



Gamma-delta T-cell engagers for the development of next- generation cancer therapeutics

Corporate Presentation
January 2024

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Investment Highlights



Proprietary Gammabody® platform

Platform selectively and conditionally activates Vδ2¹ T-cells upon cross-linking with a TAA.
Anticipate reduced risk of high-grade CRS with preferential tumor recognition expected to limit on-target/off-tumor toxicities.
Growing pipeline of bispecific T-cell engagers led by clinical-stage prostate cancer program



Promising Lead for mCRPC

Phase 1 data (ASCO GU 2023) showed good tolerability, with continued evidence of antitumor activity
Study update planned for Q3 2024



Growing Pipeline

IND/CTA submission for LAVA-1266 (CD123) for hematologic malignancies: expected in H1 2024



Validating Strategic Partners

Seagen/PFE worldwide license agreement for SGN-EGFRd2 (LAVA-1223) for EGFR+ tumors, in Phase 1
Janssen collaboration has selected a lead candidate, in preclinical development



Strong Team, IP and Cash Position

Experienced management team, with a strong IP portfolio and a cash balance of \$105 million², with an expected runway into 2026

Gammabody® Pipeline: Potential in Hematologic Malignancies and Solid Tumor Indications

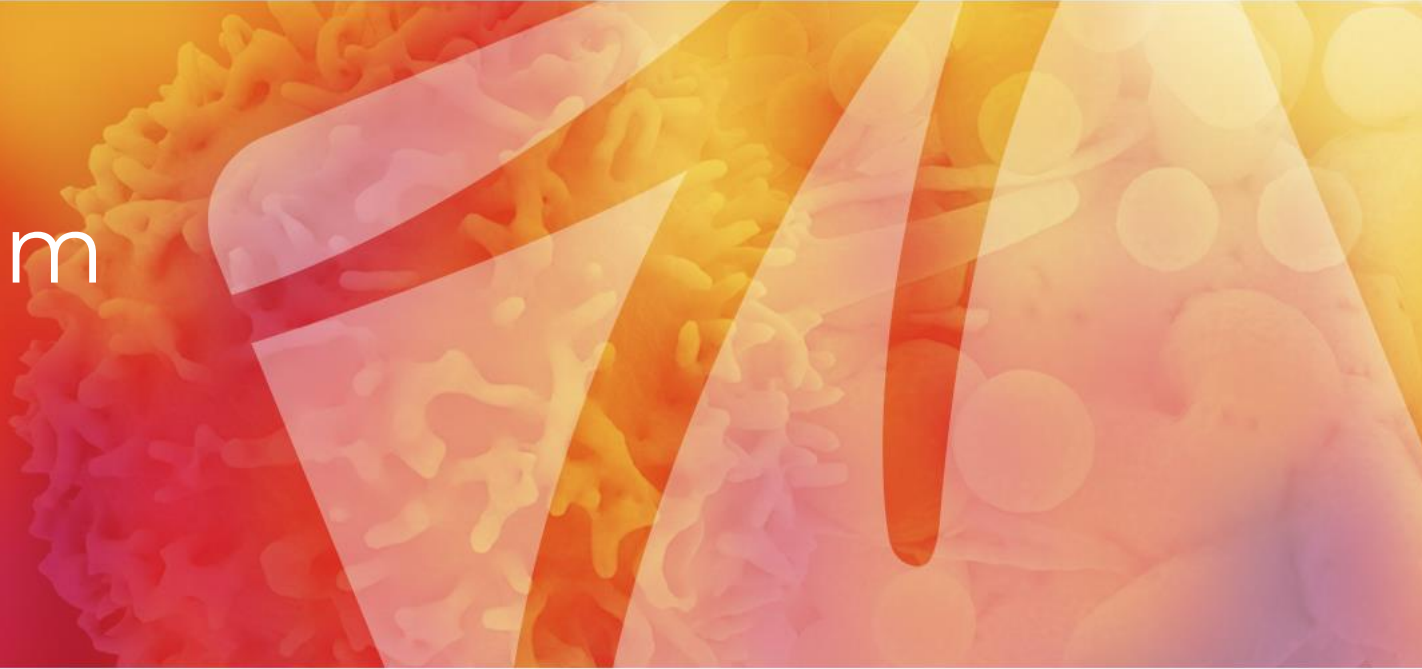
			Discovery	Preclinical	Phase 1	Phase 2
Internal Pipeline						
LAVA-1207	PSMA	mCRPC				
LAVA-1266	CD123	Hematologic Malignancies				
LAVA-1427 LAVA-1433	Undisclosed Undisclosed	Undisclosed Undisclosed				
Strategic Partnerships						
SGN-EGFRd2	EGFR	Solid Tumors				
Janssen	Undisclosed	Undisclosed				
Out-licensing Candidates						
LAVA-051	CD1d	Hematologic Malignancies				

■ Hematologic malignancy ■ Solid Tumor ■ Undisclosed

4 | 1. As of December 14, 2023, Pfizer completed its acquisition of Seagen. PSMA: prostate-specific membrane antigen; EGFR: epidermal growth factor receptor; mCRPC: metastatic castration-resistant prostate cancer

Gammabody® Platform

Overview



Proprietary Gammabody® Platform Induces Potent Killing of Tumor Cells



Targeting Vδ2 T-cells

Activated Vδ2 T-cells have a natural ability to recognize and selectively kill tumor cells and trigger a cascade of anti-cancer immune responses. Positively associated with patient outcomes



Bispecific antibodies designed to engage Vδ2 T-cells and widen the therapeutic window

Gammabody® platform selectively and conditionally activates Vδ2 T-cells upon cross-linking with a tumor-associated antigen. Anticipated reduced risk of high-grade CRS with preferential tumor recognition expected to limit on-target/off-tumor toxicities.



Designed for efficient performance

Humanized single-domain antibodies (VHHs) linked to a silent Fc for half-life extension
Safety profile could support combination therapy



Supported by strong IP

Potential patent coverage ranging from 2035 to 2041
Platform coverage as well as product-specific coverage for assets

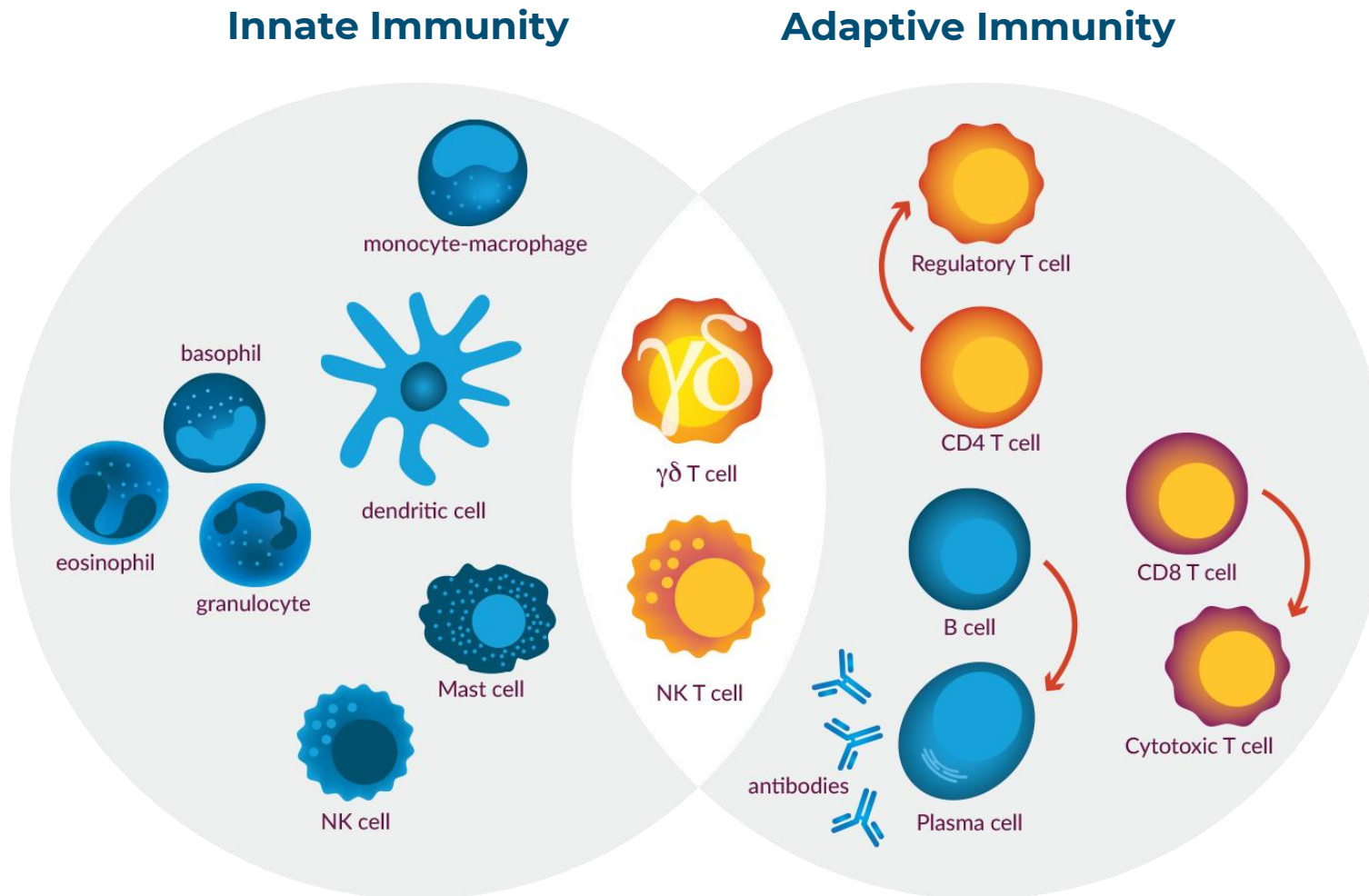


Broad opportunity in solid tumors, hematologic malignancies

T-cell engagers represent a growing therapeutic class
Analysis of Vδ2 T-cell distribution outlines a road map of target indications

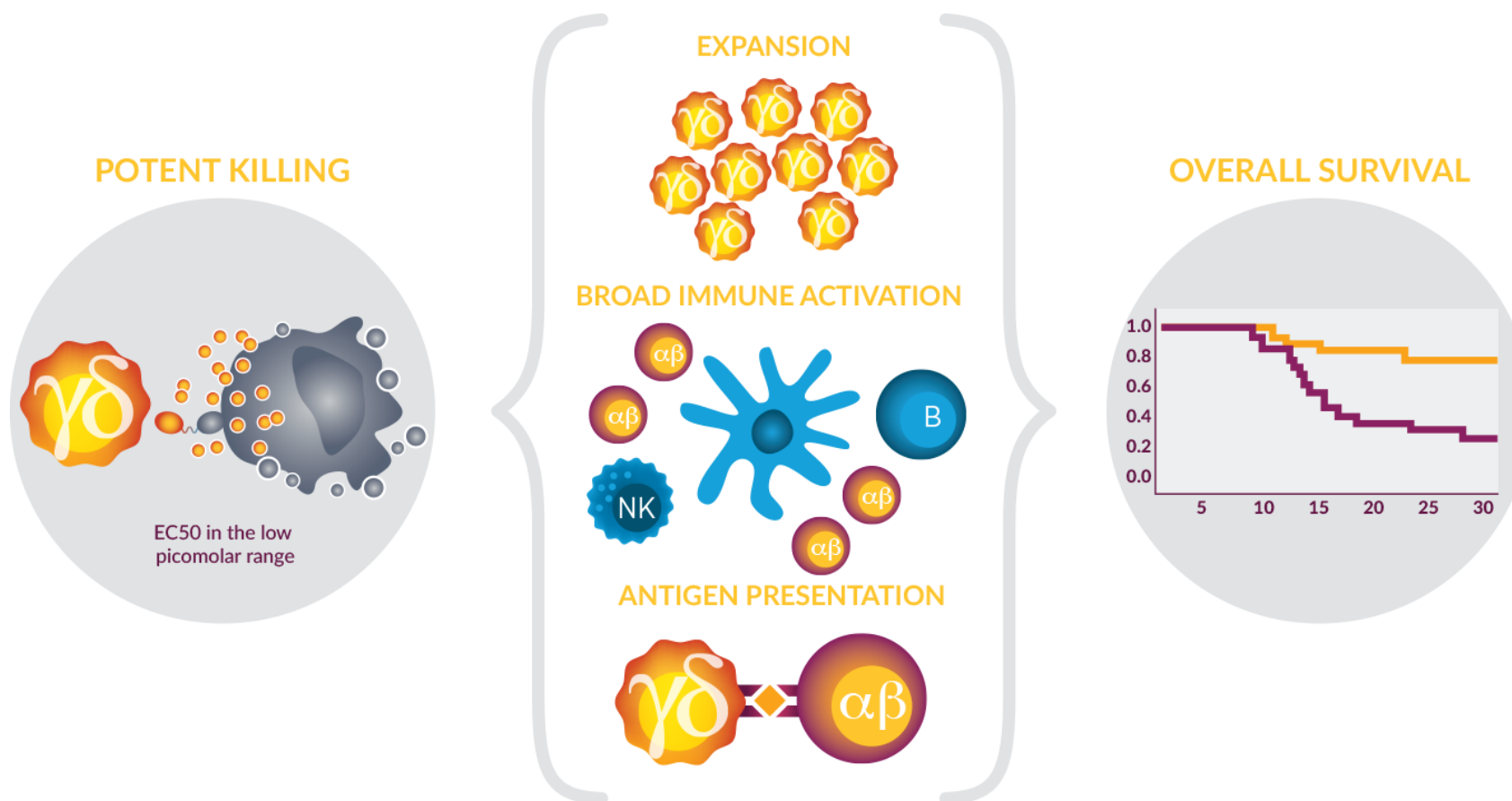
V δ 2 T-Cells

Positioned at the interface between innate and adaptive immunity



- Largest $\gamma\delta$ T-cell subset in blood: (~90-95% of total $\gamma\delta$ T- cells)
- Natural ability to recognize and kill tumor cells
- Presence of $\gamma\delta$ T-cells associated with improved outcomes in cancer patients
- Recognize tumors through phosphoantigen-BTN2A1/3A1 complex
- Consistent pro-inflammatory cytotoxic effector T-cell population

Selective Activation of V δ 2 T-Cells has the Potential to Induce Durable Tumor Responses and Improve Patient Survival



Selective activation of V δ 2 T-cells has the potential to yield potent tumor killing and durable responses through a cascade of mechanisms that may include V δ 2 T-cell expansion, broader immune activation and antigen presentation

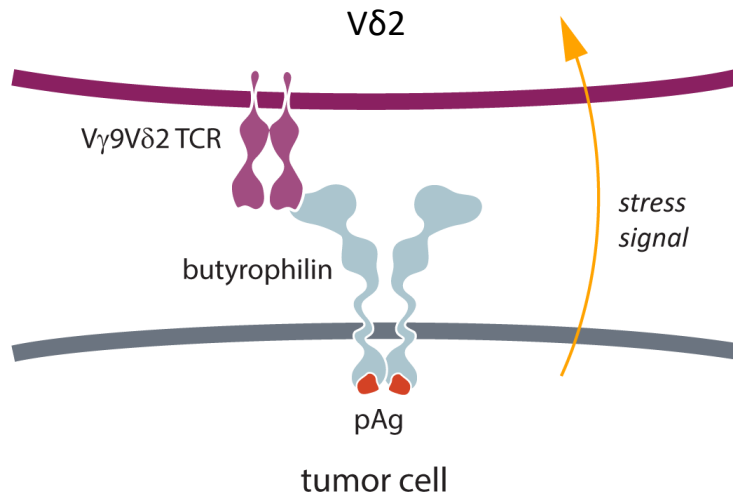
Gammabody® Platform: Specifically Directs Vδ2 T-Cells to Tumors

Highly selective platform designed to induce potent tumor killing with limited on-target off tumor toxicity

Natural Activation Mechanism¹

Vδ2 T-cells recognize stress signals

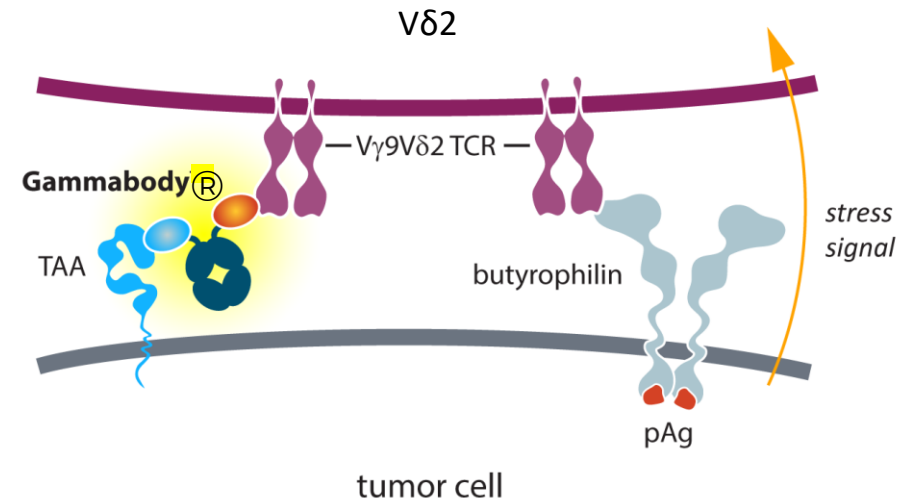
TCR interacts with pAg-butyrophilin complex



Gammabody® Provides Tumor Recognition to Trigger Vδ2 T-Cell-Mediated Immunity

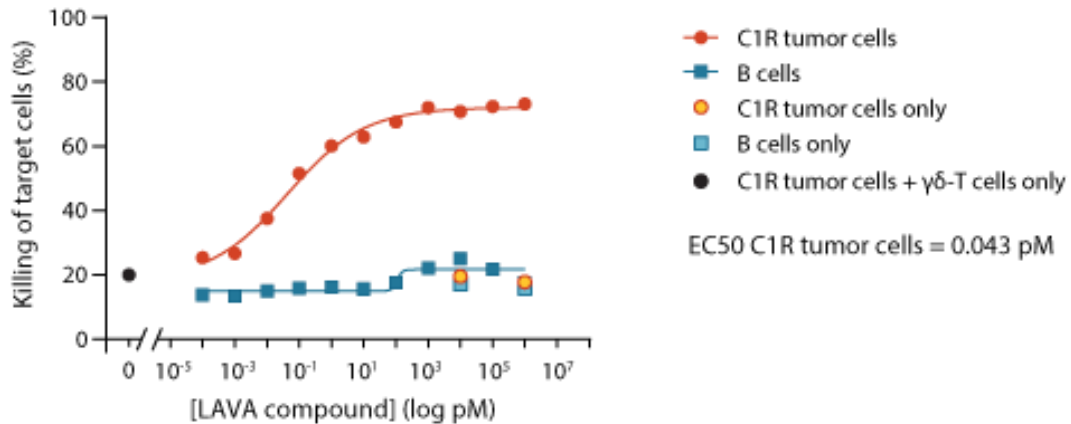
Conditionally activate Vδ2 T-cells upon crosslinking with tumor associated antigen (TAA)

Natural recognition of stress signals continues



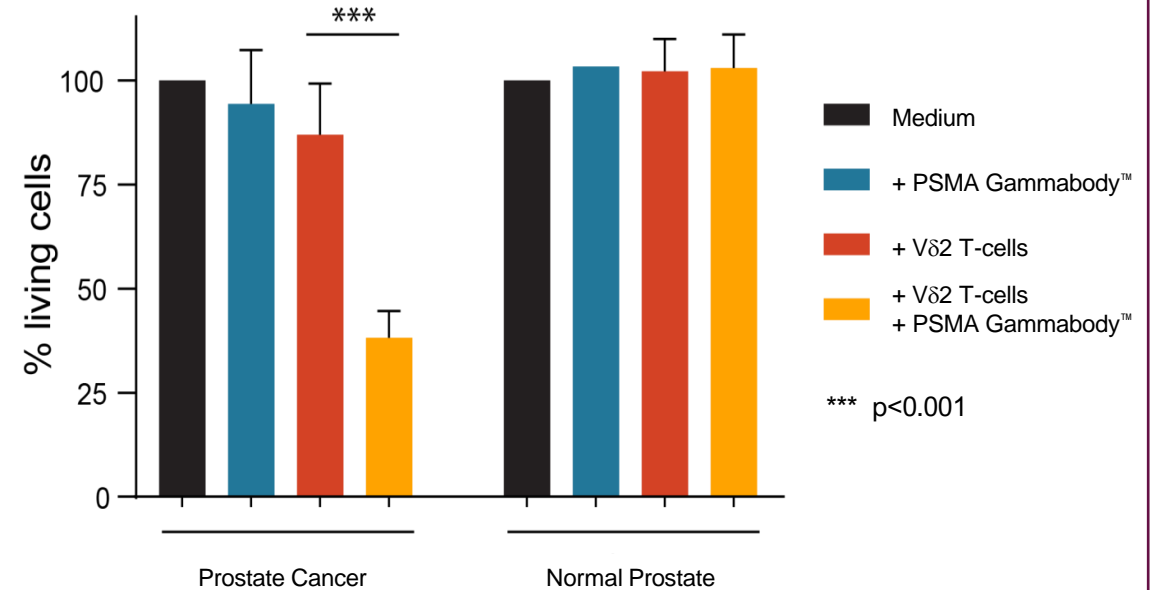
Key Features of the Gammabody® Platform

CD20 Gammabody® Mediated Killing¹



- 2:1 ratio (γδ T-cells : Target cells)
- Similar CD20 expression levels on C1R neo and B-cells

PSMA Gammabody® Mediated Killing¹



- Allow for targeting of widely expressed tumor-associated antigens
- Facilitates higher dosing potential in monotherapy
- Opportunities for combination work

LAVA-1207

Gammabody® Designed to Activate V δ 2 T-Cells by Targeting PSMA for the Treatment of mCRPC

LAVA-1207 Targets PSMA: Enrolling in Phase 1/2a Study

Update expected Q3 2024



PSMA is a clinically validated target

Highly-expressed in >90% prostate cancers¹. Higher levels negatively correlated with survival²
FDA approval of Pluvicto, a PSMA-targeted radiopharmaceutical, provides clinical validation



High unmet need

While early-stage outcomes are good, mCRPC prevalence is 50,000 in the U.S.³
With ~35,000 prostate-cancer related deaths annually in the U.S.⁴, 5-year survival for mCRPC is ~30%⁵



mCRPC is a sensible Gammabody target

Relative abundance of Vδ2s makes metastatic castration-resistant prostate cancer (mCRPC) an attractive Gammabody target



Phase 1 enrollment

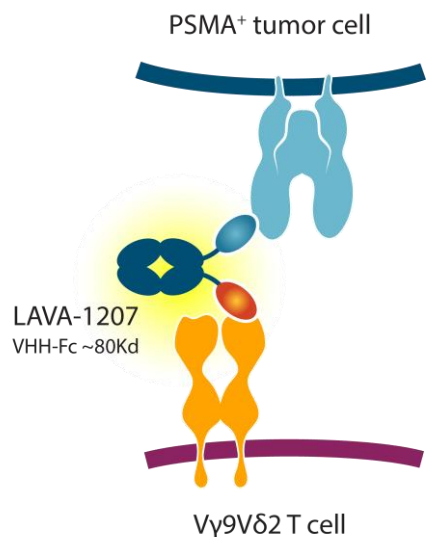
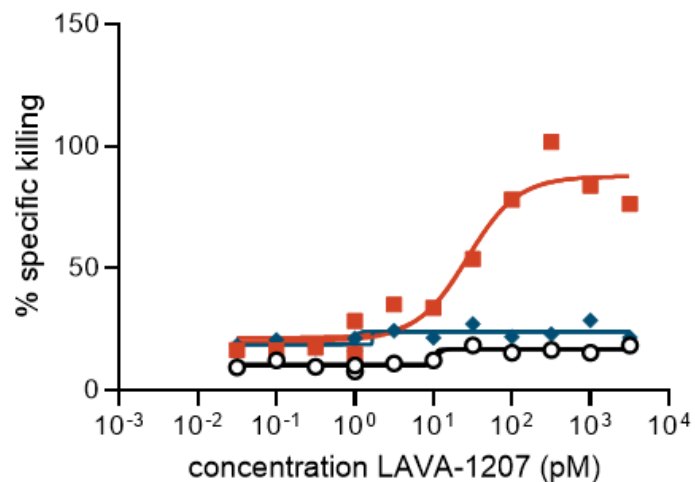
Enrollment is ongoing in the U.S. and Europe (NCT05369000)
Dosing regimen includes therapy with IL-2



Study update

Preliminary signs of clinical activity observed with disease stabilization and PSA reduction during dose escalation
Next update planned for Q3 2024

LAVA-1207: Designed to Mediate Potent Killing of PSMA-Positive Tumor Cells



Format

- Contains a Fc domain for extended plasma half-life; silenced to avoid off-target T-cell activation
- Small size (compared to regular IgG antibodies) to facilitate tumor penetration

Mechanism of Action

- Specifically directs Vδ2 T-cells to PSMA-expressing tumor cells. PSMA is a well-validated tumor target
- Mediates potent killing of PSMA-positive tumor cells
- Pre-clinical data support mechanism of action, anti-cancer activity & selectivity

Status

- Phase 1/2a trial in mCRPC; patient recruitment ongoing (NCT05369000)
- Phase 1/2a trial patient recruitment ongoing for monotherapy and IL-2 arms

Phase 1/2a Snapshot from ASCO GU 2023, from Dosing Groups 1-5¹



First 20-patients

Median patient was 68 years old, had received 4 rounds of prior therapy and was treated for 9 years
Metastasis were primarily located in bone, lymph nodes and visceral tissues



Encouraging safety profile²

No occurrence of high-grade CRS (>2). No increase in severity/frequency of TEAEs with increasing doses, or treatment discontinuations due to adverse events.



Attractive early data on pharmacokinetics

Observed linear pharmacokinetics (PK)



Attractive early data on pharmacodynamics

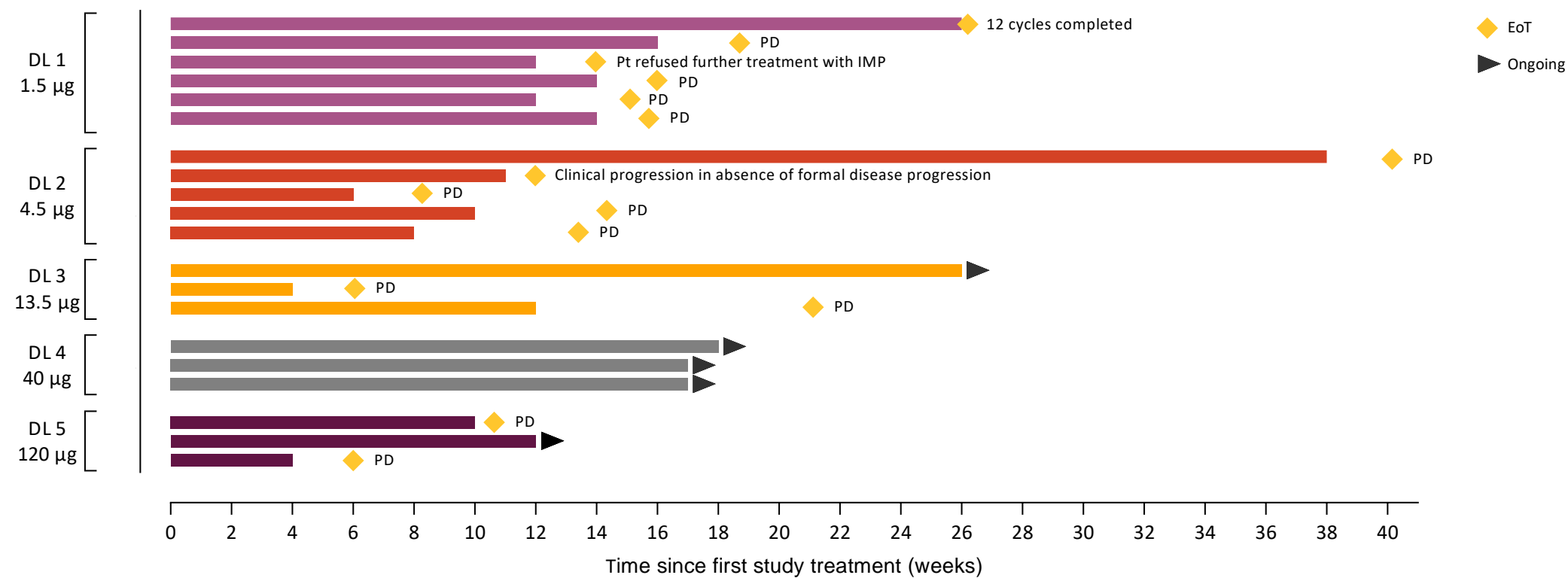
Pharmacodynamics (PD) reflects changes per MoA
Vδ2 receptor occupancy continues to increase with increasing dose



Activity and treatment duration

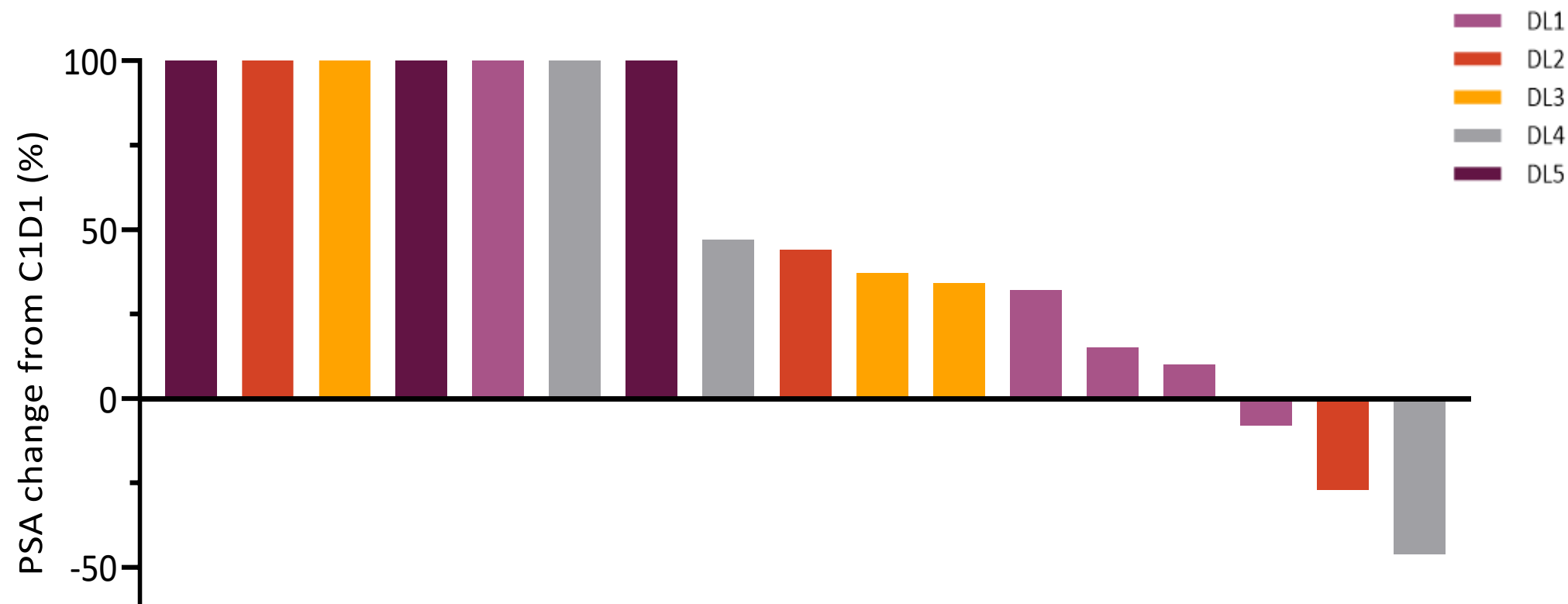
PSA reductions observed. Stable disease observed in 8 of 14 evaluable patients, as of ASCO GU 2023 data cut-off (12/8/2022)

ASCO GU 2023: Time on Treatment Shows Preliminary Signs of Antitumor Activity



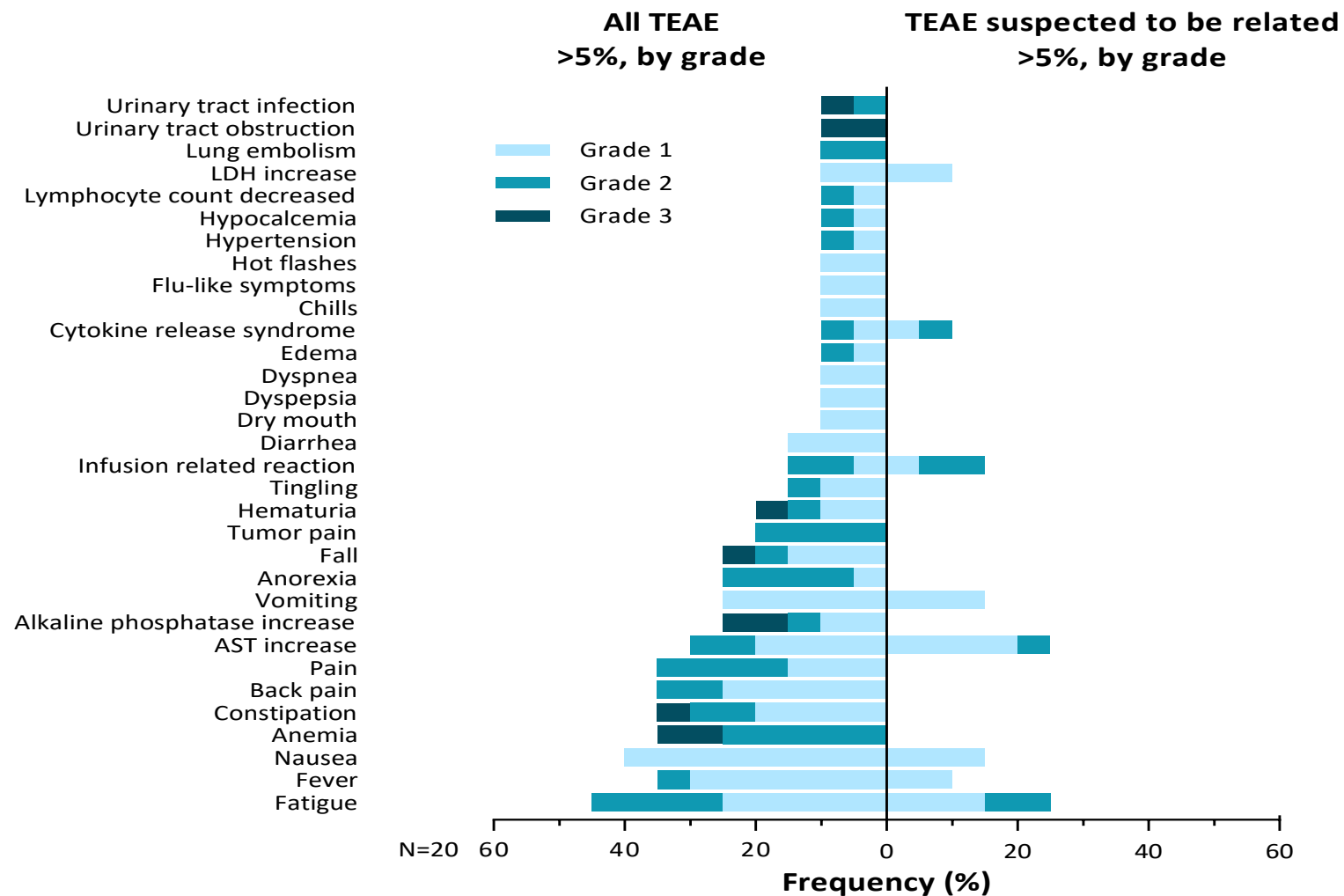
ASCO GU 2023: Best PSA Response

Continue to observe PSA reductions



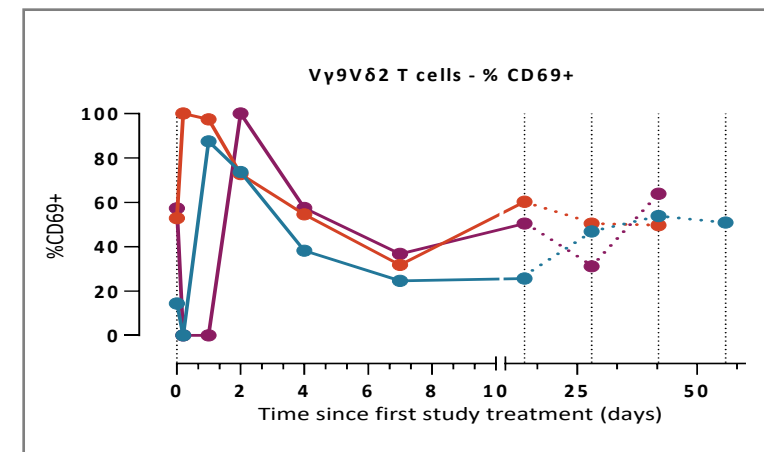
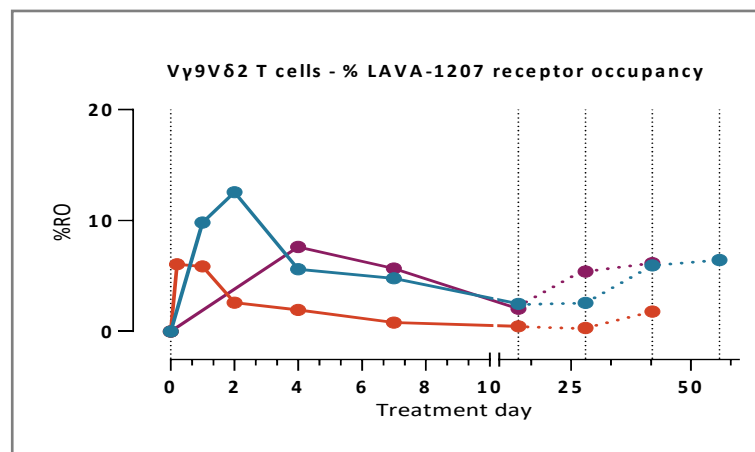
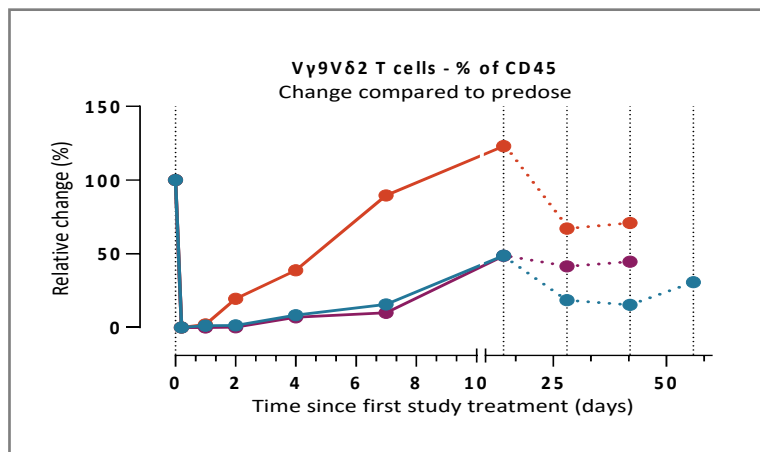
ASCO GU 2023: Initial Phase 1 Safety Data

- Favorable safety profile with no occurrence of high-grade (>2) CRS
- TEAEs that were suspected to be related were grade 1 or 2
- No increase in severity or frequency of TEAEs with increasing doses
- One grade 4 AE occurred (spinal cord compression, DL 5), which was non-related



ASCO GU 2023: Pharmacokinetics and Pharmacodynamics:

Continuing to see V δ 2 T-cell receptor occupancy with increasing doses



PK, PD Data in Keeping with MOA

- PK appears to be linear
- Pronounced drop in V δ 2 T-cell frequency 2 hr after dosing, suggesting V δ 2 T-cell re-distribution, with subsequent recovery
- V δ 2 T-cell activation markers (CD25 and CD69) upregulated following dosing
- Receptor occupancy detectable up to day 14 after EOI, with peak levels ranging from 6.1% to 12.6%

LAVA-1266

CD123 Targeting Gammabody[®] for the
Treatment of Hematologic Malignancies

LAVA-1266 Targets CD123 for AML: IND/CTA Filing Expected in H1 2024



CD123 is a clinically validated target

Over-expressed in a wide range of hematologic malignancies



Promising preclinical data

LAVA-1266 induced preferential lysis of CD123-expressing tumor cells while relatively sparing CD123-expressing normal cells



High unmet need

74,000 people living with AML in the U.S. (2020)¹
~30% 5-year mortality (adults)¹



AML is a sensible Gammabody target

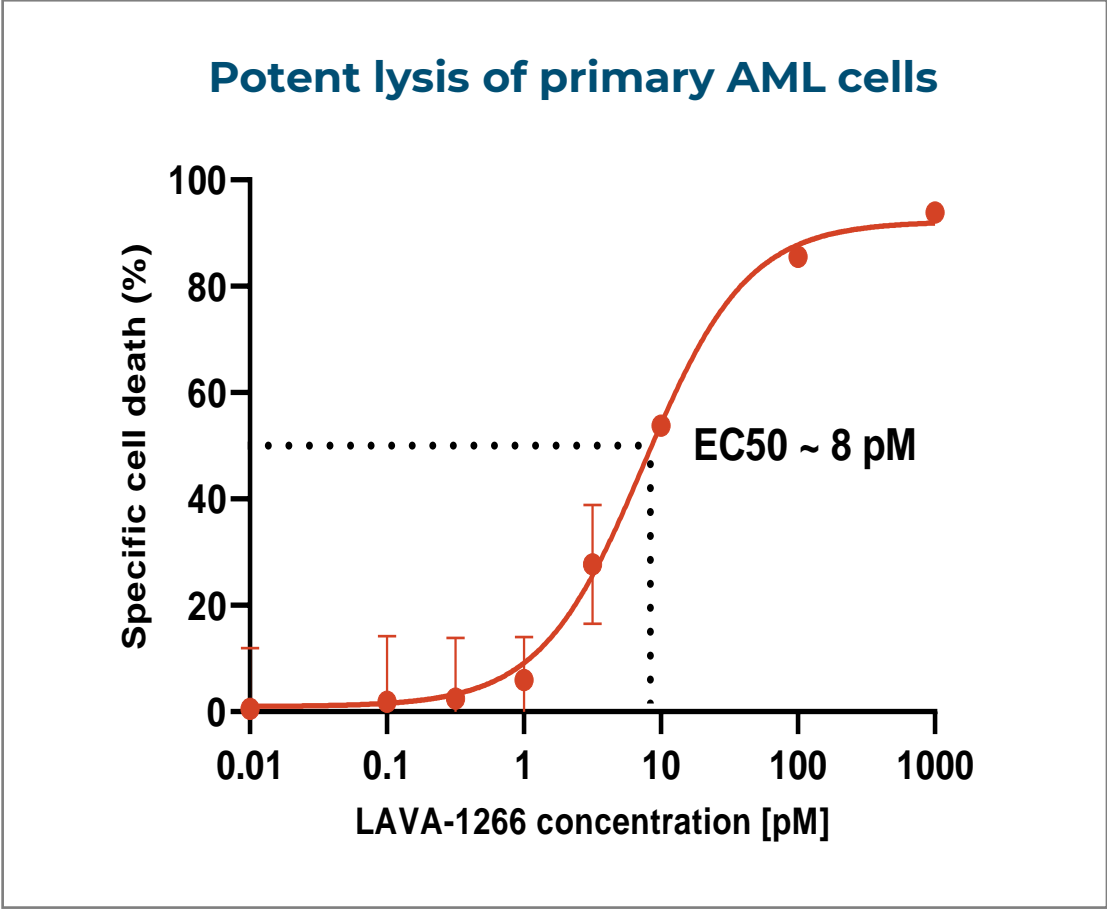
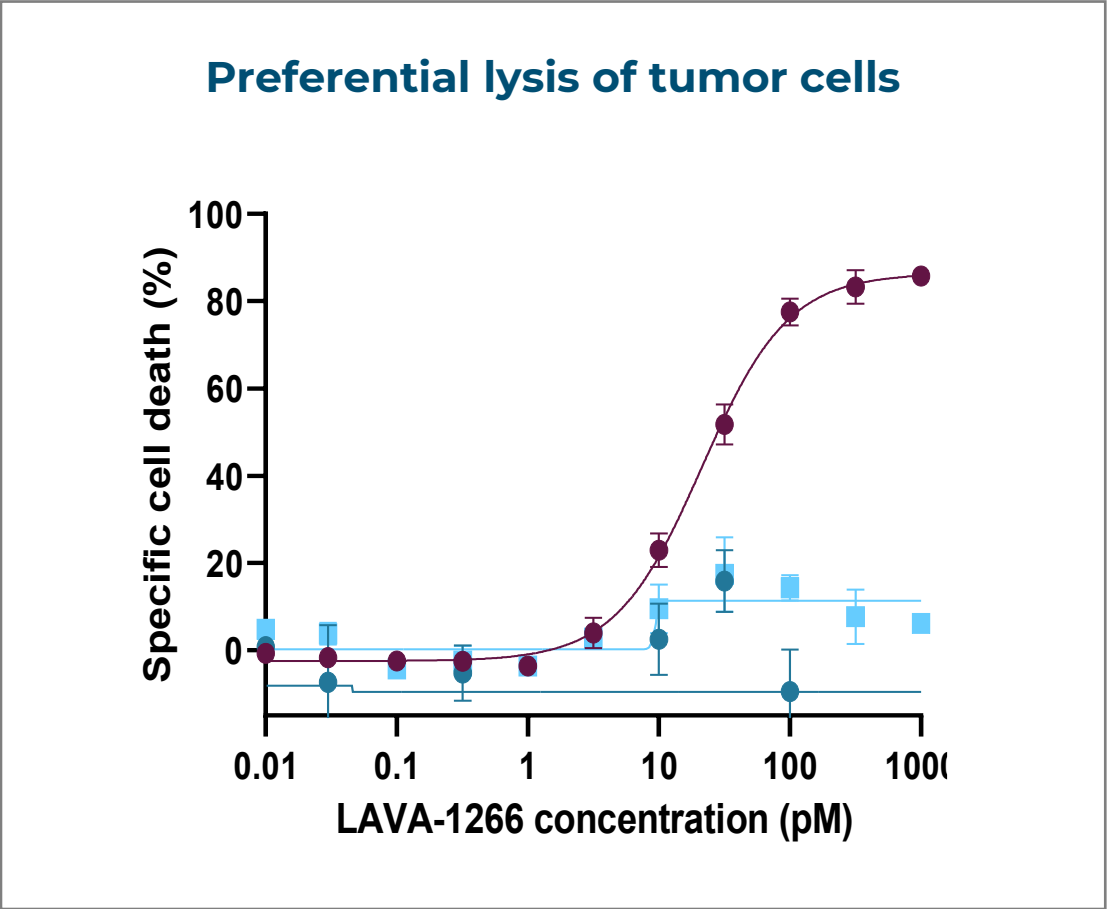
Relative abundance of Vδ2s cells in AML suggests this disease could be an attractive target for Gammabody therapies



Next update expected H1 2024

Clinical-trial enabling activities are underway, in support of an expected H1 2024 IND/CTA filing

LAVA-1266: CD123-Targeting Gammabody®



● CD123+ tumor cell line ● CD123+ healthy cells (donor 1) ■ CD123+ healthy cells (donor 2)

Strategic Partnerships

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LAVA-1223/SGN-EGFRd2 for Solid Tumors¹: Phase 1 Underway



Mechanism of action

Designed to induce preferential lysis of EGFR-expressing tumor cells while relatively sparing EGFR-expressing normal cells



Strategic partner



Agreement

Exclusive worldwide license agreement with Seagen Inc. entered into Q3 2022
Seagen to develop and commercialize SGN-EGFRd2 (LAVA-1223). Potential for milestones of up to approximately \$650 million and royalties



Payments to date

\$50MM upfront received with the signing, Sept 2022
Received licensing milestone in Q4 2023

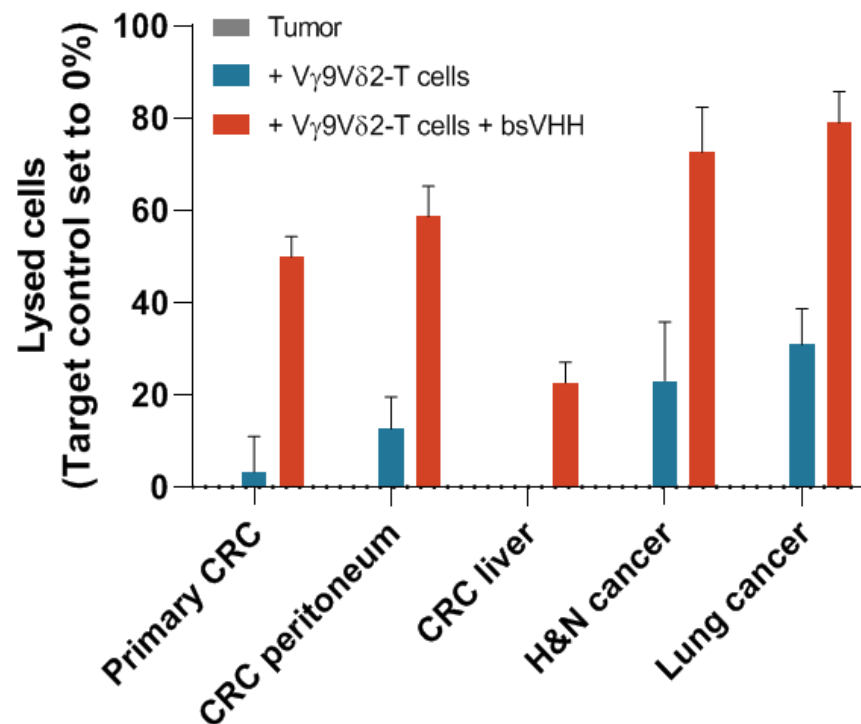


Program status

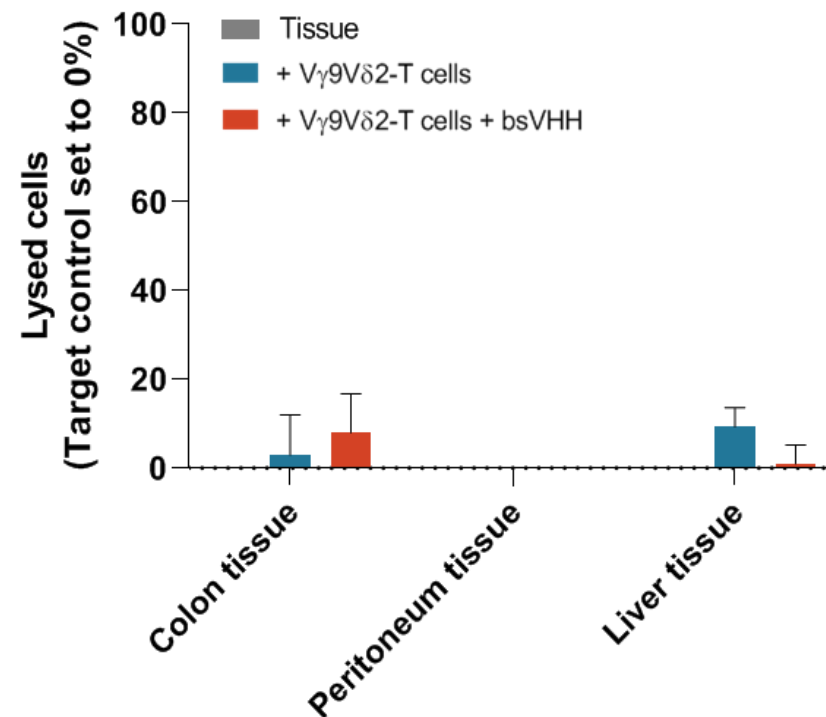
Phase 1 Clinical Trial ([NCT05983133](#)) initiated in Q4 2023

SGN-EGFRd2 (LAVA-1223) – EGFR-Targeting Gammabody®

Patient derived tumor tissue



Patient derived non-tumor tissue



Janssen Collaboration¹: Lead Candidate Selected



Mechanism of action

Undisclosed tumor associated antigen



Strategic partner



Agreement

LAVA entered into a research collaboration and license agreement with Janssen (May 2020) for the discovery and development of a novel bispecific gamma-delta T-cell engager for the treatment of cancer. Janssen is responsible for the future clinical development, manufacture, and commercialization of the candidate at Janssen's sole cost and expense



Payments

Upfront payment (undisclosed). LAVA is eligible to receive development, regulatory and commercialization milestone payments and royalties



Program status

Product candidate onboarded June 2023

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