

# A Global Leader in LAG-3 Therapeutics in Oncology and Autoimmune Disease

**Corporate Presentation – November 2022**  
(ASX: IMM, NASDAQ: IMMP)

# 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.*

*The views expressed in this presentation contain information derived from publicly available sources that have not been independently verified. No representation or warranty is made as to the accuracy, completeness or reliability of the information.*

*Any forward looking statements in this presentation have been prepared on the basis of a number of assumptions which may prove incorrect and the current intentions, plans, expectations and beliefs about future events are subject to risks, uncertainties and other factors, many of which are outside Immutep's control. Important factors that could cause actual results to differ materially from assumptions or expectations expressed or implied in this presentation include known and unknown risks. Because actual results could differ materially to assumptions made and Immutep's current intentions, plans, expectations and beliefs about the future, you are urged to view all forward looking statements contained in this presentation with caution.*

*This presentation should not be relied on as a recommendation or forecast by Immutep. Nothing in this presentation should be construed as either an offer to sell or a solicitation of an offer to buy or sell shares in any jurisdiction.*

*This presentation is authorised for release by the CEO of Immutep Limited.*

## Pioneering LAG-3 Portfolio in Oncology & Autoimmune Disease

Immutep is a pure-play LAG-3 clinical-stage company with four product candidates that address significant market opportunities in oncology & autoimmune disease

## Collaborations with Industry Leaders



Merck KGaA  
Darmstadt, Germany



## Compelling Clinical Data

Clinical trials of lead candidate efitlagimod alpha (efti) with immunotherapy & chemotherapy have shown compelling results in NSCLC, HNSCC, HR+/HER2- BC, melanoma and other solid tumors

## Global Presence



# Immutep LAG-3 Pipeline

	Program	Preclinical	Phase I	Phase II	Late Stage	Commercial Rights	
ONCOLOGY	<b>Eftilagimod Alpha (efti or IMP321)</b> APC activating soluble LAG-3 protein 	Metastatic Breast Cancer (Chemo-IO) - AIPAC					 Global Rights
		1 <sup>st</sup> Line Head and Neck Squamous Cell Carcinoma (IO-IO) - TACTI-003					
		2 <sup>nd</sup> Line Head and Neck Squamous Cell Carcinoma (IO-IO) - TACTI-002					
		1 <sup>st</sup> Line Non-Small Cell Lung Carcinoma (IO-IO) - TACTI-002					
		2 <sup>nd</sup> Line PD-X Refractory Non-Small Cell Lung Carcinoma (IO-IO) - TACTI-002					
		Solid Tumors (IO-IO-chemo) - INSIGHT-003					
		Solid Tumors (IO-IO) - INSIGHT-004				 Merck KGaA Darmstadt, Germany	
		Melanoma (IO-IO) - TACTI-mel					
		Soft Tissue Sarcoma (IO-IO-RT)					
	Metastatic Breast Cancer & Other Solid Tumors (Chemo-IO & IO-IO)					China Rights	
	<b>LAG525</b> Antagonist Antibody 	Solid Tumors + Blood Cancer (IO-IO Combo)					 Global Rights
		Triple Negative Breast Cancer (Chemo-IO Combo)					
		Melanoma (IO-IO-Small Molecule Combo)					
Solid Tumors (IO-IO Combo)							
TNBC (Chemo-IO-Small Molecule Combo)							
Small Molecule Anti-LAG-3	Undisclosed					Global Rights	
AUTOIMMUNE DISEASE	<b>GSK'781</b> Depleting Antibody 	Ulcerative Colitis					 Global Rights
		Psoriasis					
		Healthy Japanese and Caucasian Subjects					
	<b>IMP761</b> Agonist Antibody 	Undisclosed					 Global Rights


# LAG-3: Approved Checkpoint with Unique Characteristics

Immune system's role in controlling cancer has led to regulatory approval of immunotherapies targeting CTLA-4, PD-1, and now LAG-3 checkpoints





**Cytotoxic T lymphocyte  
Antigen-4 (CTLA-4)**

*Yervoy (anti-CTLA-4) approved 2011;  
commercial sales >\$2 billion in 2021*




**Programmed Cell Death  
Protein-1 (PD-1)**

*Keytruda & Opdivo (anti-PD-1) approved  
2014; combined commercial sales  
>\$24 billion in 2021*



**Lymphocyte Activating  
Gene-3 (LAG-3)\***

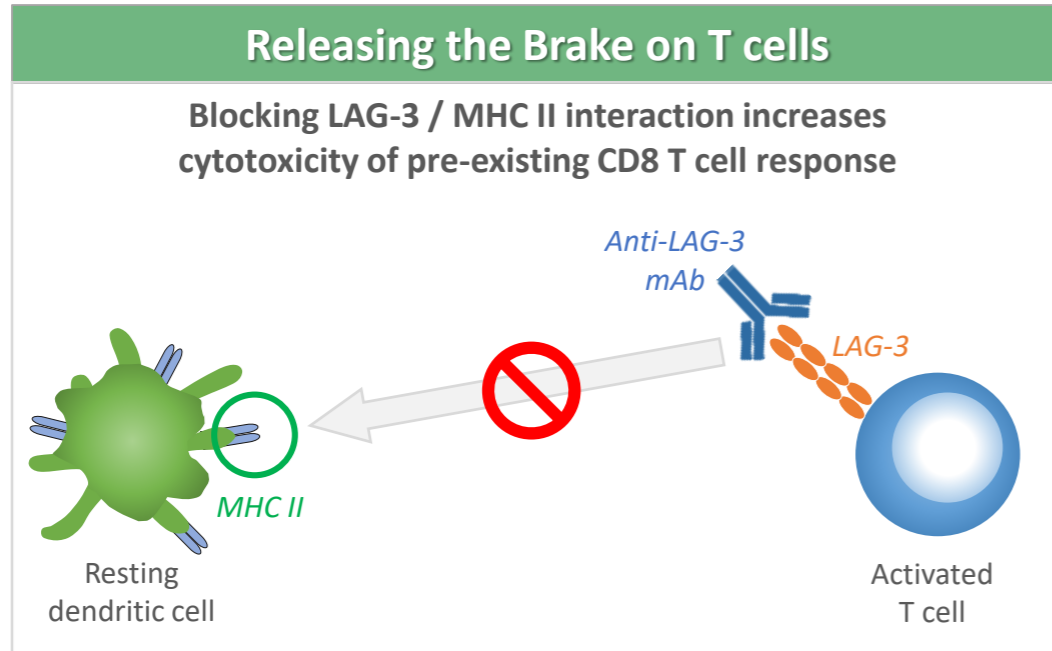
*Relatlimab (anti-LAG-3) approved 2022  
in combination with Opdivo; BMS est.  
>\$4 billion in NRA sales\*\* in 2029*



LAG-3 is unique in that its inhibition on T cells & activation of dendritic cells engages the adaptive & innate immune systems against cancer offering significant potential to: (1) improve responses to standard-of-care immunotherapy & chemotherapy, (2) limit emergence of resistance, (3) offer chemotherapy-free options in select indications.

# LAG-3 Therapeutics for Oncology

# Multiple Companies Targeting LAG-3 Inhibition

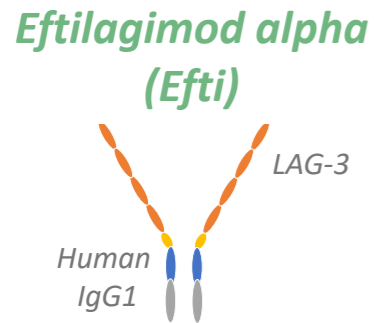


Company*	Program	Phase 1	Phase 2	Phase 3
Bristol Myers Squibb	Relatlimab	8	35	4
MERCK	Favezelimab	1	8	1
REGENERON	Fianlimab	1	1	1
NOVARTIS	Ieramilimab	1	4	
MACROGENICS	Tebotelimab	3	3	
Roche	R07247669	2	3	
Incyte	INCAGN02385	2	2	
Boehringer Ingelheim	BI754111	4	1	
Innovent	IBI110	2	1	
TESARO	TSR-033	1	1	
symphogen	SYM022	3		
F-star†	FS-118	1		

Received FDA approval in March 2022

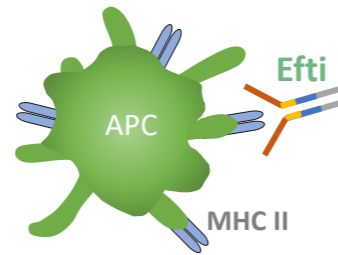
- ImmuteP designed the first anti-LAG-3 antibody and licensed it to CoStim Pharmaceuticals in 2012, which was acquired by Novartis in 2014
- Novartis' anti-LAG-3 mAb, LAG525, activates effector T cells & inhibits regulatory T cells (removing two brakes on the immune system to respond to and kill cancer cells) and has been tested in multiple clinical trials combined with spartalizumab (anti-PD-1) and chemotherapy\*\*
- IP estate for LAG525 continues to strengthen with patent grants in key markets including US, Europe, Japan and China

## Pushing the Accelerator on the Immune System

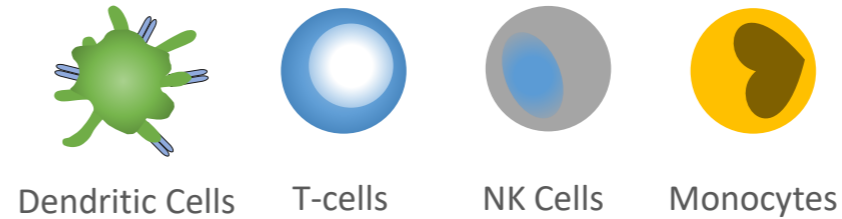


*Immutep's proprietary soluble LAG-3Ig clinical candidate is a first-in-class antigen-presenting cell (APC) agonist via MHC II that capitalizes on LAG-3's unique characteristics*

### APC activation with Efti



### Anti-tumor immune cell activation



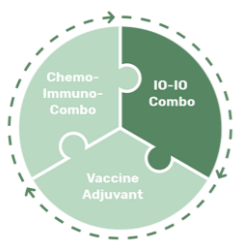
### Broad activation of immune system

- Efti capitalizes on LAG-3's unique ability to drive adaptive & innate immune systems against cancer
- Efti has high affinity for a subset of MHC II ligand on APCs and their activation drives broad stimulation of multiple anti-tumor cells

### Compelling pairing capabilities

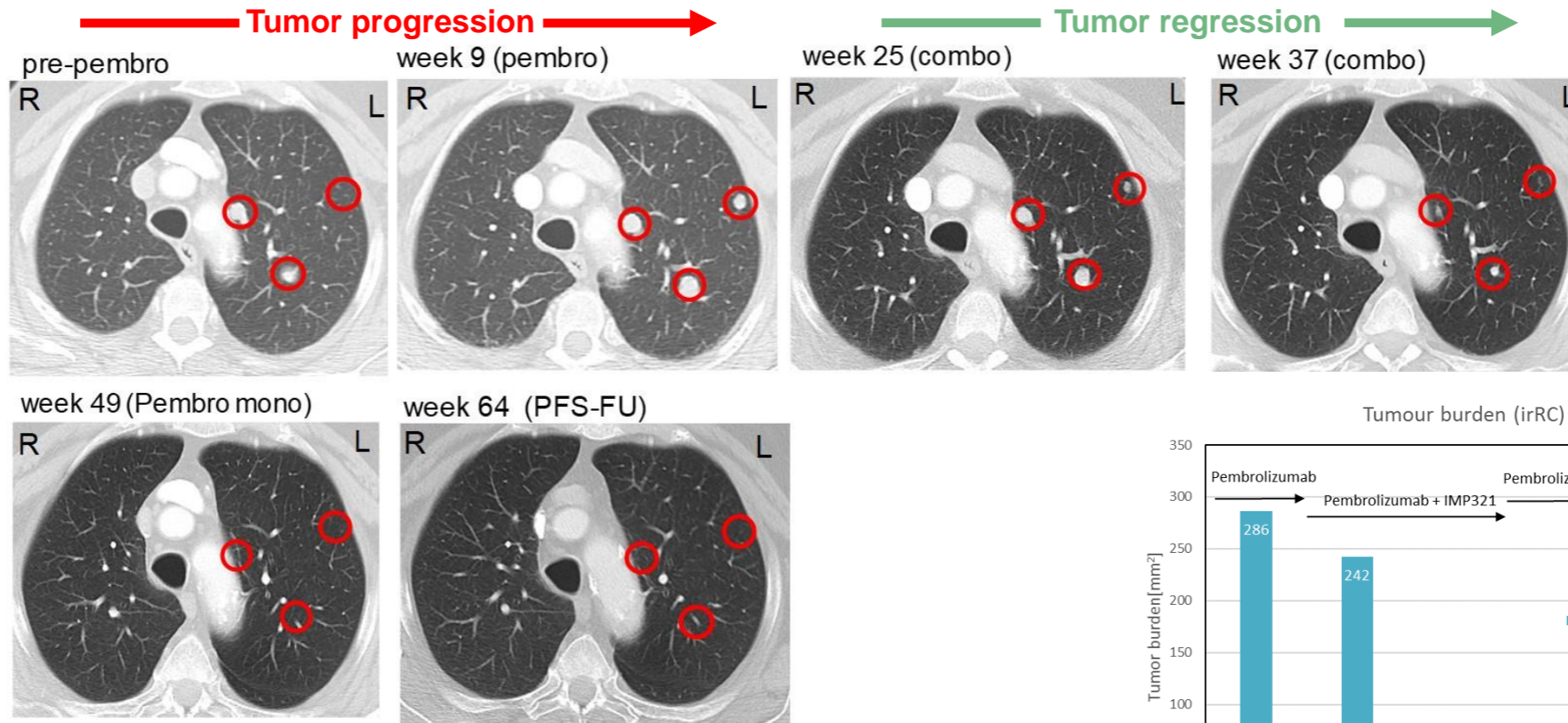
- Excellent safety profile drives high suitability for combination partnering
- Synergistic activity & encouraging clinical results with multiple agents including anti-PD-1, anti-PD-L1, and chemotherapy
- Enhances clinical activity of anti-PD-1 across PD-L1 status, including low & negative PD-L1 tumors



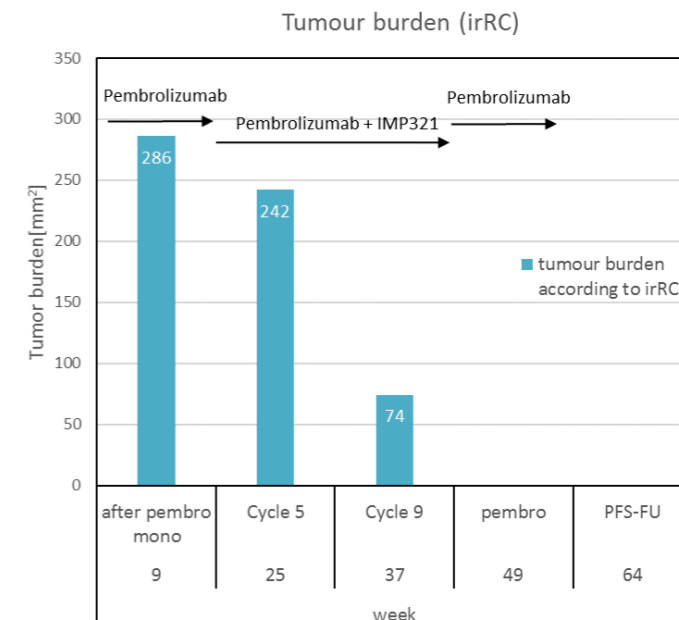


# Efti: Clinical Development TACTI-mel - Results - Single Case

## Activity: Metastatic Melanoma



- Patient progressing on Keytruda monotherapy
- At 1 yr all lesions disappeared → Complete Response (confirmed)
- Patient without treatment and disease free at last visit



# Non-Small Cell Lung Cancer

TACTI-002 trial  
Efti + Pembrolizumab

1<sup>st</sup> line Metastatic  
NSCLC

FDA

Fast Track  
Designation

ASCO 2022

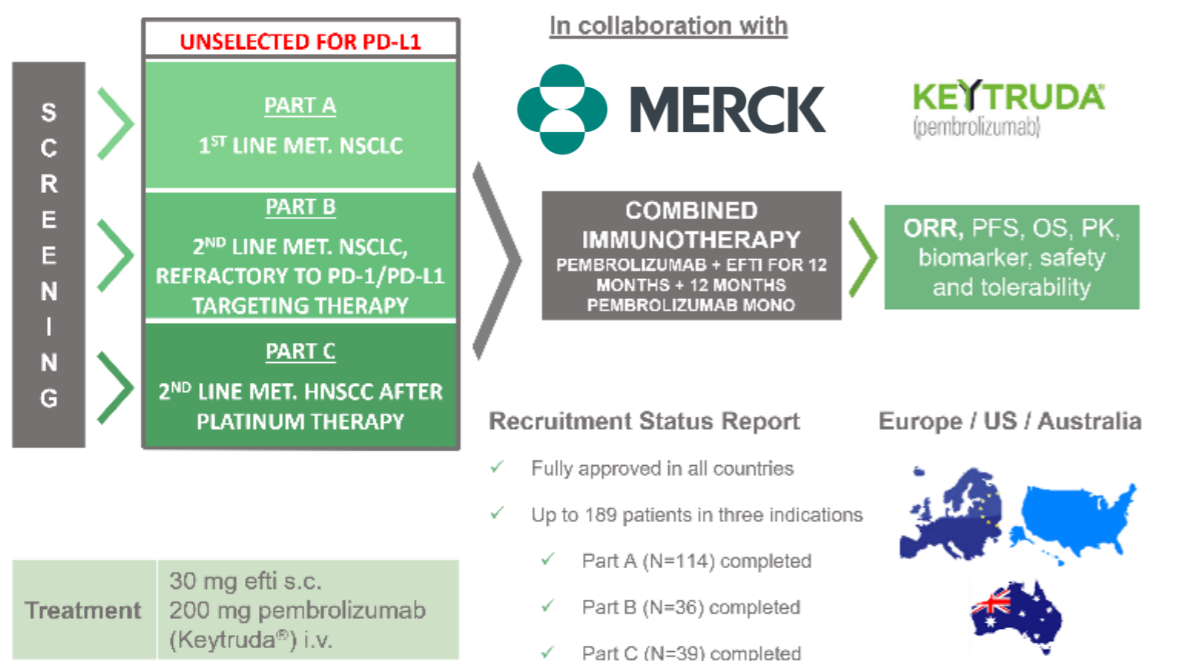


[Click here](#)

# Phase II Trial Evaluating Efti + Pembro in 1L NSCLC

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

### TACTI-002: Two ACTIVE Immunotherapeutics in NSCLC & HNSCC



Baseline characteristics for PD-L1 All Comer Trial		Part A (N=114)
Age, median (range), years		67 (44-85)
Sex, n (%)	Female / Male	30 (26.3) / 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	106 (93.0) / 8 (7.1)
PD-L1 expression TPS <sup>1</sup> , n (%)	< 1%	37 (32.5)
	1-49%	40 (35.1)
	≥ 50%	31 (27.2)
	Not evaluable	6 (5.3)
Previous therapy, n (%)	Radiotherapy	38 (33.3)
	Surgery	23 (20.2)
	Systemic therapy for non-metastatic disease	25 (21.9)

All-comer trial for patients with all levels of PD-L1 expression;  
 ~33% & ~68% of 1L NSCLC patients in TACTI-002/KEYNOTE-798 have PD-L1 TPS of <1% & <50%, respectively.

# Encouraging Clinical Results; Primary Objective Achieved; Strong ORR/PFS Across Entire PD-L1 Spectrum vs Pembrolizumab

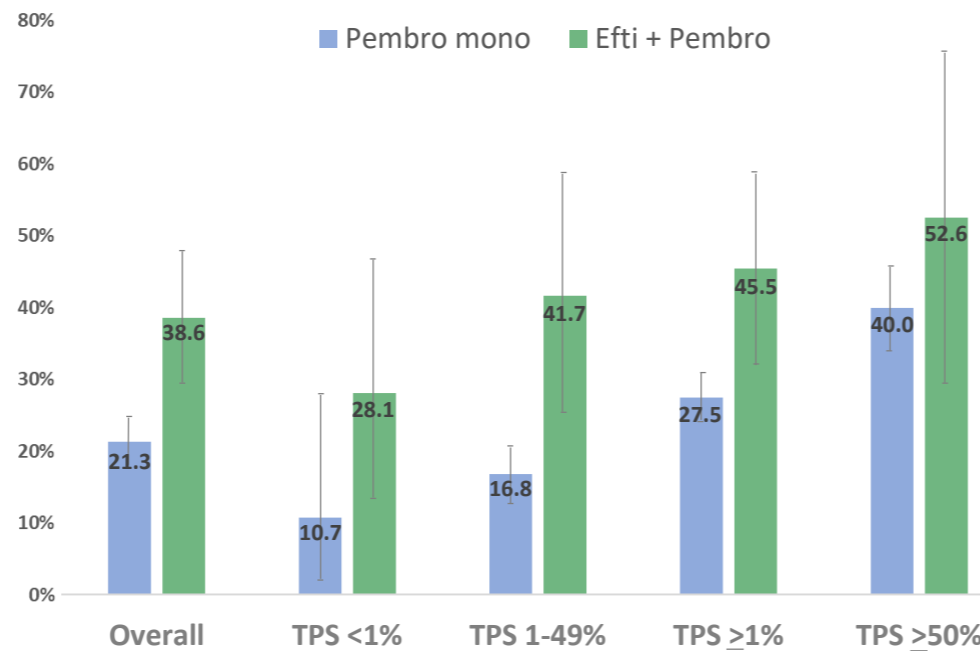
TACTI-002/KEYNOTE-798: 1L NSCLC (Part A)

## Key Takeaways

- Primary objective achieved (ORR >35%)
- Superior ORR/PFS across all PD-L1 levels
- Sustained, durable responses
- Safe, well tolerated

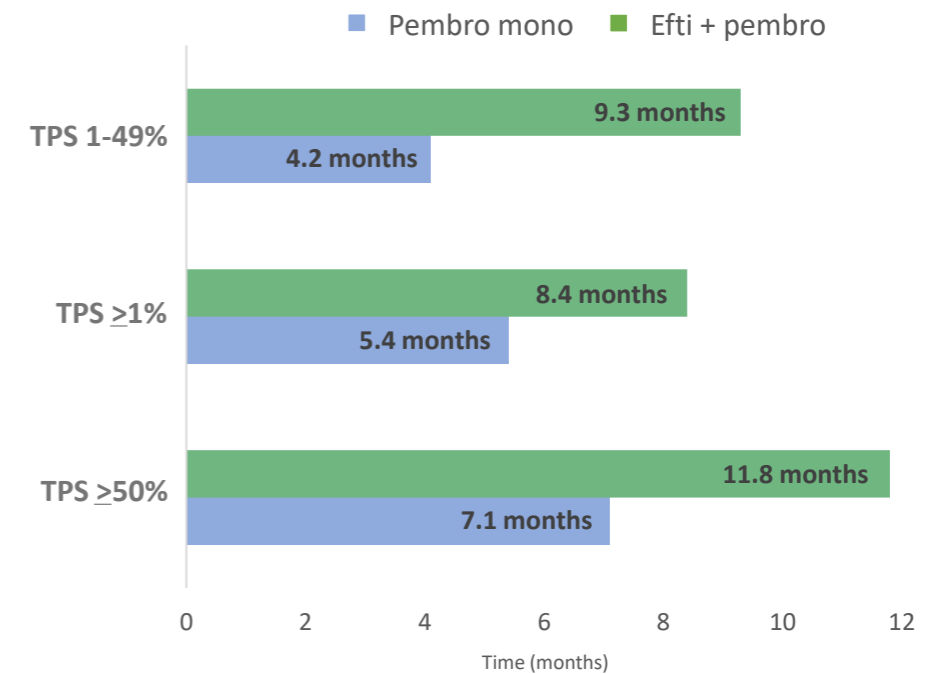
## Overall Response Rate\* (ORR)

(with 95% confidence intervals)



## Median Progression Free Survival# (PFS)

(by PD-L1 TPS Score)

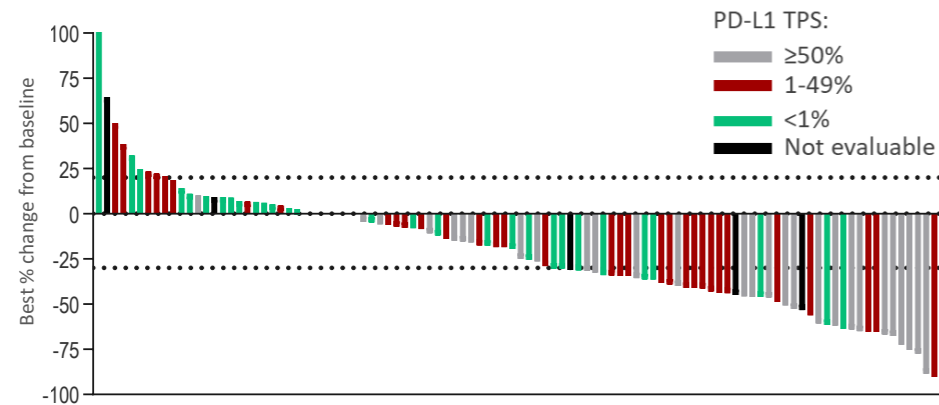


\* Efti + Pembro ORR by iRECIST, unconfirmed, Total (N=114) and PD-L1 TPS subgroups (N=87). Data cut-off April 15, 2022. Pembrolizumab ('pembro') mono efficacy used for benchmarking for ORR: Total calculation based on KN-001; KN-042 and on adjustment for the same PD-L1 subgroup distribution as in TACTI-002. < 1% TPS: calculation based on limited data set from KN-001 (1st & 2nd line altogether). 1-49%, ≥ 50%, and ≥ 1% TPS based on KN-001, KN-042. # PFS by PD-L1 status (N=108) using central assessment for 87 patients. For 21 patients, local assessment used due to non-eval central assessment results. Pembro mono efficacy used for benchmarking: 1-49%, ≥ 50%, and ≥ 1% TPS based on KN-001, KN-042. Lancet [https://doi.org/10.1016/S0140-6736\(18\)32409-7](https://doi.org/10.1016/S0140-6736(18)32409-7), Oral Presentation 2018 ASCO, EPAR assessment report, N Engl J Med 2016; 375:1823-33; KN-024 update J Clin Oncol 2019, KN-024 J Clin Oncol 2021

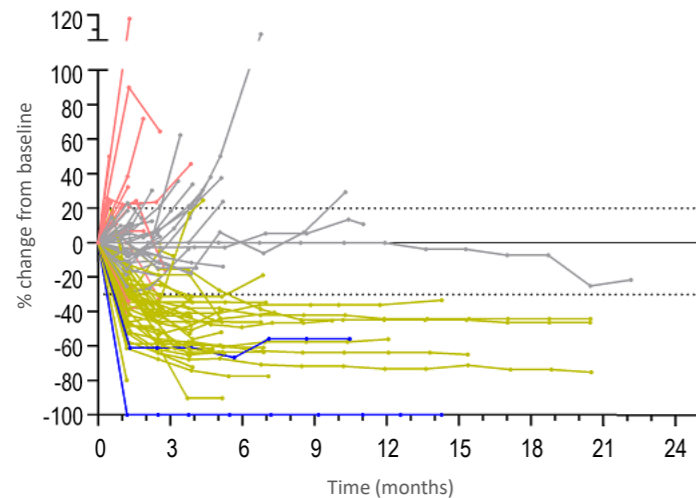
# Deep and Durable Responses

TACTI-002/KEYNOTE-798: 1L NSCLC (Part A)

## Tumor Burden Reduced in Majority of Patients



## Change in Tumor Size Over Time



- Responses are deep & long-lasting; median DoR not yet reached
- 80% (35) of responses<sup>1</sup> already confirmed & 11.4% (5) pending confirmation
- 95% of patients having a response < 4 months after study start
- Only 8.6% of patients with confirmed response<sup>2</sup> progressed ≤ 6 months until data cut-off
- 66% (68) of patients with post-baseline assessment had decrease in target lesions

# Benchmarking against IO & IO-Chemo Combinations

TACTI-002/KEYNOTE-798: 1L NSCLC (Part A)

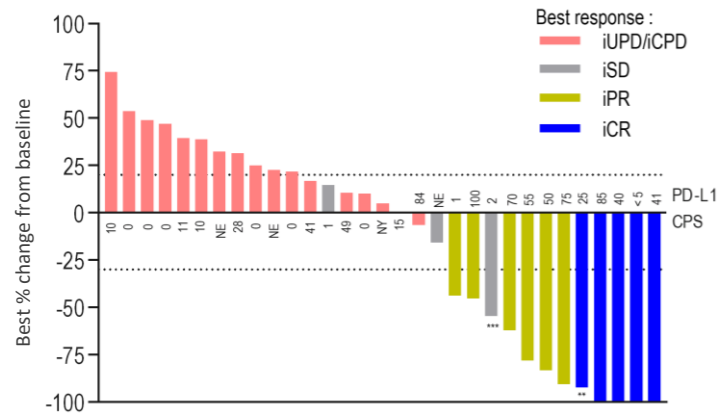
	TPS	Treatment	Efficacy <sup>(1)</sup>	Toxicity: AEs leading to disc.
1L NSCLC	≥ 1%	<b>Efti + Pembro</b>	ORR 45.5% PFS 8.4 mos	< 10%
		Pembro mono	ORR 27.5% PFS 5.4 mos	1-14%
		Ipi + Nivo <sup>(2)</sup>	ORR 36% PFS 5.1 mos	18%
	1-49%	<b>Efti + Pembro</b>	ORR 41.7% PFS 9.3 mos	< 10%
		Doublet Chemo + Pembro	ORR 49.2% (NSQ) & 50% (SQ) PFS 7.2 (SQ) & 9.2 (NSQ) mos	14%
		Pembro mono	ORR 16.8% PFS 4.1 mos	1-14%
	≥ 50%	<b>Efti + Pembro</b>	ORR 52.6% PFS 11.8 mos	< 10%
		Pembro/Atezo/Libtayo mono	ORR 35-45% PFS 7-10 mos	1-14%
	0-100%	<b>Efti + Pembro</b>	ORR 38.6% PFS 6.9 mos	< 10%
		Doublet Chemo	ORR 19-30% PFS 5-9 mos	8-22%
Doublet Chemo + Pembro		ORR 48% (NSQ) & 63% (SQ) PFS 6 (SQ) & 9 (NSQ) mos	14%	
Doublet Chemo + Atezo + Beva		ORR 56% PFS 8.4 mos	33%	
Doublet Chemo + Ipi + Nivo		ORR 38% PFS 6.7 mos	19%	

- ✓ Low efficacy of anti-PD-(L)1 monotherapy for patients with TPS <50% (~70% of total population)
- ✓ Double chemo + anti-PD-(L)1 → increased ORR & OS but shorter DoR due to chemo & more toxic; Ipi & Beva combos → high burden in terms of toxicity & high number of patients discontinuing
- ✓ Efti addresses both issues as shown with TACTI-002 results; INSIGHT-003 trial also exploring efti + pembro + chemo combination

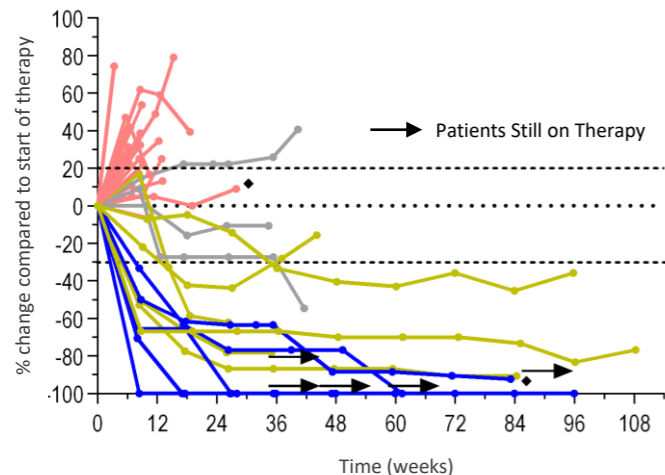
# 2L HNSCC: Robust ORR/CR with Long-Lasting Efficacy

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

## Responses at all PD-L1 levels including 5 iCRs



## Deep and durable responses



	Efti + Keytruda Combination*	Keytruda Monotherapy#
Clinical Trial & Phase	P2 (TACTI-002/KN-798)	P3 (KN-040)
Patient Number / tumor type	39 / 2L HNSCC	247 vs. 248 / 2L HNSCC
PD-L1 Status	PD-L1 All comer	PD-L1 All comer
Complete Response (CR) %	13.5%	1.6%
Partial Response (PR) %	16.2%	13%
Stable Disease (SD) %	8.1%	22.7%
ORR in evaluable patients %	35.5%	n/a
Overall Response Rate (ORR) %	29.7%	14.6%
PD-L1 CPS ≥1% group	40.7%	17.3%
PD-L1 CPS ≥20% group	64.3%	21.9%
Median PFS (months)	2.1	2.1
PD-L1 CPS ≥1% group	4.1	2.2
Median OS (months)	12.6	8.4
PD-L1 CPS ≥1% group	12.6	8.7

*Eight-fold increase in CR with efti + pembro*

*More than double ORR across all PD-L1 levels with efti + pembro*

# Fast Track Designation in 1L HNSCC

## TACTI-003: Phase IIb in 1<sup>st</sup> Line Head and Neck Squamous Cell Carcinoma (HNSCC)

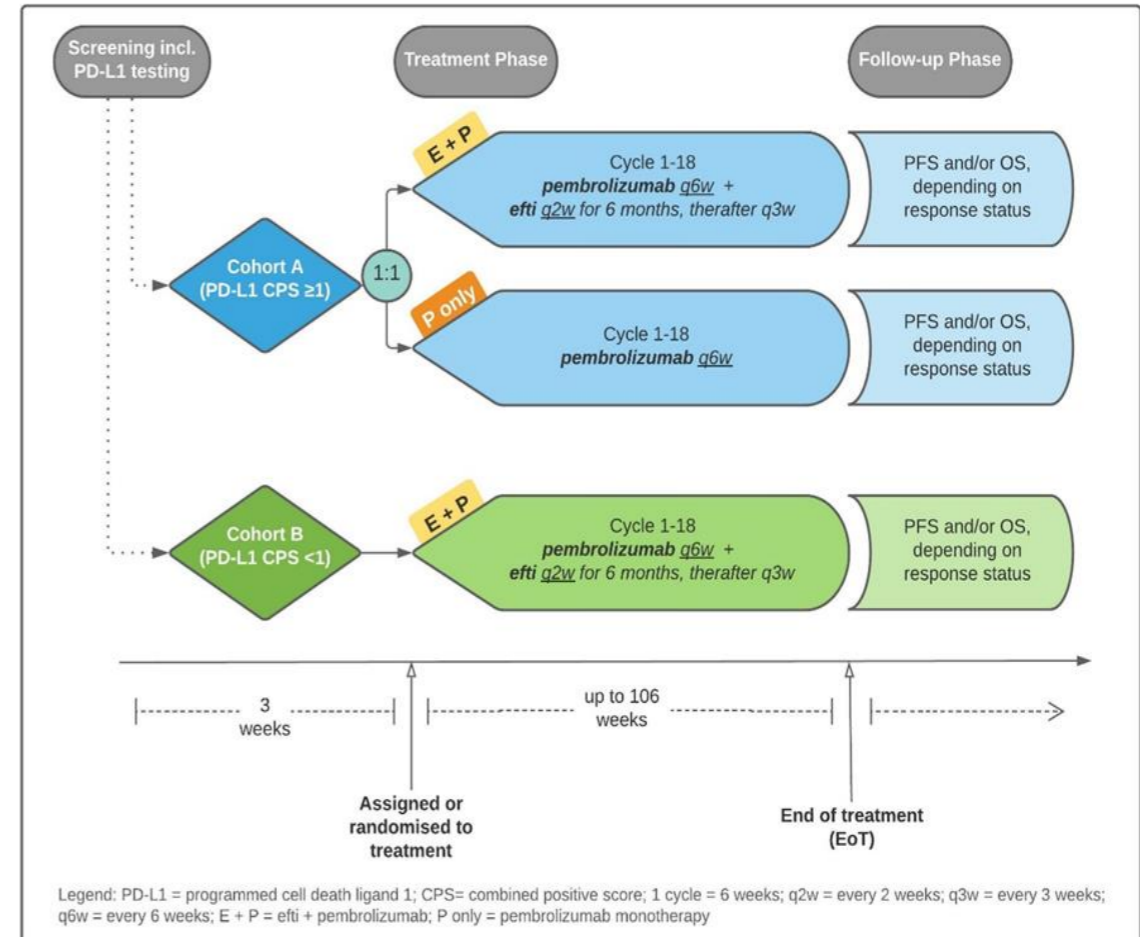


### Status:

- Recruiting (~30% enrolled; recruitment accelerating as further sites have been activated\*)
- **FDA Fast Track designation on strength of TACTI-002 data in 2L HNSCC**

### Design:

- Randomised trial with ORR as primary endpoint
- Sites worldwide (AU, US, Europe)
- Approximately 154 patients: either to be randomised to have sufficient patients in each group or in an experimental arm





## 2022 Milestones

- Year to date:
  - ✓ 1L NSCLC Oral Presentation at ASCO (TACTI-002; Part A)
  - ✓ 2L NSCLC PD-X refractory data at ELCC & WCLC 2022 (TACTI-002; Part B)
  - ✓ Fast Track Designation granted in 1L NSCLC
  - ✓ Update from TACTI-003
  - ✓ New, significant data from AIPAC study
  - ✓ IP expansion for eftilagimod alpha
- Additional clinical data updates through year end
  - New data from Phase II TACTI-002 in 1L NSCLC
  - Initial results from INSIGHT-003 (first triple-combo data)
- Expansion of existing programs (i.e., new sarcoma trial)
- Regulatory updates
- Manufacturing scale up to 2,000L
- Updates on IMP761 & partnered programs (e.g. Novartis, GSK, EOC)

## Corporate Snapshot

- Pioneering LAG-3 portfolio in oncology and autoimmune diseases with four product candidates in multiple clinical trials
- First-in-class positioning with eftilagimod alpha (efti)
- Multiple big pharma partnerships & collaborations, while retaining full control of efti (ex-China) & IMP761
- Well funded with ~A\$73.9 million\* in cash
- Cash runway to early CY2024\*
- Listings
  - ASX (primary) IMM
  - NASDAQ (ADS @ 1:10 ratio) IMMP
- Market cap ~A\$277M / \$176M US\*\*
- 7.41% owned by Fidelity (FIL)
- Total institutional ownership of ~57%



Thank You