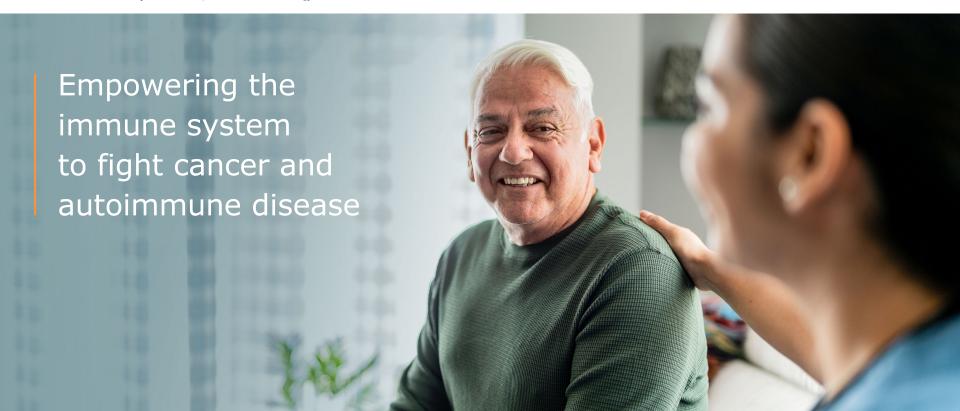
## **Immutep Corporate Presentation**



October 2025 (IMM:ASX, IMMP:NASDAQ)



## Forward Looking Statements



The purpose of the presentation is to provide an update of the business of Immutep Limited ACN 009 237 889 (ASX:IMM; NASDAQ:IMMP). These slides have been prepared as a presentation aid only and the information they contain may require further explanation and/or clarification. Accordingly, these slides and the information they contain should be read in conjunction with past and future announcements made by Immutep and should not be relied upon as an independent source of information. Please refer to the Company's website and/or the Company's filings to the ASX and SEC for further information.

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## Immutep Highlights



# A leader in LAG-3 immunotherapy

Four clinical-stage assets designed to safely empower patients' immune systems to fight cancer and autoimmune diseases through the MHC Class II & LAG-3 pathways, including first-in-class immunotherapies eftilagimod alfa (efti) and IMP761.

# Phase 3 in 1L NSCLC: Blockbuster potential

Registrational Phase III in collaboration with MSD (Merck & Co.) with potential to establish new standard-of-care in first line non-small cell lung cancer (1L NSCLC), one of the largest oncology markets expected to reach US\$48 billion in sales in 2031.

# Validation via collaborations

Multiple partnerships and collaborations with large pharma and leading institutions.















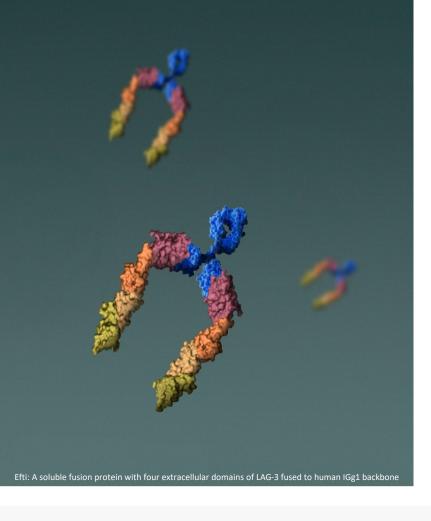
## Strong IP and balance sheet

Strong intellectual property (IP) portfolio and 12+ years of potential exclusivity for biologics like efti & IMP761. Cash & cash equivalents of ~A\$109.85 million provide runway to end of CY2026.#

## Deep Clinical Pipeline in Oncology & Autoimmune Diseases



ONCOLOGY		Preclinical	Phase I	Phase II	Phase III	Collaborations	Commercial Rights
Eftilagimod alfa MHC Class II agonist	Lung Cancer  Lung and Head & Neck Cancers  Lung Cancer <sup>1</sup>	TACTI-004: Efti + Pembroliz  TACTI-002: Efti + Pembroliz  INSIGHT-003: Efti + Pembro	umab			MSD MSD IKF	0
	Head & Neck Cancer  Metastatic Breast Cancer  Early-Stage Breast Cancer <sup>1</sup>	TACTI-003; Efti + Pembrolizumab  AIPAC-003; Efti + Paclitaxel  Neoadjuvant Efti + Chemotherapy				MSD MSD Cancer Center	Empowering the Immune System P Global Rights ex-China
	Soft Tissue Sarcoma <sup>1</sup> Urothelial Cancer <sup>1</sup>	EFTISARC-NEO: Neoadjuvar INSIGHT-005: Efti + Avelum	nt Efti + Pembrolizumab + Rad ab	iotherapy	ı	Narodeny Intervention Intervent	
AUTOIMMUNE DISEASE							
IMP761 (LAG-3 Agonist mAb)  IMP731 (Depleting LAG-3 mAb)	Healthy Volunteers  Psoriasis & Ulcerative Colitis <sup>2</sup>	IMP761 IMP731					immutep Empowering the Immune System
OUTLICENSED - Oncology							
Eftilagimod alfa	Metastatic Breast Cancer (China) <sup>3</sup>					<b></b> ♦EDC	<b>♦ □ □ Efti China Rights</b>
LAG525 (Anti-LAG-3 mAb)	Solid Tumours, Blood Cancers, TNBC, Melanoma <sup>4</sup>					<b>U</b> NOVARTIS	NOVARTIS





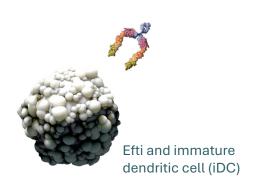
# Oncology

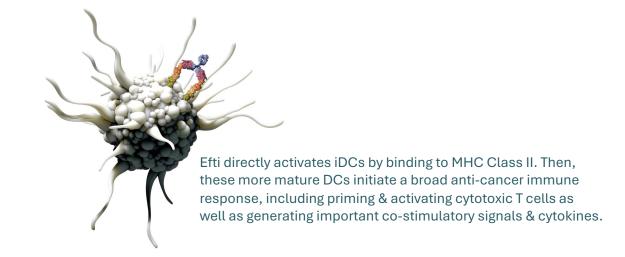
### **Eftilagimod Alfa (Efti):**

Only immunotherapy in clinical development that activates antigen-presenting cells or APCs (e.g. dendritic cells, monocytes) directly via the MHC Class II pathway to fight cancer

## Activating a Broad Anti-Cancer Immune Response with Efti







#### **COMPLEMENTARY EFFECT**

Efti initiates a complementary immune response to other cancer therapies incl. immune checkpoint inhibitors (ICIs)

#### **FAVORABLE SAFETY**

Efti combined w/ other cancer therapies (e.g. ICIs, chemo, etc.) has a safety profile in line with underlying therapy

#### **EXPANDS/ENHANCES EFFICACY**

Efti expands responders & enhances efficacy in combination with other therapies including ICIs like KEYTRUDA®

## Efti's Key Attributes and Encouraging KOL Feedback







#### **Efti Key Attributes**

- Real innovation Efti is a first-in-class asset unlike any in the immunotherapy landscape
- Pipeline in a product Can revolutionize treatment landscape for many solid tumours
- Low cost of goods Allows for reasonable pricing with strong margins
- Excellent safety profile Both as monotherapy and in multiple combination settings
- Subcutaneous administration Convenient and easy to administer

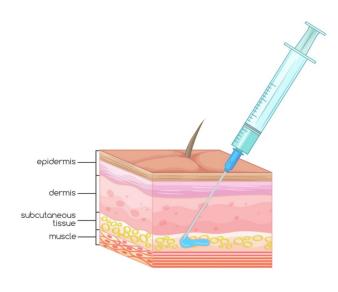
#### **KOL Feedback on Efti & 1L NSCLC Phase III**<sup>1,2,3</sup>

- Robust fundament in NSCLC Positive on efti driving higher responses & survival in previous 1L NSCLC trials
- Easy to administer and safe Do not need see high toxicity associated with other therapies
- Add-on strategy Simple add-on to standard-of-care therapy; no change to current practice
- ➤ **Easy to enrol** All-comer PD-L1 trial design allows for easy enrolment with no PD-L1 sub-group exclusions
- > **Truly first-in-class** Efti is not a "me too product" that are often seen in combination trials

## Advantages of Subcutaneous Delivery



Subcutaneous administration of efti positions it well in current & future cancer treatment landscape



#### **Efti's subcutaneous delivery:**

- Generates systemic anti-cancer immune response
- Improves patient experience vs. intravenous (IV) administration used with other therapeutics:
  - ✓ Less invasive and easier to administer
  - ✓ More flexible leading to increased patient treatment access
- Positions efti well as a combination therapy especially with anti-PD-(L)1 therapies administered either via IV or subcutaneously:
  - ✓ Move underway to subcutaneous administration among top-selling anti-PD-(L)1 therapies.¹ By 2028, MSD & Bristol Myers estimate ~30-40% KEYTRUDA & OPDIVO sales (~\$12-16 billion in sales) will shift from IV to subcutaneous delivery.²,³,⁴





## Immutep's Key Value Driver

# First Line Non-Small Cell Lung Cancer (1L NSCLC)

- Lung cancer is the leading cause of cancer death and non-small cell lung cancer (NSCLC) comprises 80 to 85% of all lung cancers<sup>1,2</sup>
- ~2.0 million NSCLC diagnoses annually
- Total addressable NSCLC drug market expected to reach US\$48 billion in 2031 with over 50% sales from immune checkpoint inhibitors including anti-PD-1<sup>3</sup>

## High Unmet Need in 1L NSCLC



Need for IO-combination therapies that maximize the immune system's ability to fight cancer

## **Evolution of Immunotherapy (IO) in 1L NSCLC**

Immune Checkpoint Inhibitors (ICI) alone

ICIs as monotherapies drive responses in up to 20% of 1L NSCLC patients **ICIs + Chemotherapy** 

ICIs + chemo expands responses yet room for much improvement and Overall Survival still below 2 years for most patients Next-generation IO + ICIs + Chemotherapy

Next-gen IO approaches like **efti** may further expand 1L NSCLC patients who respond to ICIs & extend patients' survival

## TACTI-002/KEYNOTE-798: Strong Efficacy with Efti + KEYTRUDA



TACTI-002 (Part A) Phase II: Overview & Trial Design

Overview Screening Indication: 1L NSCLC Advanced/metastatic Status: Completed

Locations: 18 centers across

six countries

- Non-squamous/squamous Enrolment: 114 patients 0-100% PD-L1 expression
  - · EGFR/ALK negative

**Treatment Phase** 

30mg efti Q2W + 200 mg pembrolizumab Q3W for 8 cycles, followed by 30mg efti + 200mg pembrolizumab for 9 cycles Up to 52 weeks

Combination Therapy

Monotherapy 200 mg pembrolizumab Q3W for 16 cycles Up to 52 weeks

Primary: ORR (iRECIST) Secondary: ORR (RECIST 1.1), PFS, OS, DOR, Safety, PK/PD

Follow Up

In collaboration with



#### Tumour Response by PD-L1 Expression Level<sup>1</sup>

	TPS <1% N=32	TPS 1-49% N=38	TPS ≥50% <sub>N=20</sub>
ORR <sup>1,2,3,4</sup>	31.3%	44.7%	55.0%
mPFS, months <sup>1,2</sup>	4.2	9.3	16.5
mOS, months	15.5	23.4	Not Reached

- Results offer strong evidence of efti's complementary effect with PD-1 inhibitors and its unique stimulation of patients' anti-cancer immune response
- Efficacy across all PD-L1 levels differentiates efti and anti-PD-1 from other chemo-free combinations
- ORR & PFS translate into excellent Overall Survival (OS)

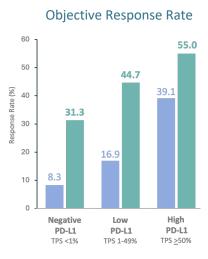


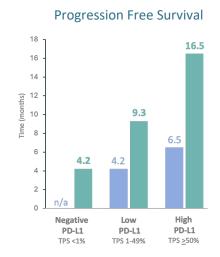


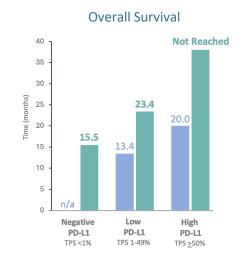


## TACTI-002: Benchmarking Efti + KEYTRUDA to KEYTRUDA Alone









Compelling efficacy from efti + KEYTRUDA across all PD-L1 levels including 1L NSCLC patients with negative & low PD-L1, for whom KEYTRUDA monotherapy works sub-optimally\*

\* KEYTRUDA monotherapy in 1L NSCLC was not approved by EMA in TPS 1-49% and was not tested after KEYNOTE-001 in TPS <1%<sup>3</sup>

Efti + KEYTRUDA<sup>1</sup> KEYTRUDA monotherapy<sup>2</sup>

## TACTI-002: Favourable Safety Profile



Differentiated overall survival from Efti + KEYTRUDA achieved without foregoing safety

Efti + KEYTRUDA and Standard-of-Care Therapies in 1L NSCLC TPS <u>&gt;</u> 1%	Drug-related Adverse Events Leading to Discontinuation <sup>2</sup>	Median Overall Survival <sup>3</sup>
Efti + KEYTRUDA (pembrolizumab)	9.6%	35.5 months
Pembrolizumab monotherapy <sup>1</sup>	9.9%	16.4 months
Pembrolizumab + Doublet Chemo (SQ)	16.8%	18.9 months
Ipilimumab + Nivolumab¹	18.1%	17.1 months
Pembrolizumab + Doublet Chemo (NSQ)	20.5%	23.3 months
Ipi + Nivo + 2 cycles of Doublet Chemo	22.1%	15.8 months

NSQ = Non-squamous; SQ = Squamous

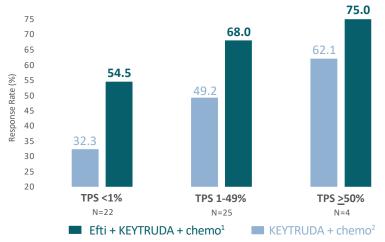
## INSIGHT-003: High Response Rates Across All PD-L1 Levels



INSIGHT-003 (Stratum C) Phase I: Overview & Trial Design

motori oos (stratam e) i nase n ov	CIVICW & ITIAI DESIGN								
Overview	Screening	Treatment Phase		Treatment Phase		Treatment Phase		Follow Up	
Indication: 1L NSCLC     Status: Ongoing     Recruitment: Completed     Enrolled: ~50 pts     Locations: Multi-center study in Germany	Advanced/metastatic Non-squamous O-100% PD-L1 expression EGFR/ALK negative	Induction 30mg efti Q2W + 200 mg pembrolizumab + platinum doublet chemo Up to 24 weeks	Maintenance 30mg efti Q2W/Q3W* + 200 mg pembrolizumab + platinum doublet chemo Max. total study treatment: 52 weeks	ORR, PFS, DOR, OS, and Safety	INSIGHT-003 is an investigator-initiated, multi-centre Phase I trial led by Frankfurt Institute of Clinical Cancer Research (IKF)				

# Benchmarking **Efti + KEYTRUDA + Chemo**Objective Response Rate to **KEYTRUDA + Chemo**



- Promising efficacy from efti + KEYTRUDA + chemo in non-squamous 1L NSCLC across all PD-L1 levels
- 61.7% ORR in TPS <50% (N=47) compares to 40.8% from KN-189. These patients with TPS <1% and TPS 1-49% have unmet need and represent ~70% of 1L NSCLC population.
- Combination with efti is feasible and safe to administer, and addition of efti does not increase toxicity of standard-of-care KEYTRUDA + chemo<sup>3</sup>

## INSIGHT-003 Case Study: Patient with PD-L1 TPS 0%



# Case study of a 75-year-old male with partial response (PR):

- PD-L1 TPS 0%, no actionable genetic alterations
- TTF1 pos. Adeno-Carcinoma, G3
- FCOG PS 1
- cT1c pN2 cM1c, Stage IVb
- Partial Response achieved after treatment cycle 3; maintained until planned end of treatment (week 52)

Baseline September 2024



TL2 Pleura: 24 mm

After 1 Year of Therapy September 2025



TL1 Lung: 7 mm

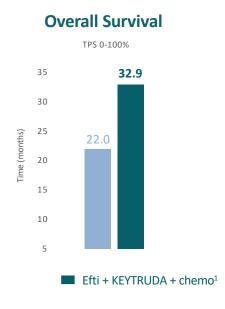


TL2 Pleura: 0 mm

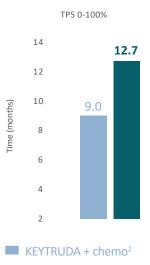
#### INSIGHT-003: Excellent PFS & Overall Survival



Complementary effect from the combination of Efti + KEYTRUDA + Chemo



#### **Progression Free Survival**



- Excellent OS & PFS in patients with a minimum follow-up of 22 months (N=21), well above historical controls
- Relative OS/PFS outperformance despite only ~19% patients having high PD-L1 TPS ≥50% who typically respond best to anti-PD- therapy vs. ~32% patients in KN-189 trial

## Efti with KEYTRUDA Expands ORR & Enhances PFS/OS



Efti + KEYTRUDA without chemo and with chemo has safely led to compelling ORR, PFS, and OS well above KEYTRUDA +/- chemo in two 1L NSCLC trials that together have enrolled >165 patients

TACTI-002 (N=114)	TPS <1%	TPS 1-49%	TPS <u>&gt;</u> 50%
Efti + KEYTRUDA ORR (%)	31.3%	44.7%	55.0%
KEYTRUDA alone	8.3%	16.9%	39.1%
Efti + KEYTRUDA PFS (mos)	4.2	9.3	16.5
KEYTRUDA alone	n/a	4.2	6.5
Efti + KEYTRUDA OS (mos)	15.5	23.4	NR
KEYTRUDA alone	n/a	13.4	20.0

INSIGHT-003 (N=51)	TPS <1%	TPS 1-49%	TPS <u>&gt;</u> 50%		
Efti + KEYTRUDA + Chemo ORR (%)	54.5%	68.0%	75.0%		
KEYTRUDA + Chemo	32.3% 49.2% 62.				
Efti + KEYTRUDA + Chemo PFS (mos)	12.7 —				
KEYTRUDA + Chemo		— 9.0 —			
Efti + KEYTRUDA + Chemo OS (mos)		<b>32.9</b>			
KEYTRUDA + Chemo		— <i>22.0</i> —			

Addition of chemo enhances strong efficacy achieved in TACTI-002 with Efti + KEYTRUDA (N=114) in 1L NSCLC

## TACTI-004/KEYNOTE-F91 Phase III Trial in 1L NSCLC



#### TACTI-004 / KEYNOTE-F91 Phase III: Overview & Trial Design

TACTION / KETHOTE 1321 Hase III.	overview & Trial Design				
Overview	Screening		Treatment Phase	Follow Up	In collaboration with
<ul> <li>Indication: 1L NSCLC</li> <li>Status: Ongoing</li> <li>Recruitment: Ongoing</li> <li>Enrolment target: ~756 pts</li> <li>Locations: +150 sites in +25 countries globally</li> </ul>	Advanced/metastatic     Non-squamous/Squamous     0-100% PD-L1 expression     EGFR/ALK/ROS negative     ECOG 0-1	Randomisation 1:1	Treatment Arm  Efti - Q2W for 6 months; Q3W thereafter Pembrolizumab - Q3W Platinum doublet chemo  Control Arm Placebo - Q2W for 6 months; Q3W thereafter Pembrolizumab - Q3W Platinum doublet chemo	Dual primary endpoints: OS & PFS	MSD MSD

- Third collaboration & supply agreement with MSD who jointly designed registrational TACTI-004 trial with Immutep announced in June 2024.
- 1:1 randomized, double-blind, global trial enrolling ~756 patients in +150 sites across +25 countries with PFS & OS dual primary endpoints.
- Immutep conducting trial & MSD supplying KEYTRUDA (typical ICI supply for trial this size is ~US\$100mm). Immutep retains efti's commercial rights with freedom to operate.
- FDA and other regulatory agencies<sup>1</sup> provided positive feedback on study design.
- +100 clinical sites across 24 countries have been activated and over 170 patients enrolled/randomised as of October 2025.<sup>2</sup>

## TACTI-004 Milestones and Multiple Potential Paths to Success



#### TACTI-004/KN-F91 Milestones

1 <sup>st</sup> Patient in: 25 March 2025				tility alysis			<b>←</b>	Interim Analysis: PFS* -	-		
<b>V</b>	Q2′25	Q3′25	YE'25	Q1′26	Q2′26	Q3′26	YE'26	Q1′27	Q2′27		
•							* Potential filing if positive				

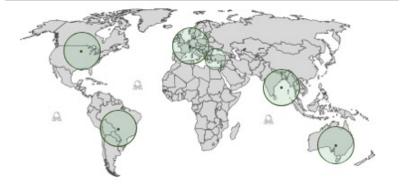
#### **Dual Primary Endpoints:**

- Progression-free Survival
- Overall Survival
  - Multiple pre-specified analyses are planned for these dual endpoints, each of which could potentially lead to a BLA and/or MMA filing opportunity

#### **Patient Stratification:**

 Important prognostics and predictive markers are used for stratification (e.g. PD-L1 levels and tumour subtypes)

#### TACTI-004 is a global trial with sites in North & South America, Europe, APAC



## Global Phase III Landscape for Advanced/Metastatic 1L NSCLC



Companies that have a CTCSA with MSD for Phase III trials in advanced/metastatic 1L NSCLC <sup>1</sup>	No PD-L1 TPS <1%	Low PD-L1 TPS 1-49%	High PD-L1 TPS <u>&gt;</u> 50%	Non- Squamous	Squamous	% of 1L NSCLC Population
% of 1L NSCLC patient population by segment $^2$ $\Rightarrow$	~35%	~35%	~30%	~70%	~30%	→ Up to 100%
Immutep TACTI-004/KEYNOTE-F91 (Efti + KEYTRUDA + chemo)	<b>~</b>	<b>~</b>	<b>~</b>	<b>~</b>	<b>~</b>	100%
Daiichi Sankyo TROPION-Lung07 (DatoDXd + KEYTRUDA)	<b>~</b>	<b>~</b>	×	<b>~</b>	×	49%
Gilead EVOKE-03 (Sacituzumab Govitecan + KEYTRUDA)	×	×	<b>~</b>	<b>~</b>	<b>~</b>	30%
Daiichi Sankyo TROPION-Lung08 (DatoDXd + KEYTRUDA)	×	×	<b>~</b>	<b>~</b>	×	21%

#### **Key aspects of TACTI-004/KEYNOTE-F91:**

- Efti a simple add-on to KEYTRUDA & chemo, the dominant standard-of-care most often chosen in 1L NSCLC
- Addressing entire PD-L1 population (TPS 0-100%) and both non-squamous/squamous patient populations
- Initial primary read-out in late 2026 thru mid-2027 followed by additional pre-specified analyses<sup>3</sup>
- Strong ORR & PFS in efti's prior 1L NSCLC trials translate to compelling Overall Survival

#### Other approaches in Phase III trials:

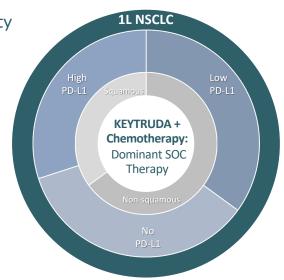
- ADCs: Often have high toxicity profiles, particularly when used in combination, that may limit broad usage in 1L NSCLC
- PD-1/VEGF: Strong ORR & PFS; unknowns to date include durability of effect and translation to Overall Survival
- TIGIT: Trial failures (e.g. SKYSCRAPER-01/06, ARC-10) & program terminations (e.g. MSD) suggest limited impact<sup>4</sup>
- Anti-LAG-3: Program terminations (e.g. MSD) and a Phase III for only TPS 1-49/NSQ suggest limited impact<sup>4,5,6</sup>

## Potential Blockbuster Commercial Opportunity



If TACTI-004 is successful it presents a potential multi-billion US\$ opportunity for efti as it will be a safe add-on to KEYTRUDA & chemo in 1L NSCLC:

- KEYTRUDA has revolutionized treatment landscape and MSD captures
  7 to 8 of every 10 metastatic lung cancer patients.<sup>1</sup> Estimates are ~27%
  of KEYTRUDA's \$29.5 billion in 2024 sales are from lung cancer.<sup>2</sup>
- Potential peak sales for efti can be reached faster vs. a typical drug launch given KEYTRUDA + chemo is the standard-of-care therapy most often used in 1L NSCLC.
- 1L NSCLC may be first of many indications that efti can improve efficacy when combined with KEYTRUDA (over 40 approvals in 18 cancer types).



#### Efti's Potential to Extend IP for PD-1 Inhibitors



BAVENCIO® JEMPERLI®

2035

LOE

2034

LOE

- Efti's comprehensive patent portfolio provides an opportunity to enhance and substantially extend established or new PD-(L)1 franchises
- Two leading PD-1 inhibitors, KEYTRUDA & OPDIVO (over \$38 billion in 2024 sales), face key patent expirations and loss of exclusivity (LOE) in 2028

# 9.3

IMFINZI° TECENTRIQ° LIBTAYO°

2030

LOE

2035

LOF

2024 Sales of Anti-PD-(L)1 Therapies (\$ Billions)

**KEYTRUDA®** 

2028

LOE

OPDIVO'

2028

LOF

2031

LOE





## Additional Oncology Indications

Efti's favorable safety profile & unique activation of immune system allows for multiple combinations in various cancers

- Metastatic Settings:
  - Head and Neck Cancer: Efti + Anti-PD-1
  - Breast Cancer: Efti + Chemotherapy
  - Urothelial Carcinoma: Efti + Anti-PD-L1
- Neoadjuvant Settings (Prior to Surgery):
  - Soft Tissue Sarcoma: Efti + Anti-PD-1 + Radiotherapy
  - Breast Cancer: Efti + Chemotherapy

## TACTI-003: Strong Chemo-Free Results in 1L HNSCC w/ CPS <1



TACTI-003 / KEYNOTE-C34 Cohort B Phase IIb: Overview & Trial Design

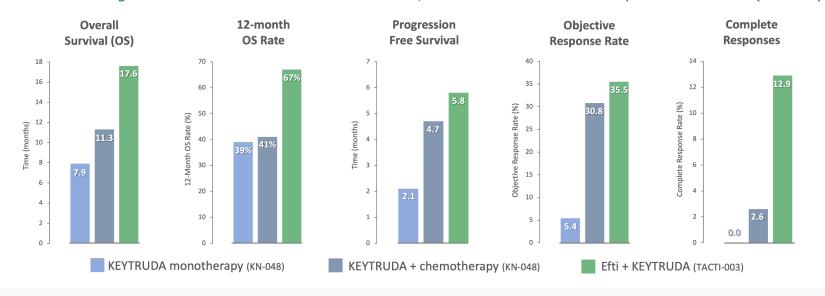
larynx

	Overview	Screening	Treatment Phase	Follow Up	
•	Indication: 1L HNSCC Status: Ongoing Recruitment: Completed / 33 pts (31 eval)	<ul> <li>Recurrent/metastatic</li> <li>PD-L1 below 1 (CPS &lt;1)</li> <li>ECOG 0-1</li> </ul>	<u>Efti + Pembrolizumab</u> 30mg efti Q2W subcutaneous +	ORR, PFS, OS, PK, Biomarker, Tolerability,	
	Landiana, Milatana, and an annual and an annual and an	<ul> <li>Oral oronharuny hypopharuny</li> </ul>	400 mg namhralinumah OGW intravanava	and Safety	

In collaboration with

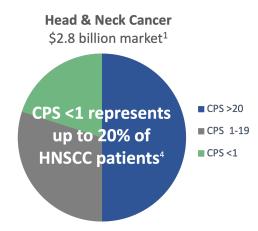


#### Benchmarking to Efti + KEYTRUDA to KEYTRUDA with/without chemo in PD-L1 expression below 1 (CPS <1)

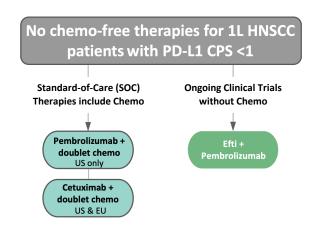


## Limited Competition in 1L HNSCC CPS <1: A Valuable Market





>890,000 HNSCC diagnoses per annum worldwide with ~100,000 patients who develop metastatic disease. 1,2,3



Efti + KEYTRUDA shows superior OS, PFS, and durability with less toxicity as compared to SOC therapies & generates high response rates.

#### **Next Steps in 1L HNSCC CPS <1**

In August 2025, Immutep announced positive FDA feedback on late-stage clinical development of efti in 1L HNSCC patients with CPS <1.

Potential paths for collaborative clinical development and accelerated approval in 1L HNSCC CPS <1 in light of FDA's Project FrontRunner include:

- A randomised trial evaluating efti and KEYTRUDA against standard-of-care therapy
- A single-arm study (e.g. 70-90 patients) with response rates and duration of response as key endpoints, followed by a confirmatory randomized study

## Primary Endpoint Met in Phase II Trial in Soft Tissue Sarcoma



#### EFTISARC-NEO Phase II: Overview & Trial Design

Overview	Screening	Treatment Phase	>	Follow Up	Onkologii im. Marii Sktodowskiel-Curie
<ul> <li>Indication: Soft Tissue Sarcoma</li> <li>Status: Ongoing</li> <li>Recruitment: Completed</li> <li>Enrolled: 40 pts</li> <li>Location: Poland's national reference centre</li> </ul>	Primary or locally recurrent STS     Grade 2 or 3; primary tumour size     Scm or locally recurrent of any     size     Measurable disease (RECIST1.1)	Efti + Pembrolizumab + Radiotherapy  30mg efti Q2W (W1, W3, W5, W7, W9)  200 mg pembrolizumab Q3W (W1, W4, W7)  Radiotherapy (50 Gy) for 5 weeks between W2 & W7	Surger	Primary Endpoint: tumour hyalinization Secondary Endpoints: DFS, LRFS, DMFS, OS & Safety*	Pathtenowy Institute Badanczy   EFTISARC-NEO is an investigator-initiated Phase II trial conducted at Poland's national reference centre for sarcoma, the Maria Skłodowska-Curie National Research Institute of Oncology (MSCNRIO)

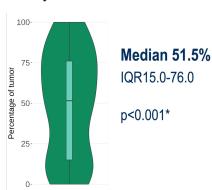
#### ESMO Congress 2025 - Proffered Paper Oral Presentation

Neoadjuvant efti + KEYTRUDA + radiotherapy met primary endpoint with median 51.5% tumour hyalinization/fibrosis (p<0.001)¹ in patients with soft tissue sarcoma (STS)

Results over 3-fold higher than median 15% from standard-of-care radiotherapy based on historical data<sup>2</sup>

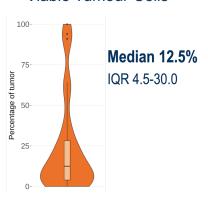
Tumour hyalinization/fibrosis may serve as early surrogate endpoint correlated with enhanced overall and recurrence-free survival in STS<sup>3</sup>

#### Hyalinization/Fibrosis



#### Viable Tumour Cells

Narodowy



#### AIPAC-003 Trial in Metastatic Breast Cancer



#### AIPAC-003: Overview & Trial Design

#### Overview

- · Indication: Metastatic Breast Cancer
- Status: Activ
- Recruitment: Phase II complete
- · Enrolled: 65 pts in PII; 6 in lead in
- Locations: 22 centers across five countries

#### Screening

- HR+/HER2-neg
- Triple negative
- Progressed on or after ≥1 line of endocrine based therapy and indicated to receive chemo

#### Treatment Phase (Dose Optimization)

#### Lead in Component

6 pts received 90mg efti + paclitaxel

#### Phase

~58 evaluable pts randomised for either 30mg or 90mg efti + paclitaxel



- HR+/- HER2-negative/low and triple negative metastatic breast cancer (MBC) patients represent ~78% breast cancer cases¹
- Patients receive efti + paclitaxel on same day; IO-chemo treatment can continue until disease progression
- Trial design incorporates feedback from FDA & EMA and provides risk-balanced approach
- Randomised Phase II dose optimization to find optimal biological efti dosing (e.g. 30mg or 90mg). 30mg dosing has been selected as optimal biological dose.
- Data update at medical conference in 2H'CY2025

## Neoadjuvant Efti in HR+/HER2-neg Breast Cancer Phase II Trial



Phase II: Overview & Trial Design

#### Overview Screening **Treatment Phase** Follow Up Indication: Breast Cancer, Stages 1-3 HR+/HER2-neg Neoadjuvant Efti + Chemotherapy Primary: Pathological Cancer Center Stages 1-3 and candidate for Status: Active Complete Response (pCR) Recruitment: Not yet recruiting neoadiuvant chemo (NAC) 30mg efti subcutaneous as monotherapy once a week for Secondary: Response rates 3 weeks and then in combination with NAC every 2 weeks Enrolment target: 50 pts Simon's two-stage design: 30 pts & tumour's Ki67 levels starting at week 4 in stage one; 20 pts in stage two Locations: Multi-centre study. United States



- George Washington (GW) Cancer Center initiating Phase II evaluating neoadjuvant efti as monotherapy and with chemo prior to surgery in HR+/HER2-neg breast cancer patients
- Study primarily funded by grants and GW Cancer Center
- Second investigator-initiated trial (IIT) to evaluate neoadjuvant efti in earlier stage disease where its unique activation of anti-cancer immune response may drive optimal benefit
  - Neoadjuvant immuno-oncology increasingly validated setting, leveraging intact tumor to prime durable immune responses and treat potential micro-metastases early
  - Neoadjuvant setting significantly increases the addressable patient population

## Efti + Anti-PD-L1 (Avelumab) in Metastatic Urothelial Cancer



INSIGHT-005 (Stratum E) Phase I: Overview & Trial Design

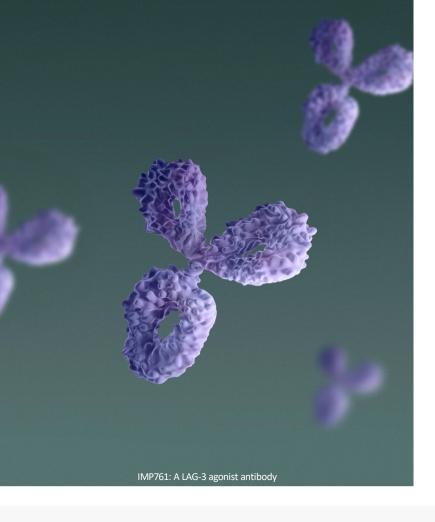
Overview	Screening	Treatment Phase	Follow Up	ik E
<ul> <li>Indication: Urothelial Cancer</li> <li>Status: Active</li> <li>Recruitment: Ongoing</li> <li>Enrolment target: 30 pts</li> <li>Locations: Multi-center study in Germany</li> </ul>	Metastatic     0-100% PD-L1 expression     Enrolment in 3 subgroups	30mg efti s.c. + 800mg avelumab i.v. on same day Q2W for a maximum of 24 cycles	Feasibility, Safety, ORR, DOR, PFS, and OS	INSIGHT-003 is an investigator-initiated, multi-centre Phase I trial led by Frankfurt Institute of Clinical Cancer Research (IKF)

In collaboration with

Merck KGaA

Darmstadt, Germany

- INSIGHT-005 evaluating safety & efficacy of efti and avelumab (BAVENCIO®), which has previously shown promising efficacy in solid tumours in Phase I trial
- Jointly funded by Immutep & Merck KGaA, Darmstadt, Germany
- Targeting area of high unmet need including patients with more metastatic cases
- Announced first patient enrolled and safely dosed in Jan 2024





## **Autoimmune Diseases**

#### **IMP761:**

First-in-class immunotherapy is a LAG-3 agonist antibody designed to target the root cause of many autoimmune disorders

# Targeting Autoimmune Diseases with a LAG-3 Checkpoint Agonist immutep



New paradigm to treat the cause - as opposed to the symptoms - of autoimmune disorders



"Findings further support the potential clinical benefits of a LAG-3 agonist in the treatment of human autoimmunity"1

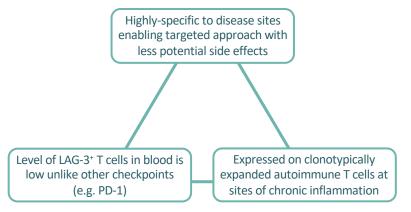


"LAG-3 agonism could be a potential target for future treatment in rheumatoid arthritis"2



"Manipulation of the **LAG-3 pathway** can serve as a promising therapeutic strategy"3

#### Unique advantages of LAG-3 allow for a more tissue-targeted approach using an agonist antibody to treat autoimmune diseases



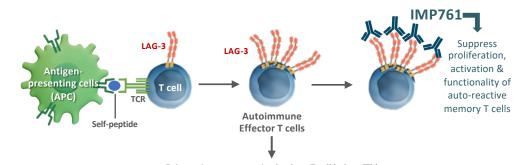
## IMP761: First-in-Class LAG-3 Agonist a Potential Game-Changer



IMP761 increases "brake" function of the LAG-3 immune checkpoint and its natural down-regulation of auto-reactive memory T cells, which represent the root cause of many disorders.

As a LAG-3 agonist, IMP761 may treat many autoimmune diseases including:

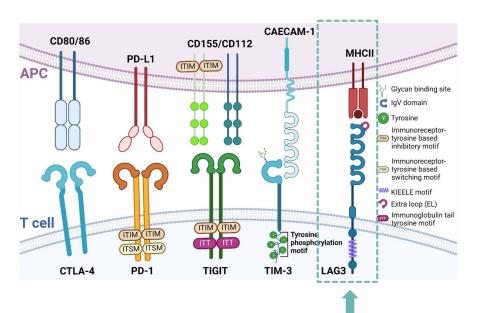
- √ Rheumatoid arthritis: market size est. \$29.6 billion\*
- ✓ Type 1 diabetes: market size est. \$9.9 billion\*
- ✓ Multiple sclerosis: market size est. \$32.9 billion\*



Epigenetic reprogramming leads to T cell helper (Th) induced Autoimmune Diseases such as Rheumatoid Arthritis (Th1), Allergic Asthma (Th2), IBS (Th17), etc.

## IMP761: Strong Inhibition of TCR Signalling & T Cell Activation





- Unlike 100+ inhibitory receptors including PD-1, LAG-3 has no tyrosine-based ITIM motif in its cytoplasmic domain<sup>1</sup>
- Inhibitory motifs unique to LAG-3 explain in part clear & rapid inhibition of T cell receptor (TCR) signalling induced by IMP761 in preclinical studies<sup>2</sup>
- IMP761 strongly blocks T cell activation via TCR in preclinical studies and has shown encouraging T cell suppression in a Phase I trial

Unique LAG-3 signalling pathway with no tyrosine-based ITIM motif

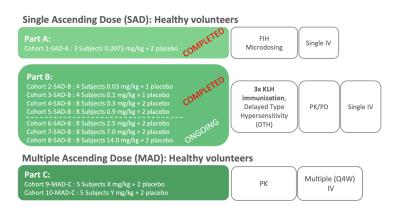
#### First-in-Human Phase I Trial of IMP761



World-class research institute, CHDR, appointed to conduct first-in-human study

#### **Overview / Key Milestones:**

- Placebo-controlled, double-blind Phase I (N = 49)
- Centre for Human Drug Research (CHDR) in Leiden, the Netherlands, conducting study in healthy volunteers
- Initial pharmacological data shows significant T cell suppression and favourable safety profile at 0.9 mg/kg dosing level<sup>1</sup>
- Single ascending dose levels continuing with 2.5, 7, 14 mg/kg
- Additional data expected in CY2025





CHDR offers a unique Keyhole Limpet Hemocyanin (KLH) challenge model allowing for evaluation of IMP761's pharmacological activity at early stages of development



## **Upcoming Milestones**



- Non-Small Cell Lung Cancer TACTI-004 futility analysis in Q1 2026
- **Soft Tissue Sarcoma** Update from EFTISARC-NEO Phase II in November at CTOS 2025
- Head and Neck Squamous Cell Carcinoma Ongoing evaluation of potential options for collaborative clinical development & paths for accelerated approval in 1L HNSCC CPS <1</li>
- Metastatic Breast Cancer Update from AIPAC-003 trial at medical conference in 2H 2025
- Early-Stage Breast Cancer Initiation of the new IIT Phase II evaluating neoadjuvant efti
- Autoimmune Diseases Update from IMP761 first-in-human Phase I trial in 2H 2025 and beyond
- Additional Updates From ongoing clinical trials, partnered programs, and potential expansion of clinical trial pipeline

## Summary



Phase III in 1L NSCLC with multi-billion US\$ opportunity

First-in-class assets in oncology and autoimmune diseases

> Well-funded with cash runway to the end of CY2026<sup>1</sup>

- Immutep is a leader in immunotherapy with substantial opportunity in both oncology & autoimmune disease
- Efti is the only immunotherapy that directly activates APCs via MHC Class II pathway and has significant potential to redefine cancer treatment landscape
  - ✓ Expands responders and enhances efficacy when combined with anti-PD-(L)1 therapy (e.g. KEYTRUDA)
  - √ Safety profile and easy subcutaneous delivery key attributes in current & future treatment landscape
  - ✓ Well positioned in terms of CMC (2,000L commercial scale production, 3-year shelf life, etc.), clinical development (e.g. no further toxicology studies needed for BLA/MAA), regulatory interactions, and IP
  - ✓ Lead indication (1L NSCLC) expected to reach US\$48 billion in drug sales in 2031 with >50% from IO
- IMP761, the world's first LAG-3 agonist antibody, is potential game-changer in treatment of autoimmune diseases
  - ✓ Encouraging safety & efficacy to date in Phase 1 study
  - ✓ May treat very large indications including rheumatoid arthritis, Type 1 diabetes & multiple sclerosis
- Multiple collaborations with large pharma and academia globally
- Strong IP portfolio; 12+ years of potential exclusivity for efti/IMP761 as biologics
- Cash & cash equivalents of ~A\$109.85 million provide runway to end of CY2026.1

# Thank You

