

ASPEN-06
Phase 2 Interim
Gastric/GEJ Cancer Data
Conference Call

October 03, 2023

Forward-looking statements

Certain information set forth in this presentation contains “forward-looking information”, under applicable laws collectively referred to herein as forward-looking statements. Except for statements of historical fact, information contained herein constitutes forward-looking statements and includes, but is not limited to the (i) results and cost and timing of our product development activities and clinical trials; (ii) completion of the Company’s clinical trials that are currently underway, in development or otherwise under consideration; (iii) our expectations about the timing of achieving regulatory approval and the cost of our development programs; (iv) projected financial performance of the Company; (v) the expected development of the Company’s business, projects, collaborations and joint ventures; (vi) execution of the Company’s vision and growth strategy, including with respect to future M&A activity and global growth; (vii) sources and availability of third-party financing for the Company’s research and development; (viii) future liquidity, working capital, and capital requirements; and (ix) industry trends. These and other risks are described more fully in ALX Oncology’s filings with the Securities and Exchange Commission (“SEC”), including ALX Oncology’s Annual Report on Form 10-K and other documents ALX Oncology files with the SEC from time to time.

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This presentation concerns product candidates that are under clinical investigation, and which have not yet been approved for marketing by the U.S. Food and Drug Administration. These product candidates are currently limited by federal law to investigational use, and no representation is made as to their safety or effectiveness for the purposes for which they are being investigated.

This presentation also contains estimates and other statistical data made by independent parties and by ALX Oncology relating to market size and growth and other industry data. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions, and estimates of ALX Oncology’s future performance and the future performance of the markets in which it operates are necessarily subject to a high degree of uncertainty and risk.

ALX Oncology program update call

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Welcome



Jason Lettmann
CEO, ALX Oncology

2

Evorpaccept program
overview



Dr. Sophia Randolph, MD, PhD
CMO, ALX Oncology

3

Gastric/GEJ cancer
overview



Dr. Josep Taberero, MD, PhD
Chief of Medical Oncology,
Vall d'Hebron University Hospital,
Barcelona, Spain

4

ASPEN-06
Prespecified interim analysis

5

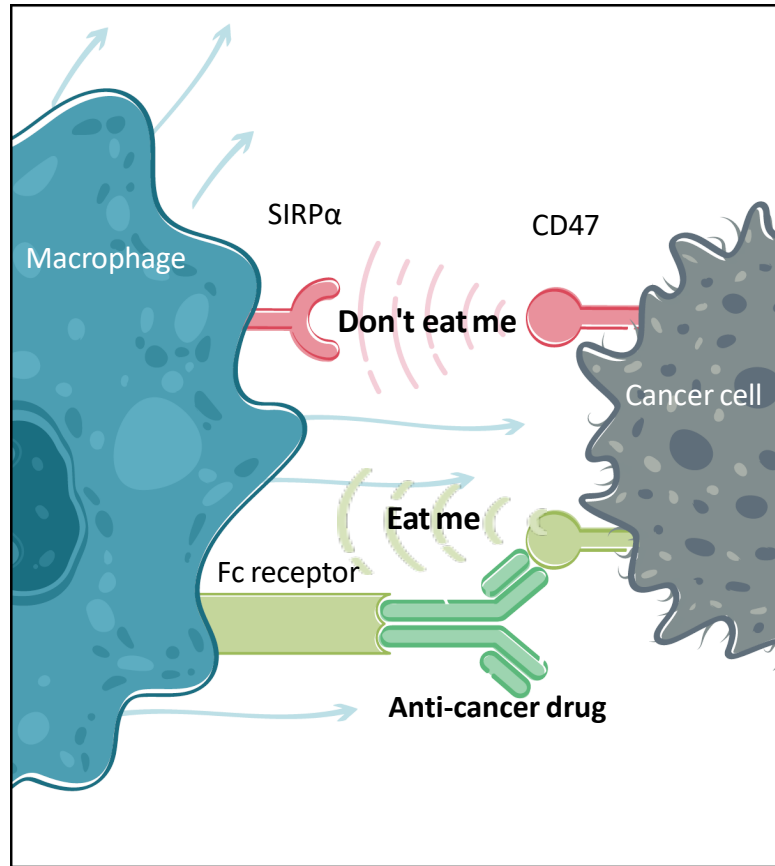
Closing remarks



Jason Lettmann
CEO, ALX Oncology

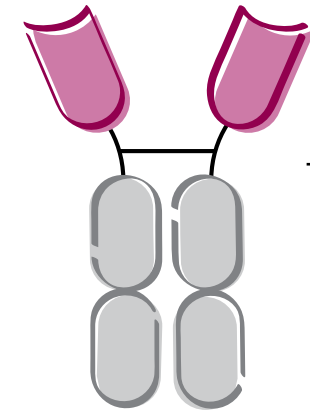
AGENDA

Evorpaccept: A first-in-class approach to targeting CD47



Target cells overexpress CD47 to evade destruction by macrophages

High affinity CD47 binding domains of SIRPα

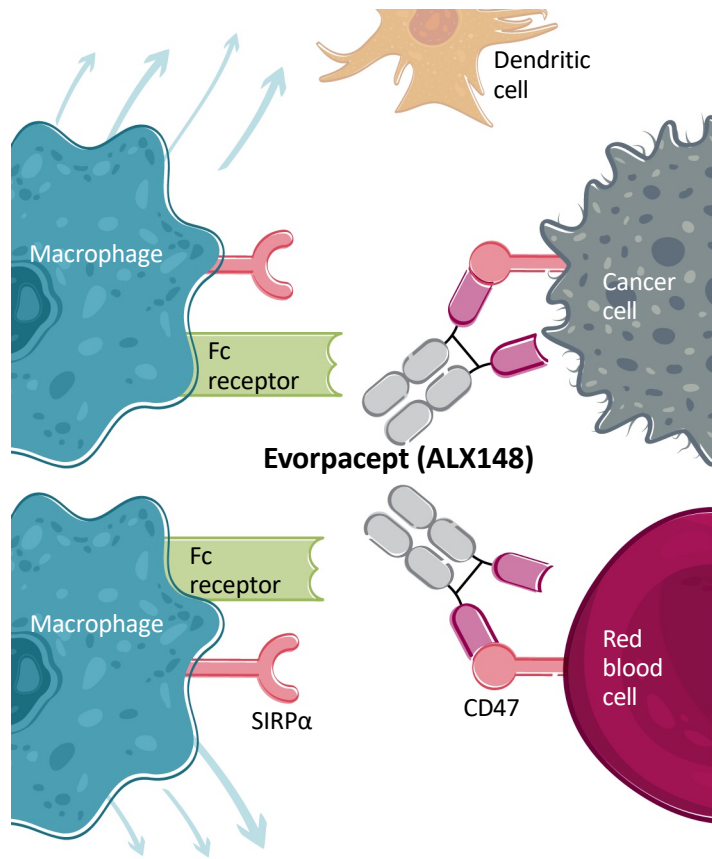


Inactive Fc domain

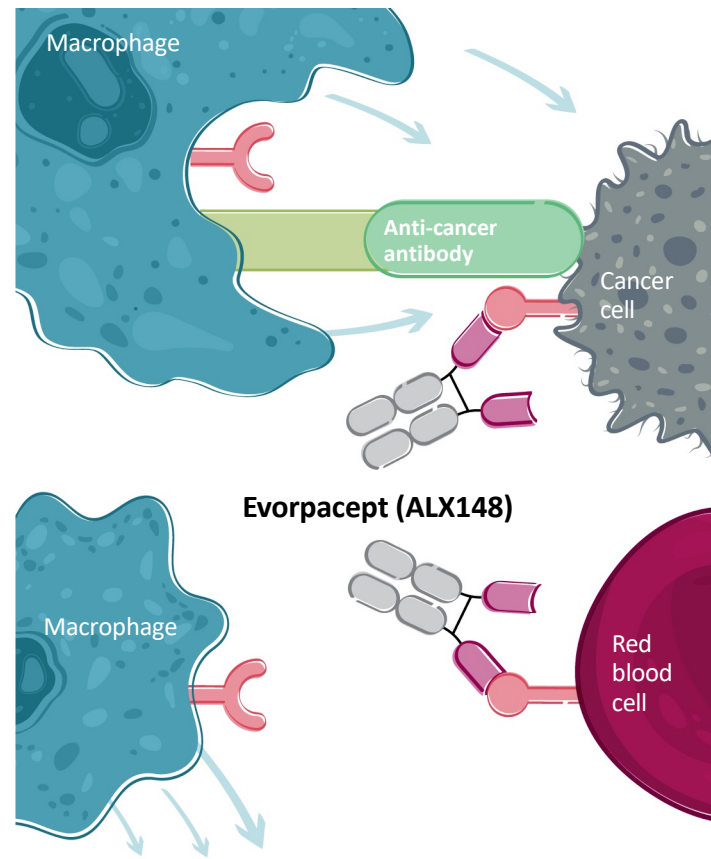
Evorpaccept

A differentiated CD47 blocker

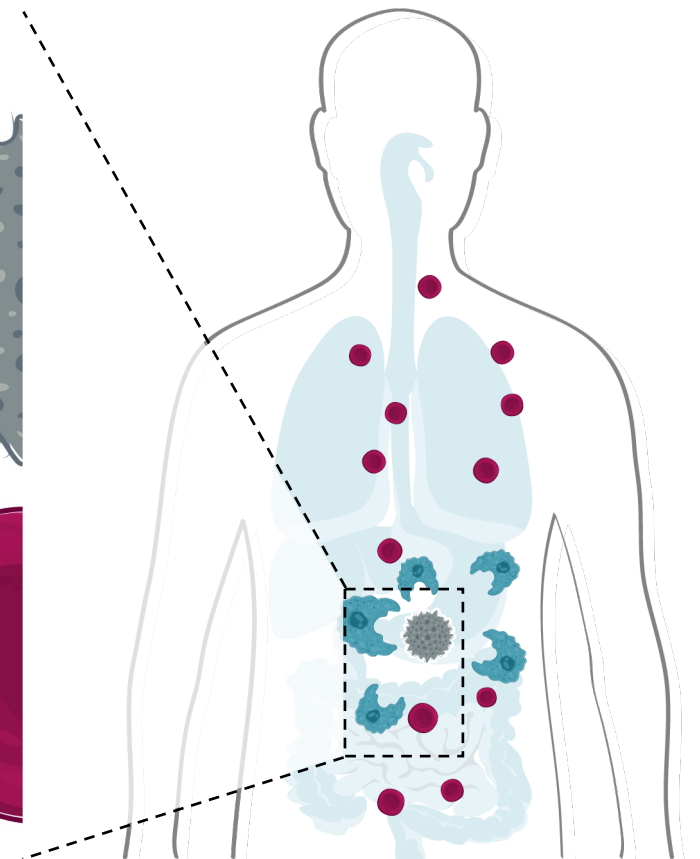
Evorpacept targets the CD47 checkpoint



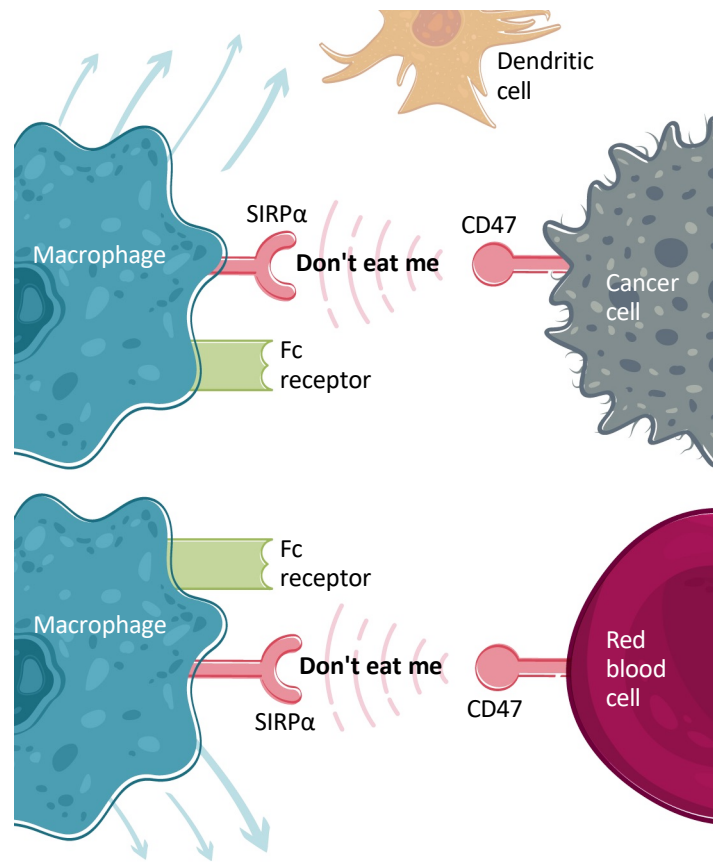
Complete CD47 blockade without targeting blood cells



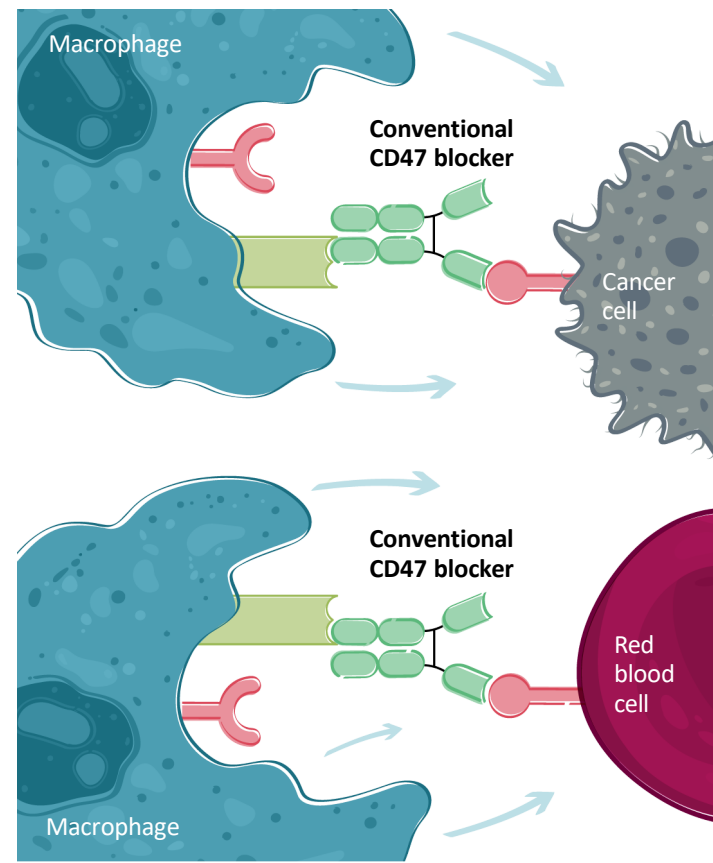
Combined with cancer therapy to specifically target cancer cells



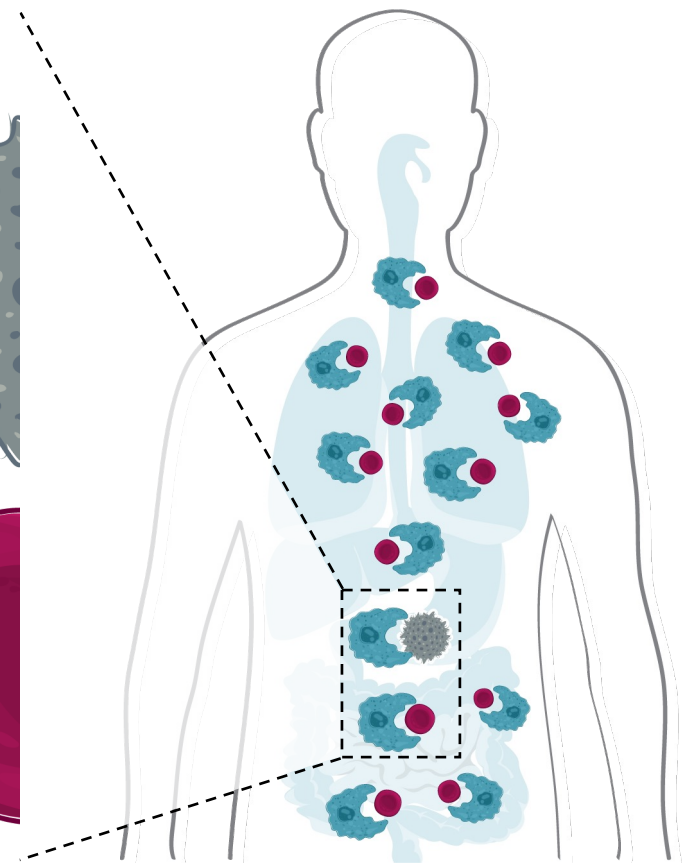
Conventional CD47 targeting is more toxic and less efficacious



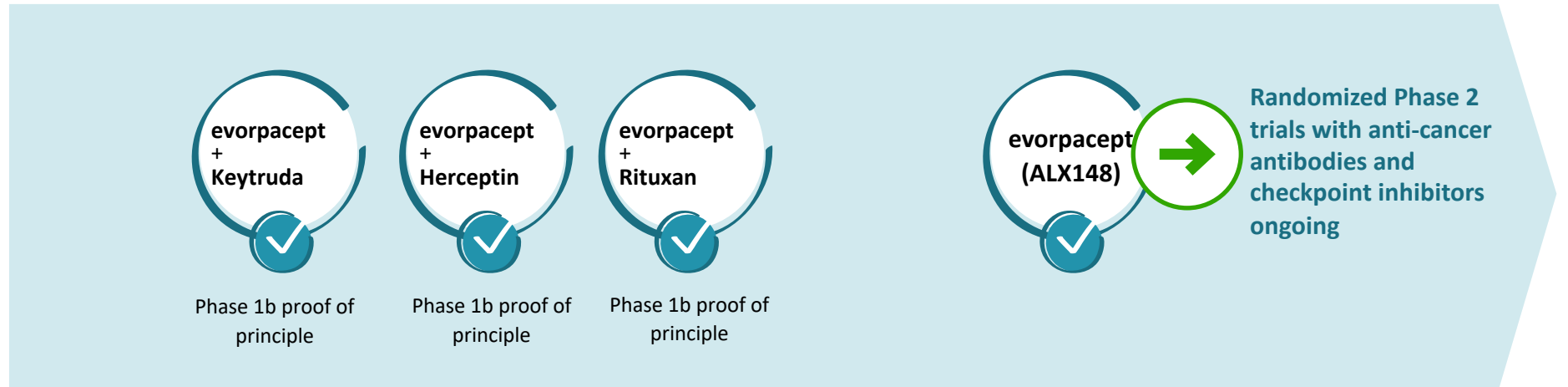
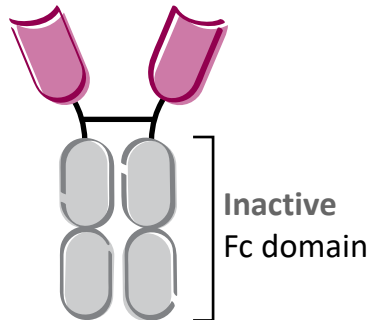
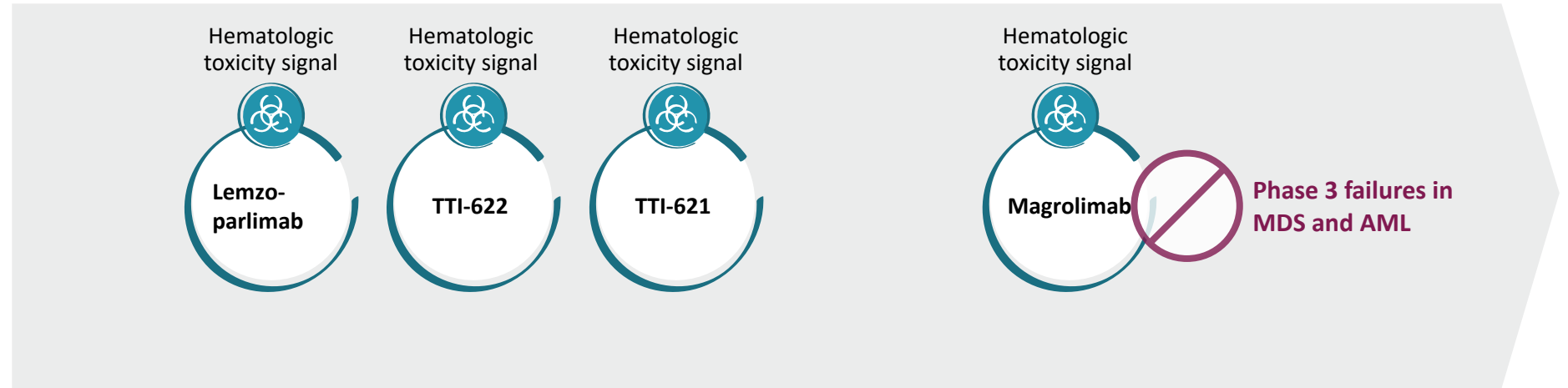
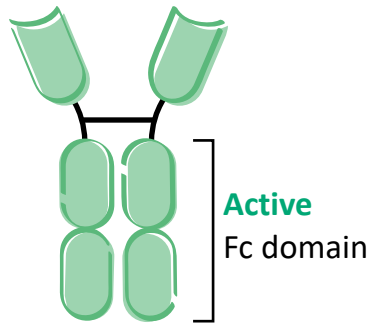
CD47 is widely expressed in both healthy and cancer cells



Indiscriminate CD47 inhibition with an active Fc will target healthy cells



Evorpaccept has demonstrated consistent tolerability and meaningful clinical activity vs. conventional approaches



Evorpacept: Pursuing a robust development plan

Indication		Evorpacept Combination Agent	Discovery	IND Enabling	Phase 1	Phase 2	Phase 3	Fast Track	Collaboration Partner	
Evorpacept Combination Studies	ANTI-CANCER ANTIBODIES AND ADCs	GC Gastric/Gastroesophageal Junction Cancer	Herceptin + Cyramza + Paclitaxel (ASPEN-06)	interim data Q4 2023				✓	Lilly	
		Urothelial Cancer	Padcev (ASPEN-07)							
		Breast Cancer	Zanidatamab							Jazz Pharmaceuticals
			Enhertu (I-SPY)							Quantum Leap Healthcare Collaborative
		MM Multiple Myeloma	Sarclisa + Dexamethasone							sanofi
CHECKPOINT INHIBITORS	HNSCC Head And Neck Squamous Cell Carcinoma	Keytruda (ASPEN-03)						✓	MERCK	
		Keytruda + 5FU + Platinum (ASPEN-04)						✓	MERCK	

Evorpaccept: Potential best-in-class CD47 blocker with consistent clinical activity and tolerability

✓ **ASPEN-06: The first prospective, randomized clinical study in the CD47 space in solid tumors**
Strong evidence that evorpaccept in combination with an anti-cancer targeted antibody improves clinical response in a population with advanced malignancy

✓ **ASPEN-06: First positive randomized data in HER2+ gastric/GEJ cancer in a population reflecting current standard of care**
The prespecified interim analysis shows evorpaccept + trastuzumab, ramucirumab, paclitaxel (Evo+TRP) compares favorably to both ramucirumab + paclitaxel (RAINBOW) as well as trastuzumab deruxtecan (DESTINY-Gastric01) in 2L and 3L gastric/GEJ cancer patients, many of whom had prior checkpoint inhibitor and trastuzumab deruxtecan (Enhertu) exposure


✓ **ASPEN-06: Interim data closely tracks both safety and efficacy data observed in the ASPEN-01 phase 1b study**
Initial randomized data shows that Evo+TRP is generally well tolerated and has improved clinical activity compared to TRP alone consistent with the positive contribution of evorpaccept to the backbone therapy


✓ **ASPEN-06: Interim data support the potential for a new standard of care for advanced gastric/GEJ cancer patients with final analysis anticipated to be completed by Q2 2024**


ASPEN-06: Registration strategy for evorpaccept in gastric/GEJ cancer

Proof of principle

ASPEN-01 Phase 1b HER2+ gastric/GEJ cancer


 South Korea, USA

 Patients: **R/R \geq 2L** with prior HER2 targeted therapy + chemotherapy
N=18

 Treatment:

Evo 10 and 15 mg/kg (QW)

+ T + R + P

 Endpoint:

Safety of combination
Anti-cancer activity


Legend:


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


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Safety of combination
Anti-cancer activity

Proof of concept

ASPEN-06 Randomized Phase 2 HER2+ gastric/GEJ cancer

 Asia, Australia, Europe and North America

 Patients: **2L/3L** with prior HER2 targeted therapy + chemotherapy
N=~122

 Treatment (1:1 randomization):

Evo 30 mg/kg (Q2W)

+ **T** + **R** + **P**

vs.

Control:

T + **R** + **P**

 Endpoint:

Anti-cancer activity:
Primary endpoint: ORR
Secondary: DOR, PFS, OS

Legend:

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

ASPEN-06: Registration strategy for evorpcept in gastric/GEJ cancer

Proof of principle

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Proof of concept

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Patients: **2L/3L** with prior HER2 targeted therapy + chemotherapy
N=~122

Treatment (1:1 randomization):

Evo 30 mg/kg (Q2W)

+ **T** + **R** + **P**

vs.

Control:

T + **R** + **P**

Endpoint:

Anti-cancer activity:
Primary endpoint: ORR
Secondary: DOR, PFS, OS

Registrational

ASPEN-06 Randomized phase 3 HER2+ gastric/GEJ cancer

Worldwide

Patients: **2L /3L** with prior HER2 targeted therapy + chemotherapy

Treatment (randomized):

Evo 30 mg/kg (Q2W)

+ **T** + **R** + **P**

vs.

Control:

R + **P**

Endpoint:

Anti-cancer activity: including OS,
PFS, ORR, DOR

Legend:

Evo Evorpcept

T Trastuzumab

R Ramucirumab

P Paclitaxel

Professor Josep Taberero, MD, PhD



Head of the Medical Oncology Department

at the Vall d'Hebron University Hospital in Barcelona, Spain



Director

of the Vall d'Hebron Institute of Oncology (VHIO)



Research interests

Development of personalized cancer medicines and identification of predictive biomarkers of anti-cancer response to therapies



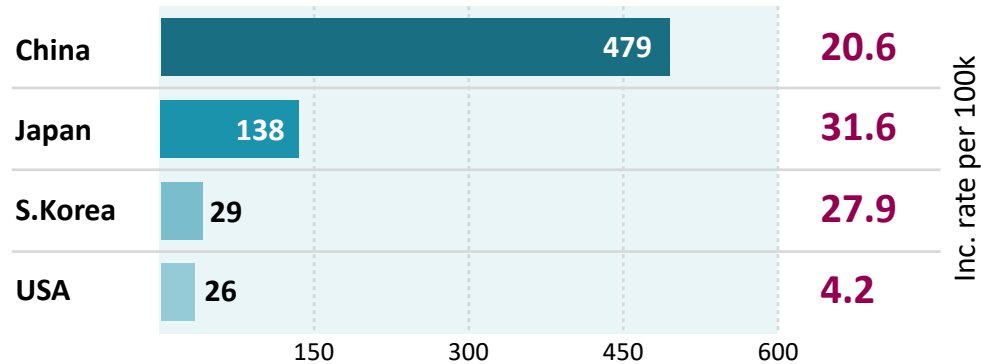
- Executive Board
- President 2018 – 2019



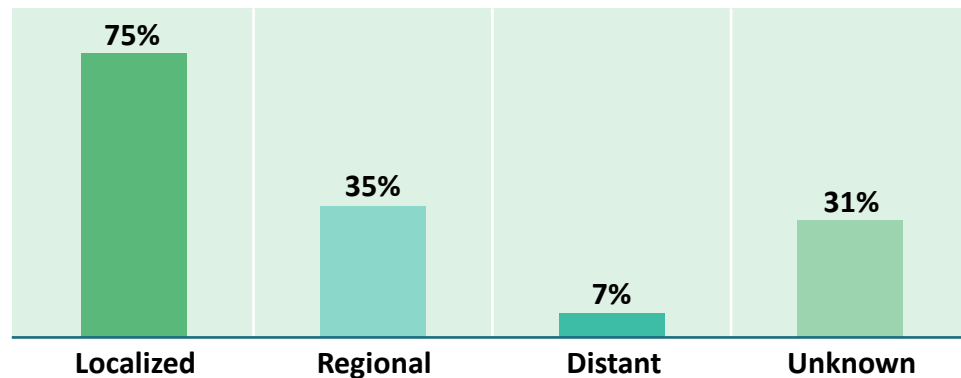
Member of several educational and scientific committees

With a global unmet need, advanced gastric/GEJ cancer provides the initial population to clinically validate evorpaccept's mechanism of action

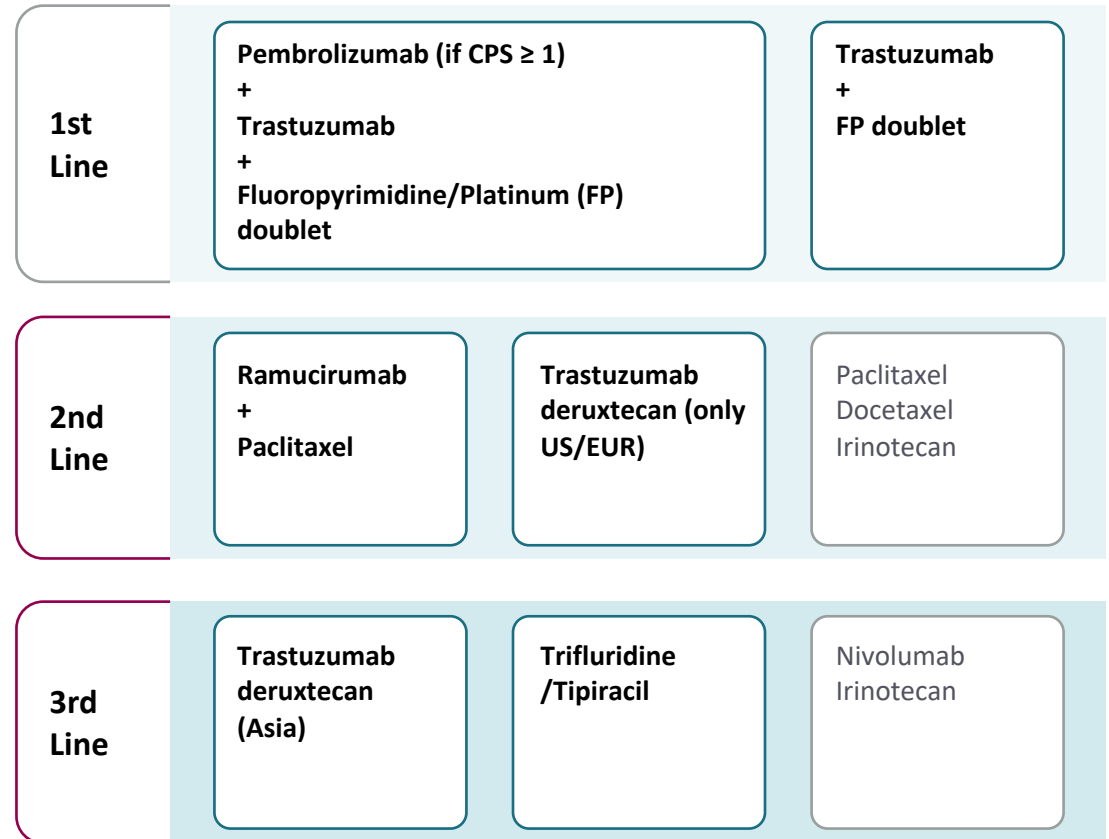
Annual new cases and ASR incidence per 100,000¹



5-Year survival by stage at diagnosis in US²



HER2+ treatment SOC by line of therapy







¹ WHO/IARC data accessed September 14, 2023 for most recent year, 2020; ASR = Age Standardized Rate;

² SEER Cancer Stats accessed September 14, 2023

Current HER2+ gastric/GEJ cancer standard of care reflects the need for novel combinations in 2L/3L

HER2+ treatment benchmarks:

RAINBOW¹ 2L





	ORR (%)	DOR	PFS	OS
Ramucirumab/Paclitaxel  N=330	 28%	4.4 months <small>IQR 2.8–7.5</small>	4.4 months <small>4.2-5.3</small>	9.6 months <small>8.5-10.8</small>
Paclitaxel  N=335	 16%	2.8 months <small>IQR 1.4-4.4</small>	2.9 months <small>2.8-3.0</small>	7.4 months <small>6.3-8.4</small>

THE LANCET
Oncology

Volume 15, ISSUE 11, P1224-1235,
October 2014

Ramucirumab plus paclitaxel versus placebo plus paclitaxel in patients with previously treated advanced gastric or gastro-oesophageal junction adenocarcinoma (RAINBOW): a double-blind, randomised phase 3 trial

DESTINY-Gastric01² 3L

Trastuzumab deruxtecan  N=126	 41%	11.3 months <small>5.6-NE</small>	5.6 months <small>4.3-6.9</small>	12.5 months <small>9.6-14.3</small>
Physicians' choice  N=62	 11%	3.9 months <small>3.0-4.9</small>	3.5 months <small>2.0-4.3</small>	8.4 months <small>6.9-10.7</small>



The NEW ENGLAND
JOURNAL of MEDICINE

Volume 382: P2419-2430
June 2020

Trastuzumab deruxtecan in previously treated HER2-positive gastric cancer – DESTINY-Gastric-01

Both large, randomized studies demonstrated modest response rates and survival benefit of ~1 year or less highlighting significant unmet medical need

¹ Wilke et al, Lancet October 2014,

² Enhertu US product insert, and Shitara et al, NEJM June 18, 2020; NE could not be estimated

ASPEN-06: Evorpaccept in combination with trastuzumab, ramucirumab, and paclitaxel in patients with advanced HER2-overexpressing gastric/GEJ adenocarcinoma


Key eligibility criteria:

HER2+ advanced or metastatic gastric or gastroesophageal junction adenocarcinoma

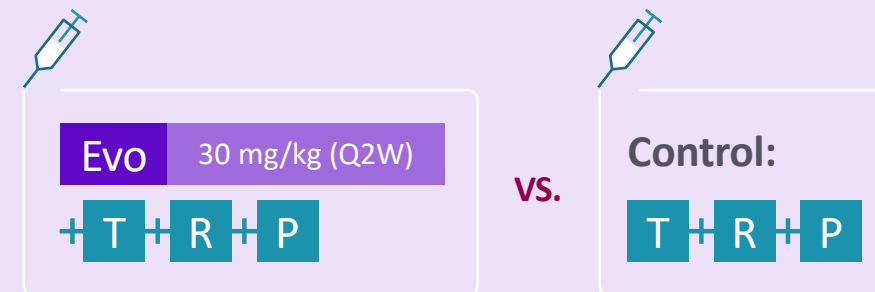
2nd line or 3rd line

- ✗ No prior treatment:
Anti-CD47 agent, an anti-SIRP agent or ramucirumab.
- ✓ Prior treatment ok:
Trastuzumab deruxtecan (Enhertu) and checkpoint inhibitors

ASPEN-06 randomized phase 2

 N=122

Treatment (1:1 randomization):



 Endpoint:

Primary: ORR
Secondary: DOR, PFS, OS

Interim analysis (N=54):

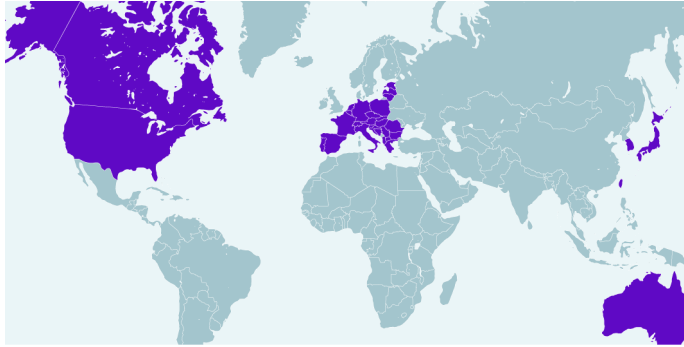
Futility met if Evo+TRP has 30% ORR or if there are more responders in TRP arm;

Final analysis (N=122):

80% power to see a 50% improvement in ORR compared to historical RP and 68% power to see 10% delta between both arms.

ASPEN-06 interim analysis: Evorpaccept administered in combination with TRP versus TRP alone

Study sites:



ASPEN-06

Patients are enrolled across 13 countries in Asia, Australia, Europe and North America.

Study regimen dose administration:

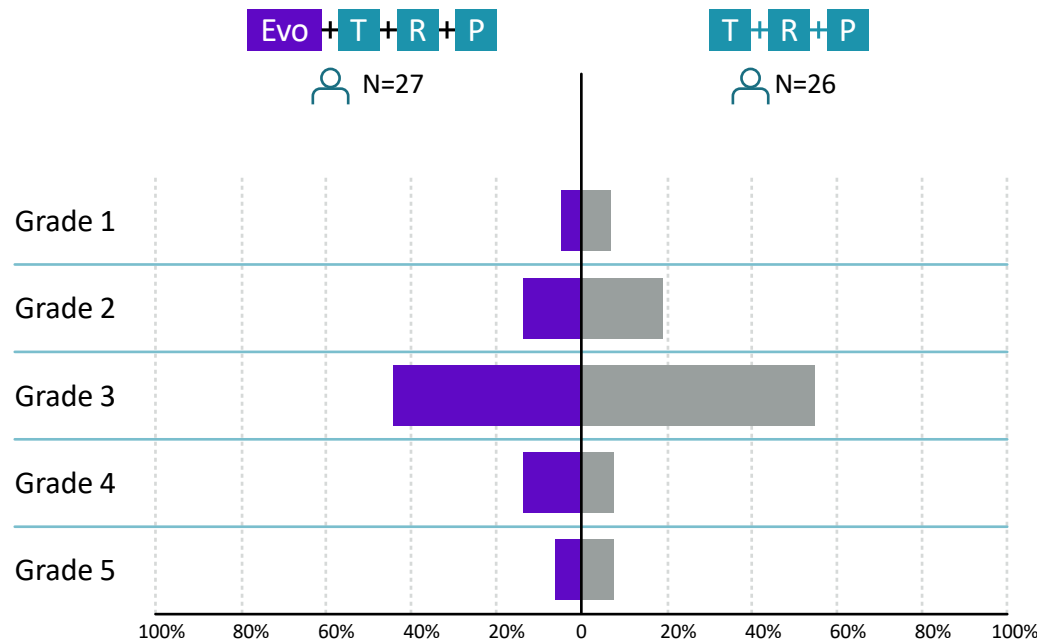
- Evo** Evorpaccept• **30 mg/kg IV Q2W**
- +
- T** Trastuzumab• **6 mg/kg > 4 mg/kg Q2W**
- +
- R** Ramucirumab• **8 mg/kg Q2W**
- +
- P** Paclitaxel• **80 mg/m²**
Days: 1, 8, 15 of 28-day cycle

Study population:

		Evo + T + R + P N=27	Control: T + R + P N=27
Median age, years (range)		65 (41-79)	57 (31-81)
Sex, n%	Male	85	70
	Female	15	30
Race, n%	Asian	52	48
	White	26	30
	Other	3.7	0
	Unknown	18.5	22
ECOG PS, n%	0	52	52
	1	48	48
GEJ, n%		15	22



Evo+TRP was generally well tolerated with a safety profile consistent with that of the backbone TRP therapy

All causality adverse events, by grade



- Evo+TRP was generally well tolerated
- The incidence of adverse events due to any cause was comparable by arm
- The incidence of cytopenias was evenly distributed by arm
- There were no on study treatment-related deaths on either arm
- Evorpcept's safety profile was consistent with its prior experience in over 400 patients that have been treated to date

ASPEN-06 interim analysis: Clinical activity of evorpaccept + TRP supports substantial contribution of evorpaccept to TRP and compares favorably to current SOC

	Evo + T + R + P  N=27	Control: T + R + P  N=27
Confirmed objective response	52%	22%
Complete response	4%	0%
Partial response	48%	22%
Duration of response	NR [3.6, NR]	7.4 [3.5, NR]

- Evo-TRP has shown substantial response activity over TRP backbone
- Initial clinical activity of evorpaccept + TRP compares favorably to ramucirumab + paclitaxel (28% ORR, 4.4 DOR)¹ as well as to trastuzumab deruxtecan (41% ORR, 11.3 DOR)²

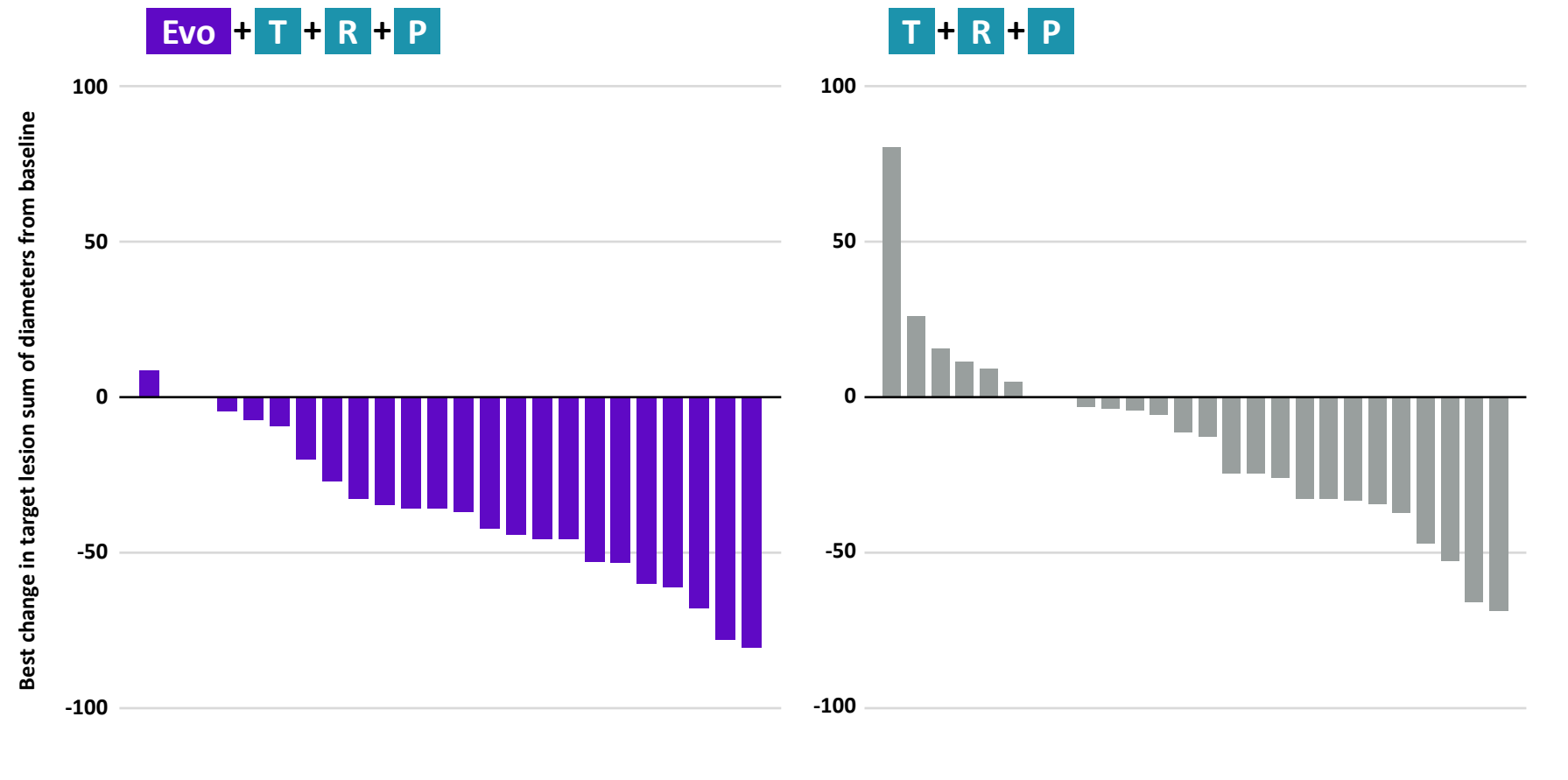
Data Cutoff as of 29 August 2023

¹ Wilke et al, Lancet October 2014,

² Enhertu US product insert, and Shitara et al, NEJM June 18, 2020; NR not reached

ASPEN-06 interim analysis: Substantial tumor shrinkage is seen in ASPEN-06 gastric/GEJ cancer patients receiving Evo-TRP compared to TRP

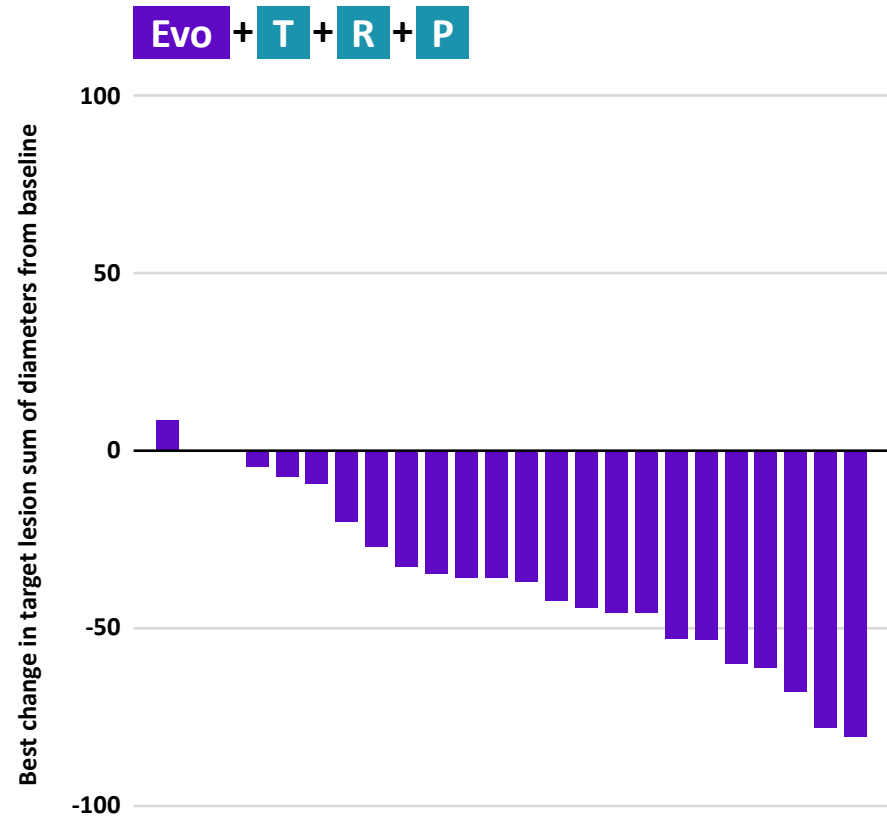
ASPEN-06 Randomized Phase 2



• Best percentage-change in target lesions from baseline reflects anti-cancer activity in most patients

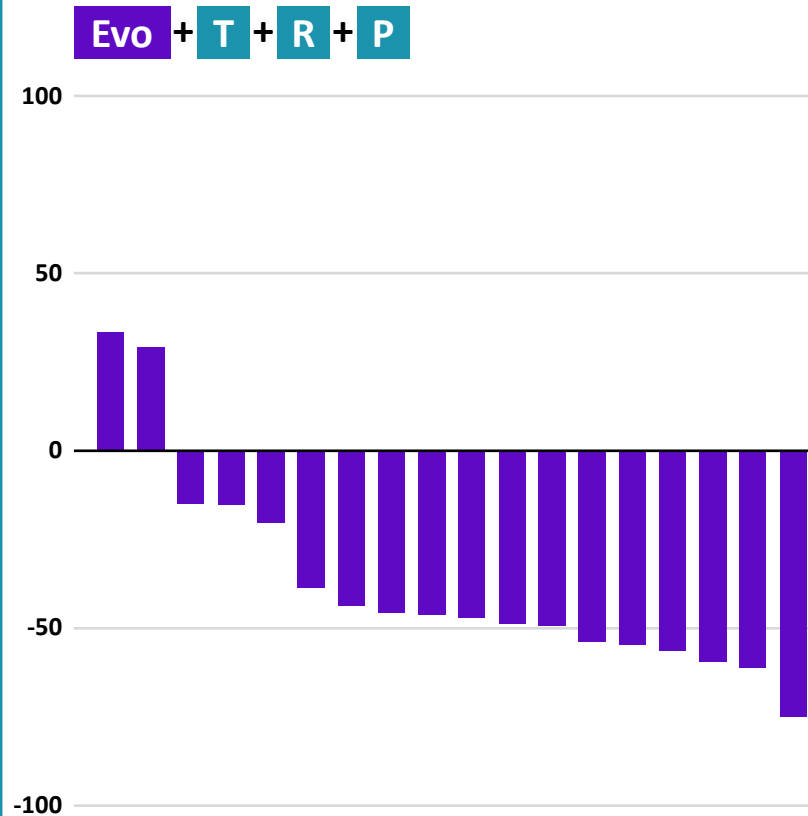
Robust tumor shrinkage is consistently seen in gastric/GEJ cancer patients receiving Evo-TRP across both ASPEN-06 and ASPEN-01

ASPEN-06 Randomized Phase 2 Study



Data Cutoff as of 29 August 2023

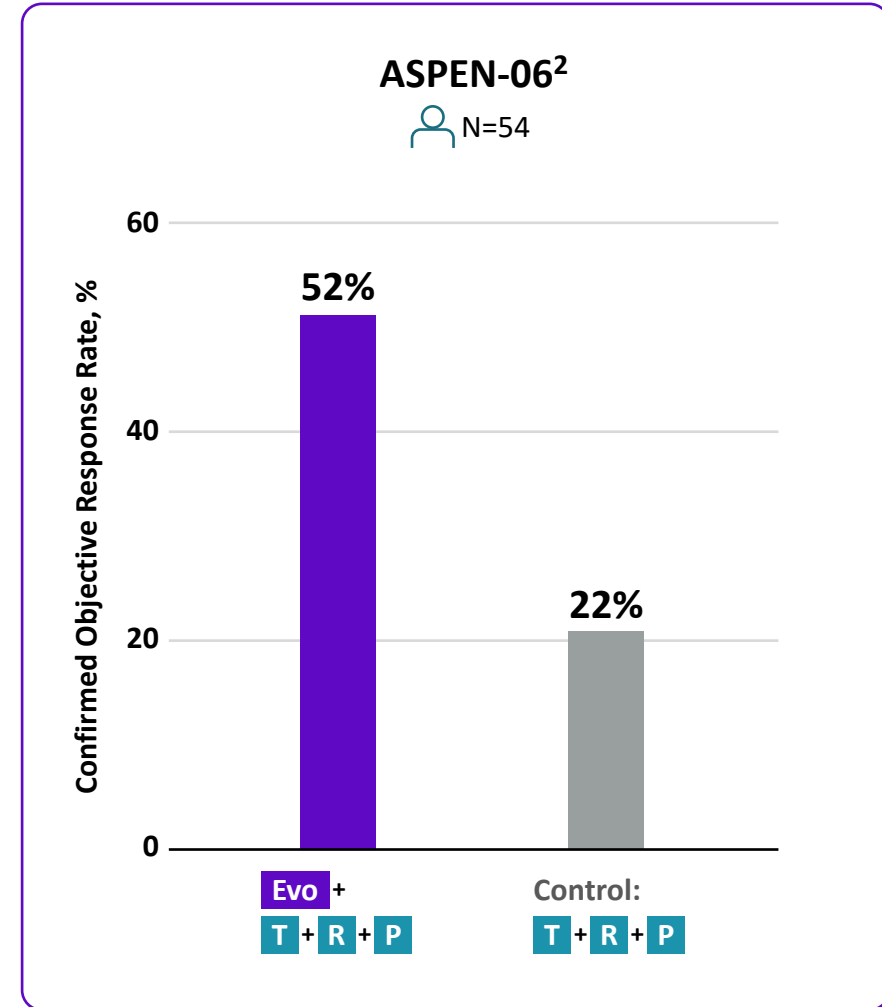
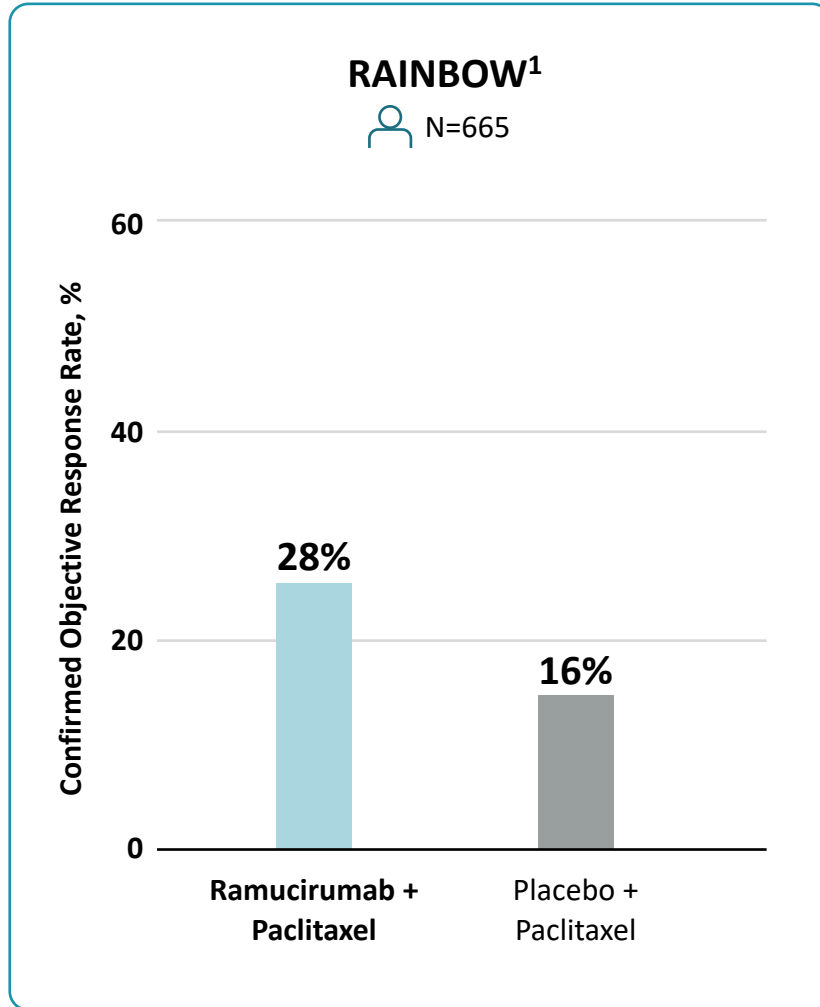
ASPEN-01 Phase 1b HER2+ GC Cohort



Data Cutoff September 1, 2021

• Responses observed across ASPEN-06 Ph2 study and ASPEN-01 Ph1b study were similar

ASPEN-06 in the context of the regulatory benchmark RAINBOW study



¹ Wilke et al, Lancet October 2014,

² ASPEN-06 IA Data as of 29 August 2023

Summary: Evorpacept demonstrates the power of engaging the innate immune response in combination with TRP anti-cancer targeted therapy in patients with gastric/GEJ cancer

Robust Clinical Activity

At the interim analysis, evorpacept demonstrates an **ORR of 52%** with an **unreached mDOR** in patients with HER2+ gastric/GEJ cancer in combination with TRP in a **contemporary 2L and 3L global population with substantial checkpoint inhibitor and trastuzumab deruxtecan (Enhertu) exposure**

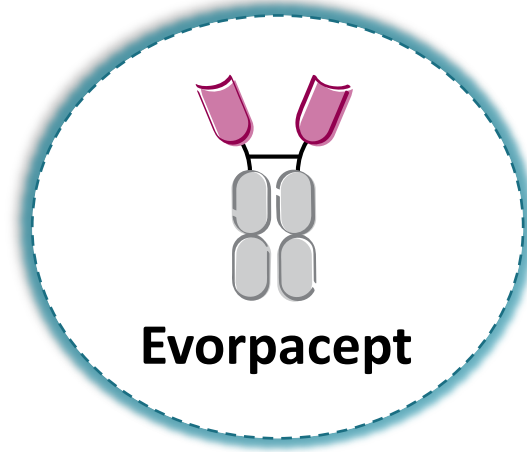
Well-Tolerated

ASPEN-06 interim data confirms that **evorpacept can be combined with TRP** with a favorable safety profile that was consistent with data from the **>400 patients treated to date**

Consistent Results

As the **first randomized data in the solid tumor setting in the CD47 space**, the interim data from ASPEN-06 further demonstrates evorpacept's encouraging safety profile and clinical activity and is in line with earlier data readouts

Evorpacept's differentiated design results in differentiated safety and clinical activity



Higher affinity
CD47 binding



More potently blocks CD47 signal on cancer cells

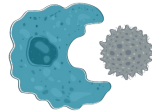
Inactive Fc
domain



Less "sink effect" = more targeted

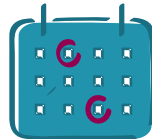
No known dose dependent cytopenia = higher dosing

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

Robust clinical
activity

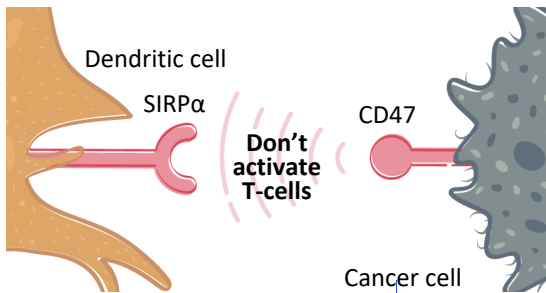
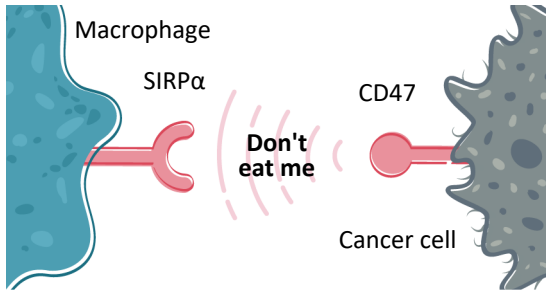
Best-in-class
safety profile

Strong solid tumor
activity

Broad combination
potential

Validated approach and our path to success

2 potential “First-In-Class” mechanisms of action



5 positive clinical readouts across multiple studies

- ✓ Ph2 Gastric/GEJ cancer randomized interim data with TRP
- ✓ Ph1b NHL data with Rituxan
- ✓ Ph1b Gastric/GEJ cancer data with TRP

- ✓ Ph1b ≥2L Head and Neck cancer (HNSCC) data with Keytruda
- ✓ Ph1b 1L HNSCC data with Keytruda + chemotherapy

9 ongoing studies in new indications and combinations



- Ph2 Gastric/GEJ cancer randomized study with TRP
- Ph1b Multiple myeloma study with Sarclisa
- Ph1b Non-Hodgkin lymphoma IST
- Ph1b Breast cancer study with zanidatamab

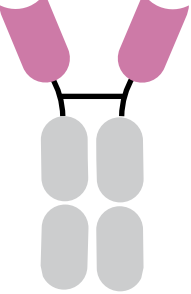


- Ph1b Urothelial carcinoma study with Padcev
- Ph1b Breast cancer study (I-Spy) with Enhertu



- Ph2 1L HNSCC randomized study with Keytruda
- Ph2 1L HNSCC randomized study with Keytruda + chemotherapy
- Ph2a 2L Ovarian cancer study with Keytruda + chemotherapy

Anticipated upcoming milestones: Significant catalysts in 2024

	1H 2024	2H 2024
 <p>Evorpcept</p>	<p>Gastric/GEJ Cancer (Phase 2) ASPEN-06 Top line final results in gastric/ GEJ from randomized trial with TRP – Q2 2024</p>	<p>Head & Neck Cancer (Phase 2) ASPEN-03 Top line results in HNSCC from randomized trial with Keytruda</p>
	<p>Non-Hodgkin Lymphoma (NHL) Phase 1B study Data from Phase 1B IST study – Q1/ Q2 2024</p>	<p>Head & Neck Cancer (Phase 2) ASPEN-04 Top line results in HNSCC from randomized trial with Keytruda and chemotherapy</p>
		<p>Gastric/GEJ Cancer (Phase 3) ASPEN-06 Initiation of registrational randomized gastric/GEJ cancer trial</p>
		<p>Urothelial Carcinoma (Phase 1b) ASPEN-07 Top line results in urothelial carcinoma with Padcev</p>
		<p>Breast Cancer (Phase 1b) I-SPY Top line results in breast cancer with Enhertu</p>
<p>Early clinical / pipeline</p>	<p>ADC pipeline Identify clinical development candidates in Q1 2024</p>	
	<p>ALTA-002 (Phase 1) initiation File IND in Q1 2024</p>	

THANK YOU