

# Cancer Immunotherapies by design™

Nasdaq: ITOS

August 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 EOS-984 to revive T cell proliferation and offer large combination opportunity broadly across cancer therapies; the expectation that 2024 will be a defining year for iTeos; our clinical, data generation and data presentation plans for 2024, including having data readouts from GALAXIES Lung-201, TIG-006 HNSCC, A2A-005 and EOS-984; our market opportunities and number of patients potentially eligible for our product candidates; the potential benefits of our collaboration with GSK; intentions around trial enrollment and recruitment; and our expected cash runway through 2027.

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 or interim data from a clinical trial may change as more patient data become available and are subject to audit verification procedures; 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 iTeos' Quarterly Report on Form 10-Q for the period ended June 30, 2024 filed with the Securities and Exchange Commission (SEC) as well as other SEC filings made by the Company which you are encouraged to review. Statements regarding the Company's cash runway do not indicate if and when the Company may access the capital markets.

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 undue 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 2H24

#### **Unlocking Adenosine Pathway**

2

Two Data Readouts Anticipated in 2H24

#### **Funded Through 2027**

3

~\$714M in pro forma cash as of June 30, 2024

#### 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                    |             |         | GALAXIES Lu | ng-301                        | Enrolling             |
| + dostarlimab   1L NSCLC PDL1high                     |             | GALAXIE | S Lung-201  |                               | Data Anticipated 2H24 |
| + dostarlimab   1L HNSCC PDL1high/low                 |             |         | TIG-006     |                               | Data Anticipated 2H24 |
| + dostarlimab + CD96   1L HNSCC PDL1high              |             | GALAXIE | S H&N-202   |                               | Enrolling             |
| + dostarlimab + chemotherapy   1L mNSCLC              |             | TIG-006 |             |                               | Enrollment Complete   |
| + dostarlimab + CD96   Advanced Malignancies          | NCT0        | 3739710 |             |                               | Enrollment Complete   |
| + dostarlimab + PVRIG   Advanced Malignancies         | NCT0        | 5277051 |             |                               | Enrollment Complete   |
| Inupadenant: Small molecule targeting A <sub>2A</sub> | receptor    |         |             |                               | iTEOS                 |
| + chemotherapy   Post-IO Chemo-naïve NSCLC            | A2A-005     |         |             | Data Anticipated<br>Late 2024 |                       |
| EOS-984: Small molecule targeting ENT1                |             |         |             |                               | iTEOS                 |
| Monotherapy   Advanced Malignancies                   |             |         |             |                               | Data Anticipated 2H24 |



## Belrestotug

EOS-448 / GSK4428859A

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



We Hold An

### Advantageous Field Position

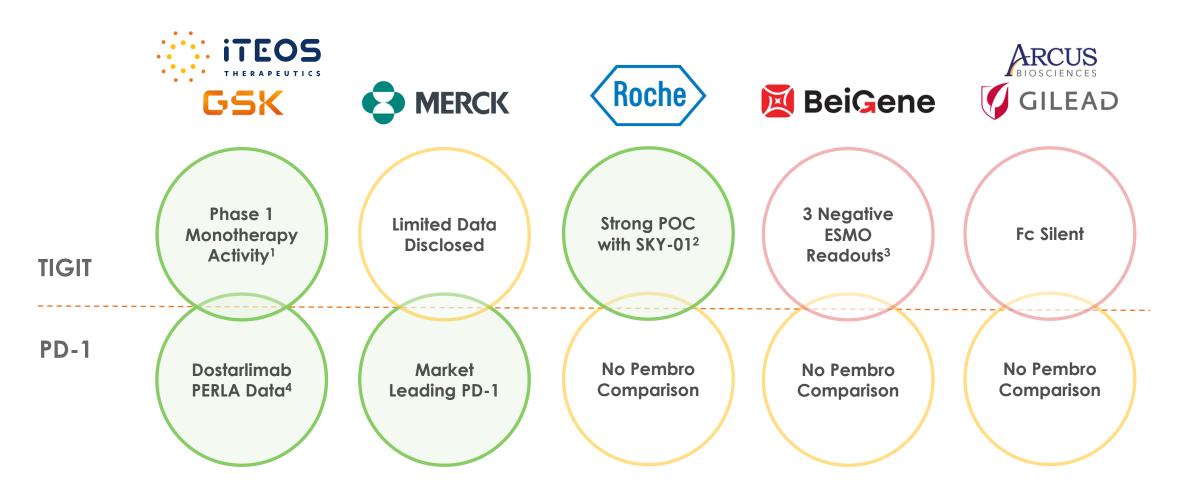
Significant momentum in 2023



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



Belrestotug + dostarlimab represent potentially 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 - AdvanTIG-203, AdvanTIG-206, AdvanTIG-202

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

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



Unique Epitope Binding

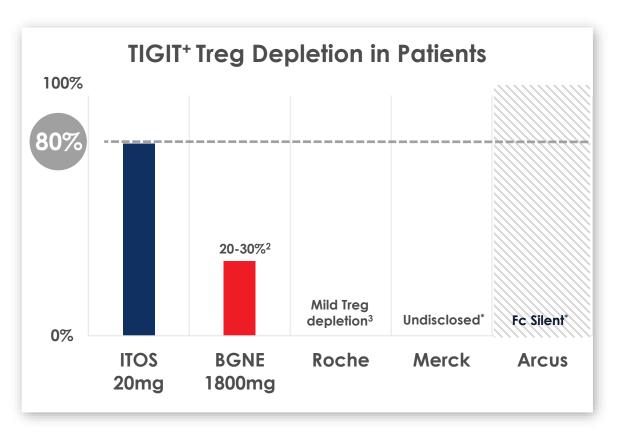
#### **High Affinity + Potency**

First and only TIGIT with proven

#### Treg depletion at all doses

Only TIGIT to Demonstrate Phase 1

Monotherapy Activity<sup>1</sup>



<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

3. doi: 10.1038/s41586-024-07121-9

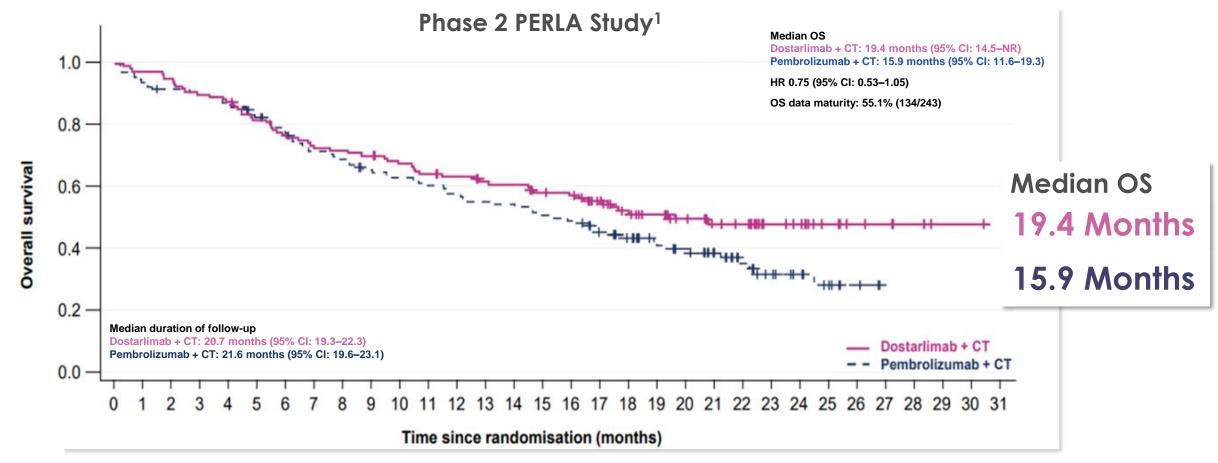
<sup>.</sup> iTeos AACR 2021

doi: 10.1136/iitc-2022-SITC2022.0768

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



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



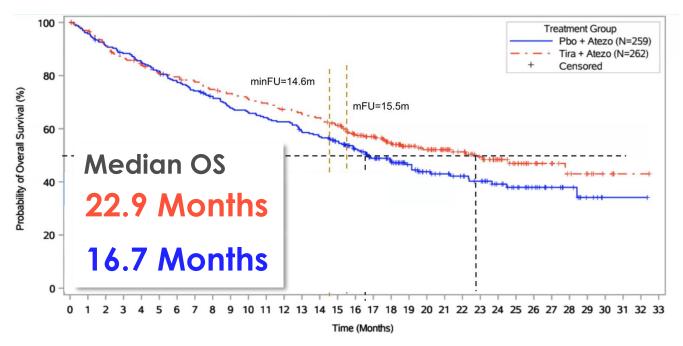
<sup>&</sup>lt;sup>1</sup>Phase 2 GSK-sponsored PERLA study in 1L NSCLC. Peters S, et al. Annals of Oncology (2023) 34 (suppl\_2): \$1254-\$1335. 10.1016/annonc/annonc1358. NB: Safety profiles in the secondary analysis of OS were similar and consistent with those previously reported in the primary analysis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

#### SKY-01: Meaningful Separation of Curves Validates TIGIT



Potential for enhancement of quality of components and clinical trial design



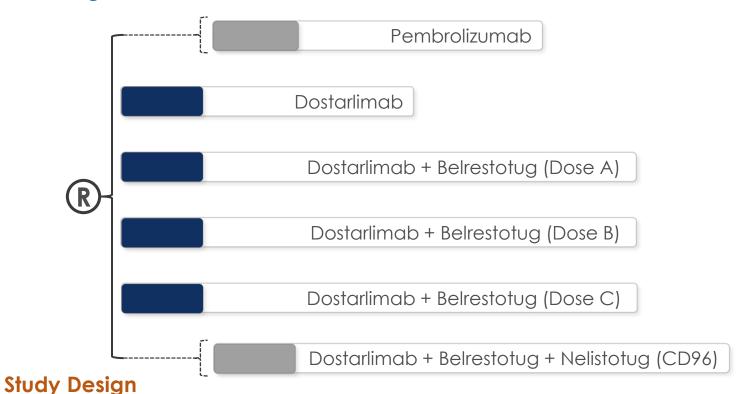


#### **Key Insights**

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

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

The largest TIGIT Phase 2 in 1L NSCLC





Key

R Subjects Randomization

**Estimated Enrollment** 

300

StatusEnrollingObjectivesEvaluate belrestotug + dostarlimab safety, efficacy, PK/PDMaskingOpen labelPrimary EndpointORR

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

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

**Delivery** IV Infusion

#### GALAXIES Lung-301 - Phase 3 in 1L NSCLC







Study Design Estimated Enrollment 1,000

Status Enrolling Objectives Evaluate

Evaluate belrestotug + dostarlimab safety, efficacy vs

placebo + pembrolizumab

**Primary Endpoint** PFS, OS

**Secondary Endpoint** ORR, MRR, DOR

PDL1 Expression ≥50%

Masking

**Lines of Therapy** No prior systemic therapy

Double-blind

**Delivery** IV Infusion

#### 1L NSCLC: Building A Meaningful Position

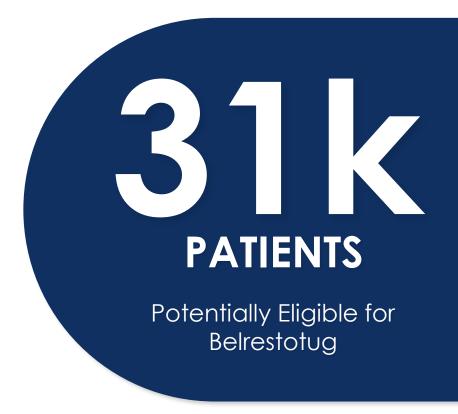


Evolving competitive landscape favoring a high-quality TIGIT:PD-1 doublet

**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 strategic launch point and clinical POC enables future exploration of other NSCLC settings and indications beyond lung



Source: Kantar, internal iTeos analysis

#### PD-1 Therapy Remains Primary Treatment for 1L NSCLC Patients

iTEOS THERAPEUTICS

No major shift in 1L NSCLC treatment trend in last two years

Patients that Receive PD-(L)1 without Chemotherapy in 1L NSCLC PD-L1 High in US

- PD-1 treatment alone remains SOC in 1L NSCLC PD-L1 high patients, followed by platinum-doublet in 2L NSCLC<sup>1,2,3</sup>
  - o PD-1 + chemo failed to improve OS vs PD-1 alone
  - PD-1 alone viewed as sufficient for most patients while reducing toxicity
  - Chemotherapy option still available in 2L NSCLC
- PD-1 + chemo typically used for high burden disease to provide rapid control/symptom relief<sup>2</sup>
- No difference in mOS or rwPFS between PD-1 alone vs chemo + PD-1 in retrospective cohort study examining 1L NSCLC treatment<sup>3</sup>

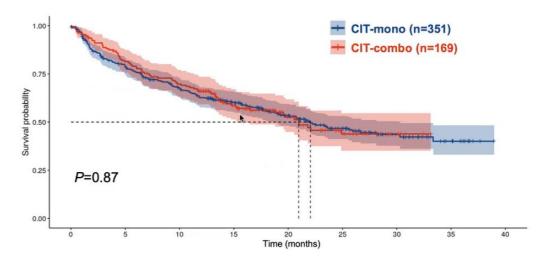
72%

**Ipsos Oncology Monitor**Chart data for 12 months ending June 2023

68%

iTeos US Oncologist Survey
(n=50: 16 academic, 34 community: 124 patient charts)

Effectiveness of PD-(L)1 Inhibitors Alone or in Combination with Platinum Doublet Chemo in 1L NSCLC with PD-L1 High Expression Using Real World Data<sup>2</sup>



<sup>1.</sup> Ipsos Oncology Monitor

<sup>2.</sup> iTeos US Oncologist Survey

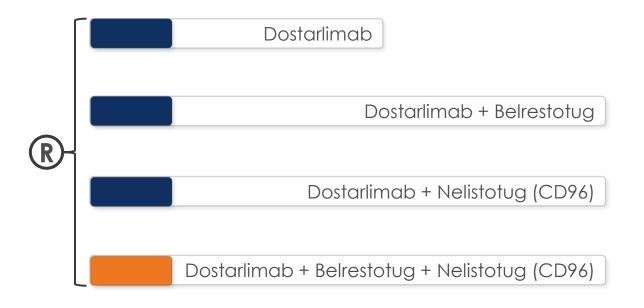
<sup>3.</sup> ESMO Virtual Plenary: Effectiveness of PD-(L)1 Inhibitors Alone or in Combination with Platinum Doublet Chemo in 1L NSCLC with PD-L1 High Expression Using Real World Data

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









Study Design Estimated Enrollment 360

Status Enrolling Objectives Evaluate antitumor activity, safety of dostarlimab + novel IOs

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 IV Infusion

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







Study Design Estimated Enrollment 40

| Status           | Enrolling                 | Objectives                     | Evaluate belrestotug + dostarlimab in two CPS populations |
|------------------|---------------------------|--------------------------------|---|
| Masking          | Open label                | <b>Primary Endpoint</b>        | ORR   |
| PDL1 Expression  | PDL1+                     | Secondary Endpoint             | PFS, OS, DOR  |
| Lines of Therapy | No prior systemic therapy | <b>Clinical Trials Listing</b> | NCT05060432   |
| Delivery         | IV Infusion               |                                |   |

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



Under-served market 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 Novel Biomarker





#### **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

#### An Empowering, Strategic Collaboration with GSK

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



#### **Success Factors**







Partner





**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

## Supraphysiological Adenosine Synthesis in TME Broadly Suppresses Immune System

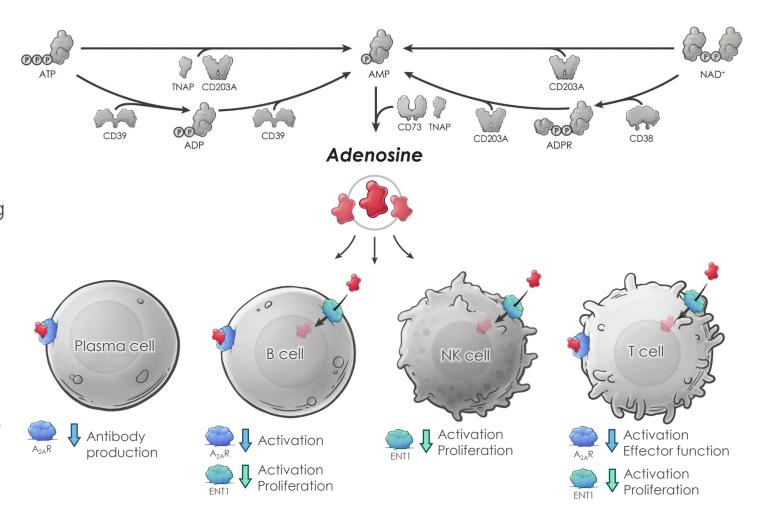


ATP/Adenosine created in response to proinflammatory stimuli, like cell stress from hypoxia and cell necrosis in the tumor

Difficult to stop adenosine production due to multiple mechanisms involved, including enzymes CD39 and CD73

**A<sub>2A</sub>R engagement** with adenosine impairs multiple immune cell activities

**ENT1 engagement** with adenosine impairs immune cell metabolism, effector function, and proliferation



#### Inupadenant: A Class of Its Own



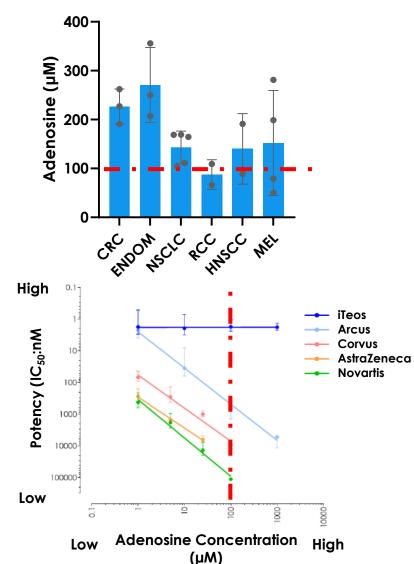
Best-in-class, highly selective  $A_{2A}R$  antagonist optimized for hostile solid TME

#### 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

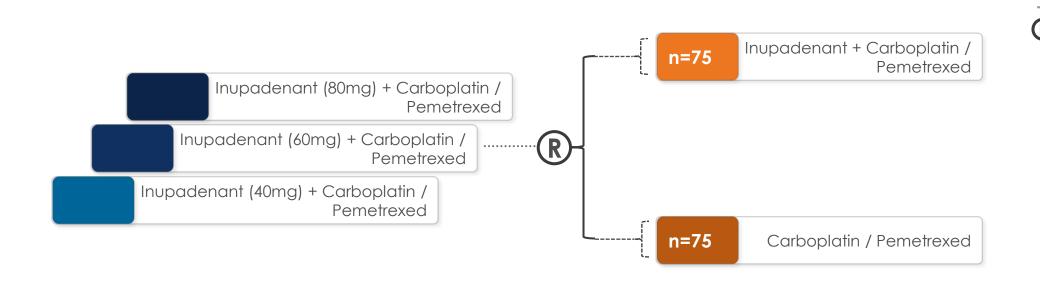
#### The Insurmountable Profile of Inupadenant

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



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





R Subjects Randomization

#### Study Design Estimated Enrollment 192

| Status           | Enrolling          | Objectives              | Evaluate clinical benefit of inupadenant + chemotherapy |
|------------------|--------------------|-------------------------|---|
| Masking          | Double Blind       | <b>Primary Endpoint</b> | ORR   |
| PDL1 Expression  | PDL1+ (all %)      | Secondary Endpoint      | PFS, OS, DOR  |
| Lines of Therapy | 1; PD-1 Inhibitors | Clinical Trials Listing | NCT05403385   |
| 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 <u>only</u> clinical trial in 2L NSCLC platinum-naïve setting



Source: Kantar, internal iTeos analysis

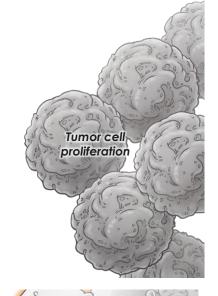
#### EOS-984: Enhancing T Cell Proliferation in the Hostile TME

One of the most meaningful discoveries in the adenosine pathway

#### The Role of ENT1

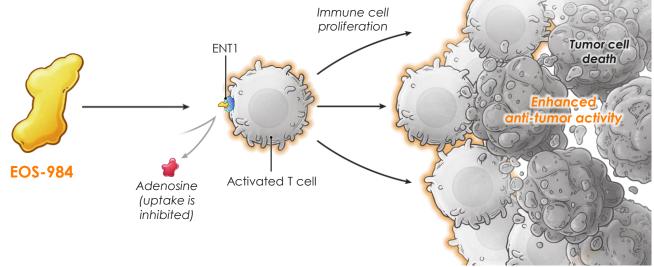
- Dominant transporter of adenosine on lymphocytes effecting:
  - T cell metabolism
- T cell effector function
- T cell expansion
- T cell survival

# Intracellular adenosine uptake ENT1 High levels of adenosine Immunosuppressed T cell



#### 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

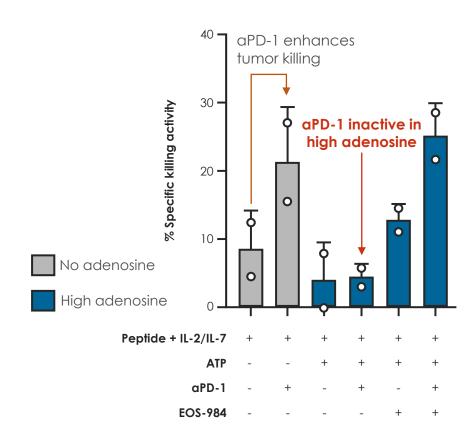


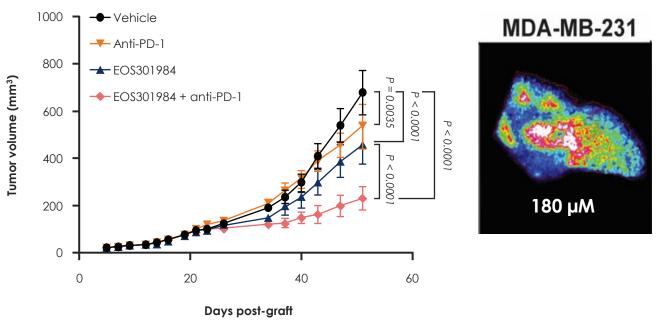
#### Anti-PD-1 Activity Enhanced by Restoration of T Cell Proliferation by EOS-984

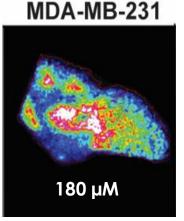


EOS-984 + aPD-1 combination maximizes tumor killing by functional memory T cells

#### **Humanized TNBC model (MDA-MB-231)** containing high adenosine



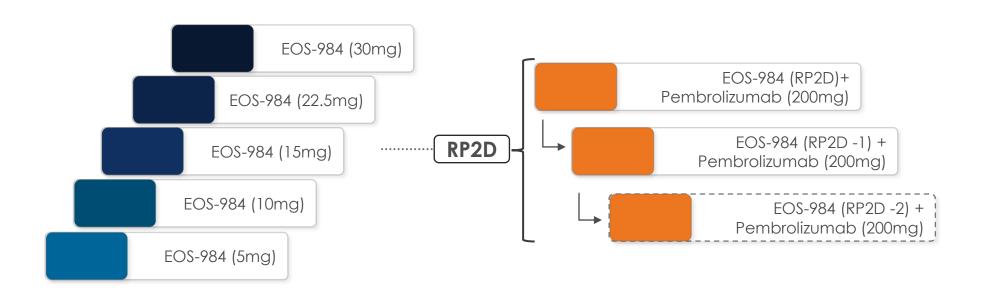




#### EOS-984: Phase 1 in Advanced Solid Tumors



Evaluation of target engagement and impact on T cells in TME



#### **Study Design**

Status **Objectives** Enrolling Evaluate safety/tolerability of EOS-984 as a monotherapy and in

combination with pembrolizumab Masking Open Label

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

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

Oral

Delivery

#### 2024: Pivotal Year of Data Generation

Innovative portfolio leveraging deep tumor immunology expertise



#### **TIGIT**

1L NSCLC

(Phase 2 GALAXIES LUNG-201)

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

1L HNSCC

(Phase 2 TIG-006)

#### **Adenosine Pathway**

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

(Phase 2 A2A-005)

ENT1 - MOA

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

(EOS-984 Preclinical)

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ENT1 - Advanced Malignancies

(EOS-984 Phase 1)

#### Funded Through Significant Milestones

As of June 30, 2024

~\$714M

Pro forma cash, cash equivalents and short-term investments

Runway through 2027



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August 2024