

Q2 2025 Results

Jason Lettmann | Chief Executive Officer
August 12, 2025



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ALX Oncology

Q2 2025 Results & Business Update

01 Overview

02 Clinical Development Update

03 Financials & Key Milestones



Jason Lettmann
Chief Executive Officer,
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Alan Sandler, MD
Chief Medical Officer,
ALX Oncology



Harish Shantharam
Chief Financial Officer,
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ALX Oncology Q2 2025 Key Accomplishments and Updates



Data from ASPEN-06 trial highlights CD47 expression as a key predictive biomarker, further demonstrating compelling clinical response of evorpacept activity in HER2+ gastric cancer



Phase 2 ASPEN-Breast evorpacept trial design updated to enable CD47 and HER2 biomarker-driven strategy in a single-arm study



Dose escalation completed in Sanofi-sponsored trial of evorpacept with SARCLISA® and dexamethasone in previously treated multiple myeloma; Sanofi moving into dose optimization



Anticipated to enroll first patient in August in the phase 1 clinical trial of novel EGFR-targeted antibody-drug conjugate (ADC), ALX2004

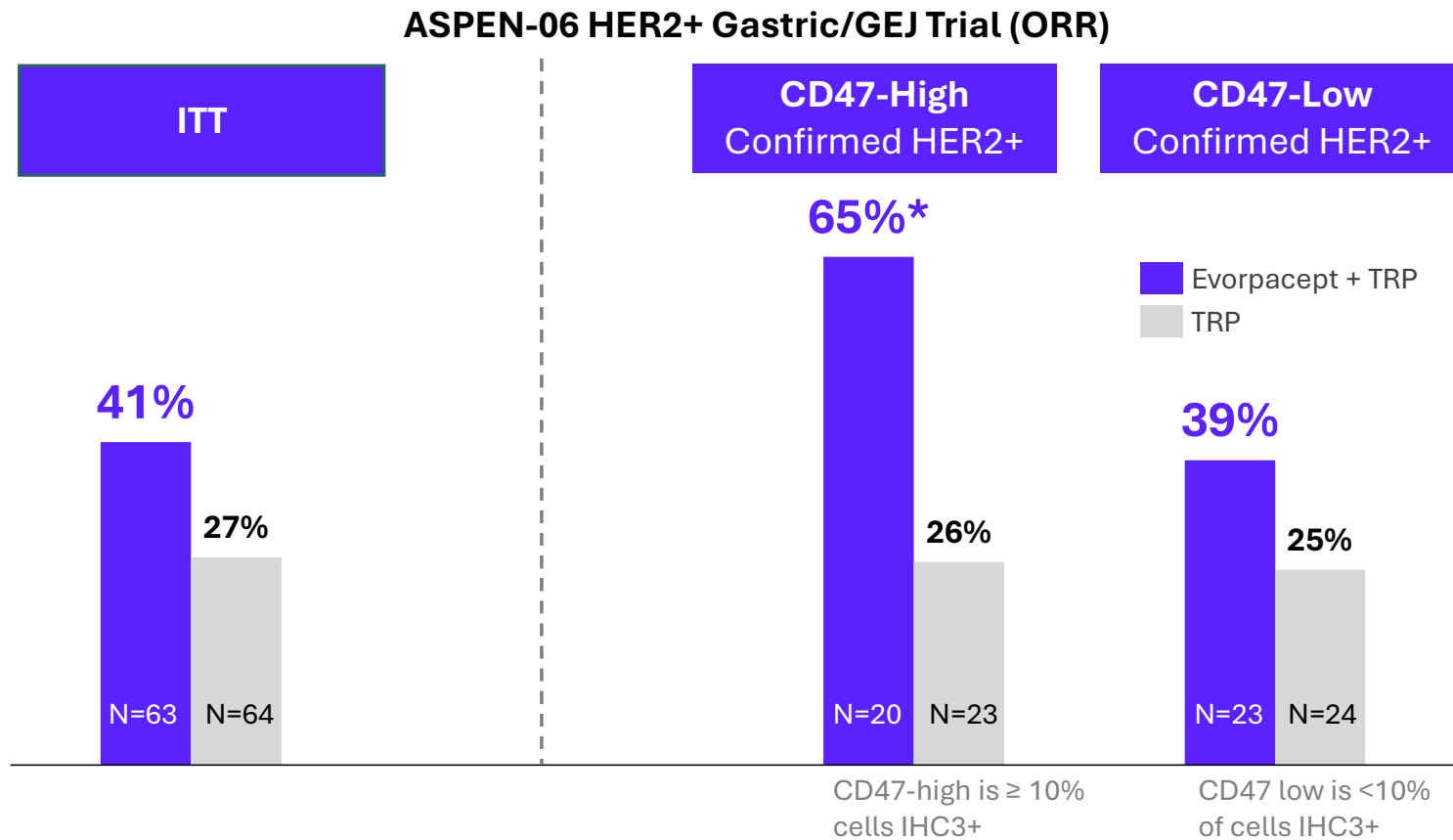


Focus on evorpacept in ASPEN-Breast and ALX2004, while pausing ASPEN-CRC, results in cash runway extended into Q1 2027



Upcoming milestones: ASPEN-06 CD47 expression data update (Q4 2025), ALX2004 initial safety (1H 2026), ASPEN-breast interim data readout (Q3 2026)

CD47 expression acts as a predictive biomarker for durable patient benefit from evorpaccept



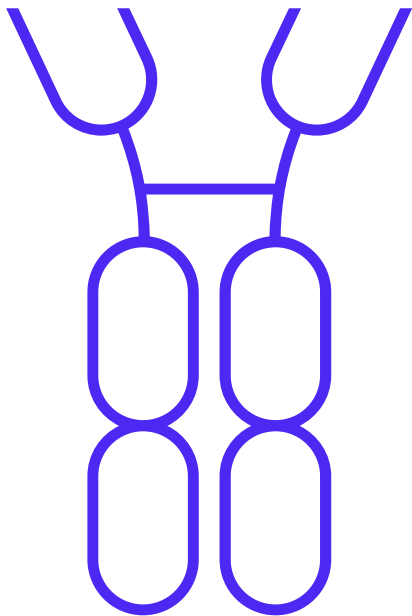
Note: Above results are based on pre-planned exploratory analysis of ASPEN-06 gastric study. HER2+ based on fresh biopsy or ctDNA amplification. Data Cutoff as of May 15, 2025. ORR per investigator. T = trastuzumab; R = ramucirumab; P = paclitaxel.

*nominal p-value < 0.05

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Evorpacept: Uniquely designed to offer a differentiated safety profile and robust clinical activity in combination with available cancer therapies

EVORPACEPT



Higher affinity CD47 binding



More potently
blocks CD47
signal on
cancer cells

Inactive Fc domain



Less “sink effect”
= more targeted

No known dose
dependent cytopenia
= higher dosing

Lower molecular weight



Increased solid
tumor penetration
and higher
effective dosing

Antibody-like pharmacokinetics

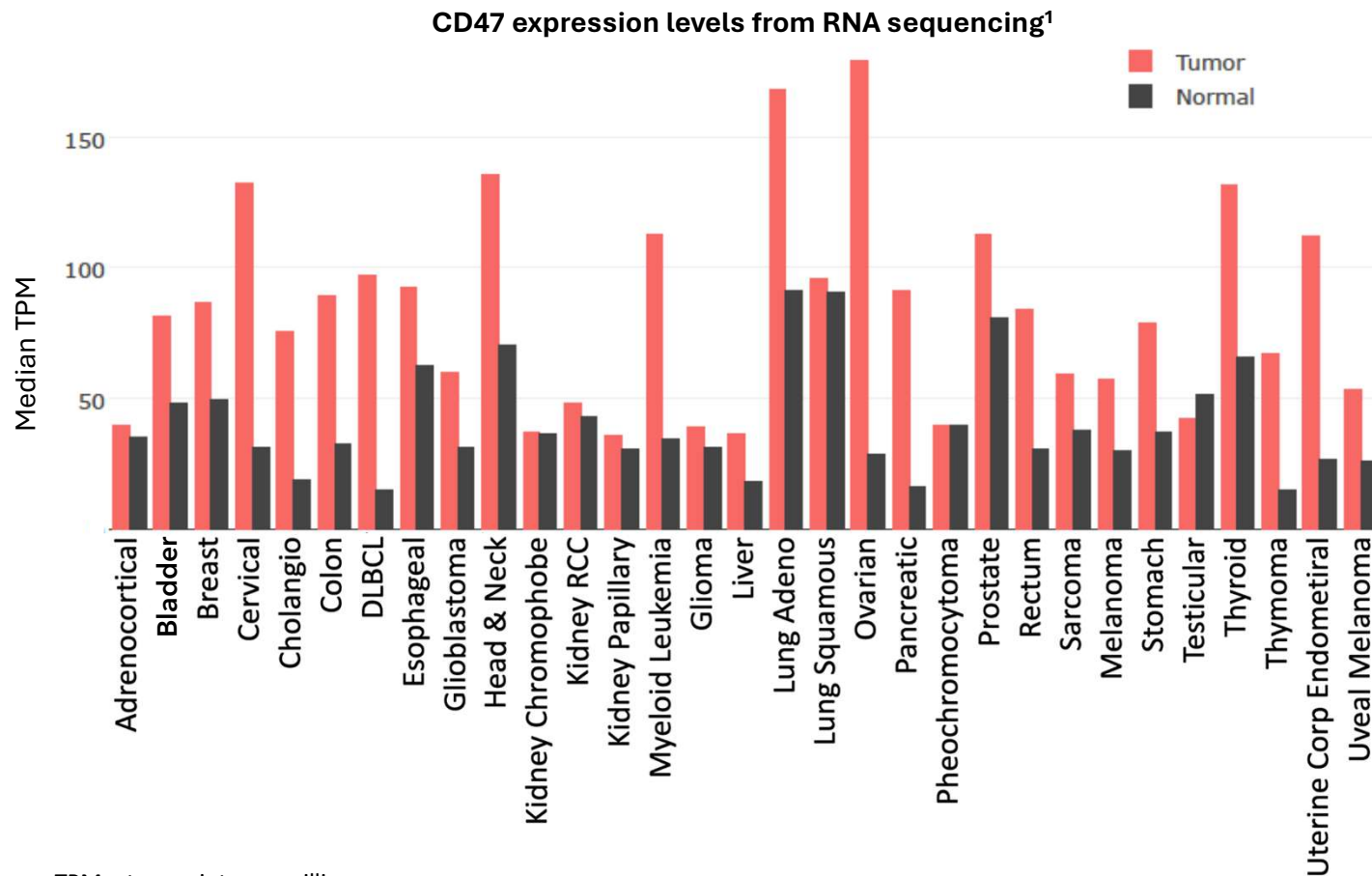


Long half
life = less
frequent dosing
and matching
regimen with
combinations

A NEW VISION FOR EVO

The first targeted immuno-oncology therapeutic to drive superior outcomes for patients with CD47 over-expressing cancers

CD47 is overexpressed across a range of solid and heme malignancies



- As a "marker of self", CD47 is expressed on all cell types²
- Cancer cells take advantage of this by overexpressing CD47
- Due to this, the vast majority of both solid and liquid tumors utilize CD47 to evade the immune system

TPM = transcripts per million

1) Tang, et al, GEPIA, 2017; 2) Dheilly, et al, Mol. Ther., 2017

Research in CD47 over the last 10+ years provides a strong foundation for utilizing CD47 as a negative prognostic biomarker

- **In a meta-analysis of 38 cohorts across 17 publications including >7,000 patients, “CD47 overexpression correlated with shorter OS in cancer patients”***

Increased CD47 expression is correlated with poor patient outcomes in many tumor types including¹:

- Oral squamous cell carcinoma²
- Nasopharyngeal carcinoma³
- Triple negative breast cancer⁴
- Ovarian cancer⁵
- Non-small cell lung cancer⁶
- Clear cell renal cell carcinoma⁷
- Hepatocellular carcinoma⁸
- Gastric adenocarcinoma⁹
- Colorectal adenocarcinoma¹⁰
- Head and neck squamous cell carcinoma¹¹
- Multiple myeloma¹²

*Yang et al, *Translational Cancer Research*, 2018

1) Huang, et al, *Scientific Reports*, 2022; 2) Pai, et al, *Cells*, 2019; 3) Wang, et al, *OncoTargets & Ther.* 2020; 4) Yuan, et al, *Oncol Lett*, 2019; 5) Li, et al, *Am J Trans Res*, 2017; 6) Barrera, et al, *Br J Cancer*, 2017; 7) Jiang, et al, *Urol Oncol*, 2022; 8) Kim, et al, *J Clin Pathol*, 2021; 9) Shi, et al, *Cancer Imm, Imm*, 2021; 10) Kim, et al, *Diagnostics*, 2021; 11) Wu, et al, *Oncoimmunology*, 2018; 12) Rastgoo, et al, *Haematologica*, 2020

Evorpacept is the leading CD47 program in development and now poised to be the next targeted immuno-oncology breakthrough



Selecting for CD47 could lead to larger magnitude of effect and increased probability of success in future studies



Predictive CD47 biomarker could facilitate smaller and faster registrational trials



Selecting for CD47-high patients who are more likely to progress enables entry into earlier lines of therapy across CD47-expressing tumor types



As data suggests 50-70% of patients overexpress CD47 in the HER2+ BC population, significant commercial opportunity in 2L+ BC



Potential of targeting the CD47 immune checkpoint is broad with high expression across multiple tumor types beyond breast cancer



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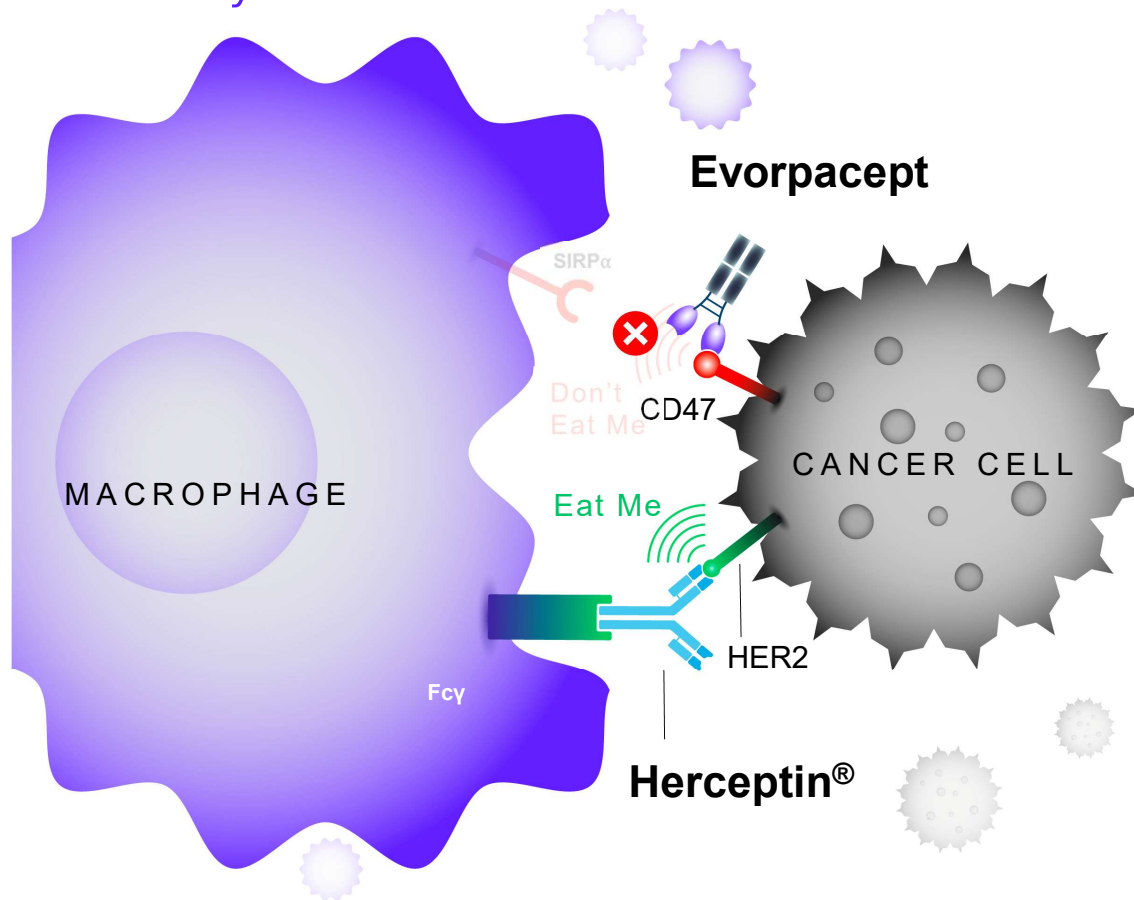
Evorpacept in patients with gastric cancer

New data from ASPEN-06 on
CD47 expression as a predictive
biomarker for evorpacept



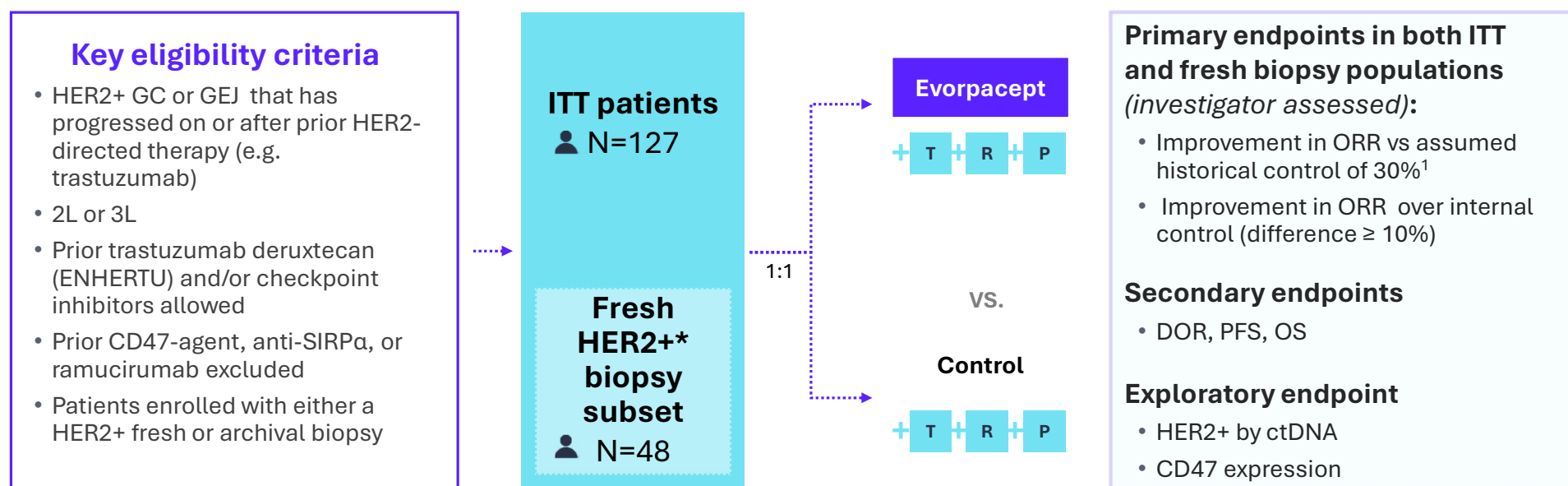
Alan Sandler, MD
CMO, ALX Oncology

Evorpacept, with its inactive Fc domain, stimulates macrophages to selectively attack cancer cells in combination with antibodies like Herceptin



- Due to CD47 expression in healthy cells driving on-target toxicity, conventional approaches utilizing Fc-active antibodies have all failed (Gilead / Forty Seven, Pfizer / Trillium, iMab, Arch, etc.)
- Evorpacept is the only CD47-blocker in development using an inactive Fc mechanism to avoid targeting healthy cells

Today: new analysis highlights CD47 expression as a key predictive biomarker further enhancing/increasing evorpaccept activity in patients with gastric/GEJ cancer



Evo Evorpaccept (30 mg/kg Q2W) **T** Trastuzumab (6 mg/kg > 4 mg/kg Q2W) **R** Ramucirumab (8 mg/kg Q2W) **P** Paclitaxel (80 mg/m² on day 1, 8, 15 of 28-day cycle)

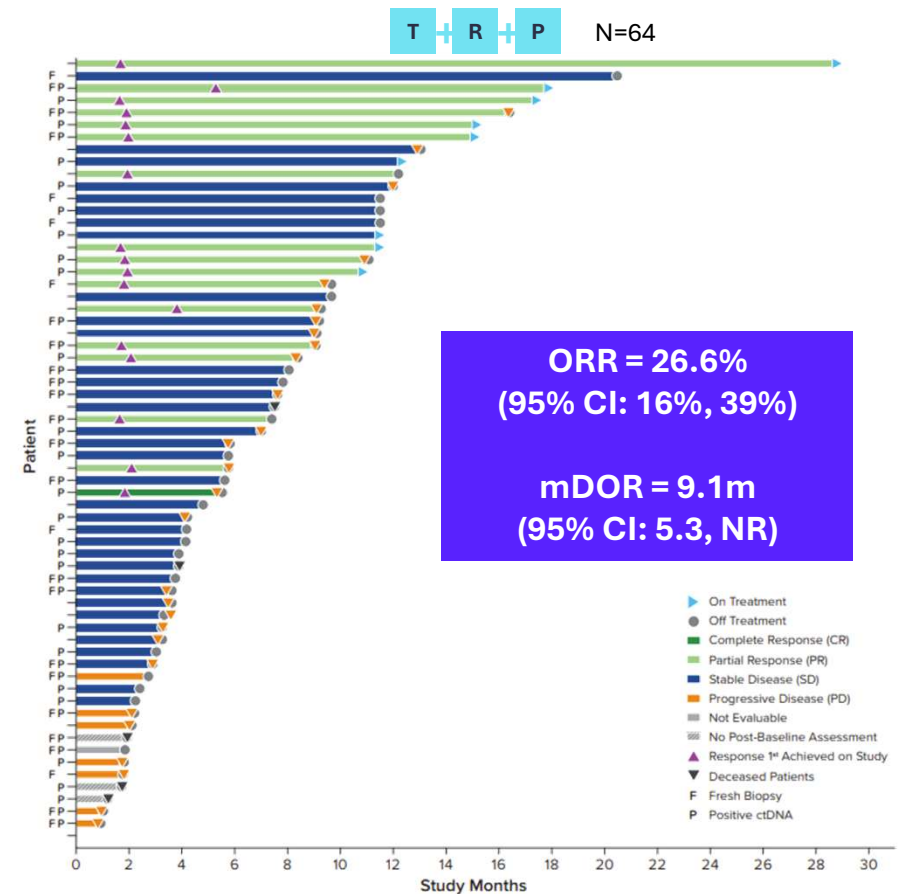
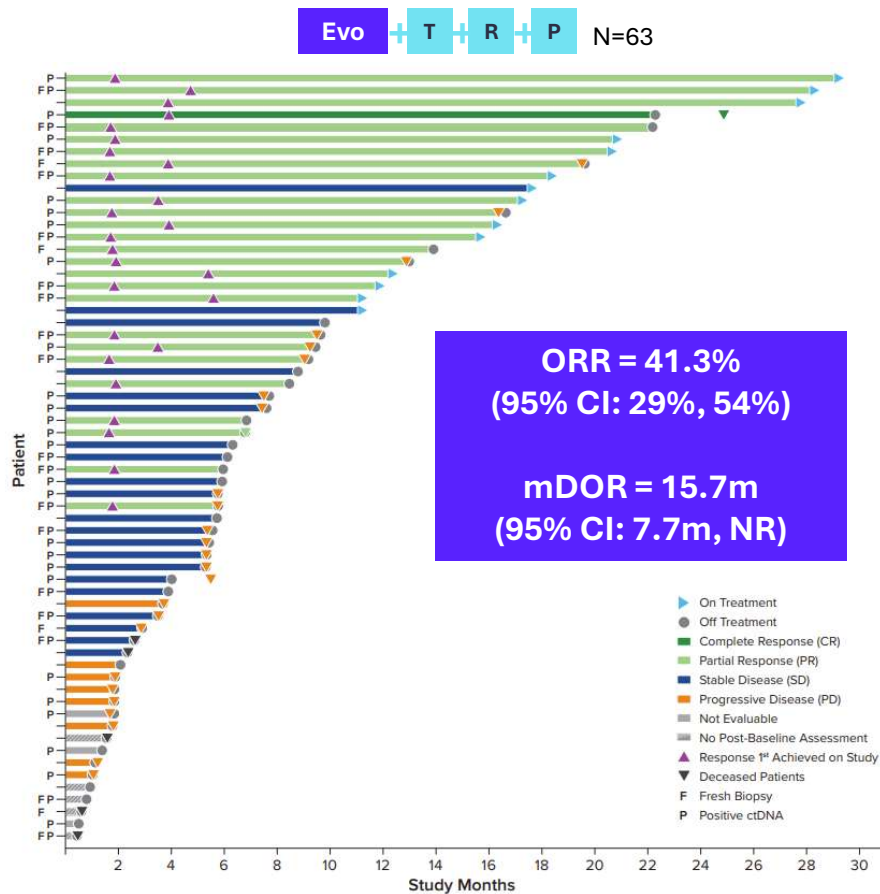
GC- gastric cancer, GEJ- gastroesophageal junction, TRP- trastuzumab, ramucirumab, paclitaxel; selected secondary and exploratory endpoints shown.

*Fresh HER2-positive is defined as biopsies that were HER2-positive after receiving prior HER2-targeted treatment

1. Wilke et al, Lancet, October 2014

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As a reminder, Evo-TRP drove a 41% ORR compared to 27% in the TRP control arm in the ITT population



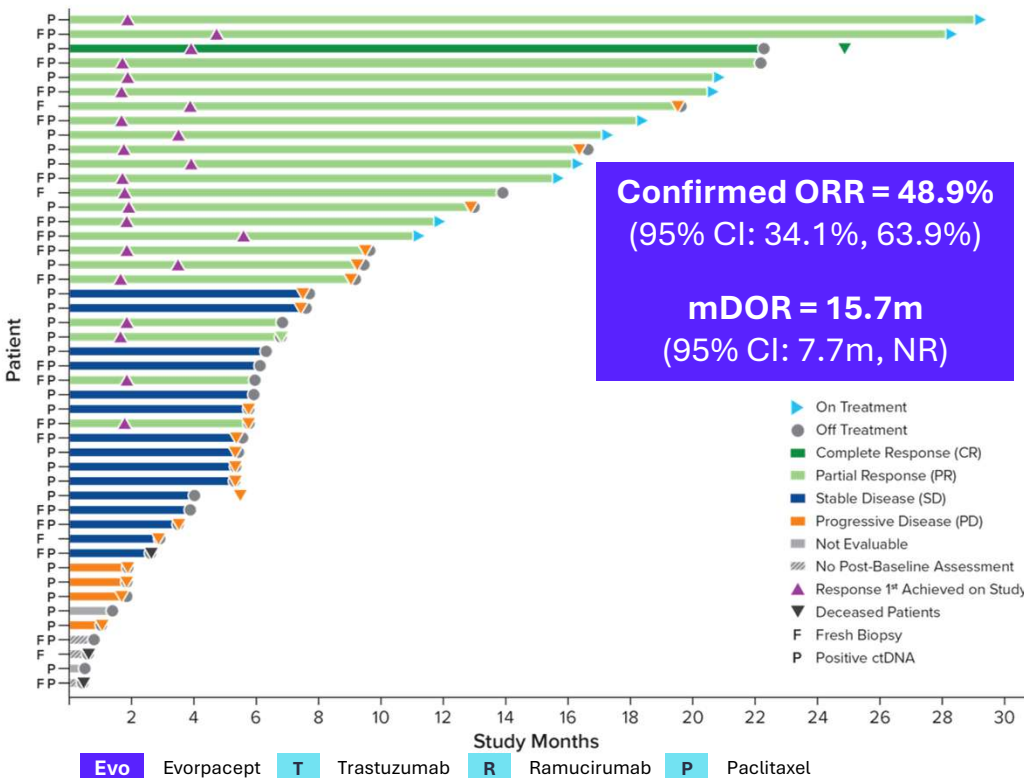
Evo Evorpacept **T** Trastuzumab **R** Ramucirumab **P** Paclitaxel

Seven patients treated with Evo+TRP and five patients treated with TRP had no post-baseline assessment or best response of NE; data cutoff as of 02 Dec 2024; NR = Not Reached

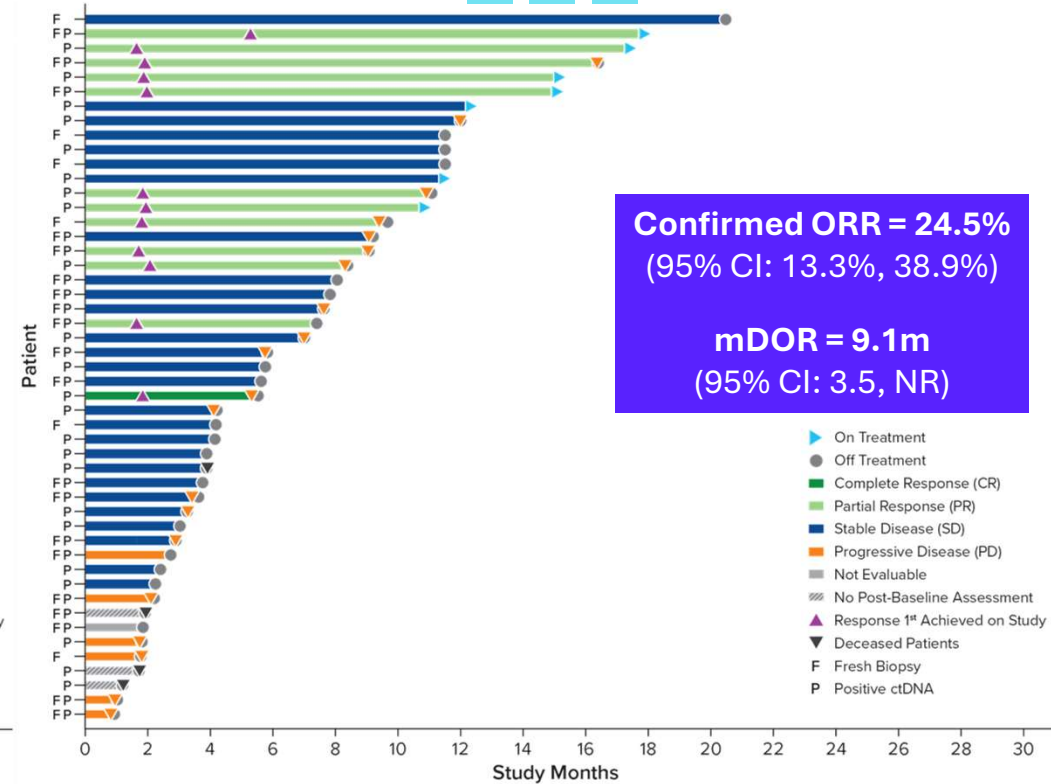
Furthermore, evorpaccept demonstrated a 49% ORR compared to 24.5% in control in patients with HER2-positivity confirmed by fresh biopsy or ctDNA

Patients with HER2+ confirmed with fresh biopsy OR ctDNA+ (n=96)

Evo + T + R + P N=47

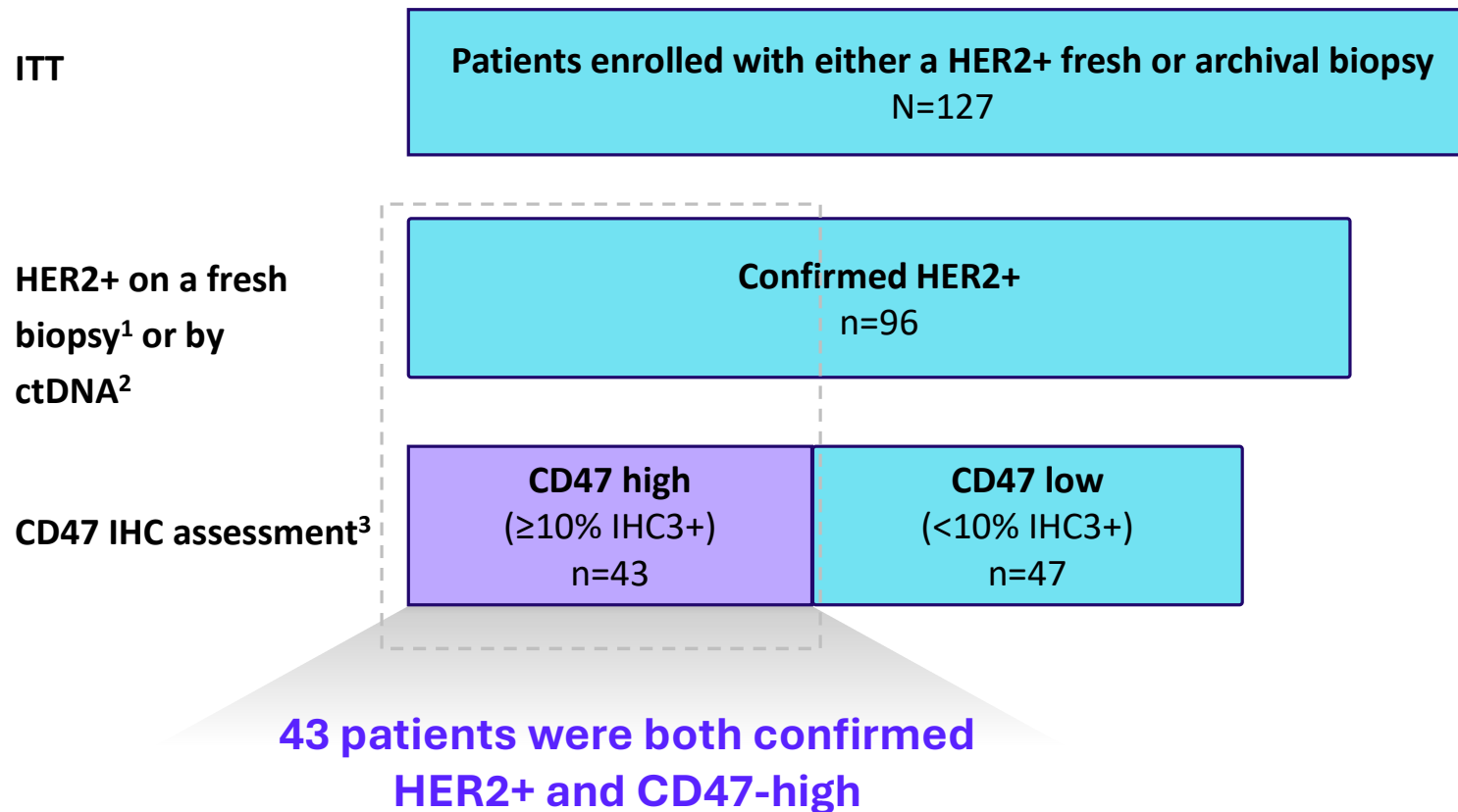


T + R + P N=49



Seven patients treated with Evo+TRP and five patients treated with TRP had no post-baseline assessment or best response of NE; data cutoff as of 02 Dec 2024; NR = Not Reached

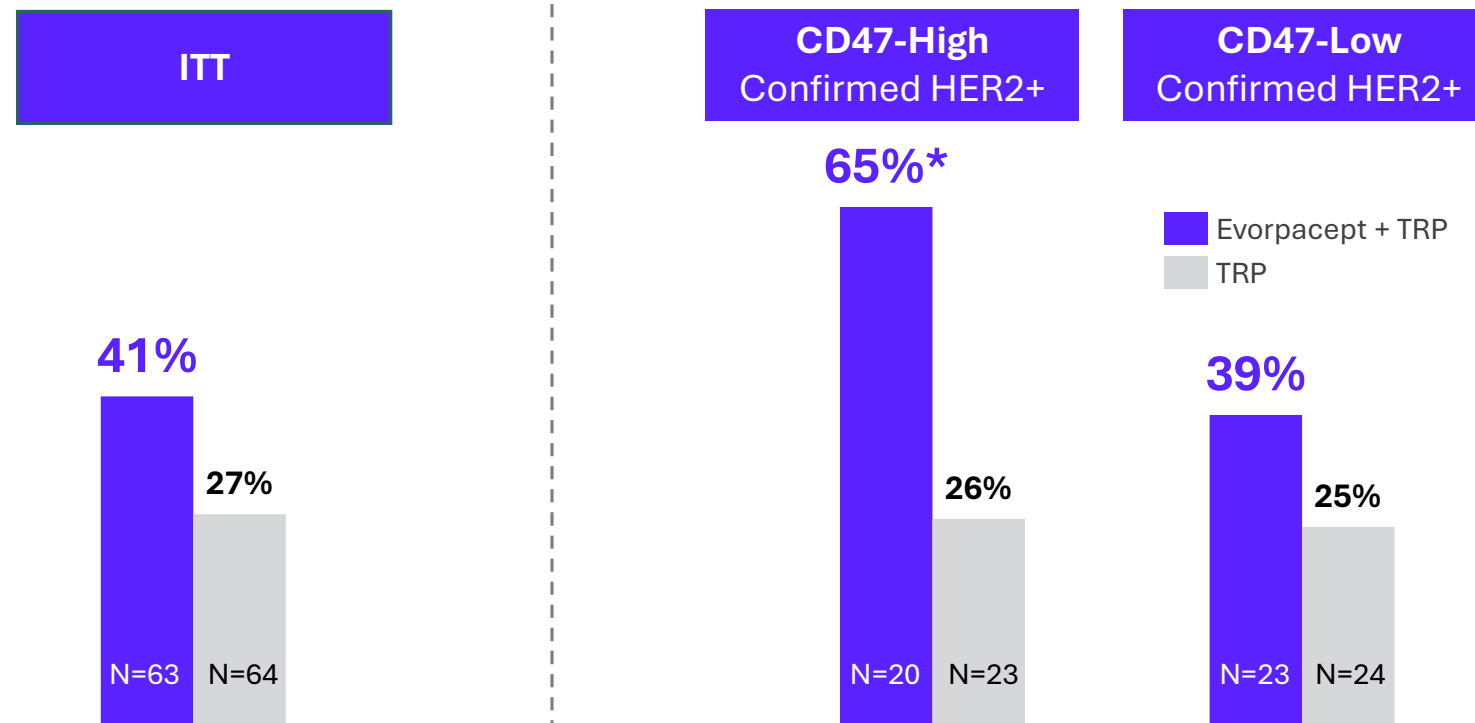
Patients in ASPEN-06 were tested for both HER2 and for CD47 expression



1) Fresh HER2-positive is defined as biopsies that were HER2-positive after receiving prior HER2-targeted treatment ; 2) HER2 (ERBB2) plasma gene amplification from Guardant360® analysis; 3) 6 patients with confirmed HER2+ had missing CD47 samples or non-evaluable samples

CD47 expression acts as a predictive biomarker for durable patient benefit from evorpaccept

ASPEN-06 HER2+ Gastric/GEJ Trial (ORR)



DOR, PFS and OS also showed strong magnitude of benefit for evorpaccept in CD47-high patients

Results consistent across multiple CD47 expression cutoffs

Full data set will be presented at an upcoming medical conference in Q4 2025

Note: Above results are based on pre-planned exploratory analysis of ASPEN-06 gastric study. HER2+ based on fresh biopsy or ctDNA amplification; CD47 low is <10% of cells IHC3+; CD47-high is ≥ 10% cells IHC3+. Data Cutoff as of May 15, 2025. ORR per investigator. T = trastuzumab; R = ramucirumab; P = paclitaxel.

*nominal p-value < 0.05

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By targeting CD47-high patients, evorpacept is now poised to be the next targeted immuno-oncology breakthrough



In patients with HER2+ and CD47-high gastric cancer (n=43), evorpacept + TRP had a 65% ORR versus 26% ORR for TRP



DOR, PFS and OS also showed strong magnitude of benefit between the CD47-high and CD47-low groups



Enables a targeted clinical development strategy and will guide our strategy in breast cancer and other tumors that overexpress CD47



Full data set will be presented at an upcoming medical conference in the fourth quarter of 2025



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ASPEN-Breast

Evorpacept in breast cancer: CD47 biomarker strategy

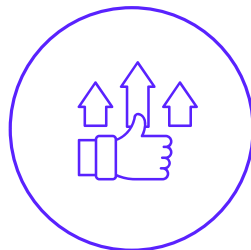


Alan Sandler, MD
CMO, ALX Oncology

The evorpacept opportunity in breast cancer

High Probability of Success

De-risked path given positive data in two HER2-positive cancers



Positive randomized data in ASPEN-06 in gastric cancer with trastuzumab, and in combination with zanidatamab (HER2-bispecific) in HER2+ breast cancer

High Unmet Need

Changing 1L SOC drives opportunity in patients who progress on ENHERTU and/or other HER2-directed therapies



Evo is active in patients who progressed on Herceptin in gastric cancer, and in breast cancer patients who progressed on ENHERTU and multiple HER2-directed drugs

Highly Targeted Approach

CD47/ HER2 biomarker-driven approach enables targeted strategy



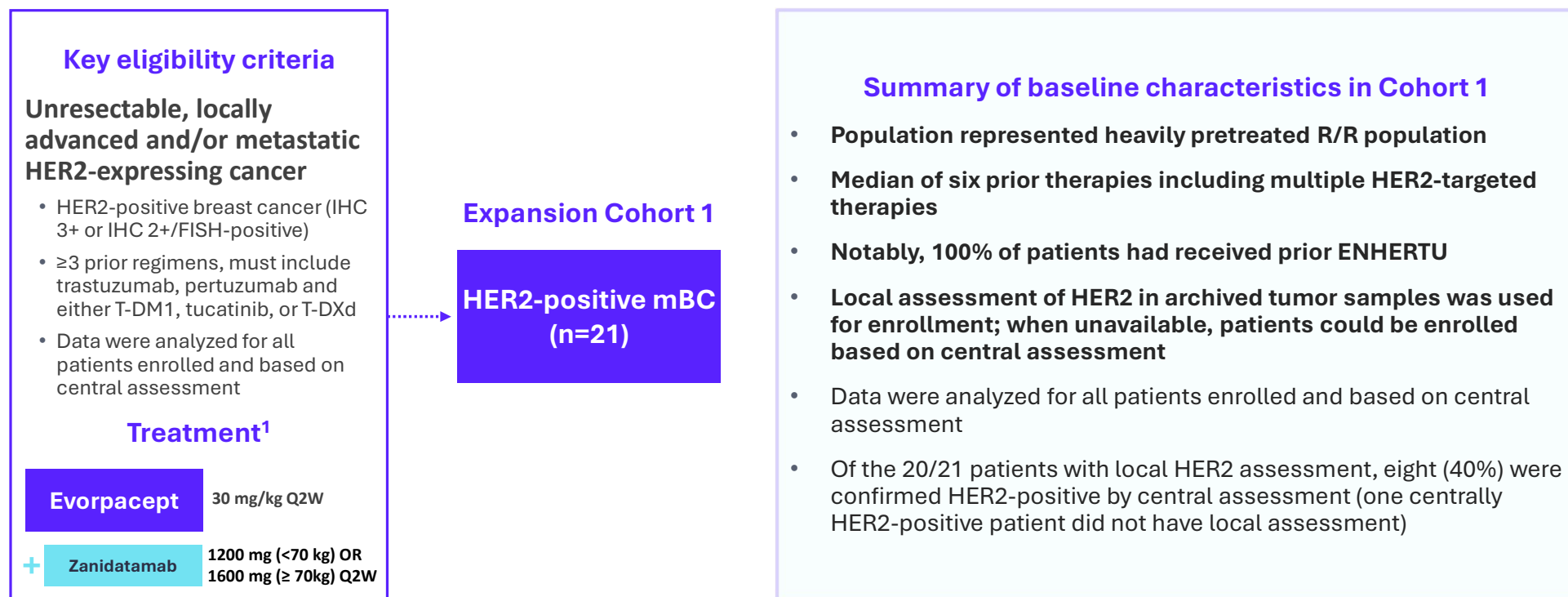
Strong scientific support for CD47/ HER2 as a key mode of resistance in metastatic BC

Trial designed to identify CD47 as a predictive biomarker potentially enabling a rapid registrational pathway

*In 7 major markets: US, EU5 (UK, Spain, Italy, Germany, France) and Japan; Source: Clarivate Market Forecast, gastroesophageal cancer, December 2024

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Phase 1b/2 trial evaluating safety and efficacy of evorpaccept plus zanidatamab in patients with breast cancer who have progressed on prior HER2-directed therapy

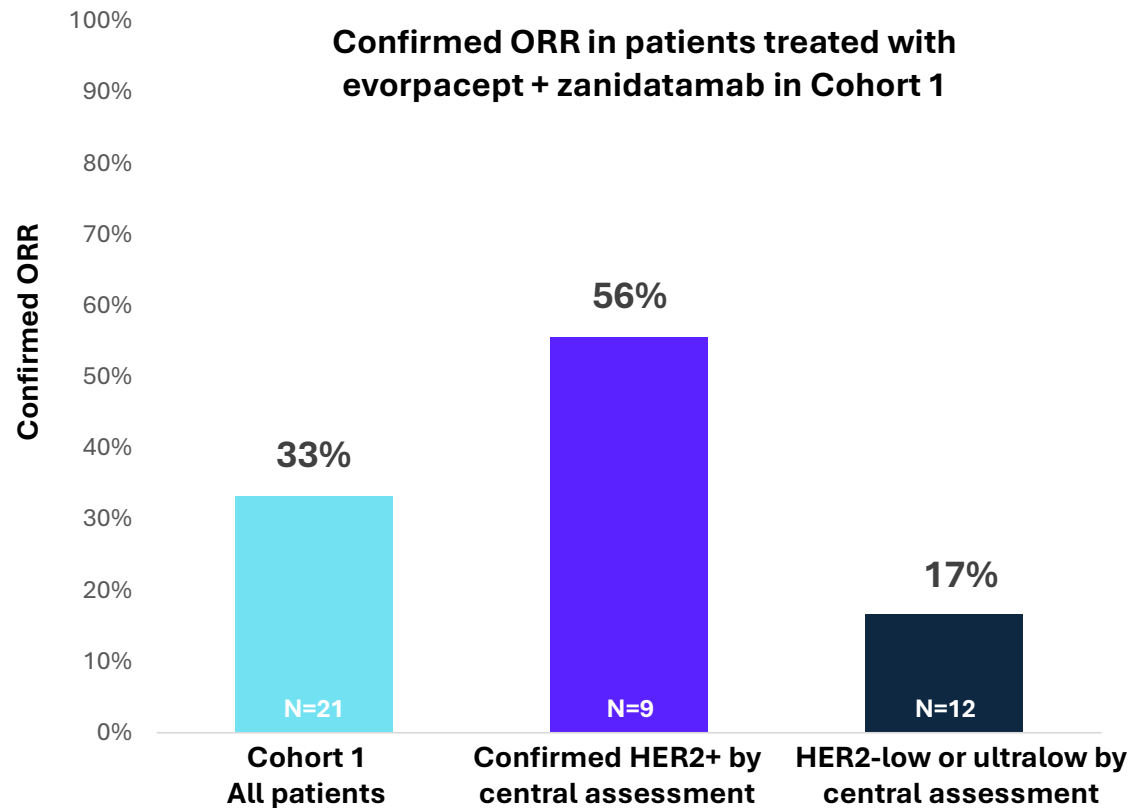


1. Mandatory IRR prophylactic treatment included corticosteroids, antihistamines, and acetaminophen. Study conducted by Jazz Pharmaceuticals

This study provides clinical data supporting further development of evorpaccept with HER2-targeted agents in patients with breast cancer

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Breast cancer patients with confirmed HER2-positivity had the greatest benefit from evorpaccept + zanidatamab



Strongest efficacy in confirmed HER2+

- ORR of 55.6% (5/9)
- mDOR NE (range: 5.5-25.9m)
- mPFS = 7.4m (95% CI: 0.6, NE)

Compares favorably to benchmark

- SOPHIA study (n=536) of margetuximab + chemo vs. trastuzumab + chemo (ORR: 22% vs. 16%)

Median follow-up (range) was 9.6 (0.6, 29.7) months, with six patients on treatment at data cutoff as of August 1, 2024; HER2-Low/Ultralow = IHC1+, IHC2+ / ISH-, IHC 0
1. *JAMA Oncol.* 2021;7(4):573-584. doi:10.1001/jamaoncol.2020.7932
Montero. et. al. SABCs 2024, Poster Spotlight Presentation. Abstr #SESS-2007

Several studies have found that CD47 protein expression in HER2+ Breast is over-expressed at time of initial diagnosis

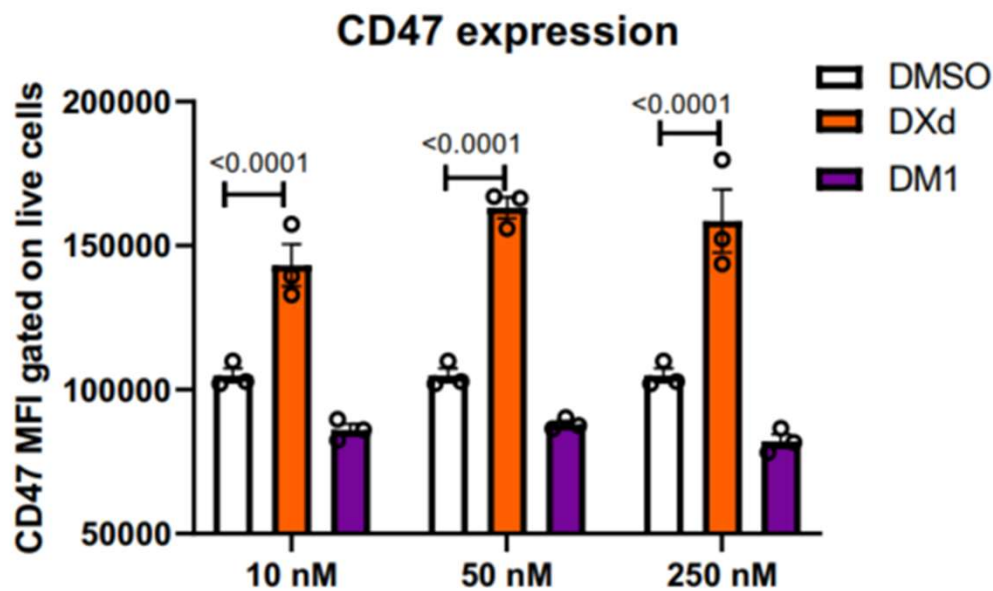
% BC CD47 High	% HER2+ BC CD47 High	Definition of High	Clone	Ref
89/200 (45%)	32/58 (55%)	IRS 7-12	A1838	Alhanafy, 2024
84/137 (61%)	16/24 (67%)	H-score ≥80	A1838	Sun, 2022
36/98 (37%)	29/82 (35%)	Mod/strong	BRIC126	Kosaka, 2021
93/282 (33%)	14/27 (52%)	IRS 6-9	ab226837	Chen, 2022
140/217 (65%)	40/54 (74%)	IRS 7-12	ab213079	Yuan, 2019
Average: 47%	Average: 54%			

- CD47 expression in breast cancer has been studied in 5 publications using different methods and clones
- CD47 is over-expressed in ~50% of breast cancer patients at diagnosis

Studies looking at CD47 protein expression in BC use varying IHC clones and scoring methods (typically incorporating both staining intensity and % positive cells)

CD47 is upregulated in response to T-DXd (Enhertu) treatment in HER2-positive breast cancer cell lines

T-DXd (Enhertu) exposure increases CD47 expression

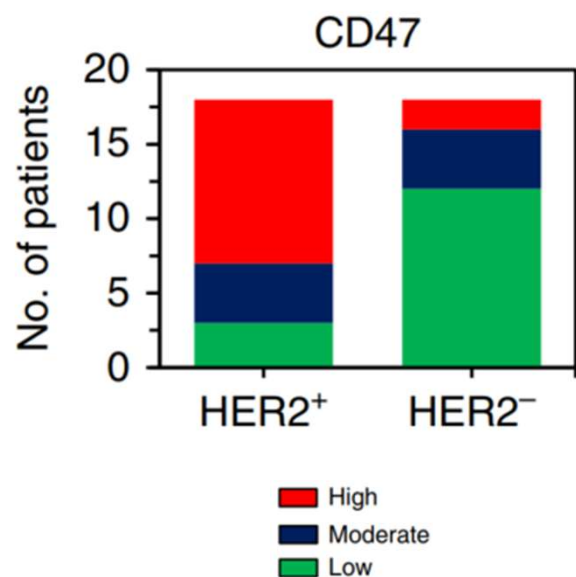


Flow cytometry assessment of surface CD47 expression on Au565 cells after 2 days of treatment with Enhertu's payload (DXd) or Kadcyła's payload (DM1) as compared to control (DMSO)

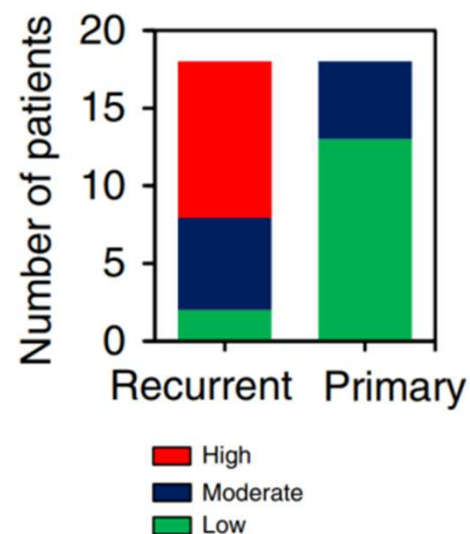
- As our ASPEN-09 breast study will target post-Enhertu patients, these data provides validation that CD47 is a key mode of ADC evasion in the relevant study population

CD47 expression in breast cancer is higher in HER2+ patients, more common in resistant cancer

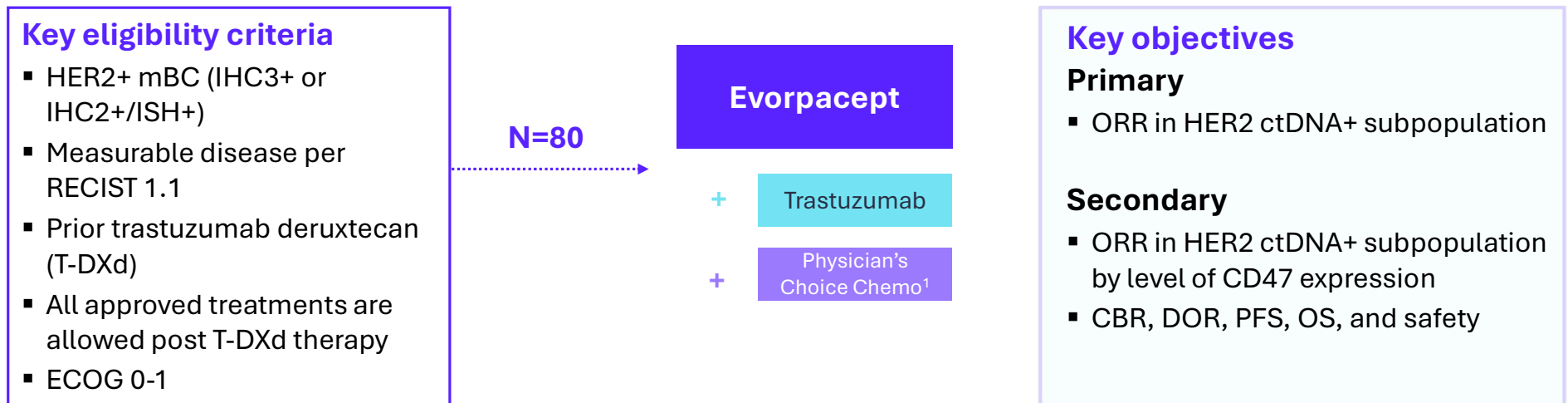
**CD47 expression is higher on
HER2+ BC cells vs HER2- and...**



**... CD47-high cells are more
common in recurrent HER2+ BC**



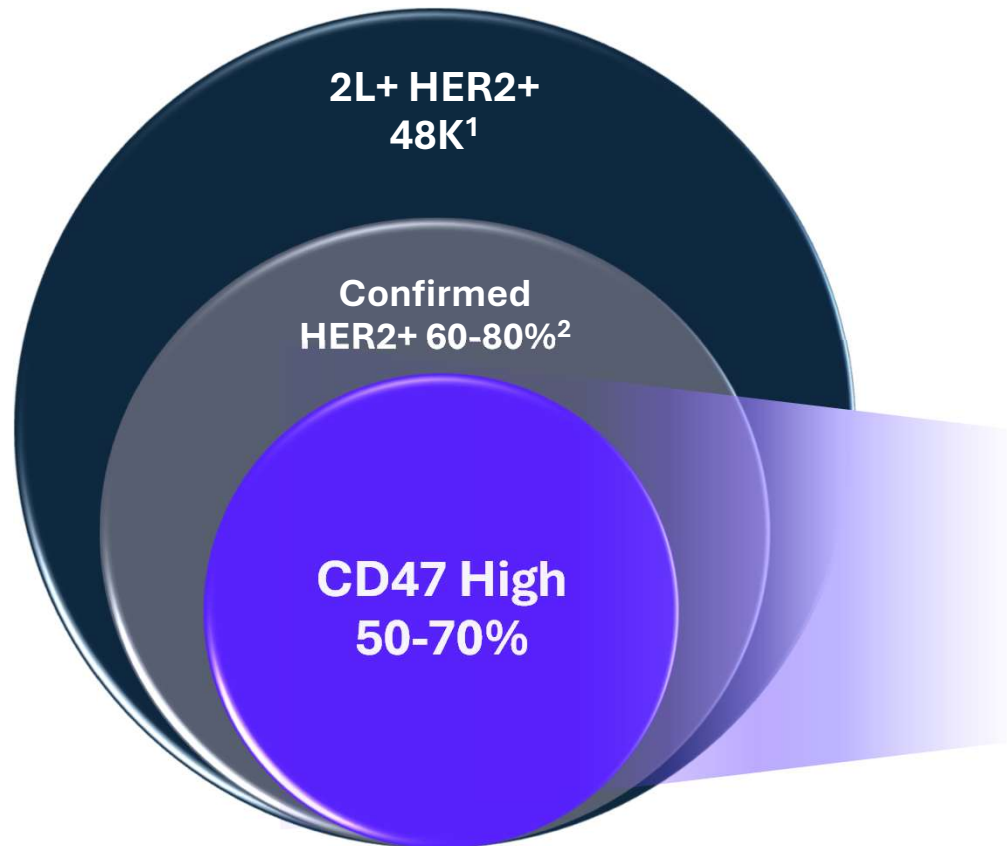
Strong expected benefit in CD47-high and HER2+ patients enables evorpaccept-attributable benefit in single-arm study



- **Inclusion of both CD47-high and CD47-low patients enables evaluation of the predictive value of CD47 as a biomarker for evorpaccept**
- **Revised study design is expected to optimize enrollment and allow for an interim data readout in Q3 2026**
- **Design is anticipated to support a biomarker-driven registrational study in HER2-positive breast cancer**

1) Capecitabine, eribulin, gemcitabine, paclitaxel, or vinorelbine

HER2+ and CD47-high 2L+ BC represents a significant initial commercial opportunity with potential to move into earlier lines of therapy



~20K addressable patients are CD47-high

Represents \$2-4B market opportunity in CD47-high, HER2+ 2L+ BC³

Annual market opportunity based on: 1) US, EU5, JPN addressable patients; ~18k patients in the US; (2) ALX advisory board feedback on breast cancer trial; (3) Monthly price estimate is based on benchmarks in US and extrapolated to core markets.

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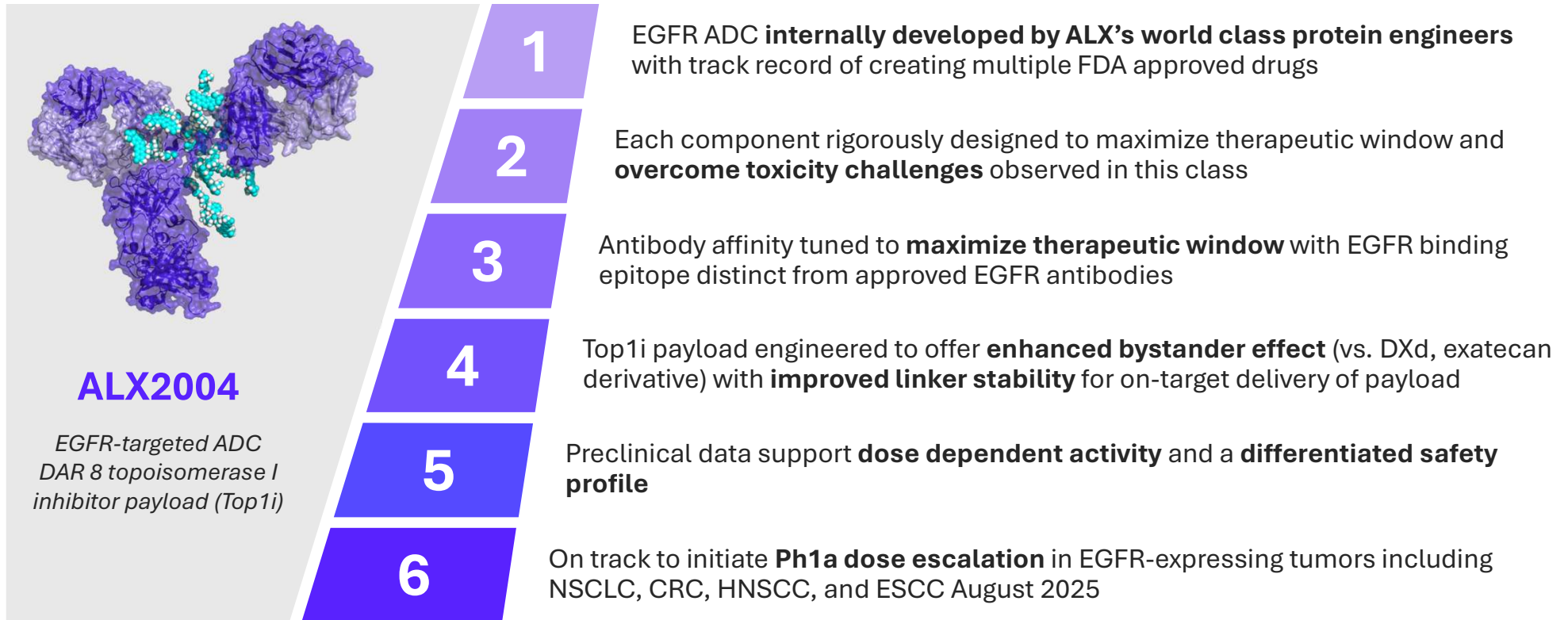


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ALX2004: potential best-in-class EGFR-targeted ADC

First ADC from ALX's linker payload platform

ALX2004 is a highly differentiated ADC in development for EGFR-expressing solid tumors now in Phase 1 trial

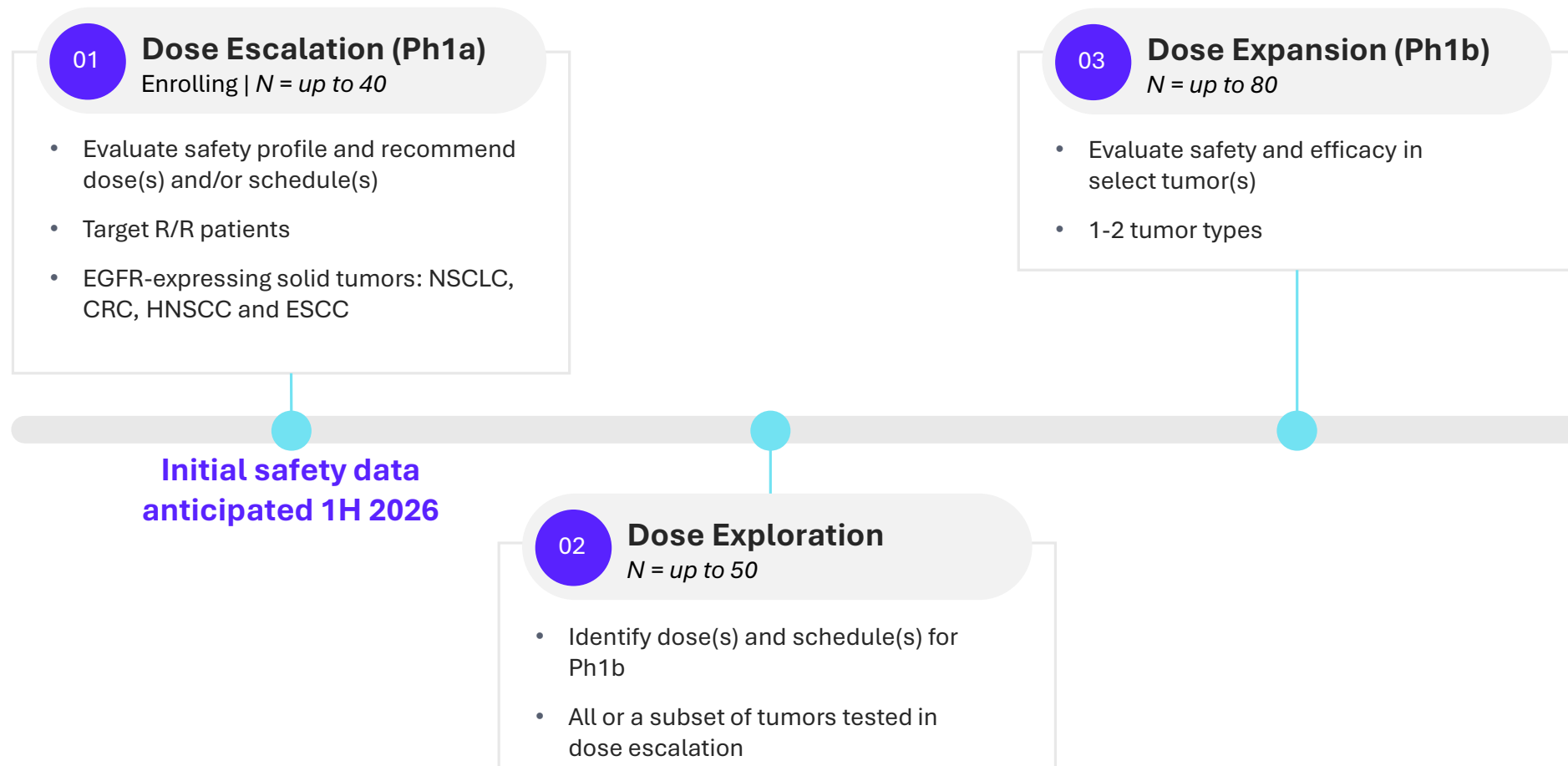


Initial safety data expected to be available in 1H2026

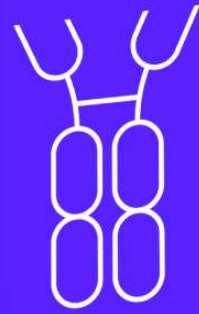
HNSCC: head and neck squamous cell carcinoma; CRC: colorectal cancer; NSCLC: non-small cell lung cancer; ESCC: esophageal squamous cell carcinoma

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Phase 1 Clinical Development Plan



HNSCC: head and neck squamous cell carcinoma (no more than 3 prior lines) ; CRC: colorectal cancer (no more than 2 prior lines of systemic chemotherapy); NSCLC: non-small cell lung cancer (no more than 2 prior lines); ESCC: esophageal squamous cell carcinoma (no more than 3 prior lines)



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FINANCIAL UPDATES AND MILESTONES

Upcoming Catalysts from Pipeline



Harish Shantharam, CFA
CFO, ALX Oncology

ALX Oncology is Pursuing a Focused Development Plan

MODALITY / TARGET	PROGRAM	INDICATION	IND ENABLING	PHASE 1	PHASE 2	PHASE 3	STATUS
EVORPACEPT PROGRAMS							
Anti-cancer Antibodies	ASPEN-Breast Evorpacept, HERCEPTIN® + chemotherapy	ENHERTU®-Experienced HER2-Positive Breast Cancer					FPI Q425
	SARCLISA® + Dexamethasone ¹ + Evorpacept	RRMM (Relapsed or Refractory Multiple Myeloma)					Completed dose escalation, moving into dose optimization
	ASPEN-06 Evorpacept, HERCEPTIN®, CYRAMZA® + Paclitaxel ²	2L or 3L Advanced HER2-Overexpressing Gastric/Gastroesophageal Junction (GEJ)					Completed, established POC, CD47 data to be presented in Q425
	Zanidatamab ³ + Evorpacept	HER2-Expressing Breast Cancer and Other Cancers					Completed, data presented at SABCS '24
ADC	ENHERTU® (I-SPY) ⁴ + Evorpacept	HER2-Positive HER2-Low Metastatic Breast Cancer					Ongoing
ALX2004 PROGRAM							
EGFR ADC	ALX2004 Dose-escalation and expansion	EGFR-Expressing Solid Tumors					FPI anticipated August '25

ALX-sponsored ongoing trial
Completed trial

ALX Oncology retains worldwide rights to evorpacept

1. Sanofi sponsors SARCLISA® clinical trial. 2. Lilly supplies CYRAMZA® for ALX Oncology's ASPEN-06 program 3. Jazz Pharmaceuticals sponsors zanidatamab clinical trial. 4. Quantum Leap Healthcare Collaborative sponsors I-SPY clinical trial.

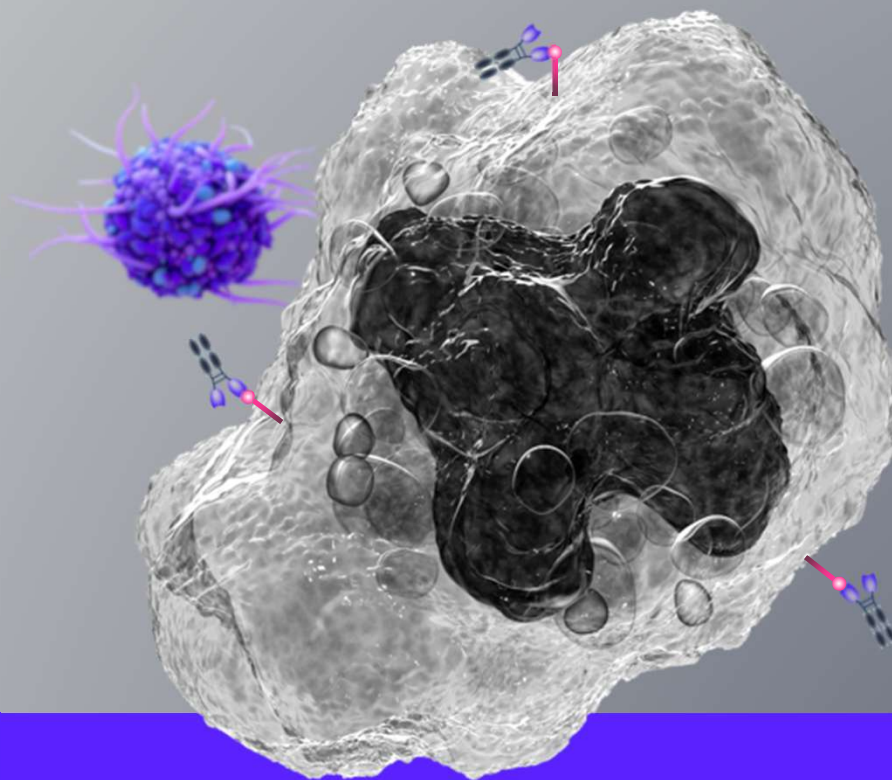


ALX is Focused on Driving Toward Two Key Inflection Points in 2026

PROGRAM	INDICATION	ANTICIPATED MILESTONES
EVORPACEPT		
ASPEN-Breast Evorpcept, HERCEPTIN® + chemotherapy	ENHERTU®-Experienced HER2-Positive Breast Cancer	FPI Q4 2025 Interim data – Q3 2026
ALX2004		
ALX2004 Dose-escalation and expansion	EGFR-Expressing Solid Tumors	FPI anticipated August 2025 Initial safety data – 1H 2026

Projected Cash Runway Now into Q1 2027

Cash, cash equivalents, and investments of \$84M as of June 30, 2025



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