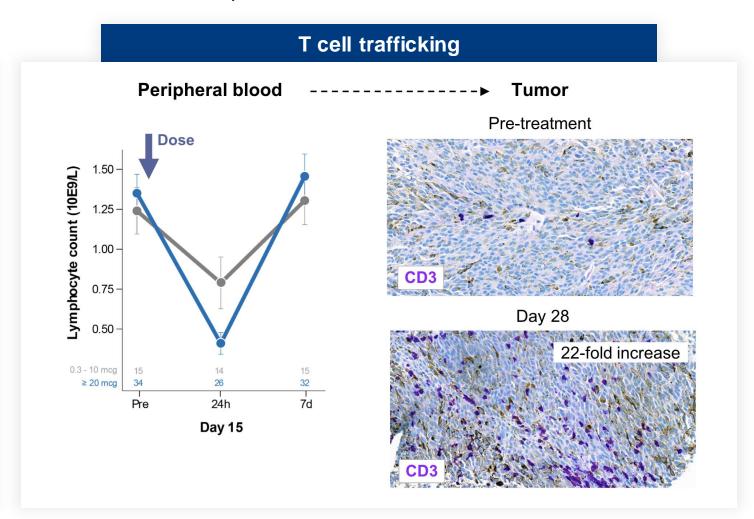
- Primary endpoint
  - Determine MTD/expansion dose
- Secondary endpoint
  - Preliminary antitumor activity
  - Pharmacokinetics
  - Pharmacodynamic markers



# Strong and consistent pharmacodynamic activity at ≥20 mcg IMC-F106C

T cell activation and re-direction into tumor seen across ImmTAC platform





# **Baseline patient characteristics**

Characteristic	Safety Population N=55	Efficacy Population N=31 <sup>†</sup>
Age - Mean (range)	60 (26, 79)	61 (36, 79)
ECOG status 0 – n (%)	30 (55%)	19 (61%)
PRAME status (IHC)		
Positive	49 (89%)	28 (90%)
Negative	2 (4%)	0
Not evaluable	4 (7%)	3 (10%)
Median H-score	195	188
Tumor type		
Melanoma	34 (62%)	17 (55%)
Uveal (UM)	26 (47%)	11 (35%)
Cutaneous (CM)*	8 (15%)	6 (19%)
Ovarian Carcinoma	10 (18%)	5 (16%)
Serous (SOC)*	7 (13%)	4 (13%)
Non-serous	3 (5%)	1 (3%)
NSCLC	4 (7%)	4 (13%)
TNBC*	3 (5%)	3 (10%)
Endometrial*	4 (7%)	2 (6%)

- Median PRAME H-score in efficacy population was high, 188; most patients enrolled regardless of PRAME testing
- Patients in efficacy population were heavily pretreated
  - Ovarian: all platinum resistant
  - CM: all received prior anti-PD1 and anti-CTLA4
  - NSCLC: all received prior anti-PD1
  - TNBC and endometrial: 2-5 prior lines of therapy

<sup>\*</sup> In efficacy population, these tumors enrolled regardless of PRAME immunohistochemistry (IHC) testing, w hich was evaluated retrospectively. NSCLC squamous also enrolled regardless of PRAME testing † Of 36 patients treated at target escalation dose of ≥20 mcg, 5 patients were excluded from efficacy analyses as they were PRAME-negative (n=2) or not yet had tumor assessment (n=3) Hamid, O., et. al, Annals of Oncology (ESMO 2022) 33 (suppl\_7): S331-S355.

#### **IMC-F106C** was well tolerated

Most frequent related AE was Grade 1/2 CRS, consistent with proposed mechanism

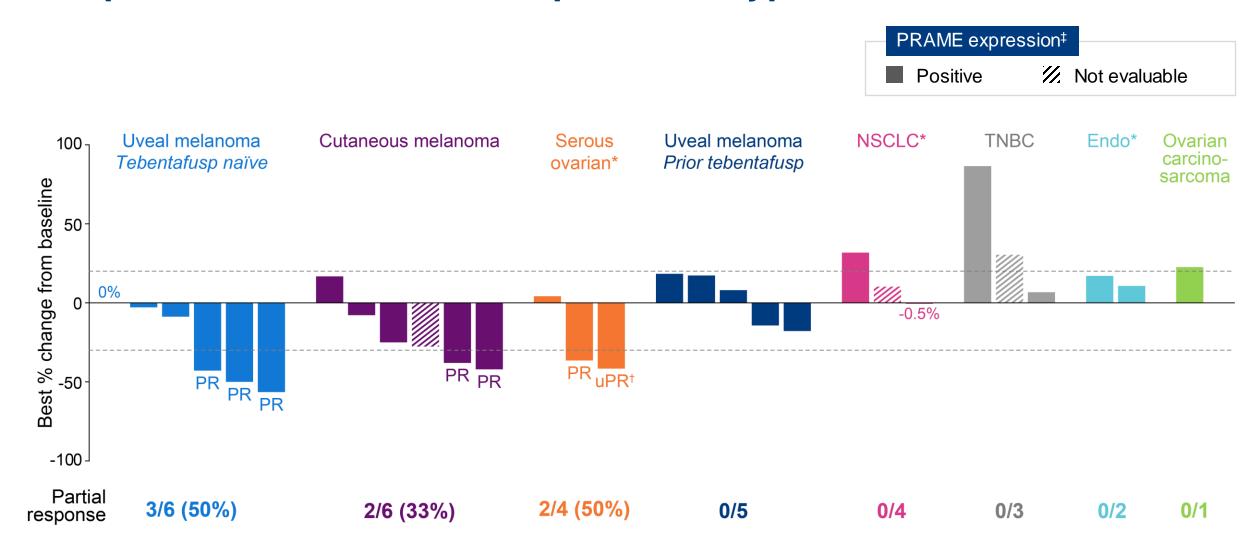
Preferred Term (MedDRA v23.1)	0.3 – 10 mcg <sup>†</sup> N=18	20 – 320 mcg <sup>†</sup> N=37	Total N=55			
All grades (events in ≥ 25% of patients), n (%)						
<b>At least one event</b> 18 (100) 34 (92) 52						
Pyrexia*	10 (56)	21 (57)	31 (56)			
Cytokine release syndrome	5 (28)	22 (59)	27 (49)			
Fatigue	6 (33)	13 (35)	19 (35)			
Hypotension*	3 (17)	15 (41)	18 (33)			
Chills	9 (50)	8 (22)	17 (31)			
Nausea	7 (39)	10 (27)	17 (31)			
Rash	3 (17)	12 (32)	15 (27)			
Grade ≥ 3 (Event	s in > 1 patient), r	n (%)				
At least one event	6 (33)	13 (35)	19 (35)			
Lymphopenia	1 (6)	7 (19)	8 (15)			
Aspartate aminotransferase increased	3 (17)	1 (3)	4 (7)			
Anemia	1 (6)	2 (5)	3 (5)			
Alanine aminotransferase increased	2 (11)	0	2 (4)			
Arthralgia	1 (6)	1 (3)	2 (4)			
Pyrexia*	0	2 (5)	2 (4)			

- MTD not reached
- No treatment-related discontinuation or Grade 5 adverse events
- CRS events were all manageable
  - Majority (77%) within first 3 doses
  - ▶ 71% Grade 1
  - ▶ 29% Grade 2
  - No Grade ≥ 3 CRS
- Adverse events attenuate over time

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

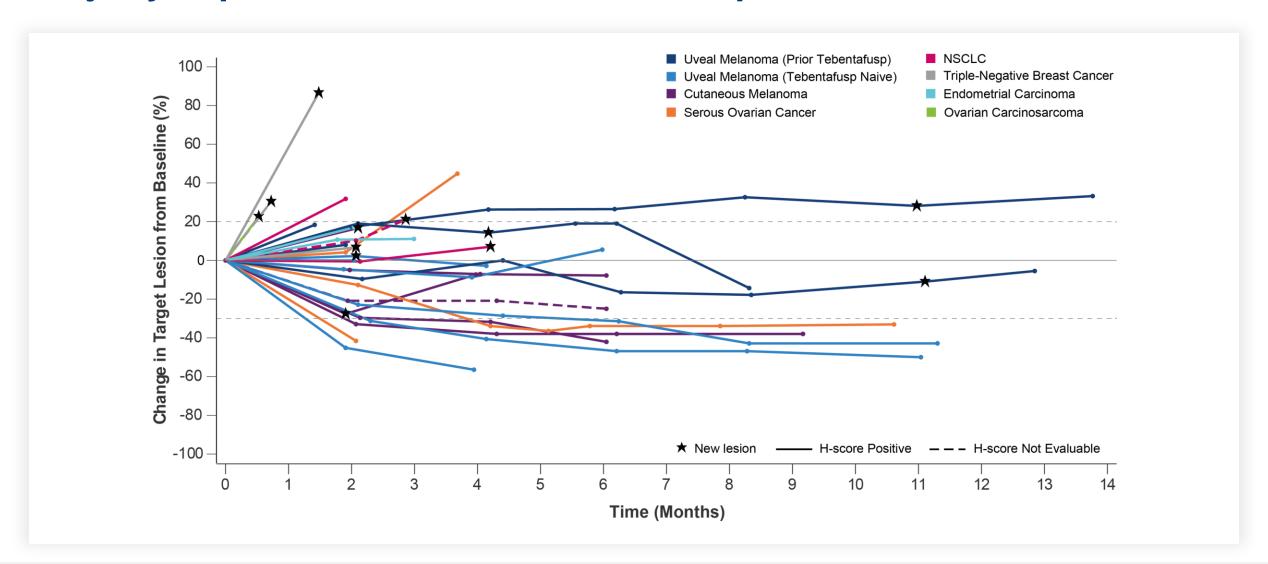
<sup>†</sup> Safety presented by intended target escalation dose on Day 15. 1/37 patients received only a single dose of 2 mcg and did not reach target dose of ≥ 20 mcg Hamid, O., et. al, Annals of Oncology (ESMO 2022) 33 (suppl\_7): S331-S355.

# Responses observed in multiple tumor types



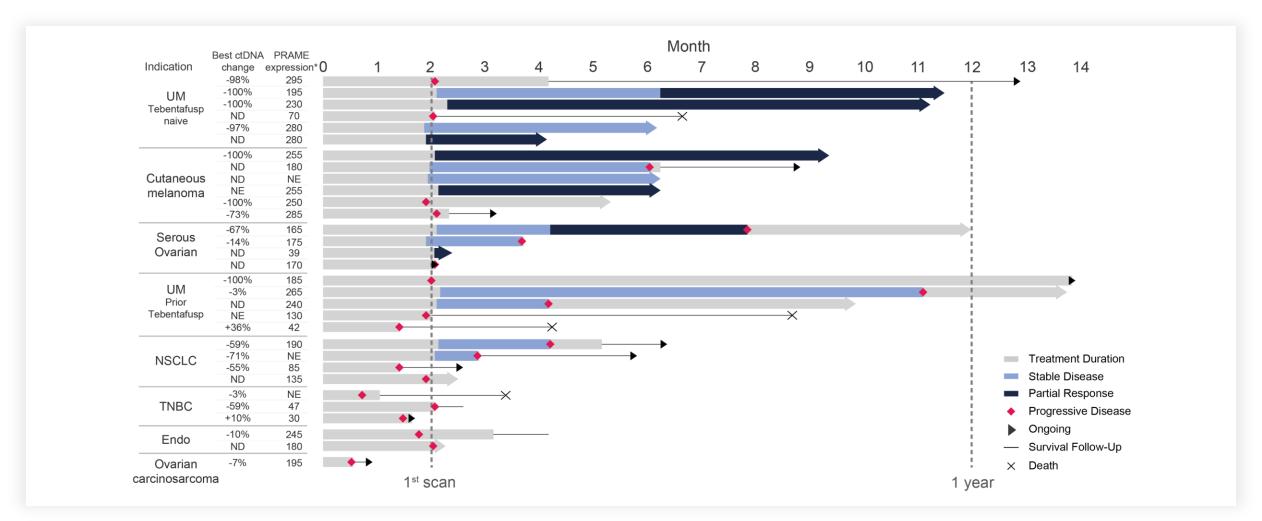
<sup>\*</sup> Two patients (1 with NSCLC, 1 serous ovarian) discontinued treatment due to PD with scan data not available at DCO; † Ovarian cancer patient with unconfirmed PR (uPR) remains on treatment and eligible for confirmation. ‡ PRAME expression assessed by IHC H-score; Two PRAME-negative patients both had PD (not shown); Endo, endometrial carcinoma; NSCLC, non small cell lung carcinoma; TNBC, triple-negative breast cancer. Hamid, O., et. al, Annals of Oncology (ESMO 2022) 33 (suppl\_7): S331-S355.

# Majority of patients have durable tumor response or disease stabilization



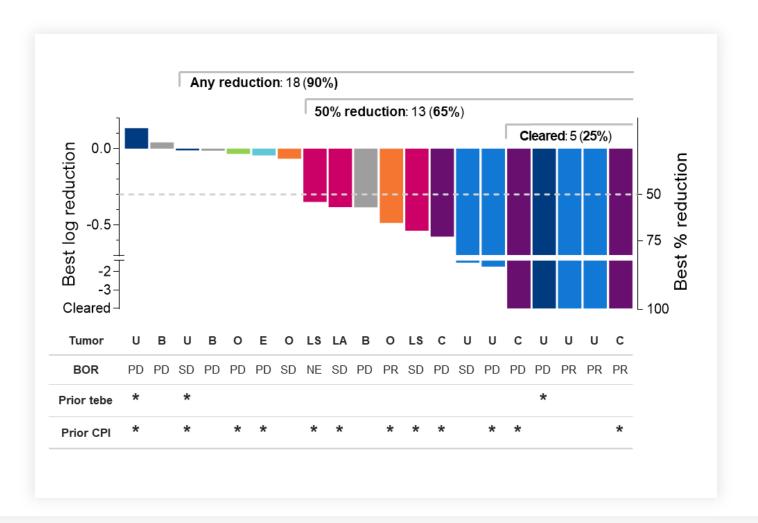
# Responses are durable, 6 of 7 PRs still ongoing

#### Two PRs ongoing for 7+ months



<sup>\*</sup> PRAME expression assessed by IHC H-score Endo, endometrial carcinoma; NSCLC, non small cell lung carcinoma; TNBC, triple-negative breast cancer; UM, uveal melanoma; ctDNA, circulating tumor DNA; ND, not yet determined (9 patients pending); NE, not evaluable; PR, partial response; Hamid, O., et. al, Annals of Oncology (ESMO 2022) 33 (suppl\_7): S331-S355.

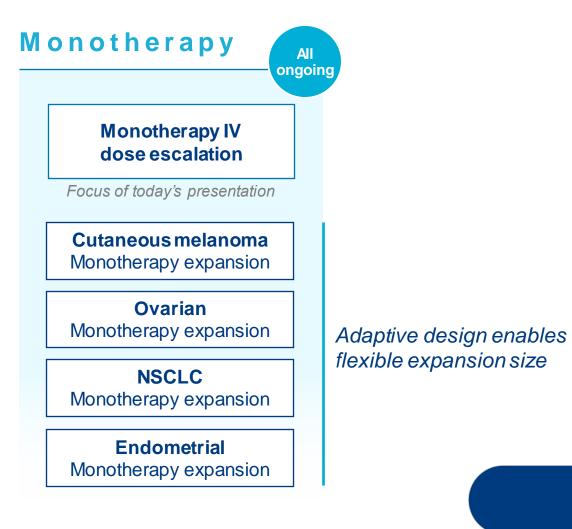
# Reduction in circulating tumor DNA observed across tumor types (n=20)<sup>†</sup>



- 4 PR patients evaluated for ctDNA had > 50% reduction, including 3 with clearance
- Two patients had ctDNA clearance despite best response of PD

ctDNA reduction correlated with OS in KIMMTRAK mUM study

# IMC-F106C-101 designed as an adaptive Phase 1/2 study



Combinations

Checkpoint inhibitor combinations

Chemotherapy combinations

ImmTAC combinations

Enables future randomized trials into earlier lines of therapy

Initial data provides optionality to develop in single arm or randomized trials

# PRAME, validated as TCR target, expressed in many solid tumors

revalence of PRAME expression <sup>1</sup>	<b>Tumor type</b>	HLA*02:01+, PRAME+ metastatic patients (G7) <sup>2</sup>		
	Endometrial	>10K		
70 4000/	Melanoma	>10K		
70-100%	Ovarian	>15K		
_	NSCLC-squamous	>30K		
50-70%	NSCLC-adeno	>40K		
	SCLC	>15K	T-4-1, 450,000	
	TNBC	>5K	Total $>150,000$	
	SCCHN		PRAME+, HLA-A2 patients/year	
	Gastric	>30K	T TO TIVIL T, TILD TO TE PATIOTION YOU	
20-50%	RCC		>30K	
	Esophageal			
	Cholangiocarcinoma			
	Cervical			

 $<sup>1.\</sup> PRAME\ prevalence\ derived\ from\ immunohistochemistry\ and\ RTqPCR\ of\ patient\ samples\ and\ analysis\ of\ TCGA$ 

# 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			

#### Clinical activity in ovarian and HNSCC

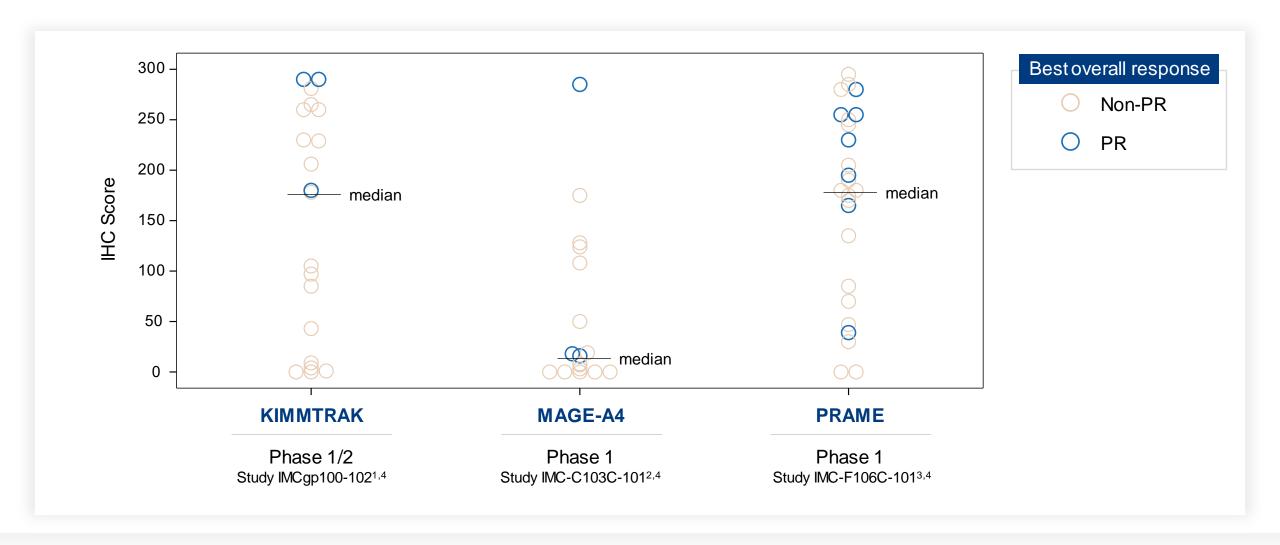
Indication	H-Score	Dose	Response	DOR
HNSCC	285	240 mcg	Confirmed PR^ (ongoing)	2+ mo
Ovarian	19	140 mcg	Overall TL reduction (-44%) but new lesions	
Ovarian	7	140 mcg	Overall TL reduction (-81%) but new lesions	
Ovarian	18	90 mcg	Confirmed PR (ongoing)	4.4+ mo
Ovarian	16	15 mcg	Confirmed PR	8.3 mo

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

<sup>\* 17</sup> 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). TL, target lesions; HINSCC, Head and neck squamous cell carcinoma; ^ confirmed after the presentation data cut-off date

D. Davar Annals of Oncology (2021) 32 (suppl 7): S1398-S1427. 10.1016/annonc/annonc786

# **RECIST** responses enriched at higher H score for PRAME



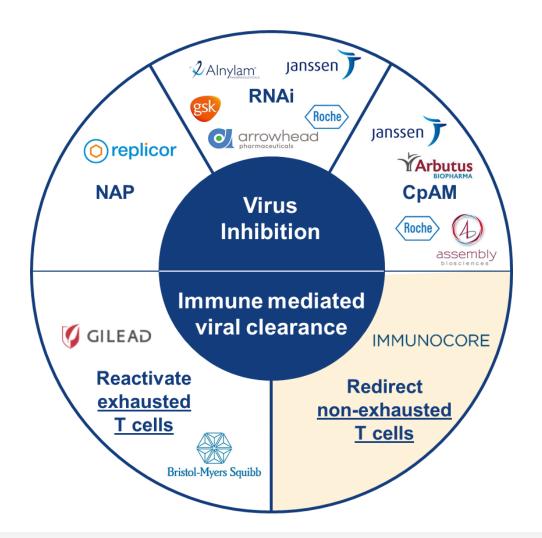
# Validation of ImmTAC platform beyond gp100

On track to present additional clinical data across all three ImmTAC programs in 2022





# Our unique approach for functional cure of chronic Hepatitis B



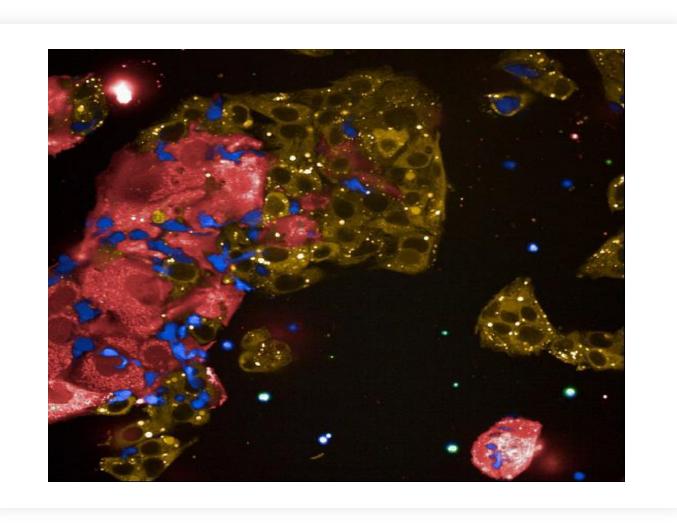
#### Key advantages of redirecting non-exhausted T cells

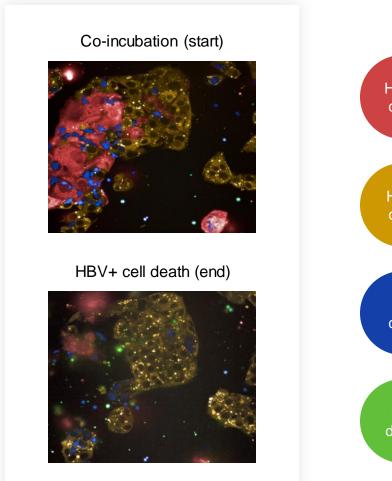
- Same CD3 MoA validated in oncology
- Independent of natural T cell reactivity to Hep B
- Goal is functional cure with finite treatment

#### 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







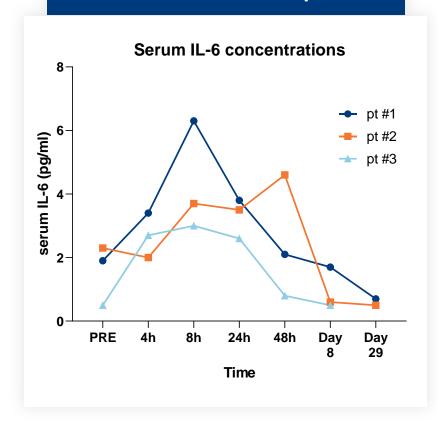




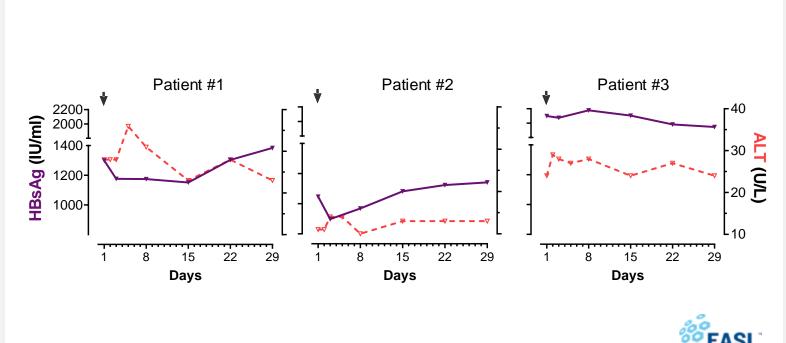


# IMC-I109V: single 0.8 mcg dose results presented at EASL 2022

#### Induction of IL-6 in all 3 patients<sup>1</sup>



#### Transient decrease in HBsAg transiently coincided with transient increase in ALT <sup>1</sup>





# 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

Initiated IMC-M113V Phase 1 2022



# We are leading the way in TCR therapeutics

Language Control of the Control of t

First and only
FDA-approved treatment
for metastatic uveal
melanoma

1 st

First T-cell engager to show Overall Survival (OS) in solid tumor

# **Executing on the global commercial launch of KIMMTRAK**

180 new to KIMMTRAK accounts in US

**+50%** of patient potential<sup>^</sup>

80 new to KIMMTRAK accounts in Germany & France

+70% of patient potential<sup>^</sup>

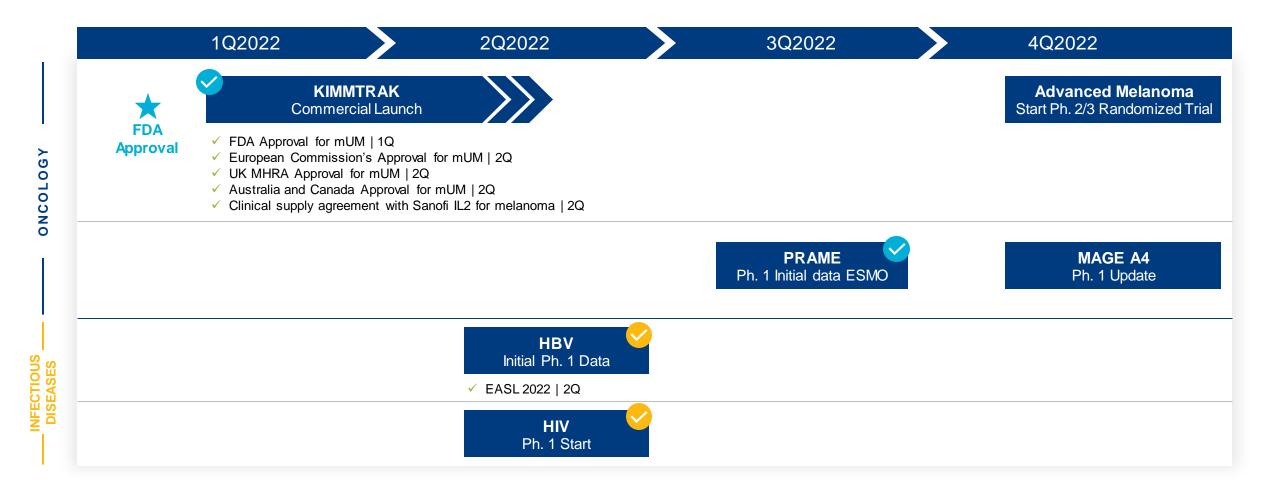
patients treated closer to home in community setting<sup>1</sup>

>90%
of estimated potential US mUM\*
lives covered for KIMMTRAK to
date with policy²

**50**% of patients are now 1L<sup>3</sup>



#### Portfolio milestones



\$387M cash and cash equivalents as of Q3 2022

# Immunocore is the most advanced TCR company



First clinically validated TCR platform with survival benefit



5 clinical-stage programs across 2 therapeutic areas



KIMMTRAK now approved in 30+ countries (incl. US, UK, EU, & Australia)



Multiple expected value inflection points over the next 12 months

