

# Cancer Immunotherapies by design™

Nasdaq: ITOS

January 2024

# Forward-Looking Statements



This presentation contains forward-looking statements. Any statements that are not solely statements of historical fact are forward-looking statements. Words such as "believe," "anticipate," "plan," "expect," "will," "may," "intend," "prepare," "look," "potential," "possible" and similar expressions are intended to identify forward-looking statements. These forward-looking statements include statements relating to the potential benefits of our product candidates and combinations, including the potential of beliestotug to be the highest quality IIGIT in the field, the potential of EOS-984's mechanism to have profound effects as a monotherapy or in combinations, and the potential of inupadenant to enhance chemotherapy therapeutic response; the expectation that 2024 will be a defining year for iTeos; our clinical and data generation plans for 2024, including initiating a IIGIT Phase 3 registrational study, having clinical data from GALAXIES Lung-201 and IIG-006 HNSCC, having clinical data from the dose escalation portion of A2A-005 in late 2024, presenting preclinical mechanism of action data from EOS-984 in the second quarter of 2024, and having topline data from the Phase 1 dose escalation trial in advanced malignancies in late 2024; our goal to gain commercial approval for belrestotug in 1L NSCLC and branch into earlier lines and potentially to a variety of IO amenable tumors; the potential of our biomarker for IIGIT in identifying indications to target and subpopulations; our market opportunities and number of patients potentially eligible for our product candidates; the potential benefits of our collaboration with GSK and the expectation that 2024 will be a year of significant momentum for this collaboration; and our expected cash runway through 2026, which contemplates the launch of multiple IIGIT Phase 3 trials.

These forward-looking statements involve risks and uncertainties, many of which are beyond iTeos' control. Actual results could materially differ from those stated or implied by these forward-looking statements as a result of such risks and uncertainties. Known risk factors include the following: success in preclinical testing and early clinical trials does not ensure that later clinical trials will be successful, and early results from a clinical trial do not necessarily predict final results; the data for our product candidates may not be sufficient to support regulatory approval; iTeos may encounter unanticipated costs or may expend cash more rapidly or more slowly than currently anticipated due to challenges and uncertainties inherent in product research and development and biologics manufacturing; the expected benefits and opportunities related to the agreement between iTeos and GSK may not be realized or may take longer to realize than expected due to a variety of reasons, including any inability of the parties to perform their commitments and obligations under the agreement, challenges and uncertainties inherent in product research and development and manufacturing limitations; iTeos may not be able to execute on its business plans, including meeting its expected or planned regulatory milestones and timelines, research and clinical development plans, and bringing its product candidates to market, for various reasons, some of which may be outside of iTeos' control, including possible limitations of company financial and other resources, manufacturing limitations that may not be anticipated or resolved for in a timely manner, negative developments in the field of immuno-oncology, such as adverse events or disappointing results, including in connection with competitor therapies, and regulatory, court or agency decisions such as decisions by the United States Patent and Trademark Office with respect to patents that cover our product candidates; and those risks identified under the heading "Risk Factors" in i

Any of the foregoing risks could materially and adversely affect iTeos' business, results of operations and the trading price of iTeos' common stock. We caution investors not to place considerable reliance on the forward-looking statements contained in this press release. iTeos does not undertake any obligation to publicly update its forward-looking statements based on events or circumstances after the date hereof.



2024

# A Defining Year for iTeos

#### **Promising TIGIT:PD-1 Doublet**

1

Two Data Readouts Anticipated in 2024

#### **Unlocking Adenosine Pathway**

2

Two Data Readouts Anticipated in 2024

#### **Funded Through 2026**

3

**\$645M** in cash as of 3Q23

# Deep Pipeline with Four Clinical Readouts in 2024



Innovative molecules and compelling combinations

	Preclinical	Phase 1	Phase 2	Phase 3	Status
Belrestotug: IgG1 antibody targeting TIGIT					iTEOS GSK
+ dostarlimab   1L NSCLC PDLI high					Planned Study
+ dostarlimab   1L NSCLC PDL1high	GALAXIES Lung-201			Data Anticipated 2024	
+ dostarlimab   1L HNSCC PDL1high/low			TIG-006		Data Anticipated 2024
+ dostarlimab + CD96   1L HNSCC PDL1high		GALAXIE	S H&N-202		Enrolling
+ dostarlimab + chemotherapy   1L mNSCLC		TIG-006			Enrolling
+ dostarlimab + CD96   Advanced Malignancies	NCT0	3739710			Enrollment Complete
+ dostarlimab + PVRIG   Advanced Malignancies	NCT0	5277051			Enrolling
Inupadenant: Small molecule targeting $A_{2A}$	receptor				iTEOS
+ chemotherapy   Post-IO Chemo-naïve NSCLC			A2A-005		Data Anticipated Late 2024
EOS-984: Small molecule targeting ENT1					iTEO5
Monotherapy   Advanced Malignancies					Data Anticipated 2024



# Belrestotug

EOS-448 / GSK4428859A

iTeos and GSK are uniquely positioned to leverage the TIGIT/CD226 axis



We Hold An

# Advantageous Field Position

Significant momentum in 2023



### There Is A Need for a Transformative TIGIT:PD-1 Doublet



# We Believe Our TIGIT:PD-1 Doublet Is Differentiating In Key Areas

Proven quality target engagement with TIGIT and FcyR

TIGIT monotherapy activity

Pembrolizumab comparison

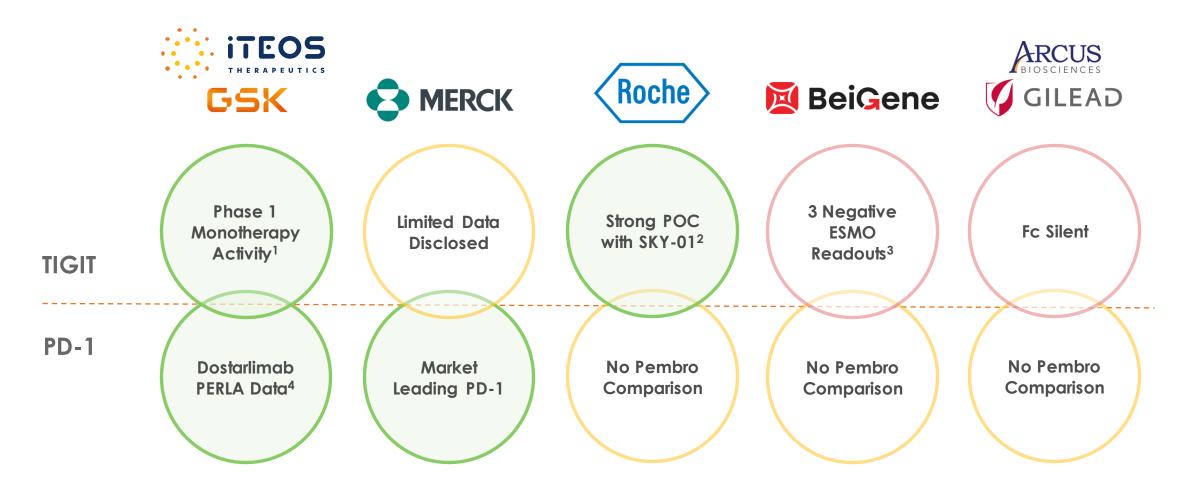


**High Quality TIGIT** 

### The Need for a Transformative TIGIT:PD-1 Doublet



Belrestotug + dostarlimab represent differentiated, high-quality therapies



<sup>1.</sup> iTeos AACR 2021

<sup>2.</sup> Genentech Phase 3 Skyscraper-01 Study-August 22, 2023 Release

<sup>3.</sup> ESMO 2023 - Adv anTIG-203, Adv anTIG-206, Adv anTIG-202

<sup>4.</sup> ESMO 2023 – Phase 2 GSK-sponsored PERLAstudy in 1L NSCLC

# Belrestotug: The First And Only TIGIT to Demonstrate Robust Target Engagement



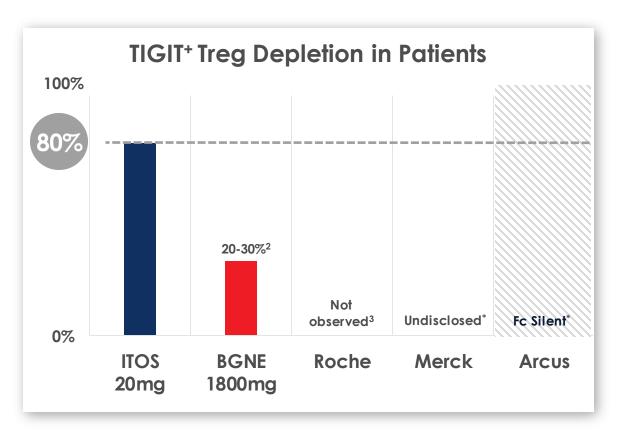
Unique Epitope Binding

**High Affinity + Potency** 

First and only TIGIT with proven

Treg depletion at all doses

Only TIGIT to Demonstrate Phase 1 Monotherapy Activity



<sup>\*</sup>Merck vibostolimab has yet to publicly show Treg depletion data; Arcus' domvanalimab is an Fc silent TIGIT and not capable of Treg depletion

<sup>3</sup>Ira Melman, Vice President, Cancer Immunology at Genentech: "We would have loved to see Treg depletion...I know that [TIGIT] is also present in fairly high abundance on regulatory T cells but neither in the mouse models nor in cancer patients can we really find much or certainly dramatic evidence that Treg compartment is diminished as a consequence of TIGIT exposure."

iTeos AACR 2021

doi: 10.1136/jitc-2022-SITC2022.0768

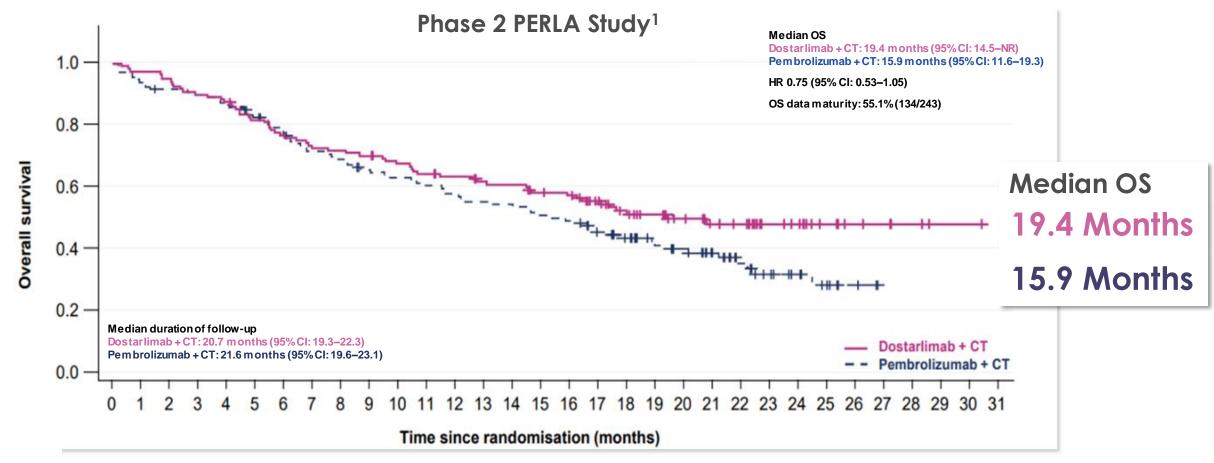
Piper Sandler Virtual Biolnsights KOL Day: Expert Call on Next Generation Cancer Immunotherapy – June 2020

# PERLA: Dostarlimab Is A Quality Anti-PD-1 Backbone



Positive numerical OS trend in 1L NSCLC favoring dostarlimab + CT vs. pembrolizumab + CT





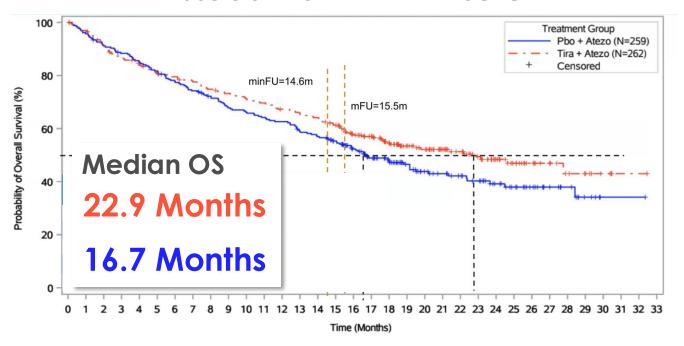
<sup>&</sup>lt;sup>1</sup>Phase 2 GSK-sponsored PERLA study in 1L NSCLC. Peters S, et al. Annals of Oncology (2023) 34 (suppl\_2): \$1254-\$1335. 10.1016/annonc/annonc1358. NB: Safety profiles in the secondary analysis of OS were similar and consistent with those previously reported in the primary analysis for the PERLA trial.

# SKY-01: Meaningful Separation of Curves Validates TIGIT



Quality of components and clinical trial design leave room for improvement





### **Key Insights**

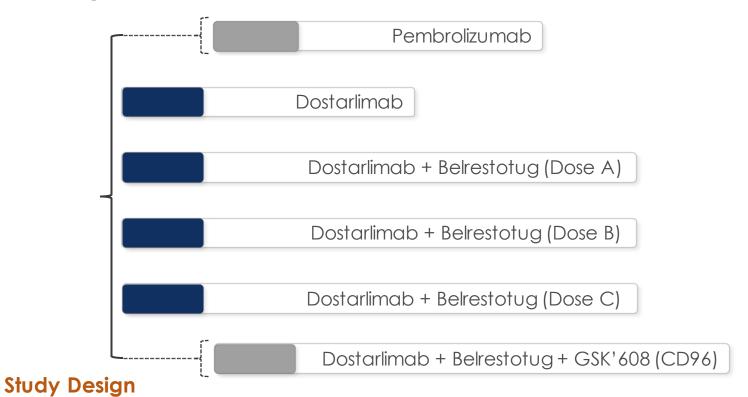
- Validated TIGIT as a target with mOS extended by ~6 months
- 2. Robust study design could provide meaningful efficacy and safety evaluation
- Incorporation of pembrolizumab as SoC control arm

# GALAXIES Lung-201 - Phase 2 in 1L NSCLC

The largest TIGIT Phase 2 in 1L NSCLC

IV Infusion

**Delivery** 





Estimated Enrollment

**Status** Enrolling **Objectives** Evaluate Belrest ot ug + Dost arlimab safety, efficacy, PK/PD

Masking Open label Primary Endpoint ORR

PDL1 Expression ≥50% Secondary Endpoint PFS, OS, DOR

**Lines of Therapy** No prior systemic therapy **Clinical Trials Listing** NCT05565378

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## 1L NSCLC: Building A Meaningful Position

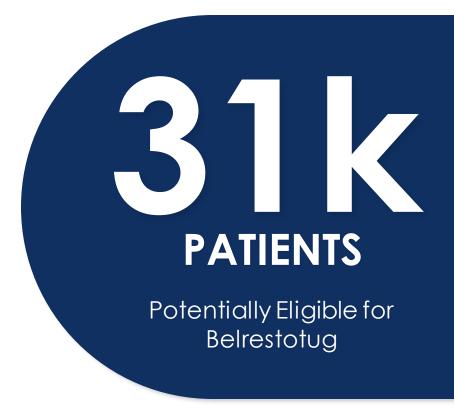
THERAPEUTICS

Evolving competitive landscape favoring a high-quality TIGIT:PD-1 doublet

**Strong scientific rationale** with high levels of TIGIT<sup>+</sup> Tregs, high infiltration of T cells, and highly amenable to IO therapies

**The right Phase 3 strategy** with right dose, right combination, right trial design, and right commercial approach

1L NSCLC launch point and clinical POC enables future exploration of other NSCLC settings and indications beyond lung



Source: Kantar, internal iTeos analysis

#### GALAXIES H&N-202: Phase 2 in 1L HNSCC





Study Design Estimated Enrollment 360

Status Enrolling Objectives Evaluate antitumor activity, safety of Dost arlimab + novel 10s

Masking Open label Primary Endpoint ORR

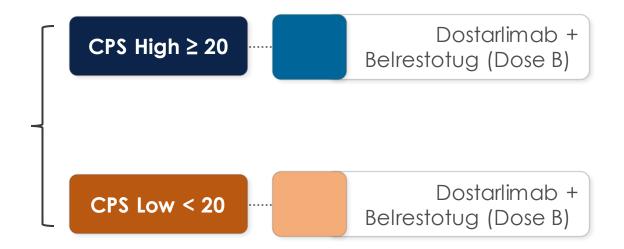
PDL1 Expression PDL1+ Secondary Endpoint PFS, OS, DOR

**Lines of Therapy** No prior systemic therapy **Clinical Trials Listing** NCT06062420

Delivery IVInfusion

### TIG-006 – Phase 2 in 1L HNSCC PDL1High/Low





Study Design	Estimated Enrollment	80
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**Objectives** Status Enrolling Evaluate Belrest otug + Dost arlimab in two CPS populations Masking Open label **Primary Endpoint** ORR PDL1 Expression Secondary Endpoint PFS, OS, DOR PDI1+ **Lines of Therapy** Clinical Trials Listing NCT05060432 No prior systemic therapy Delivery IV Infusion

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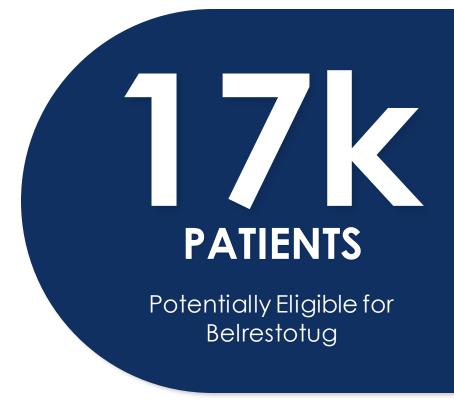
# 1L HNSCC: Potential First-to-Market Opportunity



Under-served population with strong biological rationale seeking advances

**Strong scientific rationale** with high levels of TIGIT<sup>+</sup> Tregs, high infiltration of T cells and the indication being amenable to PD-1 therapy

**Significant market opportunity** due to no ongoing Phase 3 studies, potential to be first-to-market, and the opportunity to expand to the locally advanced setting



Source: Kantar, internal iTeos analysis

# Belrestotug + Dostarlimab Are Uniquely Positioned to Fully Exploit TIGIT Pathway





# Unique Combinations







#### **Foundation**

1L NSCLC to serve as launch point for TIGIT:PD-1 doublet

#### **Expansion**

Target clinically validated indications and improve tumor activity with TIGIT:PD-1 doublet

#### Design

Utilize doublets beyond PD-1 and unique triplets to target complementary mechanisms or drugs limited by exhaustion

#### **Implementation**

Integrate promising TIGIT biomarker to precisely target sensitive indications and subpopulations

# The Right Deal & The Right Partner

Data-driven approach to unlock potential of high-quality regimens



#### **Success Factors**











**Payments** \$625M upfront, up to \$1.45B milestones



#### **Territories**

**US**: co-commercialization and **50/50 profit share** 

**Ex-US**: double digit royalties **up to 20%** 



**Developmental expenses 40%** iTeos / **60%** GSK

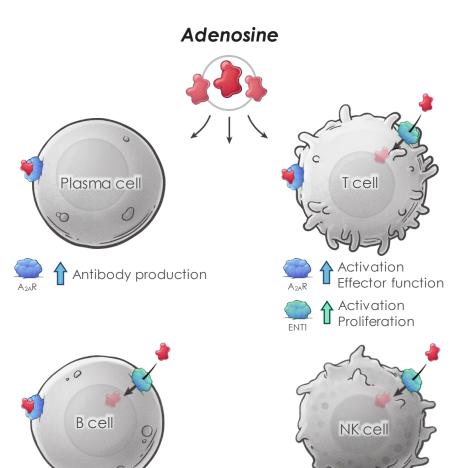


# **Adenosine Pathway**

Unlocking one of the most promising targets responsible for immunosuppression

# Addressing The Critical Adenosine Pathway Issue: Adenosine Inhibits Immune Cell Activity + Proliferation





Activation

#### **Inupadenant: Best-in-Class Approach**

- Targets  $A_{2A}R$ , restoring immune cell activity, specifically plasma cell antibody production
- First and only  $A_{2A}R$  antagonist to maintain activity at high adenosine concentrations

#### **EOS-984**: First-in-Class Approach

- Targets ENT1, a major adenosine transporter involved in T cell expansion, effector function, and survival
- Potential to restore T cell proliferation in hostile TME

# Inupadenant: A Class of Its Own

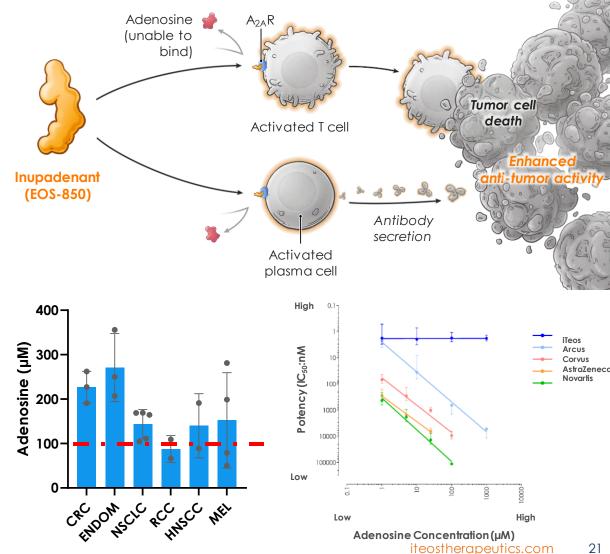
Best-in-class, highly selective  $A_{2A}R$  antagonist optimized for hostile solid TME

#### Targeting $A_{2A}R$

- A<sub>2A</sub>R activation by adenosine suppresses immune cell responses, inhibiting anti-tumor response
- Inupadenant targets A<sub>2A</sub>R, the final endpoint of the adenosine production pathway, circumventing the multiple ways adenosine is created

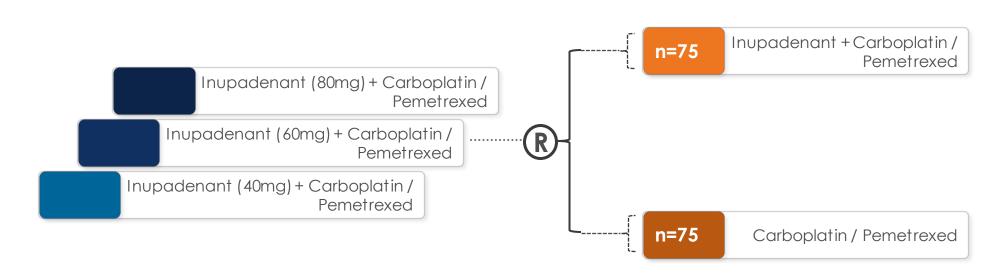
#### The Insurmountable Profile of Inupadenant

- <u>First company</u> to demonstrate TME adenosine concentration is supraphysiological and varies depending on indication
- First and only A<sub>2A</sub>R antagonist to maintain activity at high adenosine concentrations



# A2A-005: Phase 2 in 2L NSCLC (Post-IO) Chemo-Naïve





#### Key

R Subjects Randomization

#### Study Design Estimated Enrollment 192

Status **Objectives** Enrolling Evaluate Clinical Benefit of Inupadenant + Chemotherapy Masking **Primary Endpoint** Double Blind ORR PDL1 Expression **Secondary Endpoint PFS, OS, DOR** PDL1+ (all %) **Lines of Therapy** Clinical Trials Listing NCT05403385 1: PD-1 Inhibitors **Delivery** Oral

# Inupadenant Counteracts Chemotherapy's Key Downfall



2L NSCLC is an under-served population with strong biological rationale seeking advances

Chemotherapy increases adenosine levels via cell death, hindering the immune system and plasma cell activity

Inupadenant maintains potency + function at high adenosine levels, potentially enhancing chemotherapy therapeutic response

Currently only clinical trial in 2L NSCLC platinum-naïve setting



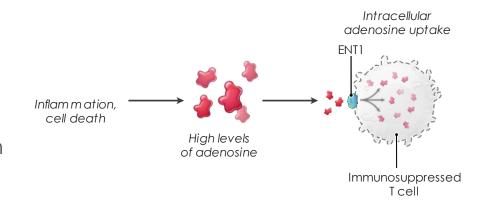
Source: Kantar, internal iTeos analysis

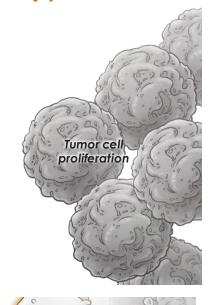
# EOS-984: Enhancing T Cell Proliferation in the Hostile TME

One of the most meaningful discoveries in the adenosine pathway

#### The Role of ENT1

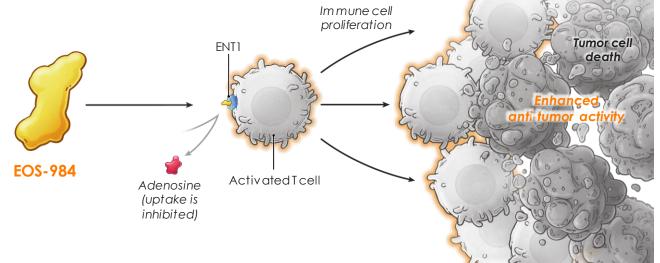
- Dominant transporter of adenosine on lymphocytes effecting:
  - T cell metabolism
- T cell effector function
- T cell expansion
- T cell survival





#### The Opportunity to Revive T Cell Proliferation

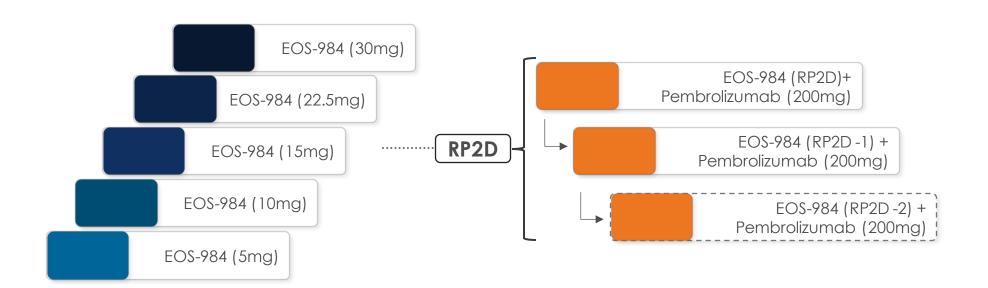
- <u>First company</u> to understand how adenosine transports into T cells and inhibits proliferation
- EOS-984 offers large combination opportunity broadly across cancer therapies



#### EOS-984: Phase 1 in Advanced Solid Tumors



Evaluation of target engagement and impact on T cells in TME



#### **Study Design**

Status Enrolling Objectives Evaluate Safety/Tolerability of EOS-984 as a Monotherapy and in

Masking Open Label Combination with Pembrolizumab

PDL1 Expression PDL1+ (all %) Primary Endpoint Safety/Tolerability, PK/PD

Lines of Therapy All-comers Secondary Endpoint ORR, PFS, OS, DOR

**Delivery** Oral

### 2024: Pivotal Year of Data Generation

Innovative portfolio leveraging deep tumorimmunology expertise



#### **TIGIT**

#### 1L NSCLC

(Phase 2 GALAXIES LUNG-201)

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#### 1L HNSCC

(Phase 2 TIG-006)

### **Adenosine Pathway**

A<sub>2A</sub>R - 2L NSCLC

(Phase 2 A2A-005)

ENT1 - MOA

............

(EOS-984 Preclinical)

............

ENT1 - Advanced Malignancies

(EOS-984 Phase 1)

## Funded Through Significant Milestones

As of 3Q23

\$645M

In cash, cash equivalents and short-term investments

Runway through 2026



# Cancer Immunotherapies by design™

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January 2024