

Corporate Presentation – June 2023 (ASX: IMM, NASDAQ: IMMP)

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#### Immutep Highlights





# Novel science and advanced pipeline

Pioneering LAG-3 immunotherapy in cancer & autoimmune diseases. Three clinical assets and two earlier stage programs.



# **Compelling** clinical data

First-in-class eftilagimod alpha (efti) has generated compelling clinical efficacy with favourable safety across several cancers.\*



# Validation through partnerships

Multiple partnerships and collaborations with large pharma.











# Global presence; strong balance sheet

Global presence and strong IP across diversified LAG-3 portfolio. Well-funded.









# Substantial market opportunity

Efti has safely improved clinical outcomes for cancer patients with anti-PD-(L)1 therapies and chemo creating large opportunity.

# Deep Pipeline



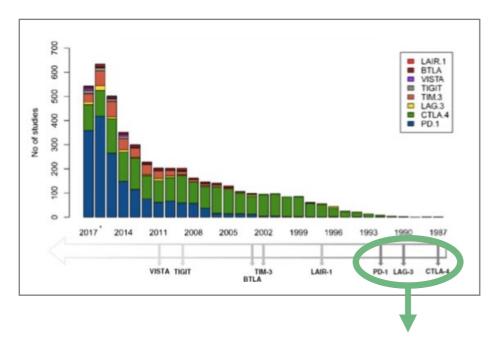
	Program	Indication	Preclinical	Phase I	Phase II	Late Stage*	Collaborations	Commercial Rights
OGY	Eftilagimod Alpha Soluble LAG-3 Protein	1L Head & Neck Squamous Cell Carcinoma (HNSCC)  1L Non-Small Cell Lung Cancer (NSCLC), 2L HNSCC, PD-X Refractory 2L NSCLC  Urothelial Cancer  1L NSCLC  Soft Tissue Sarcoma	TACTI-003   Efti+Pembr TACTI-002   Efti+Pembr INSIGHT-005   Efti+Ave INSIGHT-003   Efti+Pen EFTISARC-NEO   Efti+Pe	olizumab <sup>a</sup> Iumab <sup>§, b</sup> nbro+Chemo <sup>§</sup> embro+Radiotherapy <sup>§</sup>			MERCK MERCK Merck KGAA Darmstadt, Germany  Narodowy Instruct Onkologil Onkologil Onkologil Onkologil Onkologil Parameterstands	immutep Global Rights ex-China
ONCOTOGA	Anti-LAG-3 Small Molecule	HR+/HER2- Metastatic Breast Cancer & TNBC  Metastatic Breast Cancer & Solid Tumors  Undisclosed	AIPAC-003   Efti+Paclita: Efti+Paclitaxel and Efti+Pe				CARDIFF	Efti China Rights  immutep Global Rights
	LAG525 Anti-LAG-3 Antibody	Solid Tumors & Blood Cancer Triple Negative Breast Cancer Melanoma Solid Tumors Triple Negative Breast Cancer					U NOVARTIS	NOVARTIS Global Rights
AUTOIMIMUNE DISEASE	GSK'781 Depleting LAG-3 Antibody  IMP761	Ulcerative Colitis Psoriasis Healthy Subjects					GSK	GSK Global Rights
AUT	Agonist LAG-3 Antibody	Undisclosed						Global Rights

## Immuno-Oncology (IO) Landscape

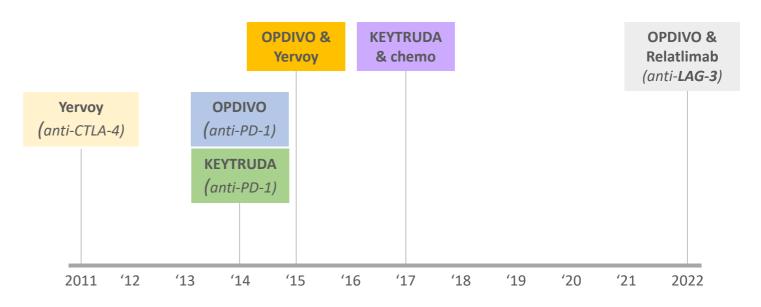




#### Timeline of Immune Checkpoint Discovery\*



#### **Evolution of Immuno-Oncology Therapies**\*\*



The immune system's role in fighting cancer has led to regulatory approval of immuno-oncology therapies targeting the immune checkpoints CTLA-4, PD-1, and LAG-3

LAG-3 is unique in that its (1) inhibition on T cell receptor signalling and (2) activation of dendritic cells both engage the immune system to fight cancer

# LAG-3 Therapeutic Landscape Overview



	Company	Program	Preclinical	Phase I	Phase II	Phase III	Total Trials	Patients
Agoniet	immutep <sup>©</sup>	Eftilagimod Alpha <sup>(5,6)</sup>		10	4	1	15	1,741
	BMS	Relatlimab		10	43	5	58	12,419
	Merck & Co. Inc.	Favezelimab		1	10	3	14	2,286
	Regeneron <sup>(1)</sup>	Fianlimab		1	1	2	4	3,932
	H-L Roche	RO7247669		3	5		8	1,489
	BeiGene	LBL-007		2	5		7	1,310
ology	U NOVARTIS	leramilimab		1	4		5	952
Oncology	Macrogenics	Tebotelimab		3	3		6	974
Softa A	Macrogenics Incyte	Tuparstobart		2	3		5	398
	B.I.	Miptenalimab		4	1		5	653
	Innovent	IBI110		3	1		4	428
	Tesaro <sup>(3)</sup>	TSR-033		1	1		2	139
	F-star <sup>(4)</sup>	FS-118		2	1		3	196
	Symphogen <sup>(2)</sup>	SYM022		3			3	97
	Jiangsu Hengr.	SHR-1802		2			2	166
Autoimmune eting Aganist	immutep®	IMP761						
Autoin Depleting	₹ gsk	GSK2831781 (IMP731)		2	1		3	207

Sources: GlobalData, Company websites, clinicaltrials.gov, and sec.gov as of Apr. 14<sup>th</sup>, 2023. Notes: The green hexagon shapes above represent programs conducted by Immutep and/or its partners. Total Trials includes all active, completed, and/or inactive trials. Patient totals are based on estimated total enrolled and/or to be enrolled. Not a complete list of currently existing LAG-3 products.

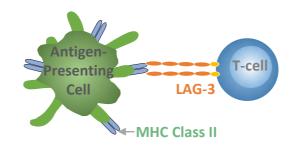
# Immutep's Pioneering Immunotherapies



Only company with four different therapeutic approaches around LAG-3 and MHC Class II interaction

# Targeting MHC Class II on APCs with Eftilagimod Alpha (Efti) Activating APC with efti (soluble LAG-3) leads to broad immune response to fight cancer, including large increase of anti-tumor cells / biomarkers\* Activated MHC Class II agonist NK Cells Dendritic Cells T Cells (CD8+, CD4+)

CXCL10



Binding of LAG-3 on T cells to MHC Class II molecules on APC leads to inhibition of T cell receptor signaling\*. Additionally, soluble LAG-3 is an efficient APC activator.

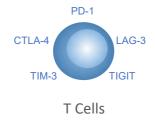
#### Targeting LAG-3 on T cells with Agonist/Antagonist Antibodies & Small Molecules **Blocking LAG-3 prevents** Targeting LAG-3 can suppress immune LAG-3-mediated co-inhibitory signaling, system's response, enabling potential allowing T cells to see and attack cancer treatment of autoimmune diseases LAG525\*\* Anti-LAG-3<sup>^</sup> **IMP761 GSK'781** Antagonist molecule mAb

Monocytes

# Efti Brings A Complementary Approach to IO-IO Combinations

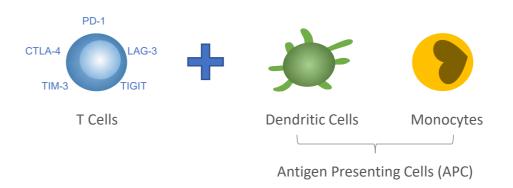


Many IO-IO combinations focus on the same immune cell (T cells) yet target different immune checkpoints on that cell. Can work well in "hot" tumour environments.



**Adaptive Immunity** 

Immutep's complementary IO-IO approach focuses on targeting different immune cells, both T cells & APC (via efti), to bring multiple facets of the immune system to fight cancer. Can work well in "hot" and "cold" tumour environments.

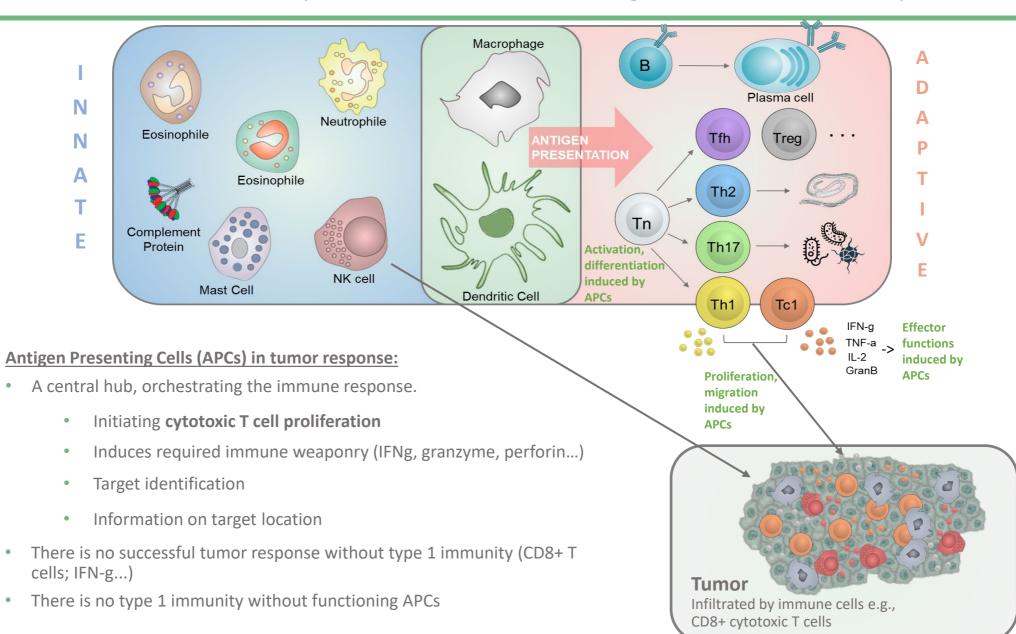


Adaptive and Innate Immunity

#### Antigen Presenting Cells



Efti Activates the 'Generals of the Immune System' via MHC Class II Leading to a Broad Immune Response



# Substantial Commercial Opportunity



Efti's ability to safely improve clinical outcomes of anti-PD-(L)1 therapies across the entire PD-L1 spectrum in multiple solid tumors\* drives substantial commercial opportunity.

#### Efti + Anti-PD-(L)1

- Doubled Overall Response Rate (ORR) of KEYTRUDA® (anti-PD-1) monotherapy in 1st line non-small cell lung cancer and in 2nd line head & neck squamous cell carcinoma in all-comer PD-L1 Phase II trial
- Complete responses (CR) in negative & low PD-L1 expressing patients with KEYTRUDA® (anti-PD-1)
- Deep, durable responses in negative & low PD-L1 expressing patients with IO insensitive cancers with BAVENCIO® (anti-PD-L1)

#### Anti-PD-11



~\$20.9 billion ~\$8.2 billion



Jemperli (dostarlimab-gxly) Injection 500 mg

~\$26 million

OPDIVO

\$29.6 Billion in 2022 sales

#### Anti-PD-L1<sup>1</sup>



~\$3.9 billion

Olimfinzio durvalumab injuridan file Intrasencias Uter 50 migrind.

~\$2.8 billion

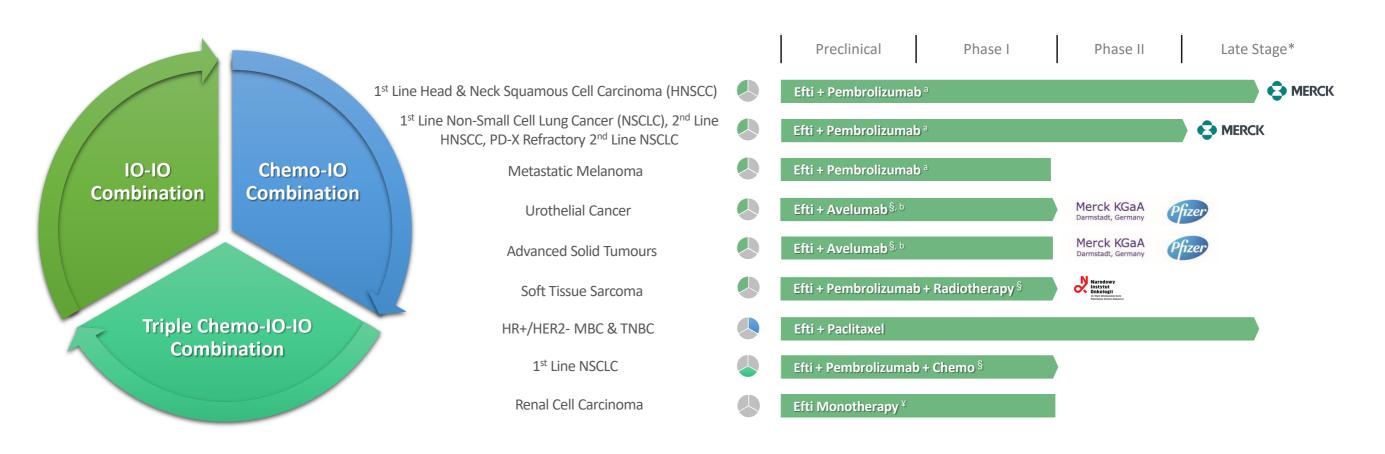


\$7.6 Billion in 2022 sales

## Efti: Pipeline in a Product



With wide-ranging clinical data showing efti's robust potential to improve anti-PD-(L)1 therapies, standard-of-care chemotherapy, and/or both together, efti defines a "pipeline in a product"



#### Late-Stage Clinical Development of Efti







#### **Late-Stage Clinical Development of Efti**

#### Non-Small Cell Lung Cancer (NSCLC) – Planning Registrational Trial in 1st line NSCLC w efti + KEYTRUDA®

- Efti + KEYTRUDA® has FDA Fast Track designation in 1st line NSCLC
- 1.87 million NSCLC diagnoses per annum; highest cause of death among all cancers<sup>1</sup>
- NSCLC drug market will nearly double to \$48 billion in 2031, and immune checkpoint inhibitors are expected to earn more than half of these sales (\$26 billion)<sup>2</sup>

#### Head & Neck Squamous Cell Carcinoma (HNSCC) – Ongoing Phase IIb evaluating efti + KEYTRUDA® in 1st line HNSCC

- Efti has FDA Fast Track designation in 1st line HNSCC
- 900K cases and >400K deaths per annum in HNSCC1
- Global head and neck cancer market size is projected to hit US\$2.99 billion by 2030<sup>3</sup>

#### Metastatic Breast Cancer (MBC) including Triple Negative Breast Cancer (TNBC) – Initiated Phase II/III AIPAC-003 Trial

- Immutep is focused on improving clinical responses for HR+/HER2-neg/low MBC and TNBC patients (~78% of breast cancer cases<sup>4</sup>)
- 2.3 million women diagnosed with breast cancer and 685,000 deaths globally in 2020<sup>5</sup>
- Metastatic breast cancer market to reach \$12.7 billion by 2024<sup>6</sup>

#### **Earlier Stage Clinical Development of Efti**

- Urothelial Cancer (Phase I), Soft Tissue Sarcoma (Phase II, investigator-initiated), and other solid tumor indications



# Non-Small Cell Lung Cancer (NSCLC)



**ASCO 2022 - Dr. Enriqueta Felip** presenting 1L NSCLC data from TACTI-002/KN-798 in Oral Presentation



**SITC 2022 – Dr. Wade lams** presenting 1L NSCLC data from TACTI-002/KN-798 in Late Breaking Abstract Oral Presentation

# 1st line Non-Small Cell Lung Cancer

**Epidemiology & Unmet Need** 





#### **1L NSCLC Epidemiology**<sup>1,2</sup>

- Lung cancer is one of the leading causes of cancer death and there are ~2.2 million cases per annum
- About 80% to 85% of lung cancers are non-small cell lung cancer (NSCLC)
- Only ~20% of patients respond to immune checkpoint inhibitor (ICI) monotherapy & median Overall Survival
   (OS) is still under 24 months for most patients
- ICI + chemo combinations have limited Duration of Response & high discontinuation rates due to toxicity
- Total addressable market (TAM) of NSCLC drug market is expected to nearly double to US\$48 billion in 2031, and ICI are expected to generate more than half of these sales<sup>3</sup>

High unmet medical need for well tolerated, efficacious and durable treatment options, preferably chemo-free

Efti could double the addressable NSCLC patient population with an effective, safe chemo-free IO regimen (i.e., patients with either 1-49% and/or >50% PD-L1 TPS)

TPS Score of ≥50% represents

~30% of 1L NSCLC patients. These patients tend to respond best to anti-PD-(L)1 therapies.

TPS Score of 1-49% represents ~35% of 1L NSCLC patients. These patients have suboptimal response to anti-PD-(L)1 therapies.

TPS Score of ≤1% represents ~35% of 1L NSCLC patients. These patients have negligible response to anti-PD-(L)1 therapies.

1L NSCLC Patient Population by PD-L1 Tumor Proportion Score (TPS)<sup>4</sup>

# Phase II All-Comer PD-L1 Trial Evaluating Efti + Pembrolizumab (KEYTRUDA®) in 1L NSCLC





TACTI-002/KEYNOTE-798: 1st Line Non-Small Cell Lung Cancer (Part A)

#### TACTI-002: Two ACTive Immunotherapeutics in NSCLC & HNSCC **UNSELECTED FOR PD-L1** In collaboration with **MERCK** 1ST LINE MET. NSCLC **PART B** COMBINED IMMUNOTHERAPY ORR, PFS, OS, PK, biomarker 2<sup>ND</sup> LINE MET. NSCLC. safety and tolerability REFRACTORY TO PD-1/PD-L1 TARGETING THERAPY Sites in Europe / US / Australia Recruitment Status Report PART C 2<sup>ND</sup> LINE MET. HNSCC AFTER ✓ Fully approved in all countries **PLATINUM THERAPY** √ Up to 189 patients in three indications ✓ Part A (N=114) completed 30 mg efti s.c. √ Part B (N=36) completed 200 mg pembrolizumab

✓ Part C (N=39) completed

Baseline characteristics for	Part A (N=114)		
Age, median (range), years		67 (4	4-85)
Sex, n (%)	Female / Male	30 (26.3) /	<sup>'</sup> 84 (73.7)
ECOG PS score, n (%)	0/1	43 (37.7) /	71 (62.3)
Smoking status, n (%)	Current or Ex-smoker / Non-smoker	108 (94.7) / 6 (5.3)	
Histology, n (%)	Squamous / Non-squamous / Unknown	40 (35.1) / 72 (63.2) / 2 (1.8	
Metastatic disease, n (%)	Yes / No	Yes / No 113 (99.1) / 1 (0.9	
PD-L1 expression TPS, n <sup>1</sup> (%)	< 1% 1-49% ≥ 50%	Central only 32 (35.6) 38 (42.2) 20 (22.2)	Central + local 37 (34.3) 42 (38.9) 29 (26.9)
Previous therapy, n (%)	Radiotherapy Surgery Systemic therapy for non-metastatic disease	38 (33.3) 23 (20.2) 26 (22.8)	

#### All-comer trial for 1L NSCLC patients with all levels of PD-L1 expression

- ~75% of patients have PD-L1 TPS of <50%</li>
- ~34% of patients have PD-L1 TPS of <1%</li>
- 99.1% had metastatic disease at study entry

(Keytruda®) i.v.

#### Compelling Clinical Results in 1L NSCLC



# TACTI-002 Phase II (1L NSCLC) clinical data and key takeaways across entire patient population, regardless of PD-L1 expression

- Primary objective achieved with 40.4% Overall Response Rate (ORR)
- Promising interim median Progression Free Survival (PFS)
- Robust interim median Duration of Response (mDoR) 21.6 months
- Efti + pembrolizumab is well tolerated and safety profile is similar to pembrolizumab monotherapy
- Efti strengthens responses to anti-PD-1 therapy across entire PD-L1 spectrum

SITC 2022 Oral Presentation - Late-Breaking Abstract was among nine abstracts, out of +1,500 submissions, to be showcased at the SITC press briefing



Combining the antigen-presenting cell activator eftilagimod alpha (soluble LAG-3) and pembrolizumab: efficacy results from the 1st line non-small cell lung cancer cohort of TACTI-002 (Phase II)

lams W<sup>1</sup>; Felip E<sup>2</sup>; Majem M<sup>3</sup>; Doger B<sup>4</sup>; Clay T<sup>5</sup>; Carcereny E<sup>6</sup>; Bondarenko I<sup>7</sup>; Peguero J<sup>8</sup>; Cobo Dols M<sup>9</sup>; Forster M<sup>10</sup>; Ursol G<sup>11</sup>; Kalinka E<sup>12</sup>; Garcia Ledo G<sup>13</sup>; Vila Martinez L<sup>14</sup>; Krebs M.G<sup>15</sup>; Campos Balea B<sup>16</sup>; Kefas J<sup>17</sup>; company authors

¹lams: Vanderbilt Ingram Cancer Center Division of Hematology/Oncology, Nashville, Tennessee, United States; ∓elip: Vall d'Hebron University Hospital, Barcelona, Spain; ¬Majem: Hospital de la Santa Creu i Sant Pau, Barcelona, Spain; ¬Majem: Hospital de la Santa Creu i Sant Pau, Barcelona, Spain; ¬Majem: Hospital de la Santa Creu i Sant Pau, Barcelona, Spain; ¬Majem: Hospital Spain; ¬Majem: Hospital Regional Universitario de Málaga, Spain; ¬Majem: ¬Ma



37th Annual Meeting and Pre-Conference Programs #SITC22

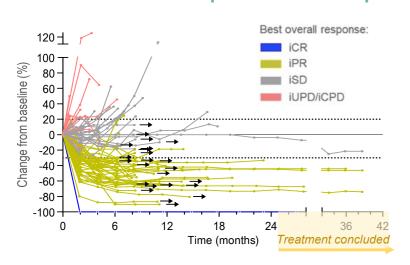
Efti + pembrolizumab received Fast Track Designation from FDA in >1% TPS in 1L NSCLC in Q4'2022 on strength of clinical results

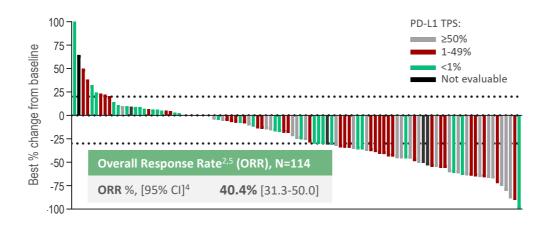
## Deep and Durable Responses Translating Into Overall Survival

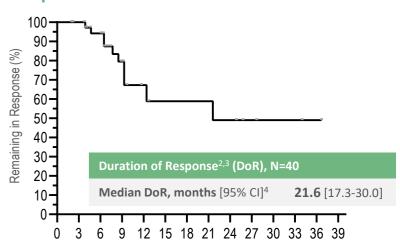


#### **PD-L1 TPS 0 - 100%**

#### Deep and durable responses across all PD-L1 expression levels<sup>1</sup>; interim median Duration of Response of 21.6 months





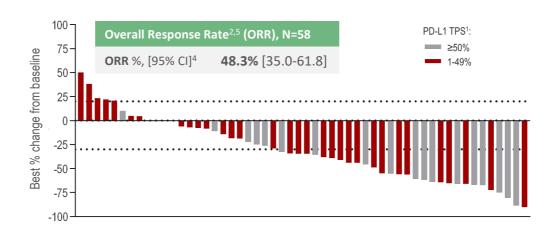


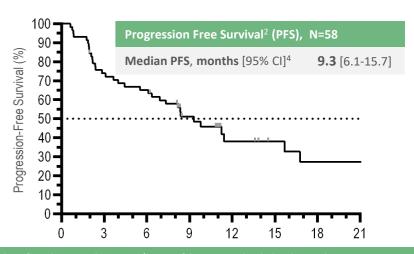
#### **PD-L1 TPS ≥1% (FDA Fast Track designation)**

Strong ORR, DoR and PFS translating into

25.0 months
Interim median Overall
Survival

new cut-off Mar 2023





# Safety



#### **General overview of AEs**

Safety parameter <sup>1</sup>	n (%)
Adverse reactions with fatal outcome <sup>2</sup>	3 (2.6)
Serious adverse reactions <sup>2</sup>	12 (10.5)
Grade ≥3 adverse reactions <sup>2</sup>	14 (12.3)
Adverse reactions leading to discontinuation of treatment <sup>2</sup>	11 (9.6)

<sup>&</sup>lt;sup>1</sup>AEs rated according to NCI CTCAE (v5.0)

#### Frequent AEs (incidence ≥10%) related to study treatment<sup>2</sup>

Adverse event (PT) <sup>1</sup>	Any grade N (%)	Grade 3 N (%)	Grade 4/5 N (%)
Pruritus	23 (20.2)	N/A	N/A
Asthenia	22 (19.3)	N/A	N/A
Rash	15 (13.2)	N/A	N/A
Diarrhoea	12 (10.5)	1 (0.9)	N/A
Fatigue	12 (10.5)	1 (0.9)	N/A

<sup>&</sup>lt;sup>1</sup> AEs rated according to NCI CTCAE (v5.0)

- Treatment with efti plus pembrolizumab is safe and very well-tolerated
- Rate of discontinuation due to drug related adverse events less than 10% and comparable to pembrolizumab monotherapy

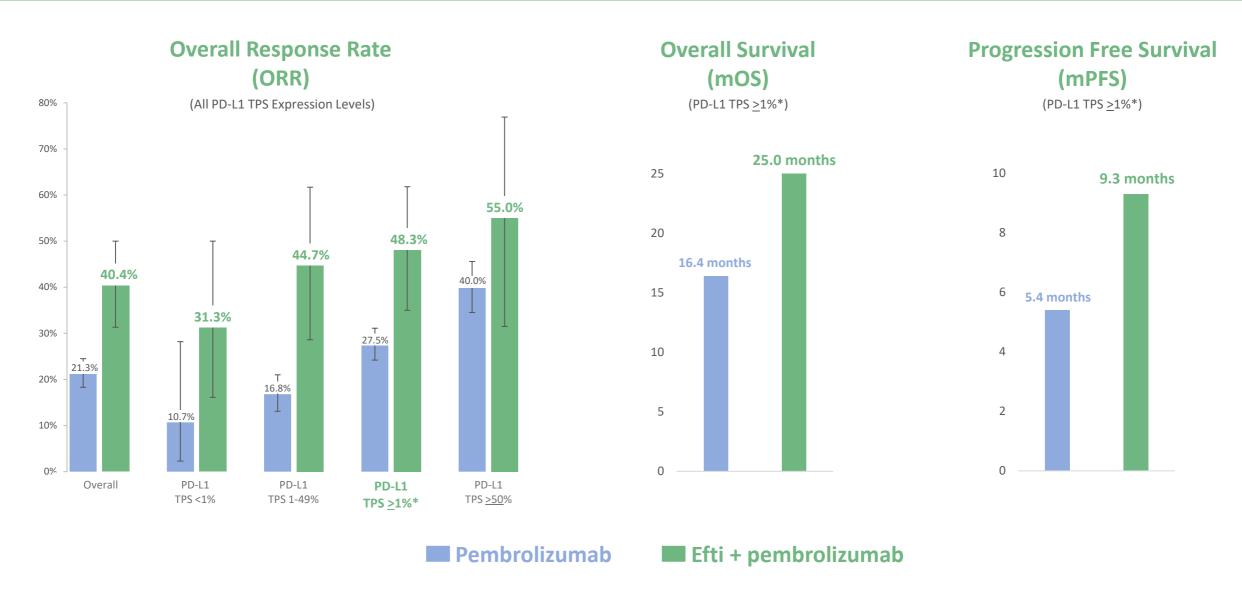
<sup>&</sup>lt;sup>2</sup>relationship to efti and/or pembrolizumab could not be ruled out

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## Benchmarking against Pembrolizumab Monotherapy

Robust Overall Response Rates, Overall Survival, and Progression-Free Survival





\*Efti + pembrolizumab has Fast Track Designation in ≥1% TPS in 1L NSCLC

## Strong Initial Overall Survival Benefit



Efficacy of efti + pembro vs. selected Standard-of-Care in patients with PD-L1 TPS ≥1% in 1<sup>st</sup> line NSCLC

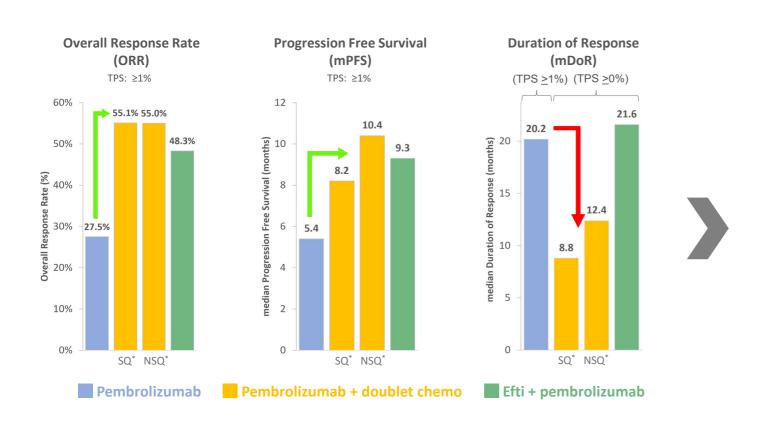
Therapy	Response Rate (RR)	Progression Free Survival (mPFS)	Duration of response (DOR)	AEs leading to disc.	Median OS <sup>2</sup>
Efti + Pembro	48.3%	9.3 months	21.6 months	9.6%	25.0 months
Pembro monotherapy <sup>(1)</sup>	27.5%	5.4 months	20.2 months	6-14%	16.4 months
lpi + Nivo <sup>(1)</sup>	36.0%	5.1 months	23.2 months	18%	17.1 months
Ipi + Nivo + limited 2 cycles of Doublet Chemo	43.3%	7.0 months	15.4 months	19%	15.8 months

**Efti + Pembro** in 1L NSCLC, TPS ≥1% population compared to other published data:

- shows strong ORR, PFS and most importantly superior OS<sup>3</sup>
- while maintaining excellent safety profile and durability of responses
- Fast Track Status has been granted

# Benchmarking against Pembrolizumab Monotherapy & Pembrolizumab-Chemotherapy Combination in 1L NSCLC





	ORR	PFS	DoR
Efti + pembrolizumab	High	High	High*
Pembrolizumab	Low	Low	High*
Pembrolizumab + chemo	High	High	Low

\*Note 34% patients in TACTI-002 have PD-L1 TPS <1% while all pembrolizumab monotherapy patients have PD-L1 TPS >1%

Efti + pembrolizumab has significant promise as a chemo-free dual immuno-oncology (IO-IO) therapy to positively impact 1L NSCLC patient outcomes.

# IO-IO-Chemo Combination Trial (INSIGHT-003) in 1L NSCLC



Promising initial efficacy & safety from first-in-human study evaluating efti + anti-PD-1 + doublet chemo<sup>1</sup>

INSIGHT-003: Phase I in 1st line Non-Small Cell Lung Cancer

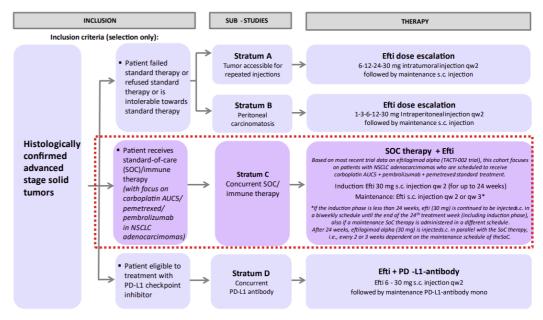
INSIGHT-003 - Third arm (Stratum C) of investigator-initiated study in metastatic 1<sup>st</sup> Line NSCLC patients evaluating triple combination therapy of efti in conjunction with doublet chemo & anti-PD-1 therapy





- Promising 67% overall response rate (ORR) and 91% disease control rate (DCR) in evaluable 1st line non-squamous NSCLC patients (N=21) despite 81% of patients having PD-L1 TPS <50%.<sup>1</sup>
- The triple combination's 67% ORR regardless of PD-L1 expression and 65% ORR in patients with PD-L1 TPS <50% (N=17) compare favourably to reported results from a registrational trial of anti-PD-1 and doublet chemo that yielded a 48% ORR regardless of PD-L1 expression and 40.8% ORR in patients with PD-L1 TPS <50%.<sup>2</sup>
- Triple combination well tolerated & appears to be safe
- Will have additional data updates in CY2023

#### **INSIGHT-003 Study Design**





# Head and Neck Cancer Data Presented at ASCO 2023

# 2nd Line Head & Neck Squamous Cell Carcinoma

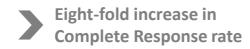


Strong, Long-Lasting Efficacy and Favorable Safety; Positive Benchmarking to Pembro Monotherapy

TACTI-002/KEYNOTE-798: 2nd Line Head & Neck Squamous Cell Carcinoma (Part C)



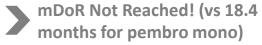


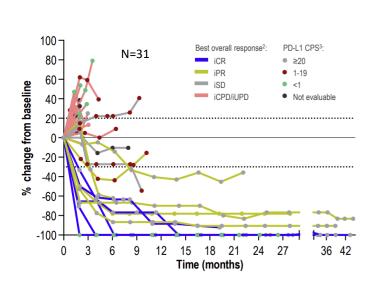


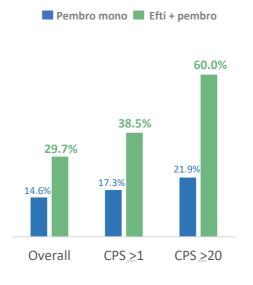


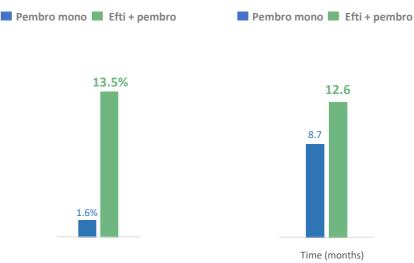
12.6

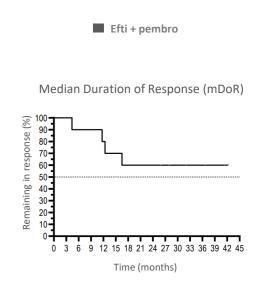
Time (months)











In addition to its impressive efficacy, this dual immuno-oncology approach continues to be safe and well tolerated with adverse reactions leading to treatment discontinuation in only two patients  $(5.1\%)^*$ , which compares favorably to pembro mono  $(6.1\%)^*$ .

# Efficacy Endpoints Across PD-L1 Subgroups in 2nd line HNSCC



Encouraging Overall Survival, Progression-Free Survival, and Duration of Response

TACTI-002/KEYNOTE-798: 2<sup>nd</sup> Line Head & Neck Squamous Cell Carcinoma (Part C)

			PD-L1	CPS <u>&gt;</u> 1
	<b>Efti + Pembro</b> Overall ITT (N=37)	<b>Efti + Pembro</b> CPS ≥20 (N=15)	Efti + Pembro CPS ≥1 (N=25)	Pembro Mono**  CPS ≥1
Overall Response Rate (ORR), %	29.7	60.0	38.5	17.3
Median Progression-Free Survival (PFS), months	2.1	13.6	2.3	2.2
6-month PFS rate, %	32.4	53.3	40.0	28.7
Median Overall Survival (OS), months	8.7*	15.5*	12.6*	8.7
12-month OS rate, %	46.0	66.7	52.0	40.0
Median Duration of Response* (DoR), months		edian follow up of 3 of Response was No	*	18.4

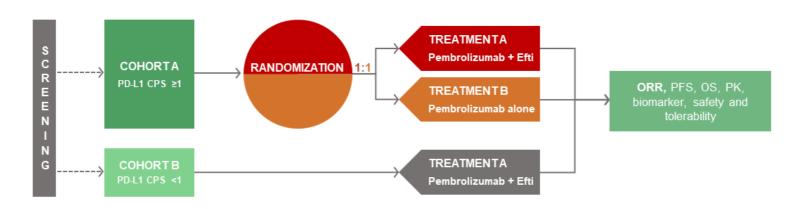
# Ongoing Phase IIb Trial in 1st Line Head & Neck Squamous Cell Carcinoma (with Fast Track Designation)



TACTI-003: Phase IIb in 1st Line Head and Neck Squamous Cell Carcinoma (1L HNSCC; app. 154 patients)

# TACTI-003 - Randomized Phase IIb Trial in 1L HNSCC patients utilizing efti + pembrolizumab versus pembrolizumab (KEYTRUDA®) monotherapy\*

- FDA Fast Track designation in 1L HNSCC on strength of the clinical results from TACTI-002 trial (Part C) in 2L HNSCC
- Clinical trial and supply agreement with Merck & Co., Inc., Kenilworth, NJ, USA (known as MSD outside the US and Canada)
- Recruiting: 75% enrolled; >25 sites activated; expect to complete enrolment by mid-2023 and have top line readout 2H of CY2023\*\*









# **Metastatic Breast Cancer**

# Efti Well Positioned to Enhance Standard-of-Care Chemotherapy in Metastatic Breast Cancer



AIPAC Phase IIb: Active Immunotherapy (Eftilagimod Alpha) and PAClitaxel (double blind, 1: 1 randomized study with 226 patients)

Efti's activation of APCs as a novel MHC Class II agonist includes significant increase in cytotoxic CD8+ T cells that can be armed with chemo-induced tumor antigens to target cancer. This synergy was demonstrated by AIPAC Phase IIb trial's encouraging results:

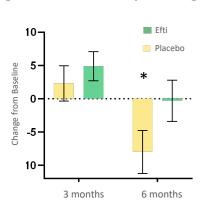
#### Positive trends in ORR, DCR and OS

	Efti + paclitaxel	Paclitaxel
Overall Response Rate	48.3%	38.4%
Disease Control Rate	85.1%	75.9%
Overall Survival	20.4 months	17.5 months

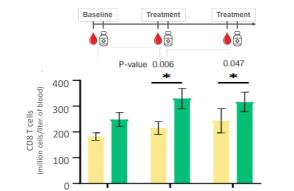
#### Significant OS improvement in three pre-specified subgroups

Pre-specified Subgroups	Median Overall Survival	Hazard Ratio	P-value
Low Monocytes	+19.6 months	HR 0.44	p=0.008
Under 65 Years	+7.5 months	HR 0.66	p=0.017
Luminal B	+4.2 months	HR 0.67	p=0.049

#### Sustained Quality of Life (QoL) vs significant decline in placebo group\*



#### **Significant increase of CD8+ T cell count**Minimal Residual Effect: samples taken just before next treatment

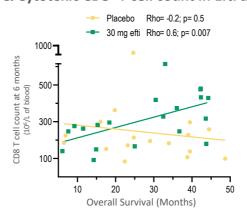


3 months

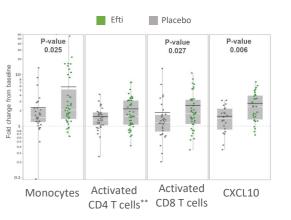
6 months

Baseline

#### Significant correlation between Overall Survival & Cytotoxic CD8<sup>+</sup> T cell count in Efti arm



#### Significant increase in anti-tumor cells and biomarkers



#### Phase II/III Trial Underway in Metastatic Breast Cancer



AIPAC-003: Active Immunotherapy (Eftilagimod Alpha) and PAClitaxel

#### AIPAC-003: Integrated Phase II/III trial in Metastatic Breast Cancer (MBC) initiated in March 2023

- Unlike previous AIPAC Phase IIb trial that administered efti and paclitaxel on different days and ceased paclitaxel at six months, AIPAC-003 patients will receive both on same day and efti + paclitaxel treatment can continue until disease progression
- Trial design provides risk-balanced approach and incorporates feedback from FDA & EMA, including expansion of HR+/HER2-neg/low MBC patient population to include triple-negative breast cancer that together account for ~78% of breast cancer cases
- First patient enrolled in May 2023\*

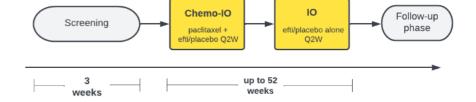
#### **Open-label lead-in component**

6 to 12 patients to test 90mg efti dosing in combination with paclitaxel driven by efti's excellent safety and FDA's Project Optimus initiative.

# Cohort 1: 80 mg/m² paclitaxel i.v. (D1,8,15) + 90 (2 x 45) mg efti s.c. (D1&15 per 4-wk cycle) Cohort 2: 80 mg/m² paclitaxel i.v. (D1,8,15) + 30 mg efti s.c. (D1&15 per 4-wk cycle) Schedule of treatments D1 D8 D15 4-week cycle Etti 30 or 90mg Paclitaxel 80 mg/m² Paclitaxel 80 mg/m² Paclitaxel 80 mg/m² Paclitaxel 80 mg/m²

→ Phase III\*\*

Randomised, double-blinded, placebo-controlled with overall survival (OS) as primary objective, which may include a specific patient population





# **Additional Oncology Indications**

# Efti + Anti-PD-L1 (Avelumab) in Urothelial Cancer & Advanced Solid Tumors





INSIGHT-004: Phase I in Various Advanced Solid Tumors & INSIGHT-005: Phase I in Metastatic Urothelial Cancer

# INSIGHT-004 – Phase I dose escalation study in advanced solid tumors\*

- Efti in combination with avelumab (BAVENCIO®) safe with promising signals of efficacy in 12 patients
- Deep & durable responses in patients with low/no PD-L1 expression and in non-immunogenic tumors
- 5/12 partial responses (42%) in different solid tumors\*\*



# INSIGHT-005 – Phase I study in metastatic urothelial cancer\*\*\*

- Investigator-initiated study evaluating safety & efficacy of efti and avelumab (BAVENCIO®) in 30 patients with metastatic urothelial cancer
- Study jointly funded by Immutep & Merck KGaA, Darmstadt, Germany
- Expansion into urothelial cancer builds on core strategy to increase target indications to exploit efti's full potential
- First patient expected to be enrolled & dosed in first half of CY2023







# Soft Tissue Sarcoma: Orphan Disease with High Unmet Need



Investigator-Initiated Trial Studying Novel Triple Combination of Efti + Radiotherapy + KEYTRUDA



EFTISARC-NEO: Open-label Triple Combination (Efti+Radiotehrapy+Anti-PD-1) Phase II trial in Soft Tissue Sarcoma (STS)



- Novel triple combination of efti with radiotherapy and anti-PD-1 therapy KEYTRUDA® (pembrolizumab)
  has potential to generate a robust anti-tumor immune response
- First time efti will be studied in neoadjuvant, non-metastatic cancer setting, which importantly will
  provide access to tumor tissue prior to and after treatment, where the impact of this novel triple
  combination on the tumor microenvironment (TME) can be assessed
- Cost-efficient Phase II study predominantly funded by an approved grant from the Polish government
- Up to 40 patients will be enrolled and dosing of first patient is anticipated in H1 of CY2023

"We are excited to begin this chemotherapy-free study combining radiotherapy with the novel immunotherapy, eftilagimod alpha, and pembrolizumab. Given efti's synergistic effects with immune checkpoint inhibitors and its ability to arm, activate, and proliferate cytotoxic T cells with radiotherapy-induced cancer antigens, this combination has a strong foundation to drive effective immunity against soft tissue sarcoma, a rare and aggressive disease in immense need of new therapeutic approaches."

- Dr. Paweł Sobczuk, Maria Skłodowska-Curie National Research Institute of Oncology



# **Preclinical Programs**

#### Novel Small Molecule Anti-LAG-3 Collaboration







Collaboration established in 2019 combining Immutep's deep LAG-3 biology experience and expertise of world leading scientists at Cardiff University

"We are delighted to collaborate with Immutep on this important project to develop a **small molecule anti-LAG-3** treatment for cancer patients that could offer the **convenience of a tablet or capsule at a fraction of the cost of existing anti-LAG-3 candidates."** 

Professor Andrew Godkin, Theme Lead in Immunology in the College of Biomedical Life Sciences, Cardiff University\*

#### Targeting the Causes of Autoimmune Diseases





#### Current Opinion in Immunology

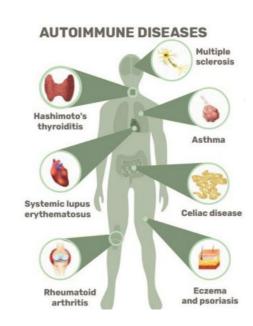
Volume 67, December 2020, Pages 1-9



#### Inhibitory receptor agonists: the future of autoimmune disease therapeutics?

Stephanie Grebinoski 12, Dario AA Vignali 1 🖂

Central and peripheral tolerance both contribute to protection against autoimmunity. The pathogenesis of autoimmunity, however, can result from critical deficits or limitations in peripheral and/or central tolerance mechanisms, presenting an opportunity for therapeutic intervention. Recent advances highlight the substantial impact of inhibitory receptors (IRs), which mediate peripheral tolerance, in autoimmunity. Deletion and blockade studies in mice, IR disruption in humans, and correlation with positive disease outcomes all highlight potential clinical benefits of enhancing IR signaling (agonism)—specifically CTLA4, PD1, LAG3, TIM3 and TIGIT—to treat autoimmune disease. Although critical questions remain, IR agonists represent an unappreciated and untapped opportunity for the treatment of autoimmune and inflammatory diseases.



#### **Present Approaches Target Symptoms** of Autoimmune Diseases

Corticoids, methotrexate, TNF & interleukin inhibitors (anti-TNF-α,-IL-6, -IL-17, -IL-23 mAbs)



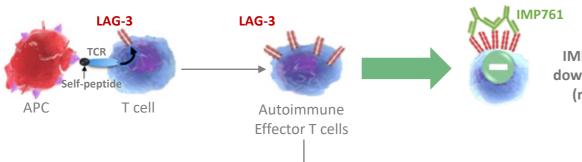
#### **Future Approaches Target Causes** of Autoimmune Diseases

Targeting autoimmune memory T cells with LAG-3 antibodies

# IMP761: First-in-Class LAG-3 Agonist is Potential Game-Changer

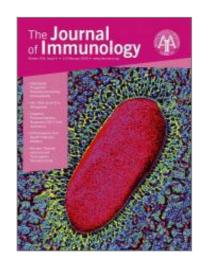


As the world's first immunosuppressive agonist antibody to LAG-3 acting upstream on activated T cells, IMP761 targets the root cause of many autoimmune diseases and represents a potential game-changer in the treatment landscape. Initiating IND-enabling studies in 1H'2023.



Epigenetic reprogramming leads to T cell helper (Th) induced AI diseases: Th1 (e.g., Rheumatoid Arthritis), Th2 (e.g., Allergic Asthma), Th17 (e.g., IBS), etc.

IMP761 increases the natural LAG-3-mediated down-regulation of auto-reactive memory T cells (root cause of many autoimmune diseases)



A LAG-3-Specific Agonist Antibody for the Treatment of T Cell-Induced Autoimmune Diseases\*

*IMP761* significantly inhibits T cell infiltration of an antigen-specific intradermal reaction in vivo in an Aqspecific delayed-type hypersensitivity (DTH) model in non-human primate study.



Juvenile idiopathic arthritis: LAG-3 is a central immune receptor in children with oligoarticular subtypes\*\*

Pre-clinical testing of IMP761 in oligoarticular juvenile idiopathic arthritis model showed decreased secretion of mostly all measured cytokines (IL-10, IL-12, IL-16, IL-4, IL-6 = p-value < 0.01)

#### **Board and Management**





Dr Russel Howard
Non-Executive Chairman

Dr Howard has over 45 years' experience in Australian & US biotech sectors, including the Walter & Eliza Hall Institute, Schering-Plough, GSK and as Maxygen CEO. He currently is Chairman of NeuClone Pty Ltd and a former Director of Circadian Technologies Ltd.



Florian Vogl, MD, PhD
Chief Medical Officer

Dr Vogl is a board-certified MD and has over 13 years in the biopharmaceutical industry with extensive clinical development expertise in the field of oncology in the Europe and the US through roles at Cellestia Biotech, Rainier Therapeutics, Novartis and Amgen.



Pete Meyers
Deputy Chairman

Mr Meyers spent 18 years in health care investment banking before taking on CFO roles in biotechnology, including Eagle Pharmaceuticals, Inc, TetraLogic Pharmaceuticals Corp, and Motif BioSciences Inc. Based in New York, he is currently CFO of Slayback Pharma.



Christian Mueller VP, Strategic Development

Mr Mueller has +10 years of clinical development experience in oncology, including at Medical Enzymes AG focusing on therapeutic enzymes for cancer treatment and at Ganymed Pharmaceuticals AG developing ideal mAbs in immune oncology.



Lis Boyce Non-Executive Director

Lis Boyce has over 30 years' experience as a corporate lawyer and is a partner at Piper Alderman. She has a strong focus on Life Sciences and Healthcare, and is deputy chair of AusBiotech's AusMedtech Advisory Group, as well as a member of AusBiotech's State Committee for NSW.



Claudia Jacoby, PhD
Director of Manufacturing

Dr Jacoby has +15 years of biotech industry experience with extensive skills in protein expression and purification as well as in analytical and preclinical development from her various positions at pre-clinical and clinical-stage pharmaceutical companies.



Marc Voigt
Executive Director & CEO

Mr Voigt has over 25 years of experience in the corporate and biotechnology sectors, including Deutsche Life Science, Revotar Biopharmaceuticals AG, Medical Enzymes AG and Allianz. He was appointed as CEO and Executive Director in July 2014. Mr Voigt is based in Berlin.



James Flinn, PhD
IP & Innovation Director

Dr Flinn is an Australian Patent Attorney with +20 years of professional experience in building and managing IP portfolios, including GSK, two US-based pharmaceutical companies, a major Australian retailer, and a Melbourne Patent Attorney firm.



Prof. Frédéric Triebel, MD, PhD
Executive Director, CSO

Prof Triebel discovered the LAG-3 gene while working at the Gustave Roussy Institute and is a pioneer in the LAG-3 field of immuno-oncology. He was the founder of Immutep S.A., which was acquired by the Company in 2014. Based in Paris, he holds a Ph.D. in immunology (Paris University).



Deanne Miller
COO, General Counsel

Ms Miller has broad commercial experience having held legal, investment banking, regulatory compliance and tax advisory positions at RBC Investor Services, Westpac Group, Macquarie Group, the Australian Securities and Investment Commission, and KPMG.



David Fang Finance Director

Joining Immutep in 2018, Mr Fang has over 12 years of accounting and auditing experience across various industries including biotechnology, manufacturing and healthcare including Group Finance Manager of Kazia Therapeutics Limited and auditor at PWC.



Chrystelle Brignone, PhD
Preclinical Development Director

Dr Brignone joined Immutep in 2004 and has more than 20 years' experience in the field of Immunology and Immune monitoring of clinical studies. As Principal Scientist since 2014, she is leading the R&D in the Immutep laboratory in France.

#### Significant Milestones Ahead in 2023



- ✓ Initiated AIPAC-003 PII/PIII trial of efti + chemo in MBC/TNBC
- ✓ Initiated cost-efficient investigator-led PII study in soft tissue sarcoma
- ✓ Final data from TACTI-002 (Part B) in 2nd line anti-PD-(L)1 refractory NSCLC
- ✓ Final data from TACTI-002 (Part C) in 2nd line HNSCC
- ✓ Received regulatory approval for initiation of jointlyfunded INSIGHT-005 with Merck KGaA, Darmstadt, Germany

- Data updates from TACTI-002 Phase II trial in 1st line NSCLC
- Complete enrolment (75% enrolled\*) and top-line readout from randomised TACTI-003 Phase IIb trial
- Data updates from triple combination INSIGHT-003 PI trial with efti + anti-PD-1 + chemotherapy in 1st line NSCLC
- Updates from INSIGHT-005 and STS studies
- IND-enabling studies of IMP761
- Updates from partnered programs
- Updates regarding expansion of clinical trial pipeline,
   e.g. Phase III trial planning in 1L NSCLC



Thank You