

**Advancing Antibody Drug Conjugates With Novel Immuno-Oncology Payloads** 

October 2025 **Corporate Overview** 

> NASDAQ: AKTX akaritx.com

### **Forward-Looking Statements**

This presentation includes expressed or implied forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended (the Exchange Act), about the Akari Therapeutics, Plc (the "Company") that involve risks and uncertainties relating to future events and the future performance of the Company. Actual events or results may differ materially from these forward-looking statements. Words such as "will," "could," "would," "should," "expect," "plan," "anticipate," "intend," "believe," "estimate," "predict," "project," "potential," "continue," "future," "opportunity" "will likely result," "target," variations of such words, and similar expressions or negatives of these words are intended to identify such forward-looking statements, although not all forward-looking statements contain these identifying words. 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Factors that may affect future results and may cause these forward-looking statements to be inaccurate include, without limitation: the risk that the Company may not realize the anticipated benefits of its merger with Peak Bio, Inc. (the "Merger") in the time frame expected, or at all; the ability to retain and hire key personnel; potential adverse reactions or changes to business relationships resulting from the Merger; the potential impact of unforeseen liabilities, future capital expenditures, revenues, costs, expenses, earnings, synergies, economic performance, indebtedness, financial condition and losses on the future prospects, business and management strategies for the management, expansion and growth of the combined business; uncertainties as to the long-term value of the Company's American Depositary Shares ("ADSs") (and the ordinary shares represented thereby), including the dilution caused by the Company's issuance of additional ADSs (and the ordinary shares represented thereby) in connection with the Merger; risks related to global as well as local political and economic conditions, including interest rate and currency exchange rate fluctuations; potential delays or failures related to research and/or development of the Company's programs or product candidates; risks related to any loss of the Company's patents or other intellectual property rights; any interruptions of the supply chain for raw materials or manufacturing for the Company's product candidates, the nature, timing, cost and possible success and therapeutic applications of product candidates being developed by the Company and/or its collaborators or licensees; the extent to which the results from the research and development programs conducted by the Company, and/or its collaborators or licensees may be replicated in other studies and/or lead to advancement of product candidates to clinical trials, therapeutic applications, or regulatory approval; uncertainty of the utilization, market acceptance, and commercial success of the Company's product candidates; unexpected breaches or terminations with respect to the Company's material contracts or arrangements; risks related to competition for the Company's product candidates; the Company's ability to successfully develop or commercialize its product candidates; potential exposure to legal proceedings and investigations; risks related to changes in governmental laws and related interpretation thereof, including on reimbursement, intellectual property protection and regulatory controls on testing, approval, manufacturing, development or commercialization of any of the Company's product candidates; the Company's ability to maintain listing of its ADSs on the Nasdaq Capital Market. While the foregoing list of factors presented here is considered representative, no list should be considered to be a complete statement of all potential risks and uncertainties. More detailed information about the Company and the risk factors that may affect the realization of forward-looking statements is set forth in the Company's filings with the SEC, copies of which may be obtained from the SEC's website at www.sec.gov. The Company assumes no, and hereby disclaims any, obligation to update the forward-looking statements contained in this press release.





### **Corporate Overview**

# Advancing Next-Generation Antibody Drug Conjugates Designed with Novel ImmunoOncology Payloads

- Proprietary PH1 spliceosome modulator payload designed to kill tumors and activate immune system
- Potential to outperform current ADC payloads with superior tumor-killing activity and favorable safety profile

#### Lead Program AKTX-101 (TROP2 PH1 ADC)

- IND/First-in-human regulatory filing in H2 2026
- Initial Focus: bladder, gastric, and lung cancer
- Strong preclinical data showing potent tumor killing
- Payload designed to evade tumor resistance mechanisms
- Payload activates multiple immune effectors

#### Second Lead Program AKTX-102 (Undisclosed Target)

- Advancing preclinical development towards first-in-class product candidate
- Initial focus: colon, gastric, and lung cancers



### Akari Is Advancing PH1 Payload In Multiple ADC Programs

PH1 payload is the cornerstone of building an ADC pipeline against wide range of tumors

ADC Programs (All PH1 Payload based)	Indication	Discovery	Preclinical	Clinical	Highlights
AKTX-101 (Trop2 Target)	Bladder, Gastric, and Lung cancers				<ul> <li>Ready for IND enabling studies supporting IND/First-in-human regulatory filing in H2 2026</li> </ul>
AKTX-102 (Undisclosed Target)	Colon, Gastric, and Lung cancers				Disclosure of target/key preclinical data Q4 2025



# ADCs Continue to Drive Cancer Therapeutic Class With Several Blockbuster Products > \$1B sales

Global ADC Market Projected to Surpass >\$34B by 2032<sup>1</sup>

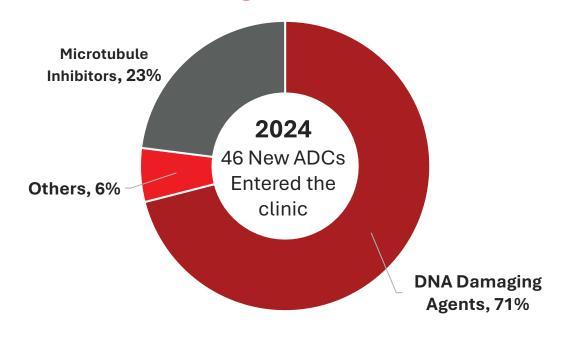
However, Current ADC Landscape Continues to Focus on Limited Payloads With Known Limitations

#### 2024 Sales from Approved ADCs

_	Product	Payload	2024 Sales		
	TRODELVY* sacituzumab govitecan-hziy 180 mg for injection	DNA Damaging Agent	\$1.3B		
	ENHERTU° trastuzumab deruxtecan	DNA Damaging Agent	\$3.8B		
	PADCEV. enfortumab vedotin-ejfv løjection for IV infusion 20 mg 8 30 mg vists	Microtubule Inhibitor	\$1.3B		
=	Ado-trastuzumab emtansine	Microtubule Inhibitor	\$2.3B		
	POLIVY* polatuzumali vedotin-piiq sucrous ros stratenous use zone; heads	Microtubule Inhibitor	\$1.3B		
_	SADCETRIS* brentuximab vedotin   for injection	Microtubule Inhibitor	\$1.9B		

15+ FDA-approved ADCs currently and dozens in late-stage development across solid tumors and hematologic tumors

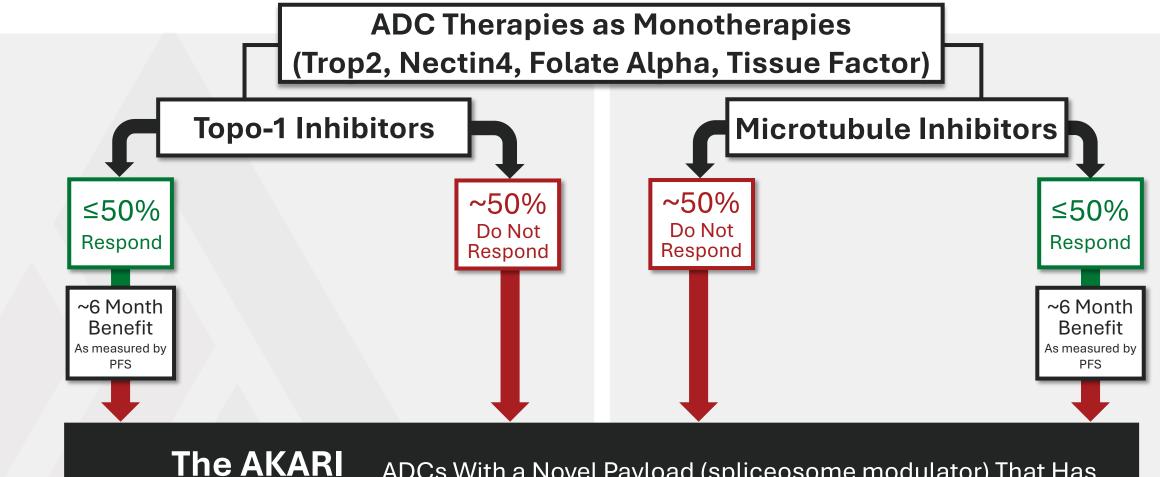
### However, Payload Innovation Is Lacking for New ADCs





**Solid Tumors** 

# Efficacy Outcomes Across A Range of Current ADCs Highlight the Need for New ADCs to Improve Response Rates and Durability





**Opportunity** 

ADCs With a Novel Payload (spliceosome modulator) That Has the Potential to Drive More Durable and Deeper Responses

# Akari Is Advancing A New Class of ADCs: Immuno-Oncology ADCs Centered Around PH1 Payload

Novel PH1 Payload is a Spliceosome Modulator That Has a Unique Efficacy and Safety Profile to Address Unmet Needs of Current ADCs

#### Potential Advantages of ADCs with PH1 Payload

## Kills Cancer Cells While Activating Immune System

Accumulation of mis-spliced proteins generates neoantigens to activate immune system while killing targeted cancer cells

### **Circumvents Traditional Resistance Mechanisms**

Current data shows PH1 is resistant to standard efflux transporters that make cancer cells resistant to ADCs

#### **Synergy With Checkpoint Inhibitors**

Preclinical data shows differential immune responses and preclinical anti-tumor response relative to current ADC + Checkpoint Inhibitor combination

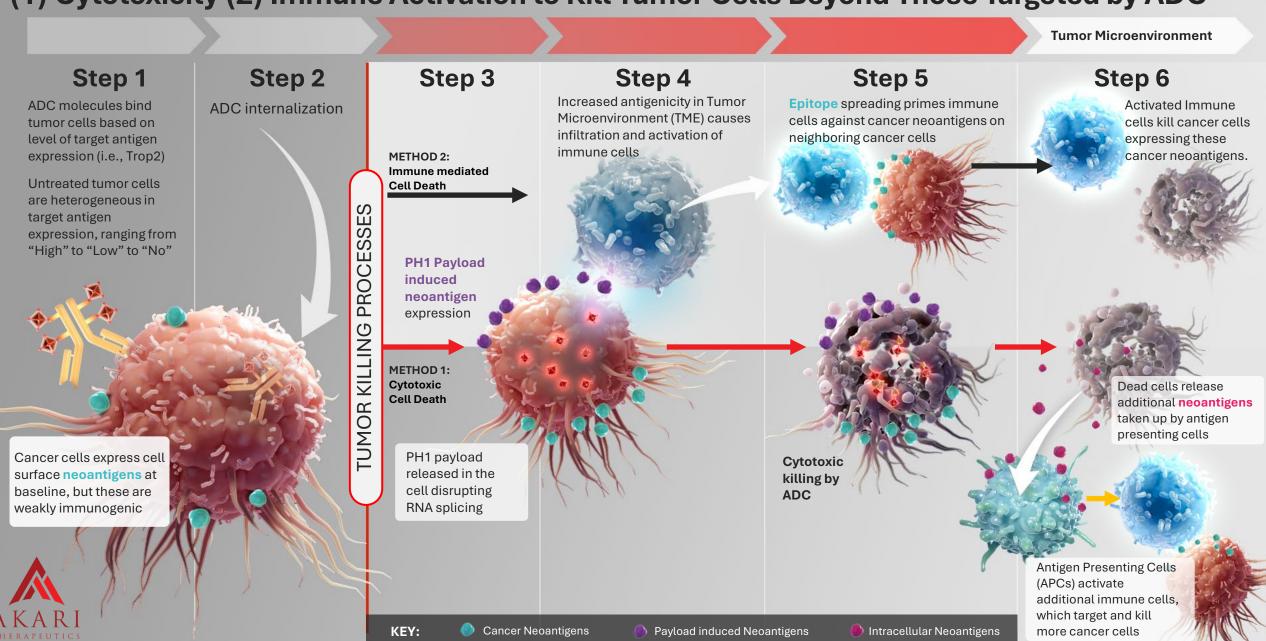
#### **Reduced Off-Target Toxicity**

Linker engineered for only intracellular release within cancer cell to mitigate off-target toxicity



#### PH1 Payload Uses a 1-2 Punch to Kill Cancer Cells:

(1) Cytotoxicity (2) Immune Activation to Kill Tumor Cells Beyond Those Targeted by ADC



Lead Program AKTX-101 (TROP2 PH1 ADC)

Targeting Trop2 Cancers With Spliceosome Modulator Payload After 1<sup>st</sup> line ADCs

IND/FIH Regulatory filing targeting H2 2026

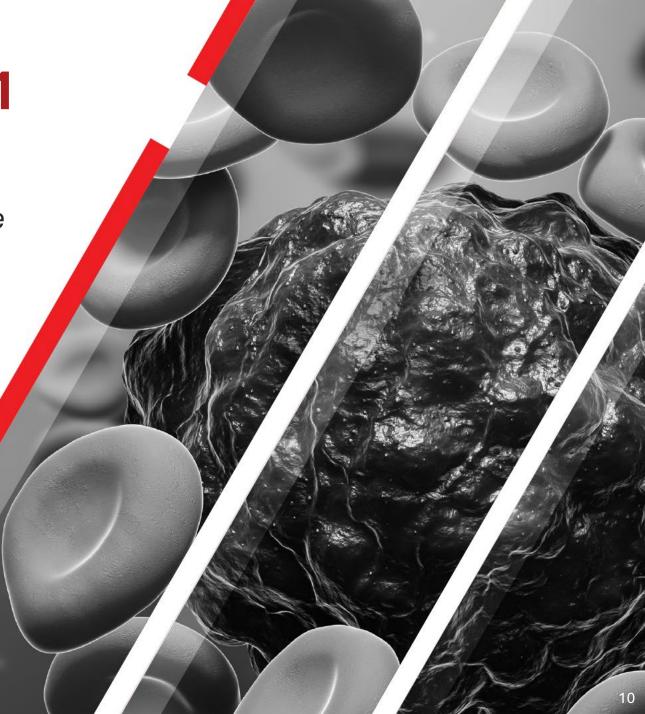
Targeting Significant Unmet Need in Bladder Cancer Post Enfortumab Vedotin (PADCEV®)

Large potential opportunities in Gastric and Lung Cancers



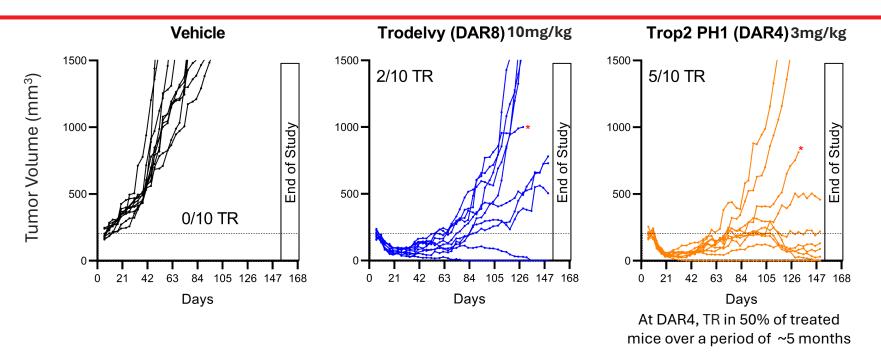
IND: Investigational New Drug

FIH: First In Human



# We Believe AKTX-101 Has Superior Activity Compared to a Marketed ADC, Trodelvy®, Based on Results from Preclinical Models

Higher Rate of Tumor Regression Compared to Trodelvy®, Even With Lower Dose and Drug/Antibody ratio (DAR)



#### Method

NCI-N87 cell xenograft model

Mice dosed IV with ADC or vehicle on day 1 and day 8 post-randomization

At DAR4, Trop2 PH1 induced TR in 50% of treated mice over a period of ~5 months

VS.

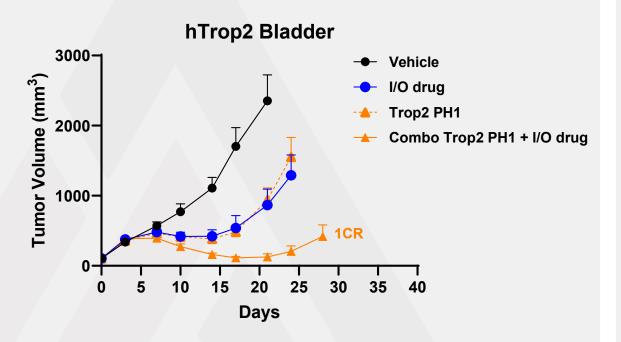
At DAR 7.6, Trodelvy induced TR in only 20% of treated mice over a period of ~5 months

Group	TGI <u>+</u> Std Err (day 20)	p Value vs Trop2 PH1 (DAR4) (day 20)	TGI <u>+</u> Std Err (Day 41)	p Value vs Trop2 PH1 (DAR4) (day 41)
Trop2 PH1 (DAR 4)	87.7 <u>+</u> 1.0		90.3 <u>+</u> 1.8	
Trop2 PH1 (DAR 2)	80.5 <u>+</u> 1.8	0.00014	78.5 <u>+</u> 3.2	0.0014
Trodelvy (DAR 7.6)	79.0 <u>+</u> 2.1	0.000028	83.3 <u>+</u> 2.4	0.035

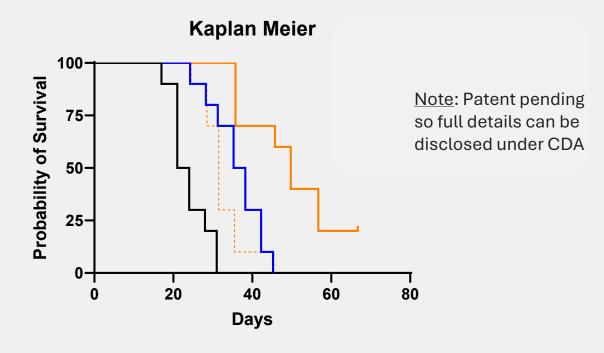


# AKTX-101 Demonstrates Compelling Activity In a Difficult Bladder Cancer Model Both as a Single Agent and Synergistically With Checkpoint Inhibitors





### Superior Overall Survival Combination With IO

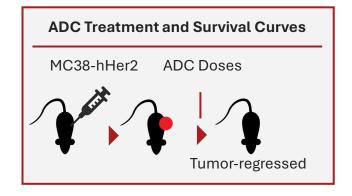




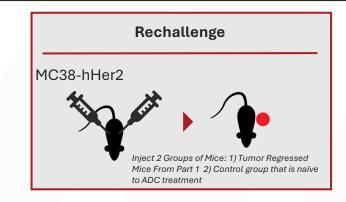
# PH1 Payload Drives Cancer Tumor Regression and Creates Immunological Memory That Prevents Tumor Recurrence

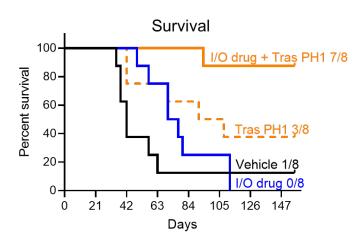
Part 1: Potent Activity of PH1 ADC As Single Agent and With Checkpoint Inhibitor

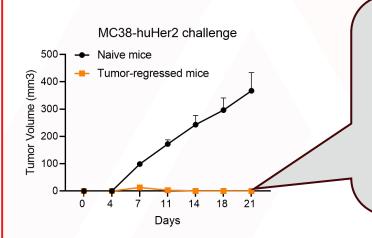
Part 2: PH1 Payload Induces Tumor-Specific Immunological Memory Against Cancer Cells



Rechallenge Regressed Mice Only (7/8 From Combination Arm)







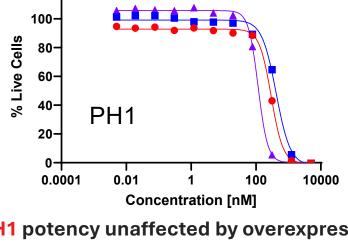
Demonstrates PH1
payload creates
Immunological
Memory That Enables
Immune System To
Prevent Tumor Growth
When Mice is
Rechallenged



# PH1 Payload Designed to Overcome Traditional Resistance Mechanisms Experienced by Currently Approved or In Development ADC Payloads

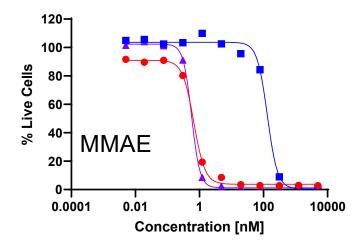
**Method:** Normal MES-SA cells & MES-SA cells selected for overexpression of MDR transporter 1/2 exposed *in vitro* to PH1 or anti-tubulin payload MMAE (monomethyl auristatin E), in presence or absence of MDR 1/2 inhibitor elacridar

- MES-SA uterine sarcoma cell line
- MES-SA cells expressing high levels of MDR transporter 1 and 2 which can pump toxins out of cells
- → MES-SA cells with high MDR expression + MDR inhibitor elacridar



120-

PH1 potency unaffected by overexpression of multidrug resistance (MDR) transporters



200X higher MMAE concentration required to kill cells overexpressing MDRs; inhibition of MDR 1/2 by elacridar restores MMAE potency



# Trop2 PH1 ADC Has Demonstrated Favorable Safety Profile in Non-Human Primate Study

No toxicities such as interstitial lung disease, neutropenia, mucosal inflammation or neuropathies were observed

### Tolerated in NHP at doses well above efficacious dose

- DAR₄ADC tolerated at 6mpk Q3W X 3 repeat doses
- DAR<sub>2</sub>ADC tolerated at 6mpk Q3W X 3 repeat doses

### Mitigatable side-effects that reset to baseline within 2 weeks

- Mild and reversible elevations in liver enzymes
- Mild and reversible reduction in platelets
- Skin rash

### Toxicity profile compatible for combination with Checkpoint Inhibitors

- No evidence of lung complications, pneumonitis
- No Colitis or Hypothyroidism

### Differentiated safety profile with other Trop2 ADCs in clinic

- No Neutropenia, Leukopenia or Diarrhea as observed with Trodelvy®
- No ILD or mucosal inflammation was observed with Dato-DXd



# AKTX-101 Path to IND/FIH Registrational Filing Based on Target Key Milestones\*

2025 2026

Topline Results of New Preclinical Studies (Lung)

Target Completion
of Initial GMP
Manufacturing Batches

Expected Final GLP Toxicology
/Nonclinical Package

Initiate an IND/FIH Regulatory
Filing for Initial Clinical Studies

Expect to Have Future Interactions with FDA Leading Towards Potential IND/First-In-Human Regulatory Filing in H2 2026





### Early Stage ADC's Continue to See Strong Interest and Deal Flow

Licensee	Licensor	Phase	Asset	Target	Date	Deal Type	Upfront Payment	Total Deal Highlights
TUBULIS		Phase 1/2	TUB-040	NaPi2b	10/2025	Financing	\$361M	\$361M (€308M) Series C to advance ADC pipeline, including TUB- 040 in Phase 1/2 for ovarian and NSCLC. Interim data expected at ESMO 2025
Roche	HANSOH P H A R M A	Phase 1	HS-20110	CDH17	10/2025	Licensing	\$80M	Global license for CDH17-targeted ADC (HS-20110) valued up to \$1.45B with \$80M upfront. Expands Roche's ADC portfolio; Phase 1 for colorectal and other solid tumors, excluding rights in China.
EVOPOINT信诺维 Blosciences	astellas	Phase 2	2 ADCs	CLDN18.2	5/2025	Licensing	\$130M	\$130M upfront + potential \$70M in near-term milestones; \$1.34B in development, regulatory and commercialization milestones
<b>₩</b> ararıs	Johnson-Johnson	Discovery	Undisclosed Research	Undisclosed	4/2025	Collaboration	Undisclosed	Research Agreement to develop ADCs
<b>₩</b> ararıs	TAIHO PHARMA	3 Preclinical	Acquired	Entire Company	3/2025	Acquisition	\$400M	\$400M upfront + \$740M in potential milestones
F普主物 LEPU BIOPHARMA	ARRIVENT	Preclinical	1 ADC	Undisclosed	1/2025	Licensing	\$47M	\$47M upfront + \$1.16B total milestones and royalties
biohaven°	Merus	Discovery	3 ADCs	Undisclosed	1/2025	Collaboration	Undisclosed	Research collaboration and license agreement to co-develop 3 ADCs
Synaffix	Mitsubishi Tanabe Pharma	Discovery	Undisclosed Research	Undisclosed	1/2025	Licensing	Undisclosed	Undisclosed upfront payment licensed.
Synaffix connect to cure	Boehringer Ingelheim	Discovery	Undisclosed Research	Undisclosed	1/2025	Collaboration	Undisclosed	Undisclosed upfront with up to \$1.3B in milestones and royalty payments
<b>₩</b> ararıs	Chugai	Discovery	Undisclosed Research	Undisclosed	1/2025	Collaboration	Undisclosed	Undisclosed upfront with up to \$780M in milestones and royalty payments
Vela <b>⊮i</b> go	AVENZO THERAPEUTICS	Preclinical	VAC-103	EGFR x HER3	1/2025	Licensing	\$50M	\$50M upfront for rights outside of China, \$1.15B total deal potential + royalties.
Dual <mark>Ť</mark> tyBio <sup>映 恩 生 物</sup>	AVENZO THERAPEUTICS	Preclinical	AVZO-1418/DB-1418	EGFR/HER3	1/2025	Licensing	\$50M	\$50 million and will be eligible to receive up to approximately \$1.15 billion in development, regulatory and commercial milestone payments
WuXi Biologics Global Solution Provider	bioscience	Preclinical	3 ADCs	PTK7-ADC, MUC16-ADC, SEZ6-ADC	12/2024	Licensing	\$44M	\$44M upfront + \$265M in development and \$540M in commercial milestones, plus single-digit royalty (all 3 assets included)
Synaffix	ELEVATION ONCOLOGY	Preclinical	EO-1022	HER3	12/2024	Licensing	\$368M	\$368 million in upfront and clinical, regulatory, and commercial milestone payments, plus tiered royalties on net sales



### **Investment Summary**

Antibody Drug Conjugates Designed With Novel Immuno-Oncology Payload, PH1 (Spliceosome Modulator)

Filling significant gap in ADC payload innovation

 >90% use DNA damaging agents (topo-1 inhibitors, others) or microtubule inhibitors

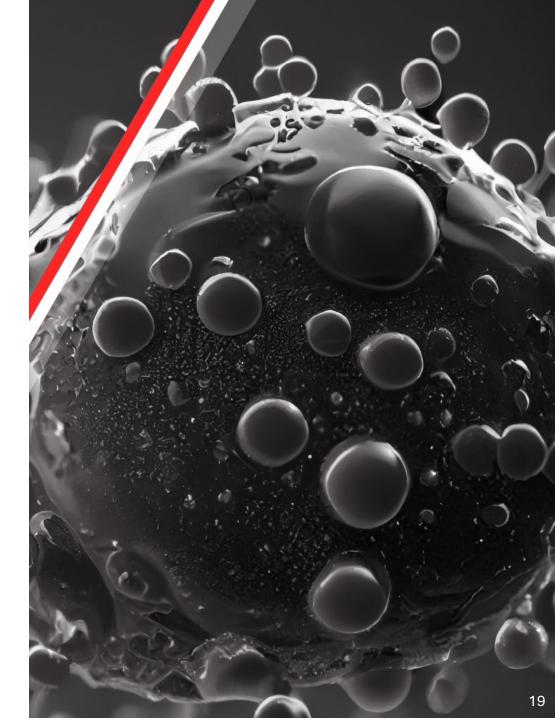
AKTX-101: Lead TROP2 ADC has clear timeline and milestones to target IND/FIH regulatory filing for initial clinical studies

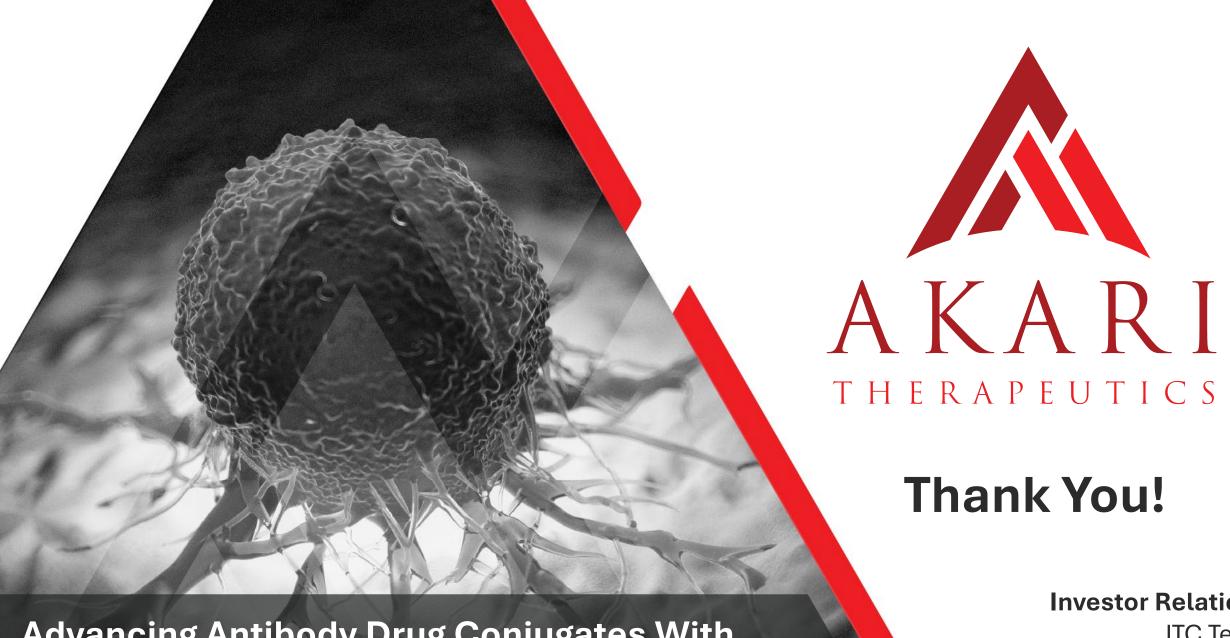
AKTX-102: Potential best-in-class antibody combined with PH1 payload entering preclinical development

 Large opportunity in colon, gastric and lung cancers, in niches where current ADCs have been limited

Continued deal flow for early-stage ADC programs underscores potential for near-term value creation







**Advancing Antibody Drug Conjugates With Novel Immuno-Oncology Payloads** 

**Thank You!** 

**Investor Relations** JTC Team aktx@jtcir.com (908) 824-0775