# **ALX Oncology**

**Corporate Presentation** 

April 2024

Don't eat me

#### **Forward-looking statements**

Certain information set forth in this presentation contains "forward-looking information", under applicable laws collectively referred to herein as forwardlooking 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 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: The CD47 Leader

ALX Oncology is advancing a highly differentiated immuno-oncology pipeline led by evorpacept, a potential best and first-in-class CD47 innate immune system checkpoint inhibitor that has been studied in over 500 patients

Evorpacept is the first CD47 inhibitor to demonstrate robust clinical activity and a differentiated safety profile across both solid and hematologic tumors highlighted by the first positive randomized data in the field in gastric cancer in Q4 '23

A prespecified interim analysis of ASPEN-06, a randomized Ph2 study for the treatment of advanced HER2+ gastric/GEJ cancer, showed a confirmed overall response rate for the evorpacept arm of 52% vs. 22% for control and encouraging early durability

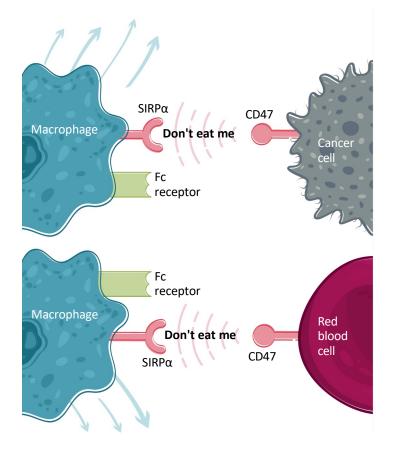
Multiple positive clinical studies across NHL, gastric, and head and neck (HNSCC) have been completed to date and currently pursuing additional studies in combination with 3 therapeutic classes: anti-cancer antibodies, checkpoint inhibitors and ADCs

Significant upcoming milestones anticipated in 2024 for evorpacept include final top line results from the Ph2 gastric/GEJ study, results from two randomized Ph2 studies in HNSCC, and new clinical data in NHL (AACR 2024), breast, and urothelial cancer

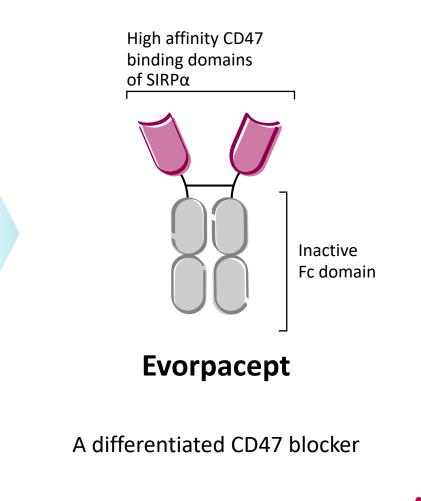
Expanding evorpacept to new indications and building a strong pipeline beyond evorpacept supported by multiple pharma partnerships and a strong balance sheet with cash runway into early 2026



# **Evorpacept: A first-in-class approach to targeting CD47**

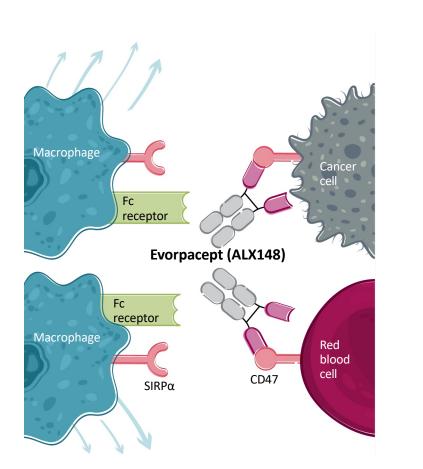


Target cells overexpress CD47 to evade destruction by macrophages

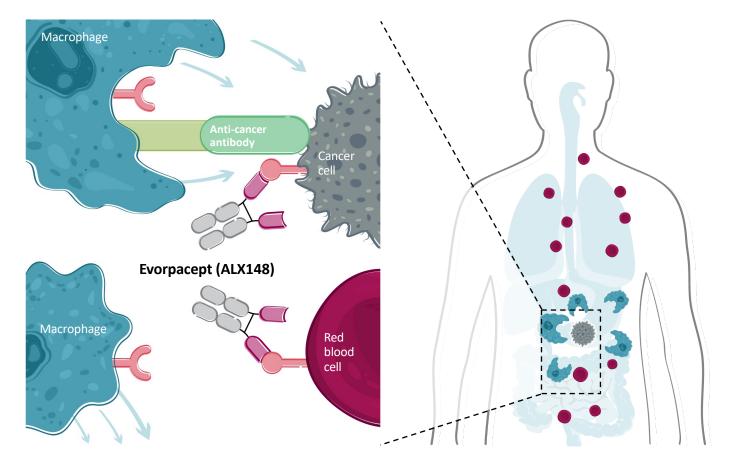


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# **Evorpacept targets the CD47 checkpoint**



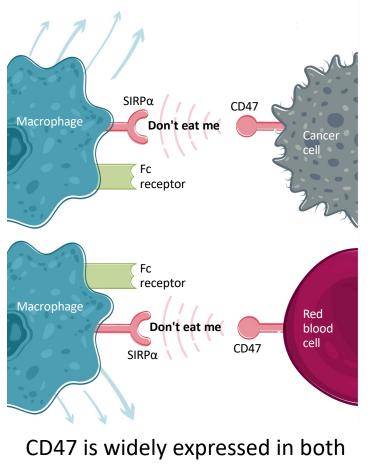
Complete CD47 blockade without targeting blood cells



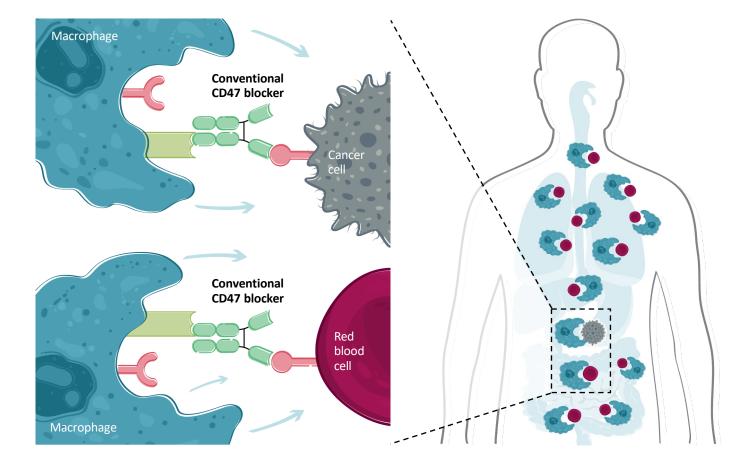
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Combined with cancer therapy to specifically target cancer cells

# **Conventional CD47 targeting is more toxic and less efficacious**



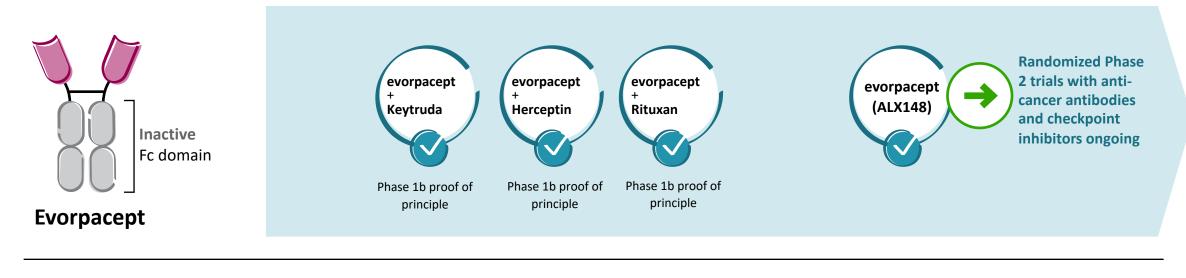
healthy and cancer cells

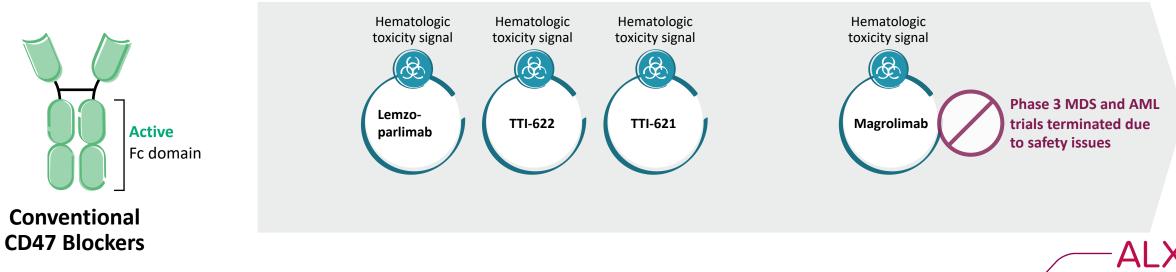


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Indiscriminate CD47 inhibition with an active Fc will target healthy cells

# Evorpacept has demonstrated consistent tolerability and robust clinical activity vs. conventional approaches

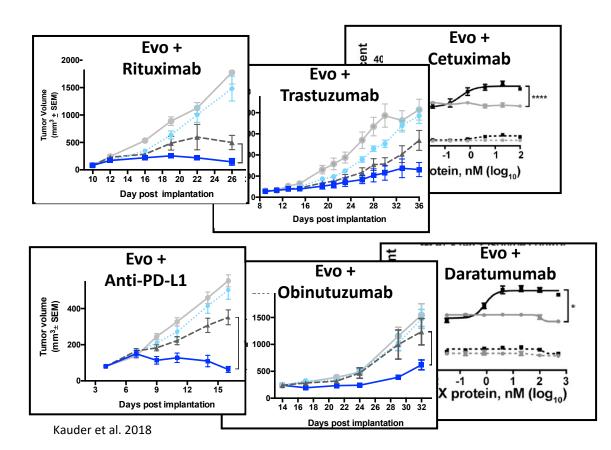




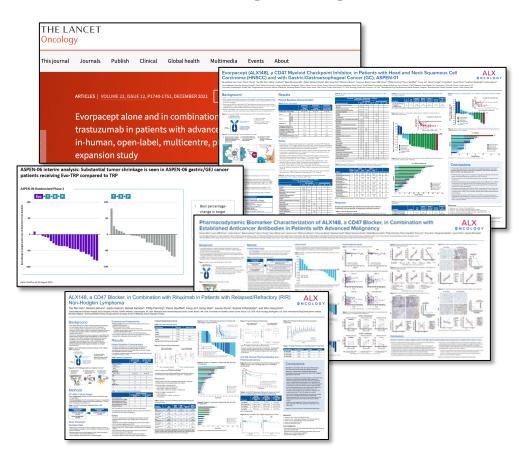
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## Evorpacept's consistent activity profile is due to its distinct molecular design

**Evorpacept enhanced preclinical antitumor activity across multiple classes of therapies...** 

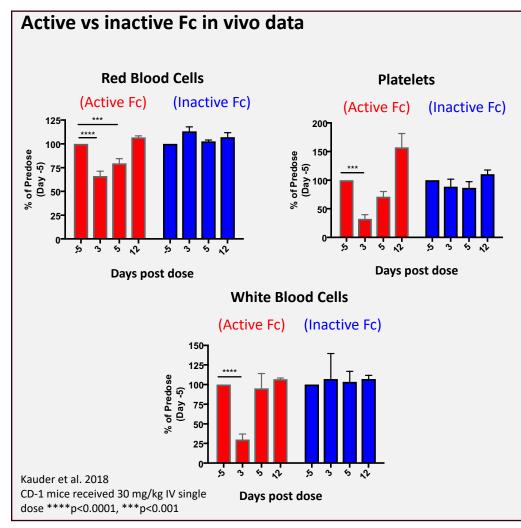


#### ...translated to 5 positive clinical studies across both solid and hematological malignancies





### Evorpacept has demonstrated a consistent tolerability profile across multiple tumors & combinations



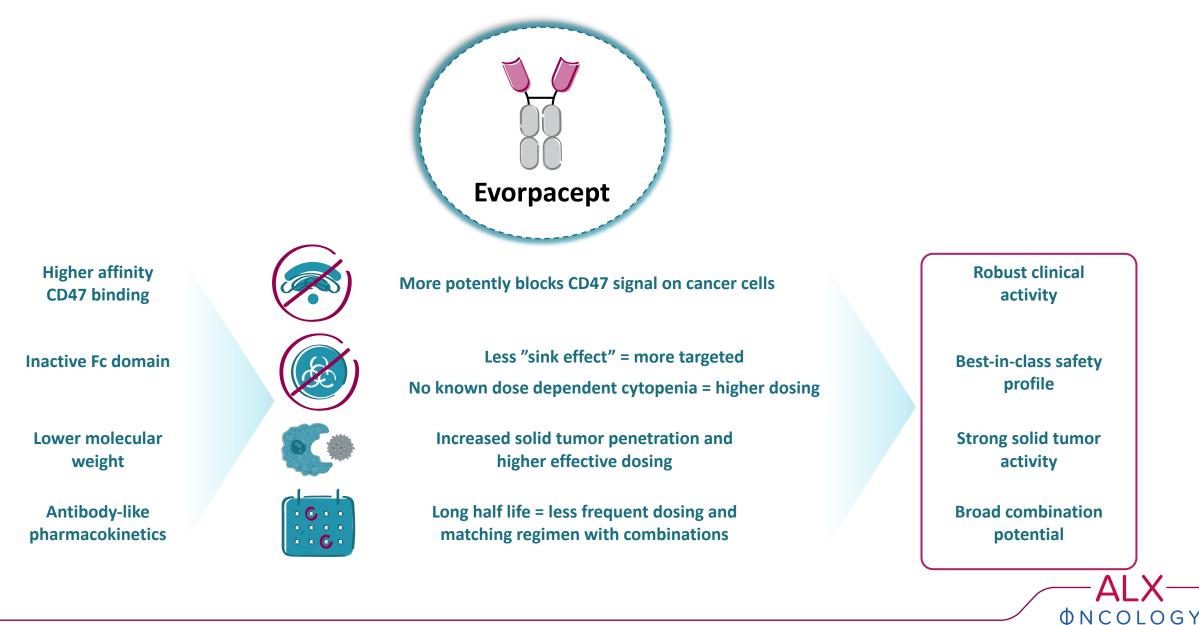
Treatment related adverse events	+ Ċyramza	+ Herceptin a + chemo =18)	Keyt + chem	evorpacept + Keytruda + chemo (N=13) Total n (%) ≥Grade 3		acept + ruda 52)
Fatigue	2 (11.1%)	≥Grade 3	1 (7.7%)	≥Grade 3	Total n (%) 6 (11.5%)	
Rash / dermatitis acneiform	4 (22.2%)	-	-	-	5 (9.6%)	-
AST increased	-	-	-	-	9 (17.3%)	-
Platelets decreased	-	-	-	-	4 (7.7%)	
ALT increased	-	-	-	-	7 (13.5%)	
Pruritus	2 (11.1%)	-	-	-	5 (9.6%)	-
Pyrexia	-	-	-	-	3 (5.8%)	-
Decreased appetite	-	-	-	-	2 (3.8%)	-
Anemia	1 (5.6%)	-	1 (7.7%)	1 (7.7%)	5 (9.6%)	1 (1.9%)
Infusion reaction	-	-	-	-	4 (7.7%)	-
Neutropenia / neutrophil count decrease	-	-	1 (7.7%)	-	2 (3.8%)	1 (1.9%)
Nausea	-	-	-	-	2 (3.8%)	-
Alkaline phosphatase incr	-	-	-	-	3 (5.8%)	-
Arthralgia	-	-	-	-	3 (5.8%)	-
WBC decreased	-	-	-	-	3 (5.8%)	-
Myalgia	-	-	-	-	2 (3.8%)	-
Diarrhea	3 (16.7%)	-	-	-	-	-
Urticaria	3 (16.7%)	-	-	-	-	-
Lymphocyte count decreased	1 (5.6%)	1 (5.6%)	-	-	-	-
Headache	1 (5.6%)	-	-	-	-	-
Stomatitis	1 (5.6%)	-	-	-	-	-
Back pain	1 (5.6%)	-	-	-	-	-
Vision blurred	1 (5.6%)	-	-	-	-	-
Abdominal pain / abdominal pain upper	1 (5.6%)	-	-	-	-	-
Hypersensitivity	-	-	1 (7.7%)	1 (7.7%)	-	-
Pneumonitis	-	-	1 (7.7%)	-	-	-
Constipation	-	-	-	-	-	-
Vomiting	-	-	-	-	-	-

#### The lack of preclinical toxicity due to the inactive Fc in vivo has translated to a well-tolerated profile in clinic

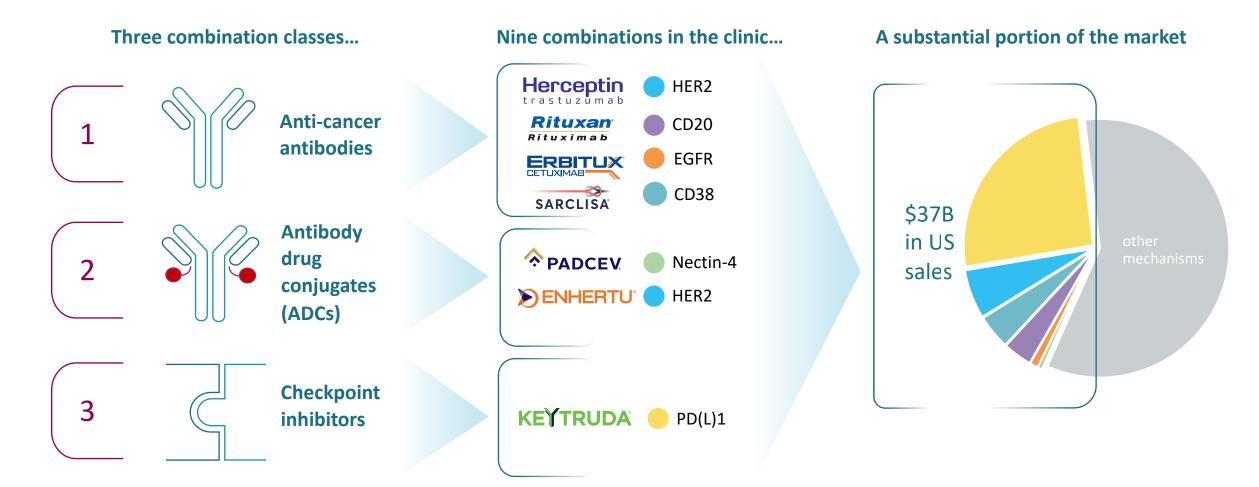
Phase 1 ASPEN-01 cohorts. For combination cohort of evorpacept plus Keytruda, treatment related adverse events occurring in >1 subject in all histologies at 10 & 15 mg/kg QW; data as of April 1, 2020. For combination cohorts of evorpacept plus Keytruda and chemotherapy (5FU, platinum) or plus Herceptin and chemotherapy (Cyramza, paclitaxel), all treatment related adverse events are reported; data as of September 01, 2021.



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



## A bold vision for evorpacept: Deliver a first-in-class, universal combination agent



#### Three distinct modalities currently being tested in the clinic... targeting nearly half of the US oncology market

US sales by drug class based on Clarivate | DRG Disease Landscape & Forecast US sales estimates for 2022 for cumulative total sales across compound classes. Total 2022 US oncology spending from 2023 IQVIA Global Oncology Trends.



# Pursuing a robust development plan

lr	dication	Evorpacept Combination Agent	Discovery	IND Enabling	Phase 1	Phase 2	Phase 3	Fast Track	Supply/ Collaboration Partner
SUC	<b>GC</b> Gastric/Gastroesophageal Junction Cancer	Herceptin + Cyramza + Paclitaxel (ASPEN-06)	Announced po	ositive interim da	ta in Q4 2023				Lilly
CINO	Urothelial Cancer	Padcev (ASPEN-07)							
tion Studies	Breast Cancer	Zanidatamab							Jazz Pharmaceuticals.
Combination		Enhertu (I-SPY)							QL Leap Healthcare Collaborative
Evorpacept Co	<b>MM</b> Multiple Myeloma	Sarclisa + Dexamethasone							sanofi
	HNSCC Head And Neck	Keytruda (ASPEN-03)							
	HNSCC Head And Neck Squamous Cell Carcinoma	Keytruda + 5FU + Platinum (ASPEN-04)							

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# **Evorpacept + anti-cancer antibodies**

## HER2+ Gastric/ GEJ Cancer

ASPEN-06 Phase 2 Study:

Evorpacept

+ Herceptin + Cyramza + paclitaxel

# Non-Hodgkin Lymphoma (NHL)

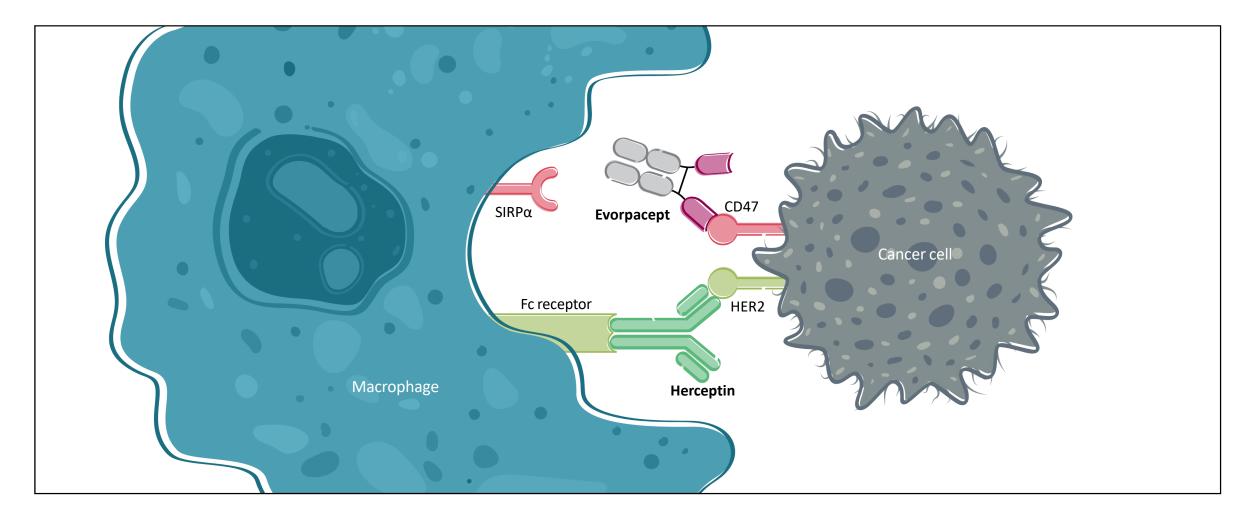
ASPEN-01 Phase 1b NHL Study:

Evorpacept

+ Rituxan

Don't eat me

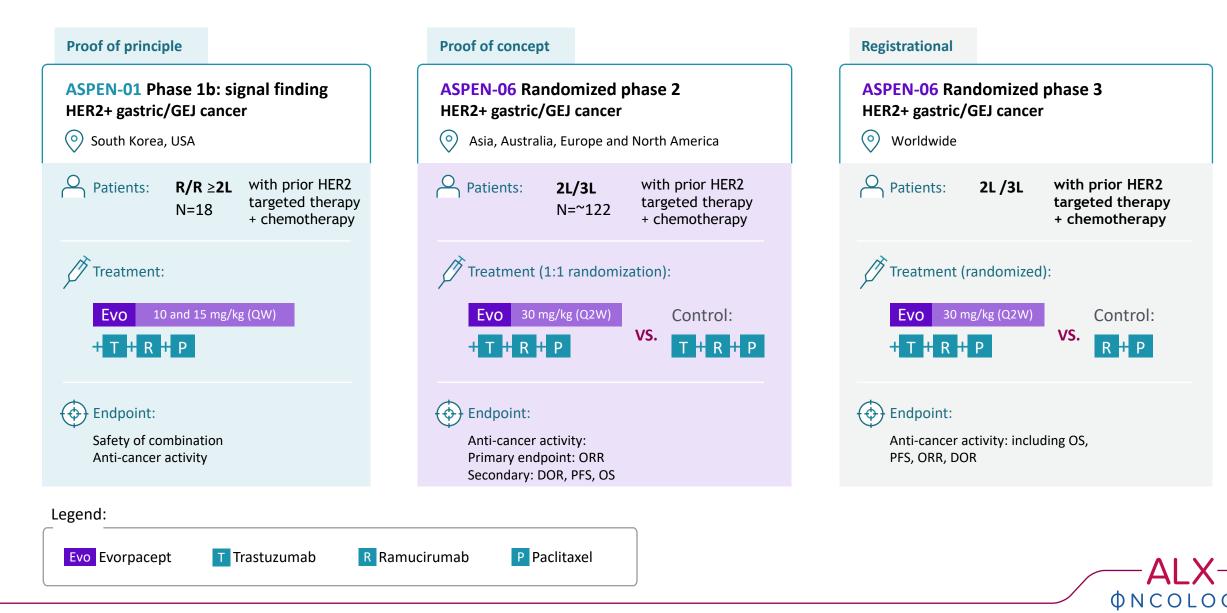
## **Evorpacept + Herceptin mechanism of action**



**Evorpacept increases antibody dependent cellular phagocytosis in combination with Herceptin** 

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



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

AINBOW <sup>1</sup> 2L	ORR (%)	DOR	PFS	OS	
Ramucirumab/Paclitaxel	<sup>0%</sup>	100% 4.4 months IQR 2.8–7.5	<b>4.4</b> months 4.2-5.3	<b>9.6</b> months 8.5-10.8	Volume 15, ISSUE 11, P1224-1235 October 2014 axel versus placebo plus paclitaxel
Paclitaxel	16%	<b>2.8</b> months IQR 1.4-4.4	<b>2.9</b> months 2.8-3.0	<b>7.4</b> months 6.3-8.4	ly treated advanced gastric or tion adenocarcinoma (RAINBOW): ted phase 3 trial

#### DESTINY-Gastric01<sup>2</sup>3L

Trastuzumab deruxtecan	41%	<b>11.3</b> months 5.6-NE	<b>5.6</b> months 4.3-6.9	<b>12.5</b> months 9.6-14.3	The NEW ENGLAND JOURNAL of MEDICINE Volume 382: P2419-24 June 20 Trastuzumab deruxtecan in previously treated HER2-positive		
Physicians' choice	11%	<b>3.9</b> months 3.0-4.9	<b>3.5</b> months 2.0-4.3	<b>8.4</b> months 6.9-10.7	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

<sup>1</sup>Wilke et al, Lancet October 2014, <sup>2</sup>Enhertu US product insert, and Shitara et al, NEJM June 18, 2020; NE could not be estimated



ASPEN-06: Evorpacept 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

#### 2<sup>nd</sup> line or 3<sup>rd</sup> line

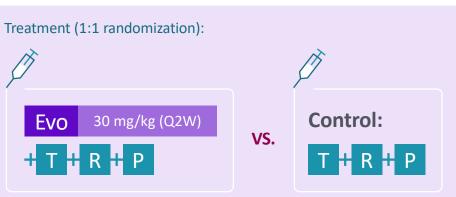
#### **X** No prior treatment:

Anti-CD47 agent, an anti-SIRP $\alpha$  agent or ramucirumab.

#### ✓ Prior treatment ok:

Trastuzumab deruxtecan (Enhertu) and checkpoint inhibitors

ASP	EN-06 randomized phase 2	•
Q	N=122	



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):

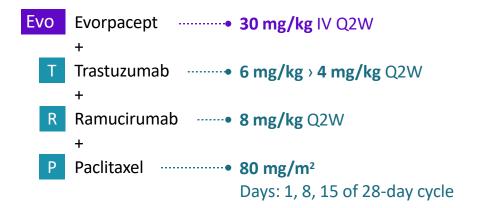
80% power to see a 50% improvement in ORR compared to historical RP and 68% power to see 10% delta between both arms.



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



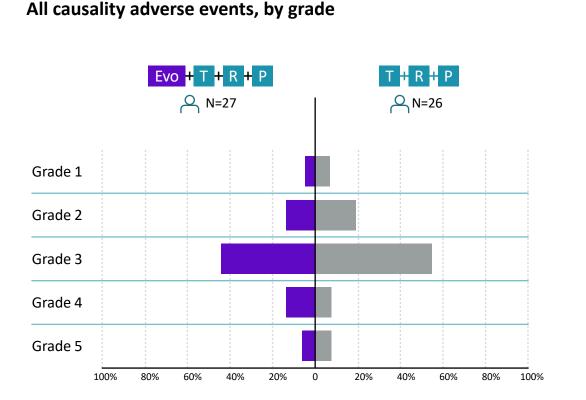
#### Study regimen dose administration:

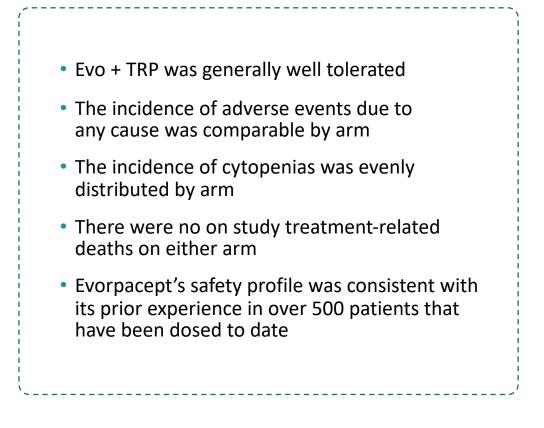


Study pop	ulation: -				
		Evo	Control:		
		+ T + R + P	T + R + P		
		⊖ N=27	○ N=27		
Median age, years (range)		65 (41-79)	57 (31-81)		
Sex,	Male	85	70		
n%	Female	15	30		
	Asian	52	48		
Race,	White	26	30		
n%	Other	3.7	0		
	Unknown	18.5	22		
ECOG PS,	0	52	52		
n%	1	48	48		
GEJ, n%		15	22		

Data Cutoff as of 29 August 2023

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







Data Cutoff as of 29 August 2023

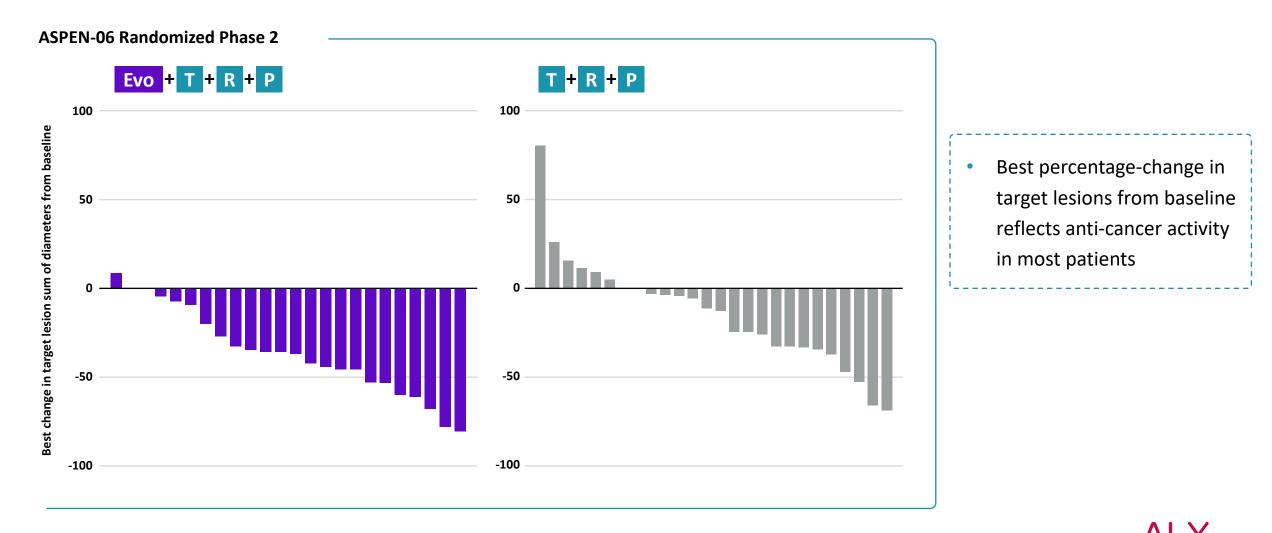
# ASPEN-06 interim analysis: Clinical activity of Evo + TRP supports substantial contribution of evorpacept to TRP and compares favorably to current SOCs

	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	<b>NR</b> [3.6, NR]	<b>7.4</b> [3.5, NR]

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

Data Cutoff as of 29 August 2023 <sup>1</sup> Wilke et al, Lancet October 2014, <sup>2</sup> Enhertu US product insert, and Shitara et al, NEJM June 18, 2020; NR not reached

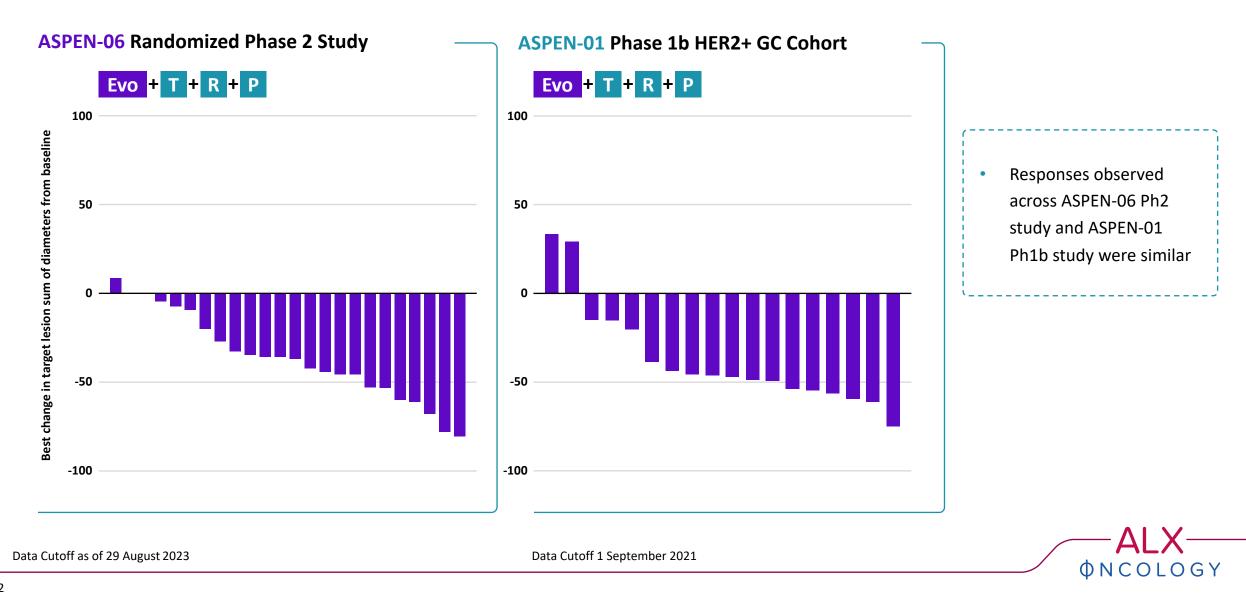
ASPEN-06 interim analysis: Substantial tumor shrinkage is seen in ASPEN-06 gastric/GEJ cancer patients receiving Evo + TRP compared to TRP



 $\Phi$ NCOLO

Data Cutoff as of 29 August 2023

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



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**

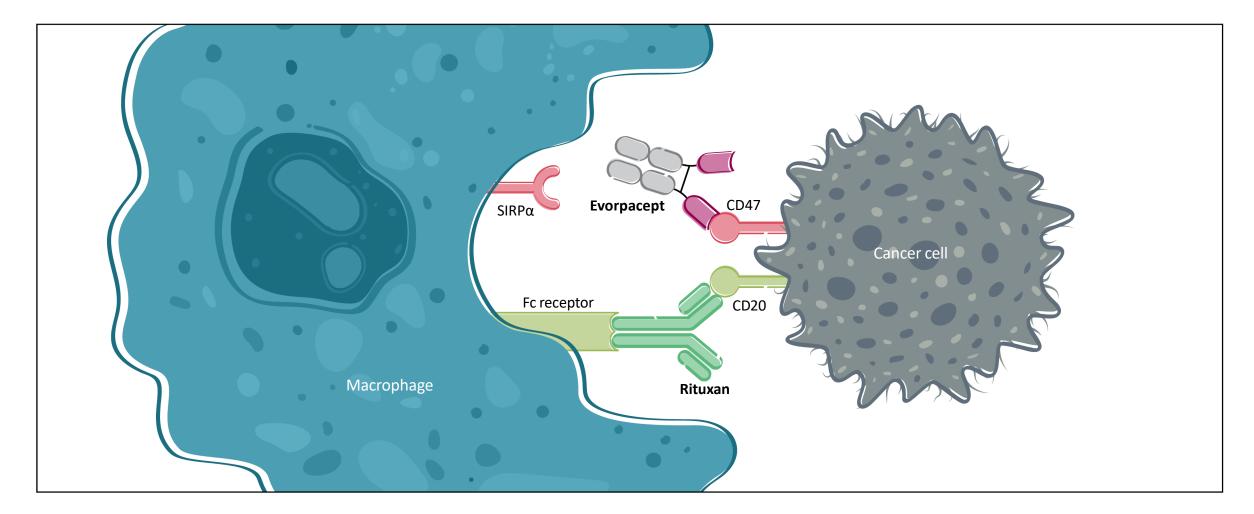
ASPEN-06 interim data confirms that **evorpacept can be combined with TRP** with a favorable safety profile was that was consistent with data from the >500 patients treated to date

#### **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 + Rituxan mechanism of action**



**Evorpacept increases antibody dependent cellular phagocytosis in combination with Rituxan** 



# Promising activity observed for evorpacept plus an anti-cancer antibody in a hematologic malignancy Phase 1b clinical trial of evorpacept + Rituximab in patients with aggressive / indolent NHL

ohorts		-	pt (10 mg/kg Rituximab	Evorpacept (15 mg/kg QW) + Rituximab		
	Population	N	ORR	N	ORR	
relapsed/refractory NHL, prior regimen with Rituximab	All	22	40.9%	10	70.0%	
Treatment:	Aggressive	15	33.3%	6	50.0%	
evorpacept 10 or 15 mg/kg once a week (QW) + Rituximab 375 mg/m <sup>2</sup> once a week for 4 weeks, once monthly for 8 months	Indolent	7	57.1%	4	100.0%	

Indolent = Follicular Lymphoma and Marginal Zone Lymphoma.

ORR = Objective Response Rate. MTD = maximum tolerated dose.

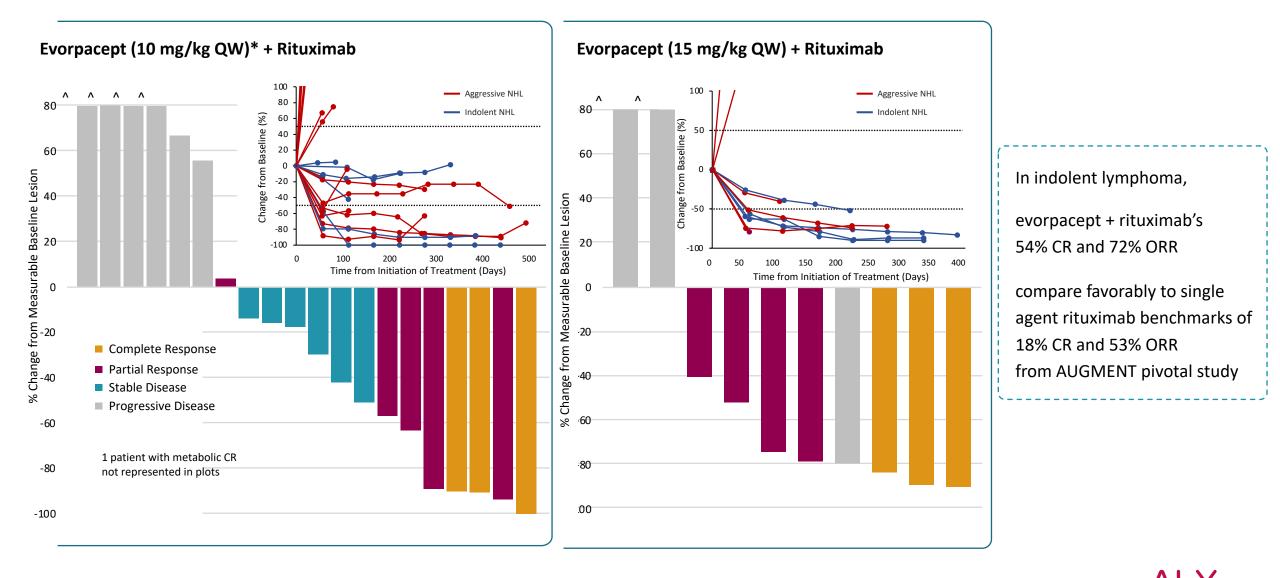
Aggressive = Diffuse Large B-cell Lymphoma and Mantle Cell Lymphoma.

#### All patients enrolled (22/22) had received prior Rituximab therapy

- Evorpacept demonstrated higher response rates at higher dosing
- No dose-limiting toxicities were reported in either the 10 or 15 mg/kg group, and the MTD was not reached



# Phase 1b clinical trial of evorpacept + Rituximab in aggressive / indolent NHL

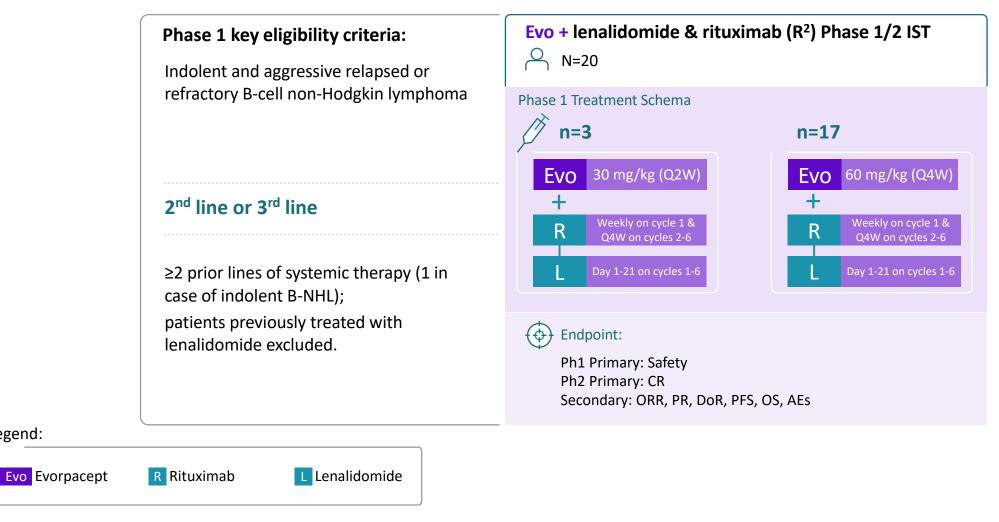


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Data Cutoff October 1, 2020; Response evaluable patients; Responses include metabolic response per Lugano Response Criteria. ^ more than 80% increase from baseline. \* 1 patient with rapid fatal progressive disease not represented in plot

# Phase 1/2 of evorpacept + $R^2$ in indolent and aggressive relapsed or refractory **B-cell non-Hodgkin lymphoma**





N = Response Evaluable Patients; Indolent = Follicular Lymphoma and Marginal Zone Lymphoma; Aggressive = Diffuse Large B-cell Lymphoma and Mantle Cell Lymphoma; CR = Complete response; PR = Partial response; ORR = Objective response rate; DoR= Duration of response; PFS = Progression free survival; AEs = Adverse events; IST = Investigator Sponsored Trial

P. Strati. AACR 2024, Oral Presentation. Abstr #CT037

Investigator Sponsored Trial

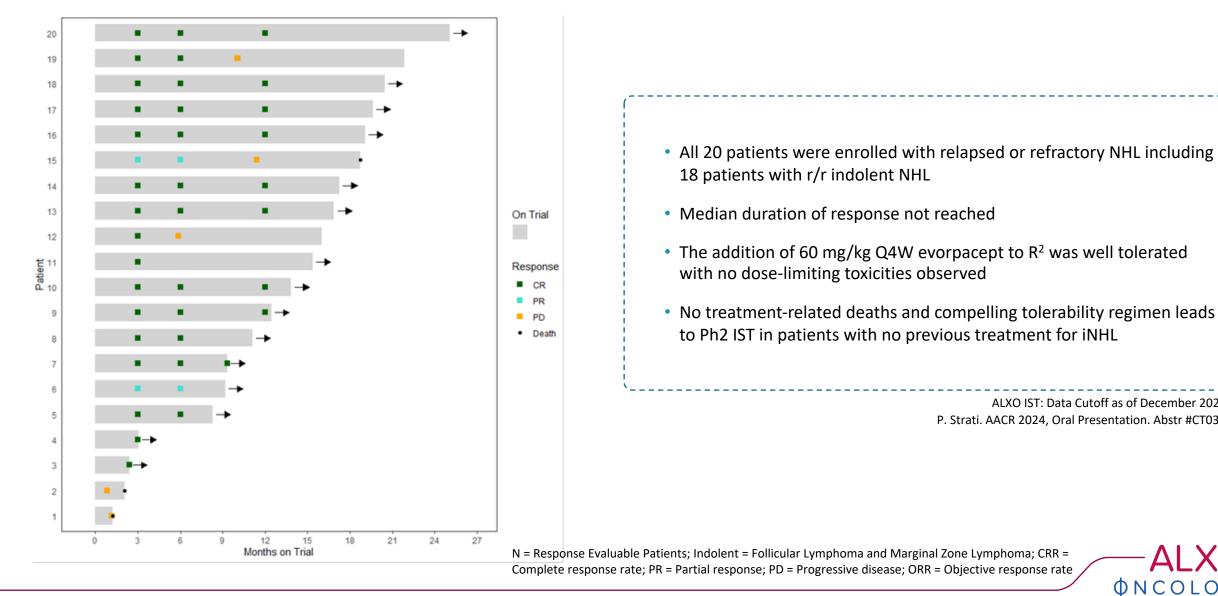


Legend:

# Encouraging initial activity of evorpacept + R<sup>2</sup> in iNHL with a favorable safety profile



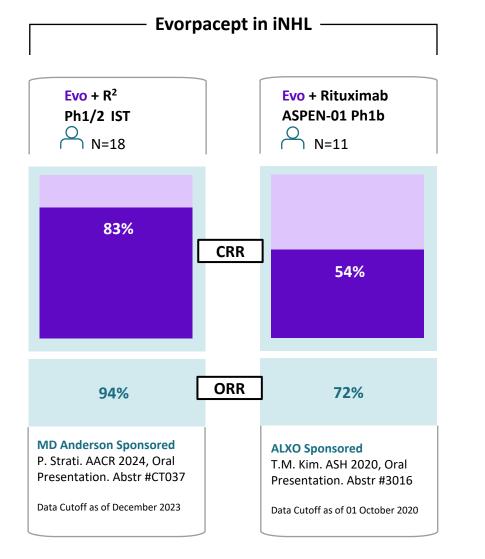
A best ORR of 94% and a CRR of 83% in patients with indolent R/R B-NHL

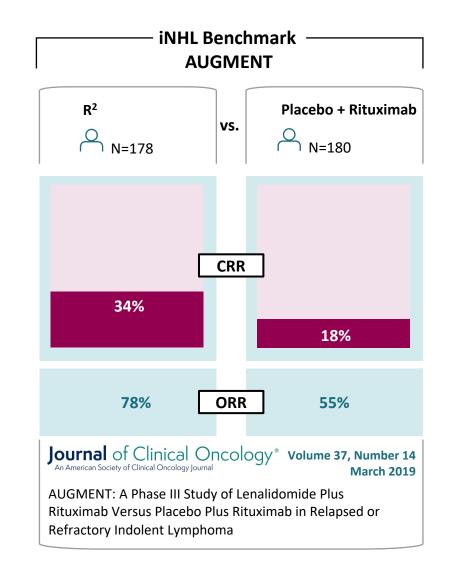


ALXO IST: Data Cutoff as of December 2023

P. Strati. AACR 2024, Oral Presentation. Abstr #CT037

### **Evorpacept-based regimens show consistent activity in indolent NHL trials**





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R<sup>2</sup> = Lenalidomide + Rituximab; N = Response Evaluable Patients; Indolent = Follicular Lymphoma and Marginal Zone Lymphoma; CRR = Complete response rate; ORR = Objective response rate; IST = Investigator Sponsored Trial

# Two ongoing studies with anticancer antibodies in hematologic malignancies

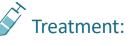


Phase 1/2 Non-Hodgkin Lymphoma IST



N=20

Relapsed or refractory B-cell NHL, 1 or more prior systemic therapies

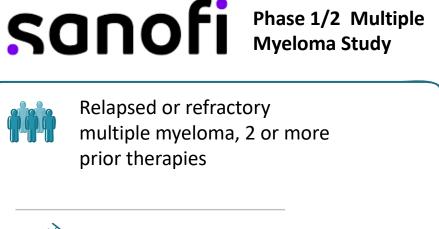


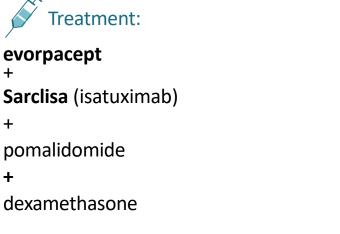
```
evorpacept 30 mg/kg every two
weeks (Q2W) or 60 mg/kg every 4
weeks (Q4W)
+
Rituxan (rituximab) weekly on cycle 1 and
Q4W on cycles 2-6
+
```

Revlimid (lenalidomide) D1-21 on cycles 1-6

# Oral presentation at AACR 2024, now dosing 1L R2-naive NHL patients

IST: Investigator-sponsored trial. Multiple myeloma trial sponsored by Sanofi with ALX collaboration





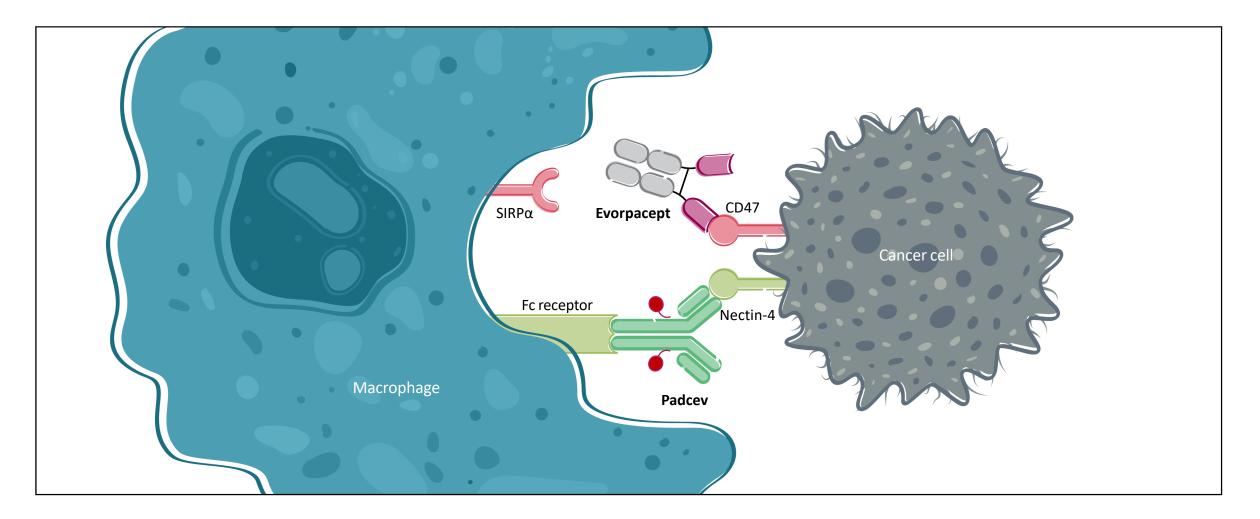


# Evorpacept + antibody-drug conjugates (ADCs)

Urothelial (Bladder) Cancer
 ASPEN-07 Phase 1b Study:
 Evorpacept + Padcev

Don't eat me

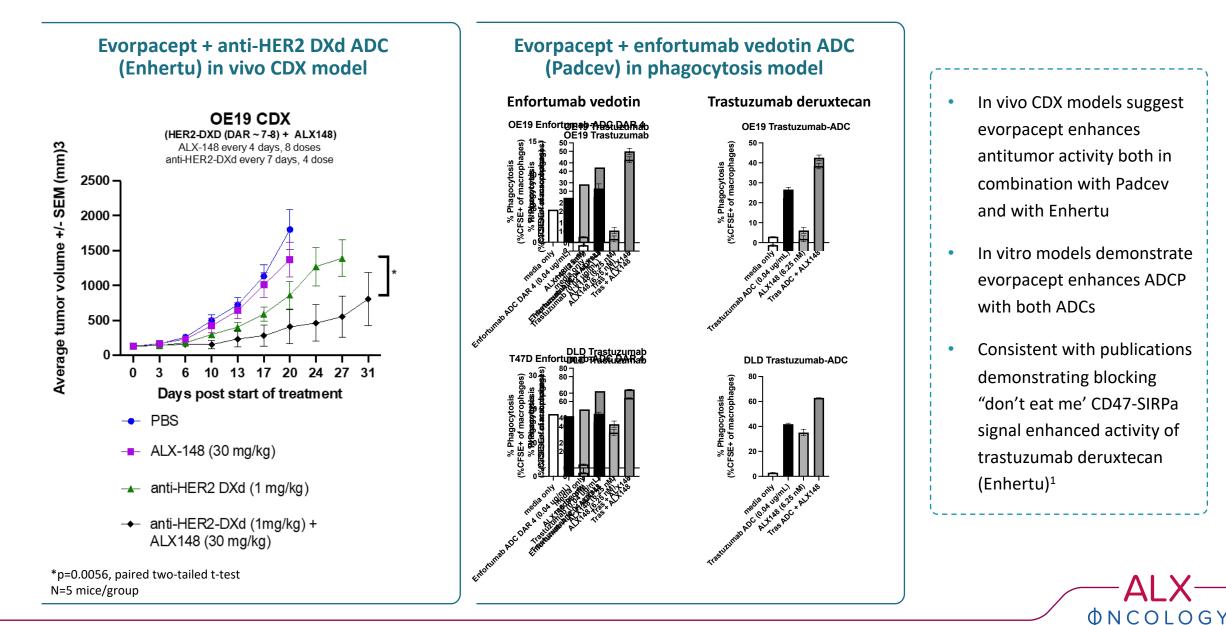
## **Evorpacept + ADCs mechanism of action**



**Evorpacept increases antibody dependent cellular phagocytosis (ADCP) in combination with Padcev** 

**ØNCOLOGY** 

### Preclinical data supports CD47 blockade enhances ADC efficacy through increased phagocytosis



# Advancing clinical studies in breast and urothelial cancer to assess evorpacept's synergistic potential with ADCs

#### ASPEN-07 - Phase 1b Urothelial Study Design



N=20

locally advanced or metastatic urothelial carcinoma, prior platinum-based chemotherapy and PD-1/L1 inhibitor



evorpacept 20 or 30 mg/kg every two weeks (Q2W) + Padcev (enfortumab vedotin) 1.25 mg/kg IV

on Days 1, 8, and 15 of each 28-day cycle

## Data update Q2-2024



Phase 1b Breast Cancer Study Design



Unresectable or metastatic HER2-positive or HER2-low breast cancer

# Treatment:

evorpacept 20 or 30 mg/kg every
two weeks (Q2W)
+
Enhertu (trastuzumab deruxtecan) 5.4
mg/kg every three weeks (Q3W)

Top line data Q4-2024



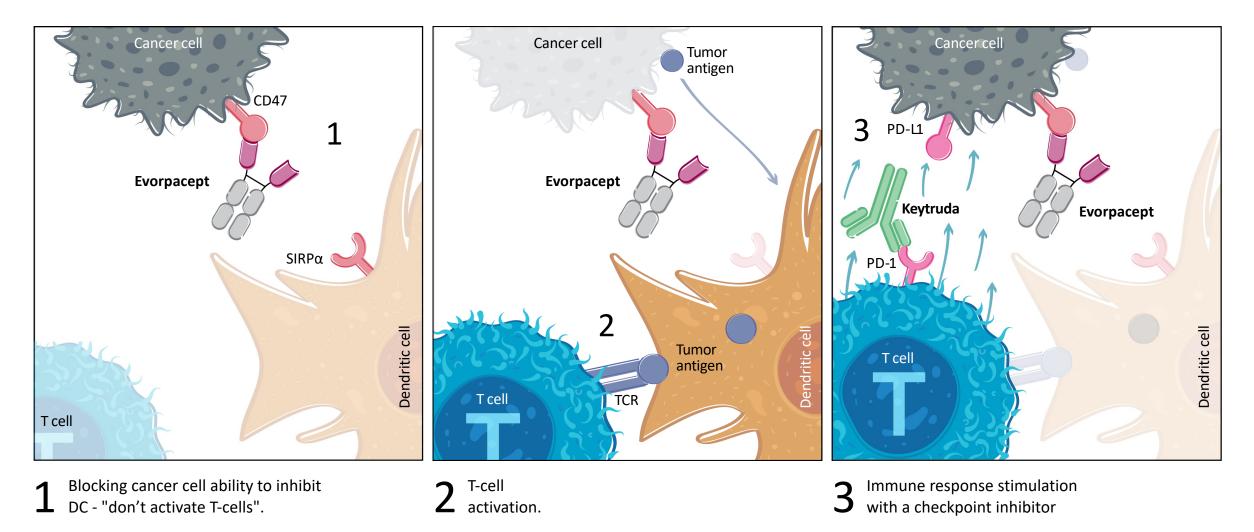
# **Evorpacept + checkpoint inhibitors**

**1L Head & Neck Squamous Cell Carcinoma (HNSCC)** ASPEN-03 Phase 2 Study: Evorpacept + Keytruda

### 1L Head & Neck Squamous Cell Carcinoma (HNSCC)

ASPEN-04 Phase 2 Study: Evorpacept + Keytruda + chemotherapy

## HNSCC trial: evorpacept + Keytruda mechanism of action



**Evorpacept activates dendritic cells and enhances cross-priming of T cells** 

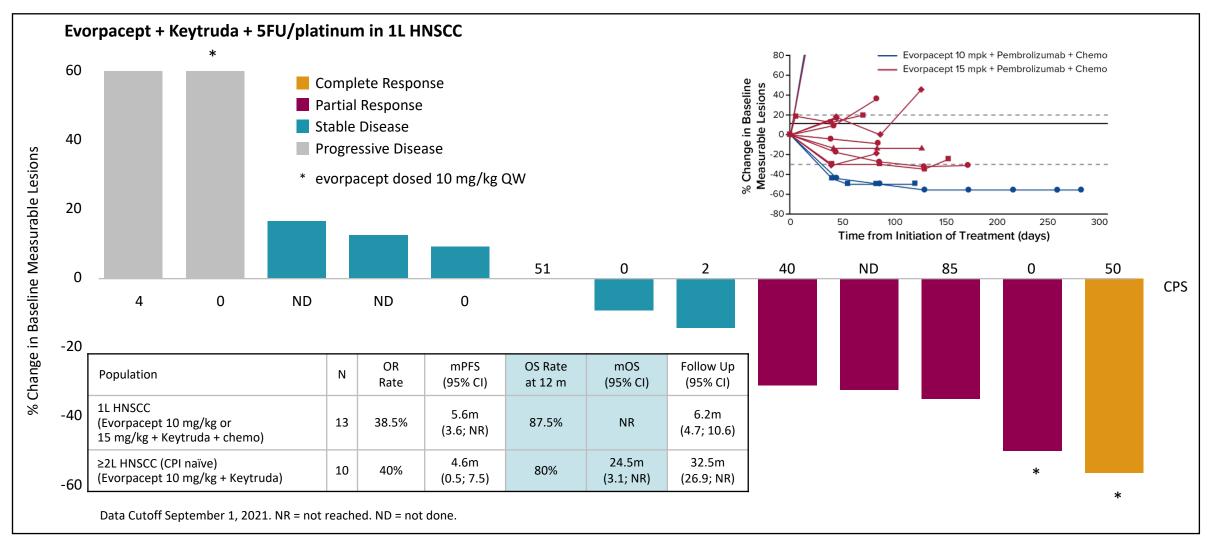
**ØNCOLOGY** 

# Current standard-of-care in 1L HNSCC is Keytruda +/- chemo and the KEYNOTE-048 studies highlight the benchmark and significant unmet need

	Population	N	ORR (%)	PFS (m) [95% Cl]	OS Rate at 12 m	OS (m) [95% Cl]	Follow Up (m) [95% Cl]	
	KEYNOTE-048: 1L HNSCC pembrolizumab + 5FU/platinum	281	36%	4.9 [4.7–6.0]	53%	13.0 [10.9–14.7]	13 [6.4–26.6]	<ul> <li>KEYNOTE-048 supported Keytruda's 1L HNSCC</li> </ul>
1L _	<b>KEYNOTE-048:</b> <b>1L HNSCC</b> cetuximab + 5FU/platinum	278	36%	5.1 [4.9–6.0]	44%	10.7 [9.3–11.7]	10.7 [6.6–19.7]	approvals and provide the benchmarks for ASPEN- 03 and ASPEN-04
	KEYNOTE-048: 1L HNSCC, CPS ≥1 pembrolizumab	257	19%	3.2 [2.2–3.4]	50%	12.3 [10.8–14.3]	11.5 [5.1–25.7]	<ul> <li>Of note, OS benefit at 12 months correlated with OS benefit.</li> </ul>
	KEYNOTE-048: 1L HNSCC, CPS ≥1 cetuximab + 5FU/platinum	255	35%	5.0 [4.8–5.8]	44%	10.3 [9.0–11.5]	10.7 [6.6–19.7]	

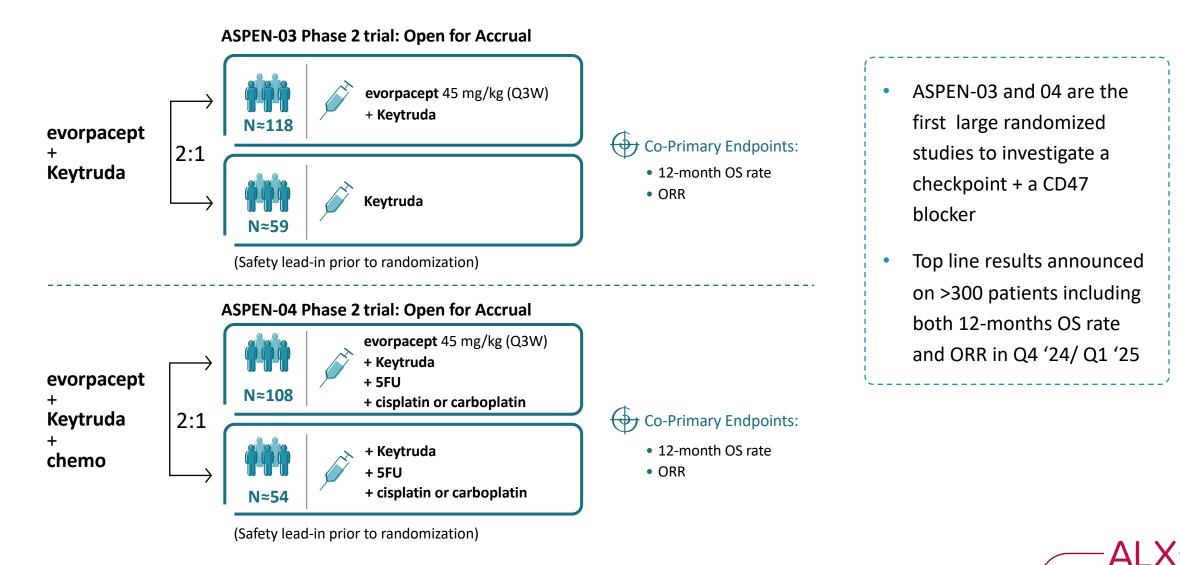
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## ASPEN-01 Phase 1b HNSCC: Evorpacept + Keytruda + 5FU/platinum first line checkpoint naive



Data as of 1 February 2022. NC = not calculable, (95% CI) 1L HNSCC: mOS not reached (CI: 5.99-NC) with median follow up of 15.8 months (CI: 5.0-17.8) ≥2L HNSCC (CPI-Naïve): mOS of 24.6 months (CI: 3.13-NC) with median follow-up of 35.3 months (CI: 27.0-41.0)

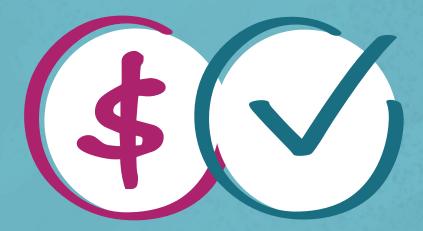
# First line head and neck cancer: Phase 2 development plan, ASPEN-03 and ASPEN-04



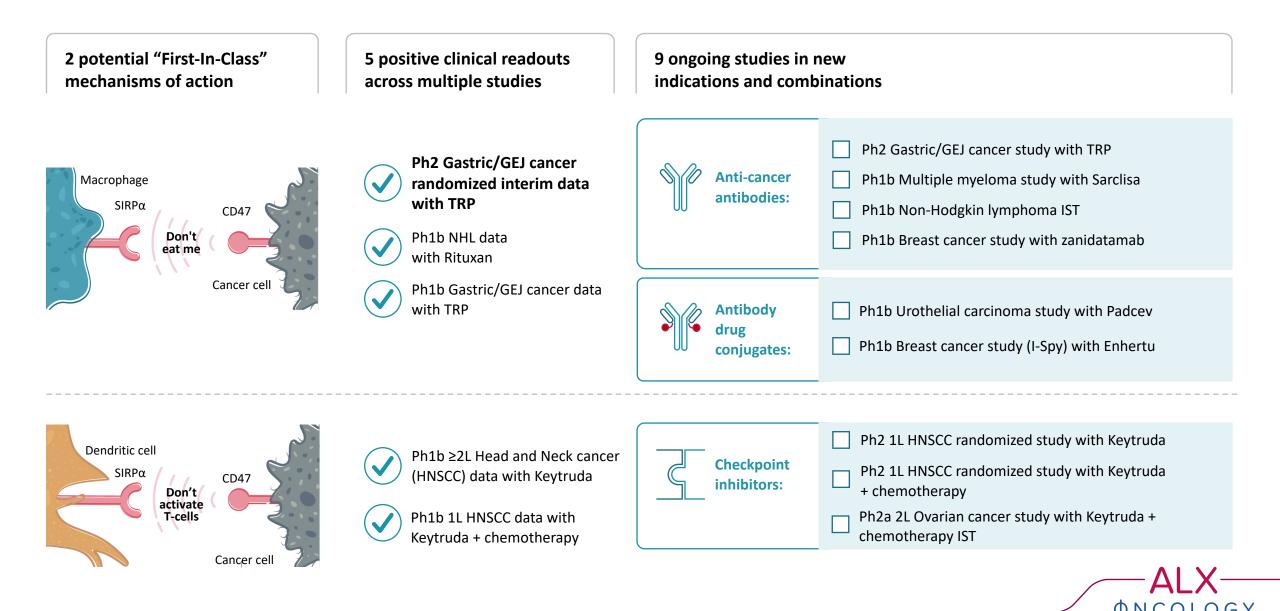
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Dosing schedules: Keytruda and chemotherapy Q3W

# Upcoming Milestones and Financials

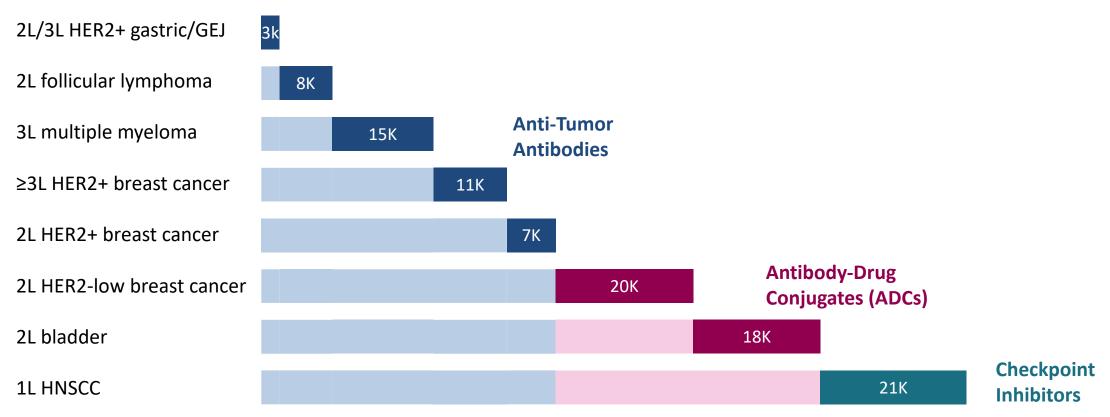


### Validated approach and our path to success



# ALX clinical trials position evorpacept to become a market leader in metastatic disease across combining with three key modalities

#### US addressable patient populations from evorpacept clinical trials



**ØNCOLOGY** 

#### Current clinical trials with evorpacept address >100,000 cancer patients in the US

Addressable patient population sources: Decision Resources Guide; Market Research; industry IR materials.

# Anticipated upcoming milestones: Significant catalysts in 2024

2024 Evorpacept Milestones
Non-Hodgkin Lymphoma (NHL) Phase 1B study Data from Phase 1B IST study – AACR '24
<b>Urothelial Carcinoma (Phase 1A)</b> ASPEN-07 data update in urothelial carcinoma with Padcev - Q2 '24
Gastric/GEJ Cancer (Phase 2) ASPEN-06 Top line results from gastric/ GEJ from randomized trial with TRP – June/ July '24
<b>Breast Cancer (Phase 1B)</b> I-SPY Top line results in breast cancer with Enhertu– Q4 '24
<b>Head &amp; Neck Cancer (Phase 2)</b> ASPEN-03 Top line results in HNSCC from randomized trial with Keytruda – Q4 '24 / Q1 '25
<b>Head &amp; Neck Cancer (Phase 2)</b> ASPEN-04 Top line results in HNSCC from randomized trial with Keytruda and chemotherapy – Q4 '24/ Q1 '25
Gastric/GEJ Cancer (Phase 3)

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ASPEN-06 Initiation of registrational randomized gastric/GEJ cancer trial – Q4 '24

### **Financial information**

### **Approximately \$605M in net proceeds raised to date including:**

- \$170M IPO in July 2020
- \$195M follow on in December 2020
- \$59M follow on in October 2023

\$90M of \$100M loan facility potentially available with \$10M drawn to date

Cash, cash equivalents and investments balance as of December 31, 2023 of approximately \$218M

**Expected cash runway into early 2026** 

