

**ATACs: Unique new mode of action to fight cancer**

## Forward looking statements

This communication contains certain forward-looking statements, relating to the Company's business, which can be identified by the use of forward-looking terminology such as "estimates", "believes", "expects", "may", "will" "should" "future", "potential" or similar expressions or by general discussion of strategy, plans or intentions of the Company. Such forward-looking statements involve known and unknown risks, uncertainties and other factors, which may cause our actual results of operations, financial condition, performance, or achievements, or industry results, to be materially different from any future results, performance or achievements expressed or implied by such forward-looking statements.

Such factors include, among others, the following: uncertainties related to results of our clinical trials, the uncertainty of regulatory approval and commercial uncertainty, reimbursement and drug price uncertainty, the absence of sales and marketing experience and limited manufacturing capabilities, attraction and retention of technologically skilled employees, dependence on licenses, patents and proprietary technology, dependence upon collaborators, future capital needs and the uncertainty of additional funding, risks of product liability and limitations of insurance, limitations of supplies, competition from other biopharmaceutical, chemical and pharmaceutical companies, environmental, health and safety matters, availability of licensing arrangements, currency fluctuations, adverse changes in governmental rules and fiscal policies, civil unrest, acts of God, acts of war, and other factors referenced in this communication.

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# Key achievements



## Differentiated ADC Technologies

- In Plug & Play mode
- 2 years from target to IND



## Strong IP

- Several IP families
- Monopoly in the Amanitin/MoA space



## GMP Manufacturing

- Fully synthetic process for Amanitin
- 5 GMP batches completed



## Strategic partnerships

- Huadong: China-focused partnership
- Takeda: ATAC technology partnership



## Clinical Stage

- 1 ATAC in ongoing Phase I
- 2 additional ATAC INDs within the next year



## Corporate & Finance

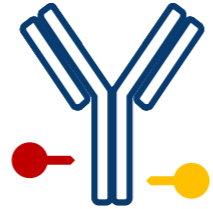
- Experienced leadership team
- Cash (runway): EUR 50.7m\* (mid-2025)

\* as per end of August 2023

# Strong in-house R&D capabilities and expertise



Synthetic chemistry



Antibody generation  
& bioconjugation



Preclinical testing



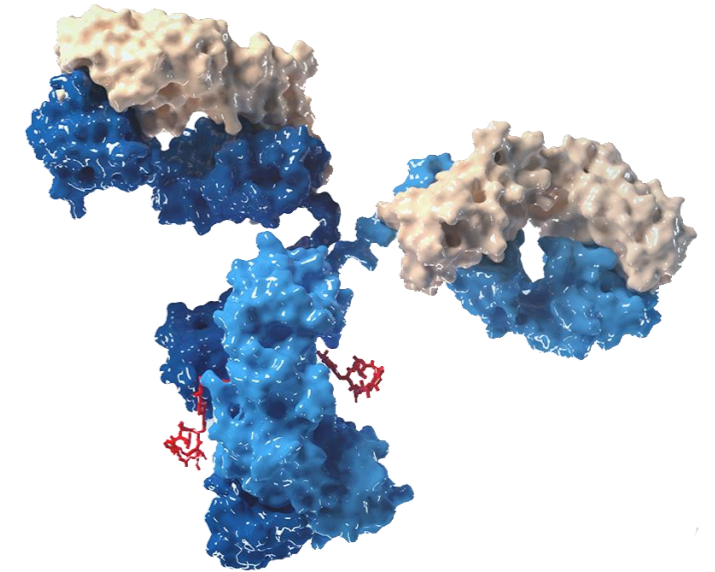
CMC



Bioanalytical sciences



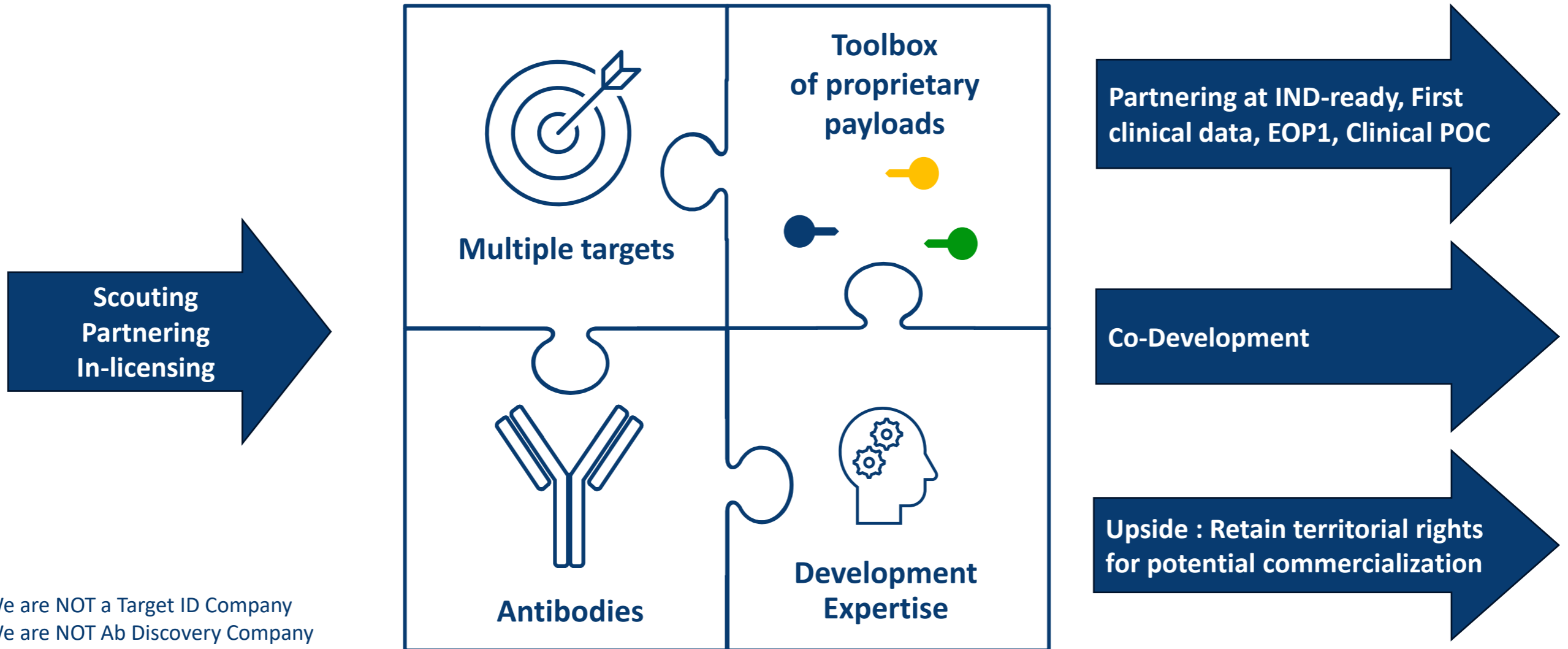
Clinical Development



Best ADC candidate in the shortest time

# Value creation through development of best-in-class ADC assets

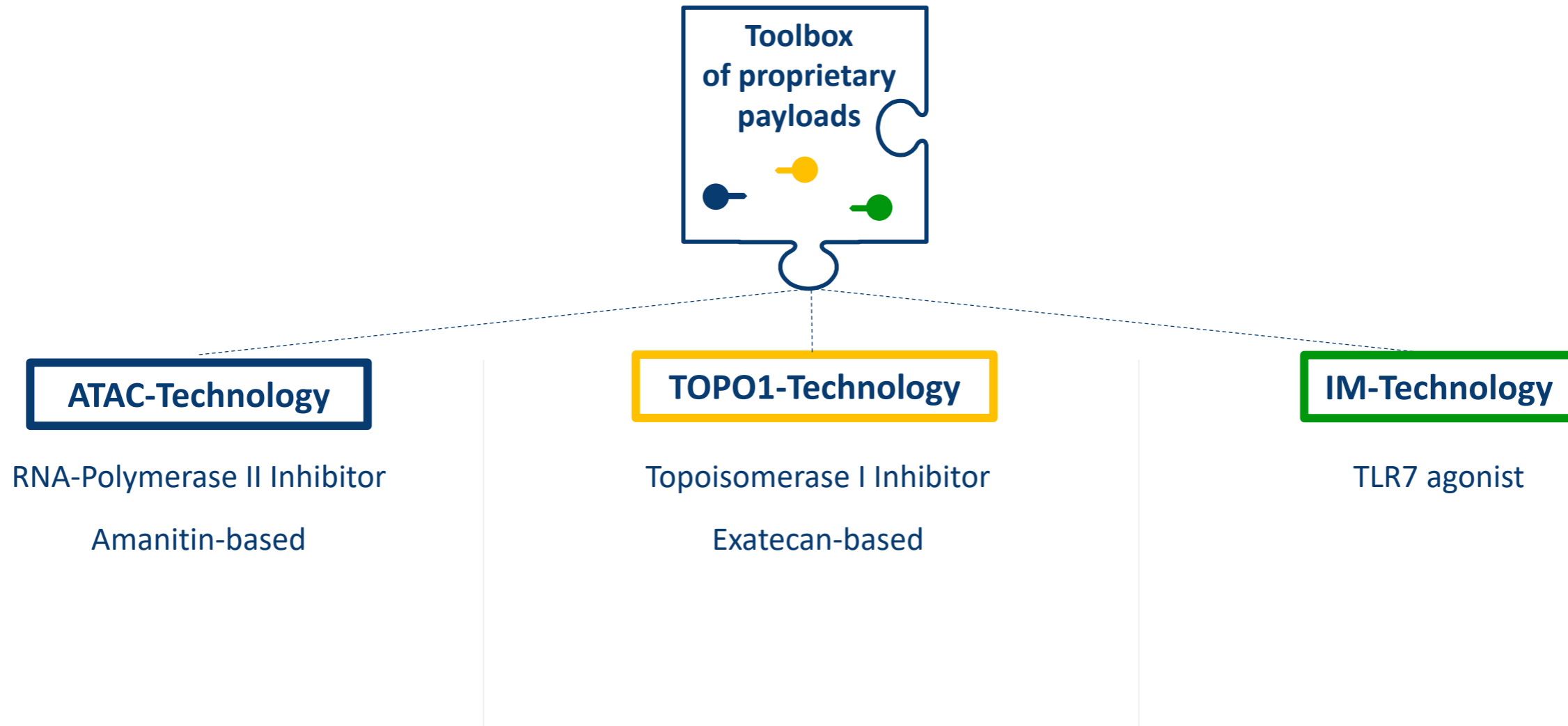
## Discovery & development engine



We are NOT a Target ID Company  
We are NOT Ab Discovery Company



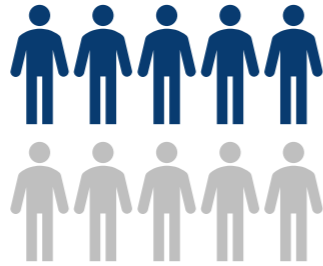
# Payload Toolbox – Multiple MOAs



# Growing pipeline of proprietary and partnered programs

	Product	Target	Indication	Research	Preclinic	Phase I	Phase II	Phase III	Partner	
ATAC pipeline	HDP-101	BCMA	Multiple Myeloma	[Progress bar]					Huadong (China+)	
	HDP-102	CD37	NHL (DLBCL/CLL)	[Progress bar]					Huadong (option China+)	
	HDP-103	PSMA	Prostate cancer	[Progress bar]					Huadong (China+)	
	HDP-104	GCC	Gastrointestinal (e.g., CRC)	[Progress bar]					Huadong (option China+)	
TOPO	HDP-201	n/a	Solid tumors	[Progress bar]					Proprietary	
ATAC partners	TAK-ATAC	n/a	Oncology	[Progress bar]					Takeda	
Legacy assets	TLX250-CDx	CA-IX	Renal Carcinoma Urothelial Carcinoma, TNBC	[Progress bar]						Telix
	TLX250	CA-IX	Renal carcinoma	[Progress bar]						Telix
	RHB-107		Oncology/GI, Covid-19	[Progress bar]						RedHill

# Resistance is one of the biggest challenges in oncology



**1 in 2**

people will be diagnosed  
with cancer in their  
lifetime



**> 90%**

of cancer deaths  
are caused by drug  
resistance



# The journey of many cancer patients

## Before Treatment



## Treatment



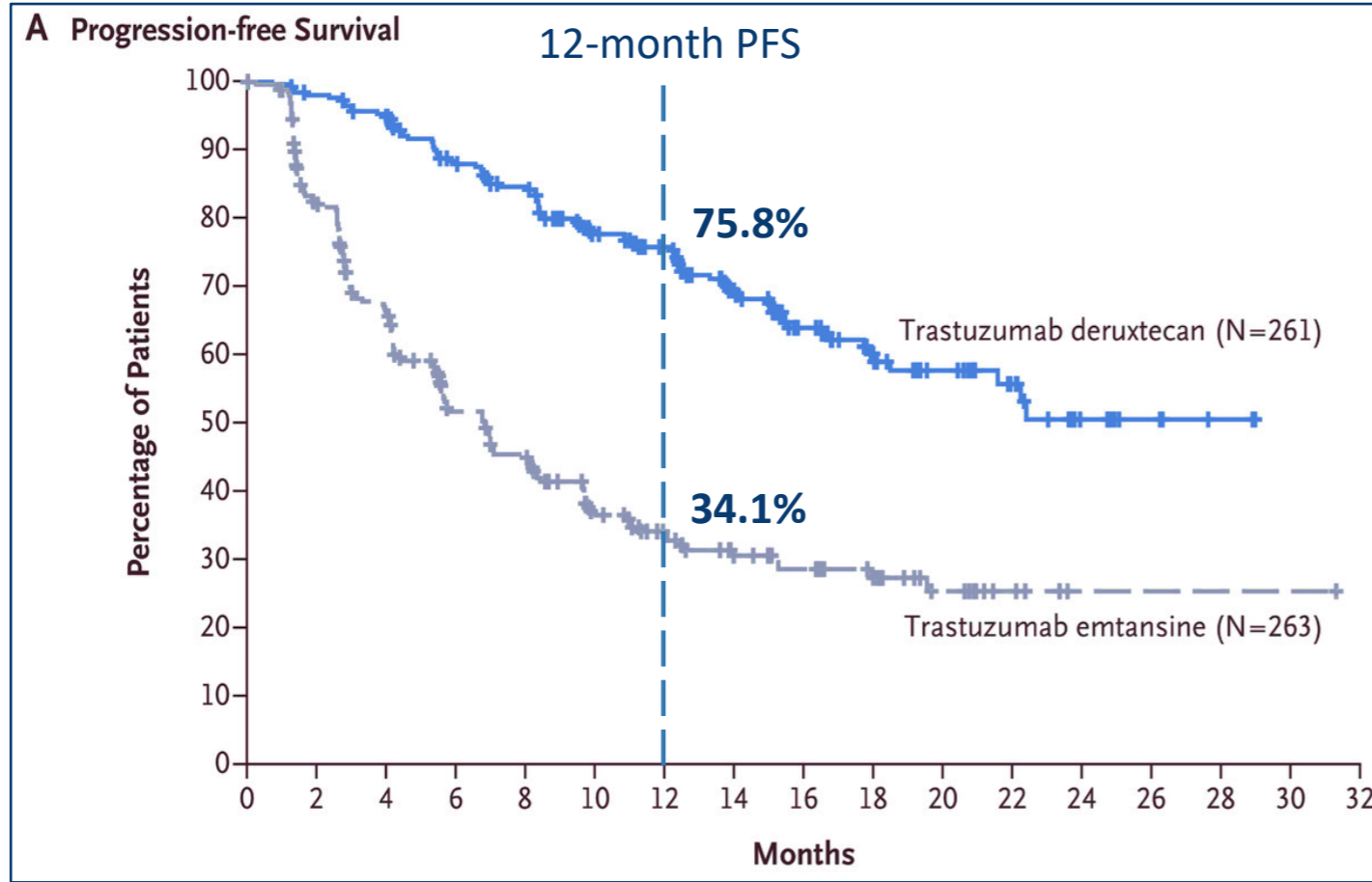
## Resistance & Relapse



Wagke, N. et al, J Clin Oncol. 2011; 29(22): 3085–3096

**We need new drugs with new mode-of-action to overcome resistance**

# The payload MOA is what makes the difference!



Cortés, J. *et al*, N Engl J Med 2022; 386:1143-1154

**Enhertu®**

Payload: deruxtecan (Topo 1 inhibitor)

**Kadcyla®**

Payload: emtansine (Tubulin inhibitor)

**Same target (Her2), same antibody (Trastuzumab), same patient population**

# Novel payloads to overcome resistance

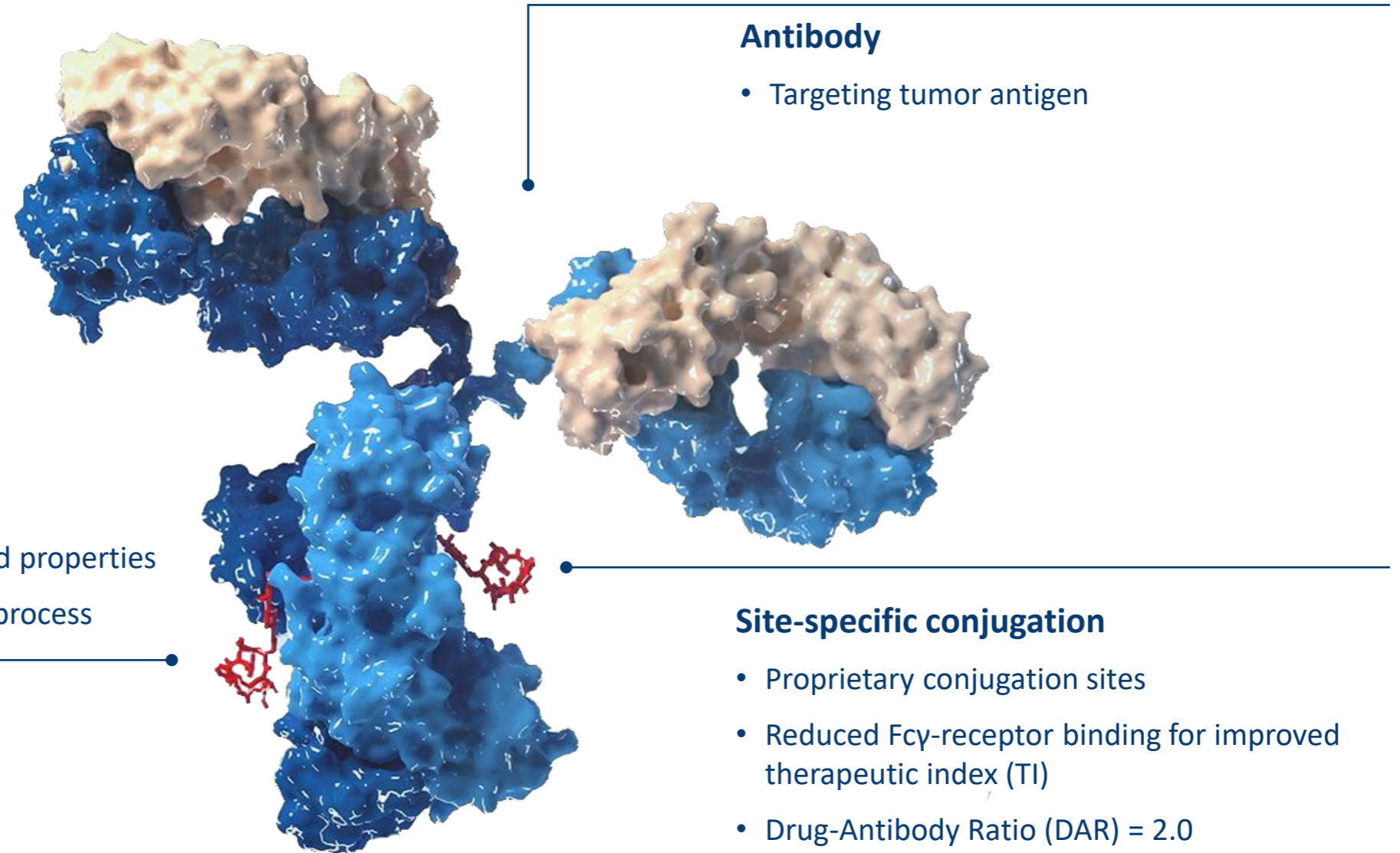
	Tubulin inhibitors <i>e.g. Maytansines &amp; Auristatines</i>	DNA-damaging agents <i>e.g. PBDs, PDDs, IGNs, Calicheamicin, Duocarmycins</i>	Topoisomerase inhibitors <i>e.g. Camptothecins, Deruxtecan, SN-38</i>	RNA polymerase inhibitors <b>Amanitin</b>
<b>Potency</b>	High	Ultra-high	Low	Medium
<b>Hydrophilicity</b>	✗	✗	✗	✓
<b>Overcome resistance</b>	✗	✗	✗	✓
<b>Active on non-dividing cells</b>	✗	✓	✗	✓
<b>Biomarker</b>	✗	✗	✗	✓
<b>Target Exclusivity / Single player / IP monopoly</b>	✗	✗	✗	✓

**Amanitin has a mechanism of cytotoxicity that is radically different from that of conventional chemotherapy**

# ATACs are ADCs with amanitin as a payload

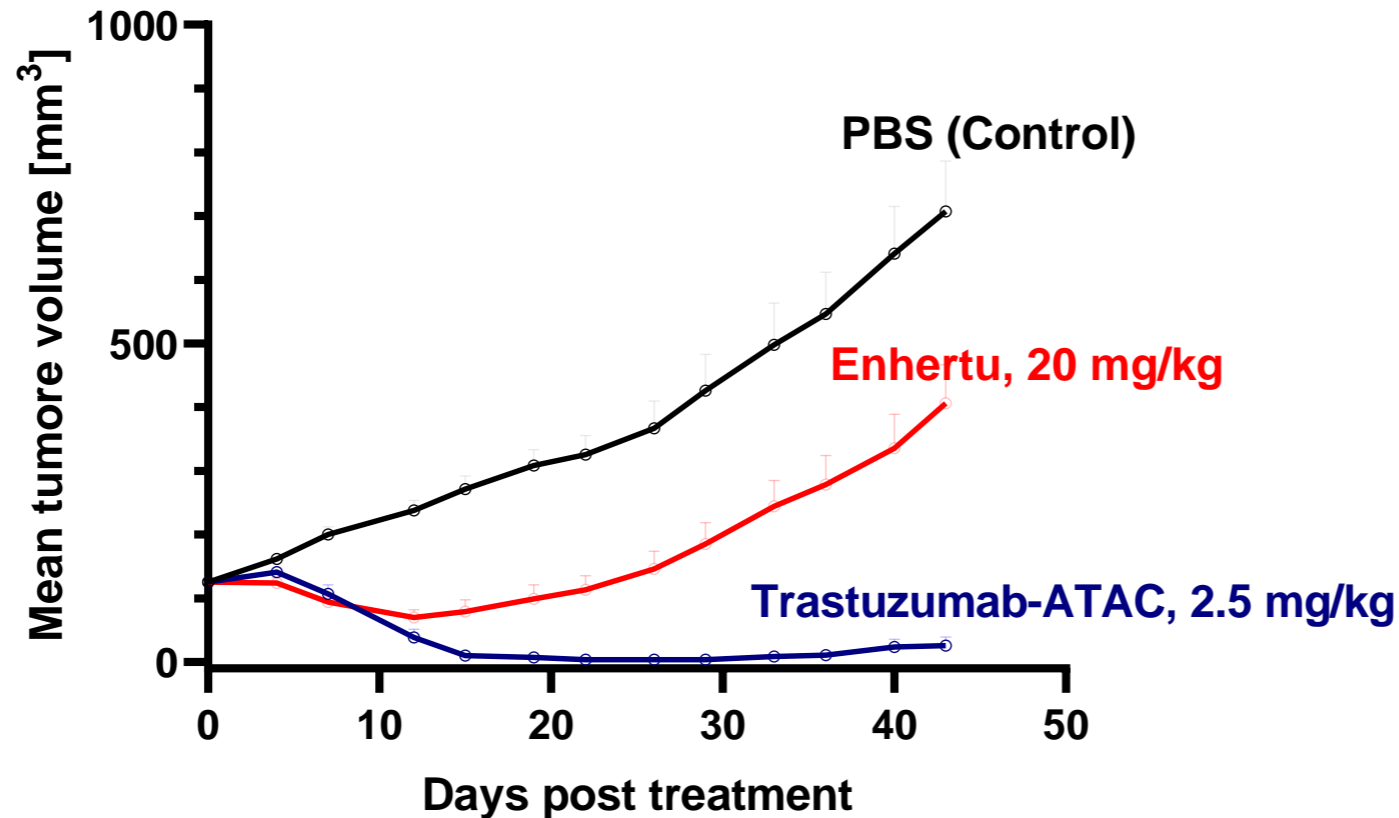
## Amanitin as warhead

- Differentiated mechanism of action:  
*inhibition of RNA Polymerase II*
  - Kills dormant tumor cells
  - Overcomes resistance
  - Predictive biomarker
- Synthetic amanitin derivatives with improved properties
- GMP manufacturing through fully synthetic process



# The Payload Makes The Difference

Breast cancer model (JIMT-1 Xenograft) is resistant to Kadcyła<sup>®</sup> and Enhertu<sup>®</sup>

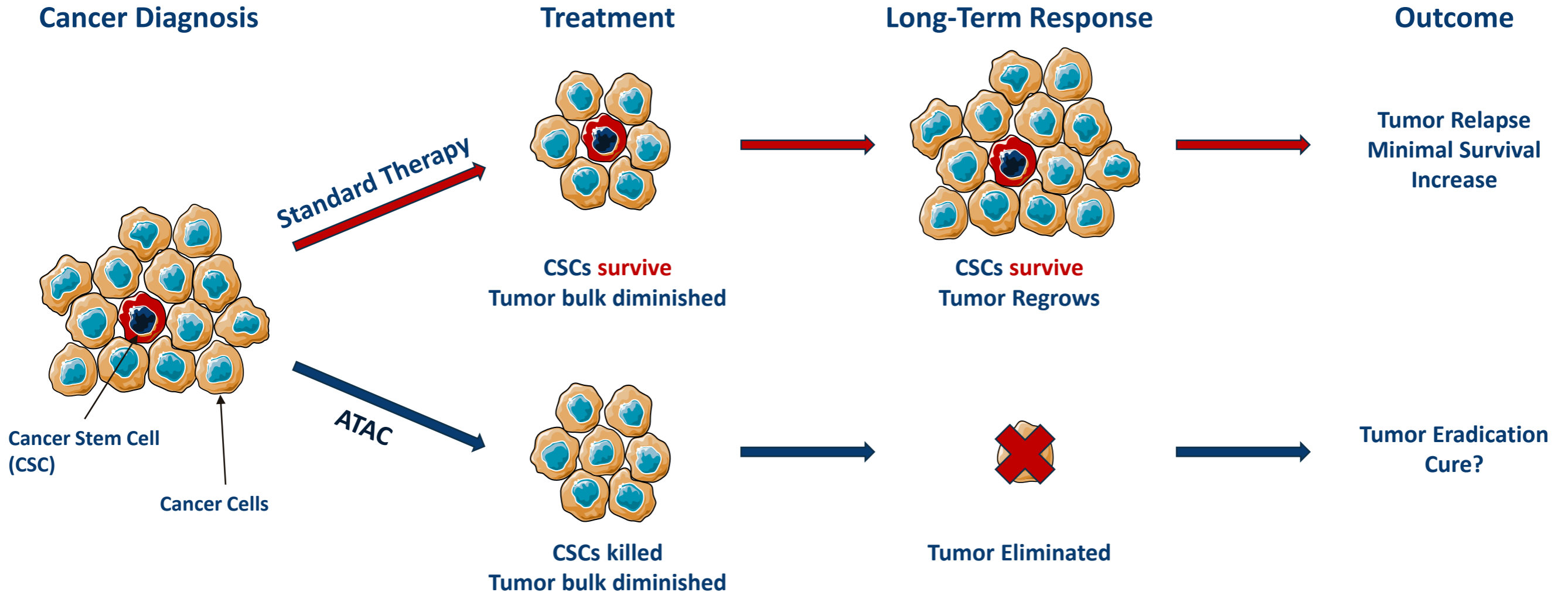


- Same antibody (Trastuzumab), different payload (amanitin vs. topoisomerase inhibitor)
- **Complete remission after single-dose application of HER2-ATAC.**

**Trastuzumab ATAC leads to complete remission in resistant model after single-dose**



# ATACs address the limitations of current cancer therapies



**Amanitin has a mechanism of cytotoxicity that is radically different from that of conventional chemotherapy**



# Del(17p): Potential platform-wide predictive biomarker

## Deletion of TP53 (tumor suppressor)

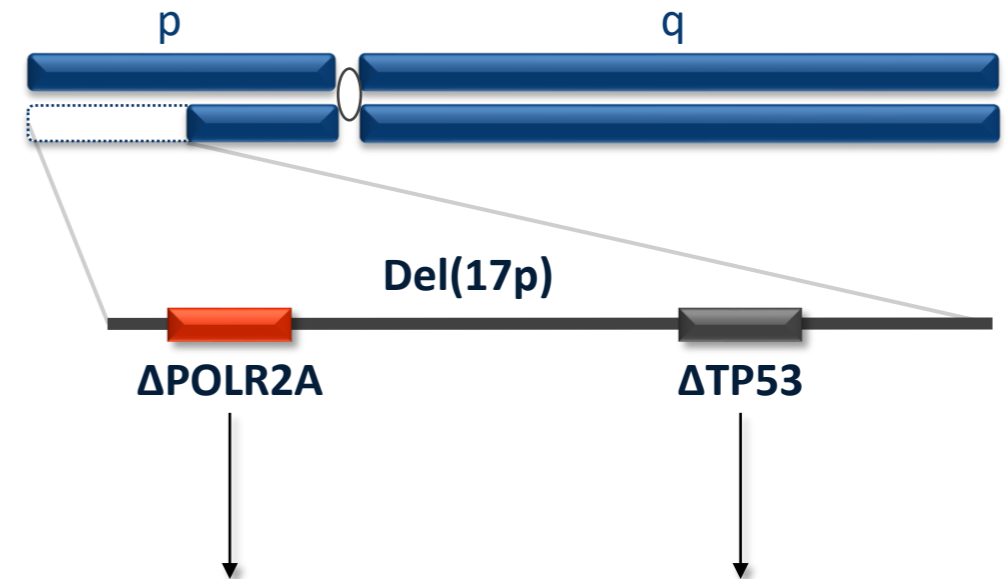
- High incidence
- More aggressive tumors **resistant** to SoC and poor prognosis

## Deletion of RNA Polymerase II (POLR2A is co-deleted)

- Higher sensitivity to ATAC treatment

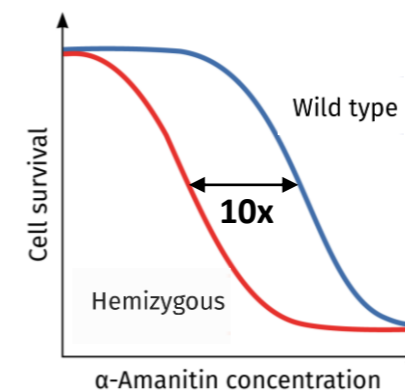
## Occurs only in tumor cells

- Wider therapeutic window in patients with del(17p) tumors
- Across cancer indications and tumor types



Intracellular target of amanitin: Increases ATAC sensitivity

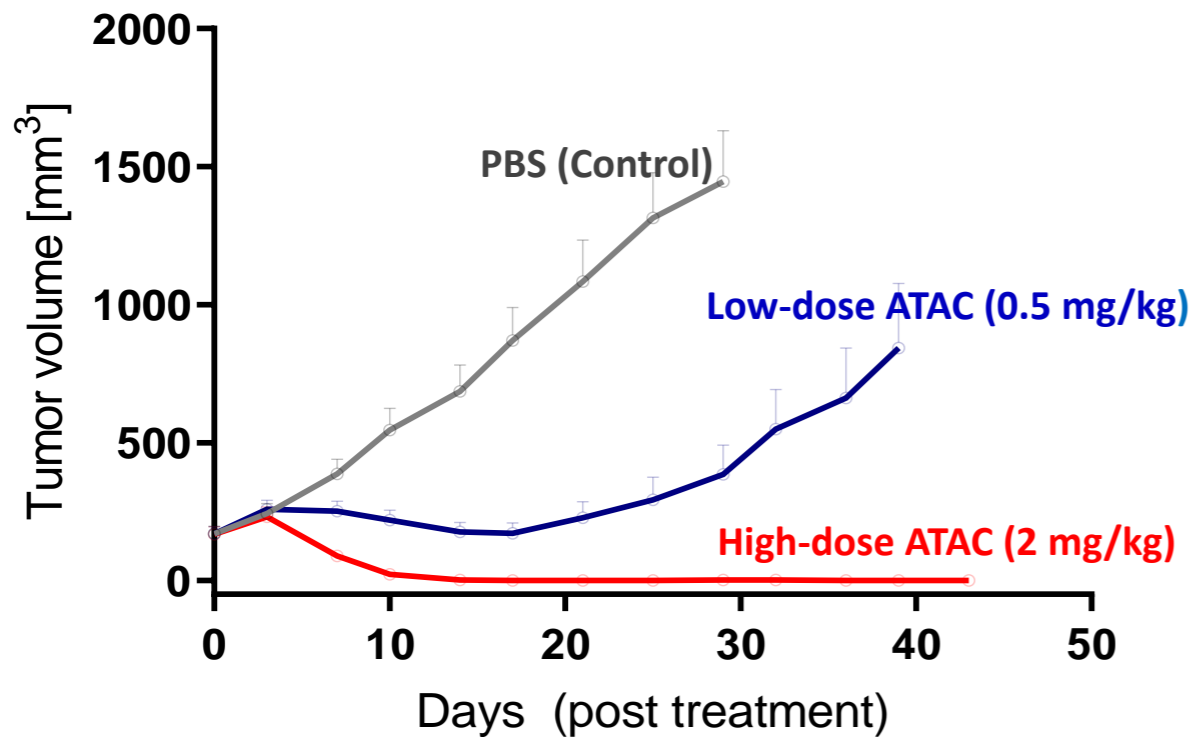
Tumor suppressor: Increases tumor aggressiveness & resistance



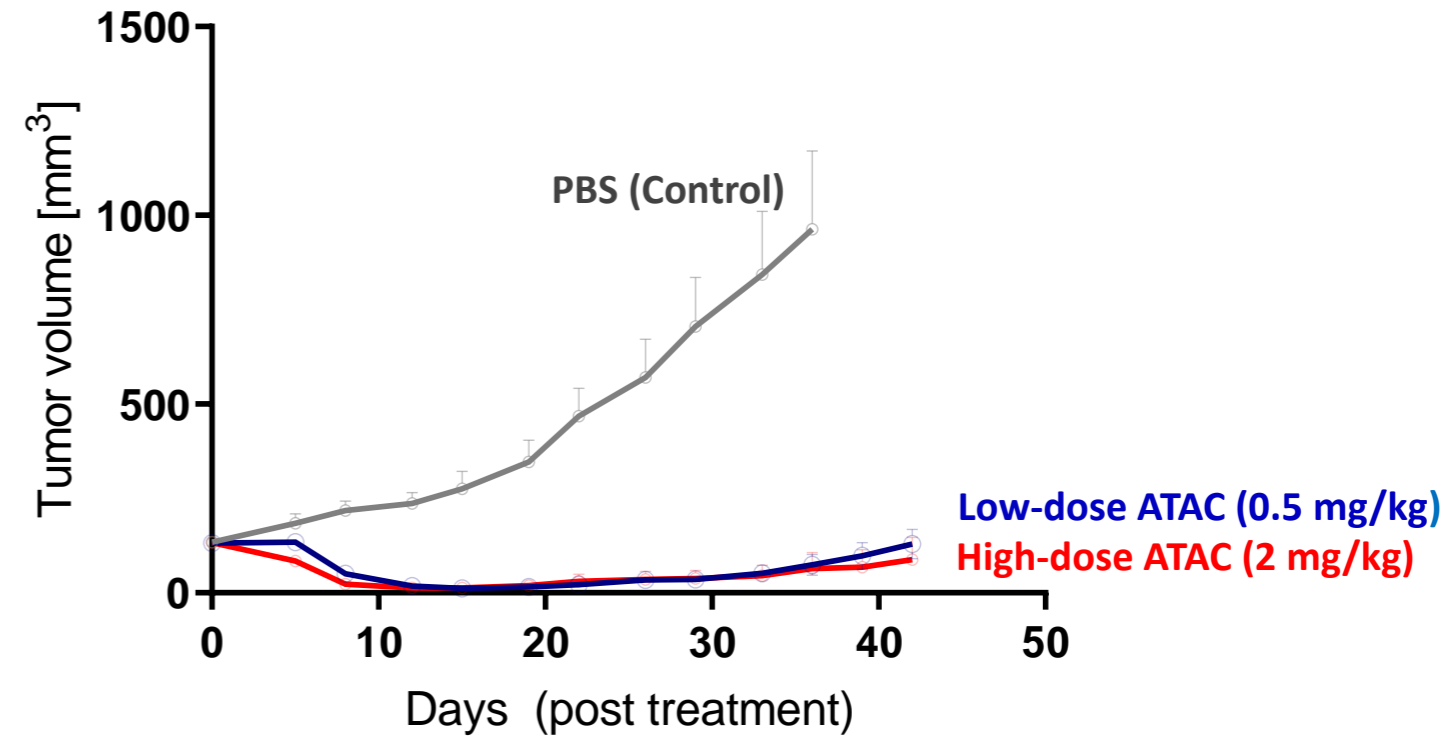
# Del(17p): Potential platform-wide predictive biomarker

Her2 1+ patient-derived xenograft models

**Wildtype** - normal RNA Pol II levels



**Del(17p)** - reduced RNA Pol II levels



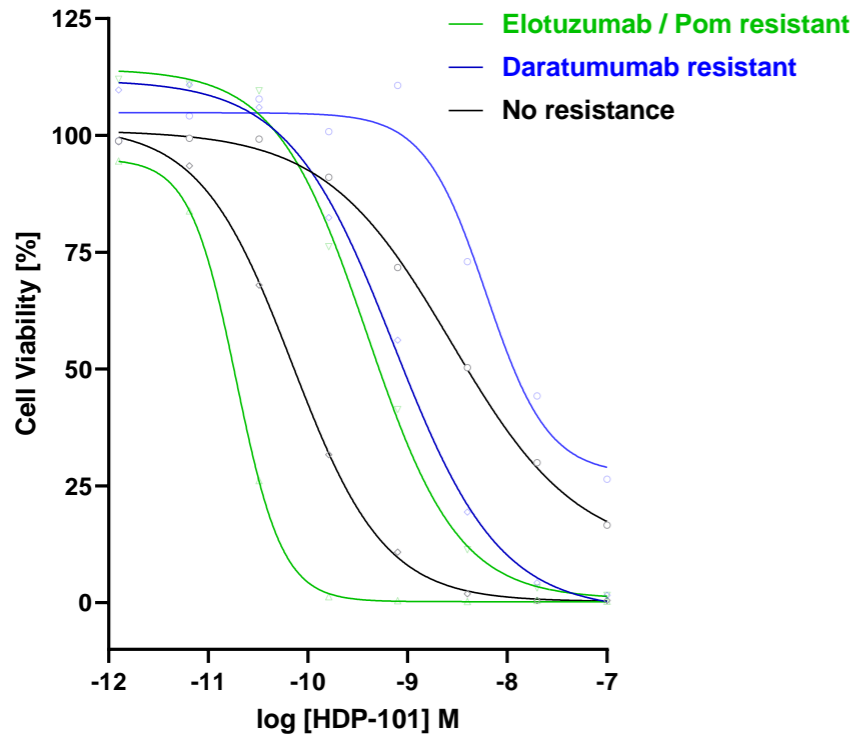
**Less amanitin is required to kill del(17p) cells**

**Wider therapeutic index in patients with del(17p) tumors**

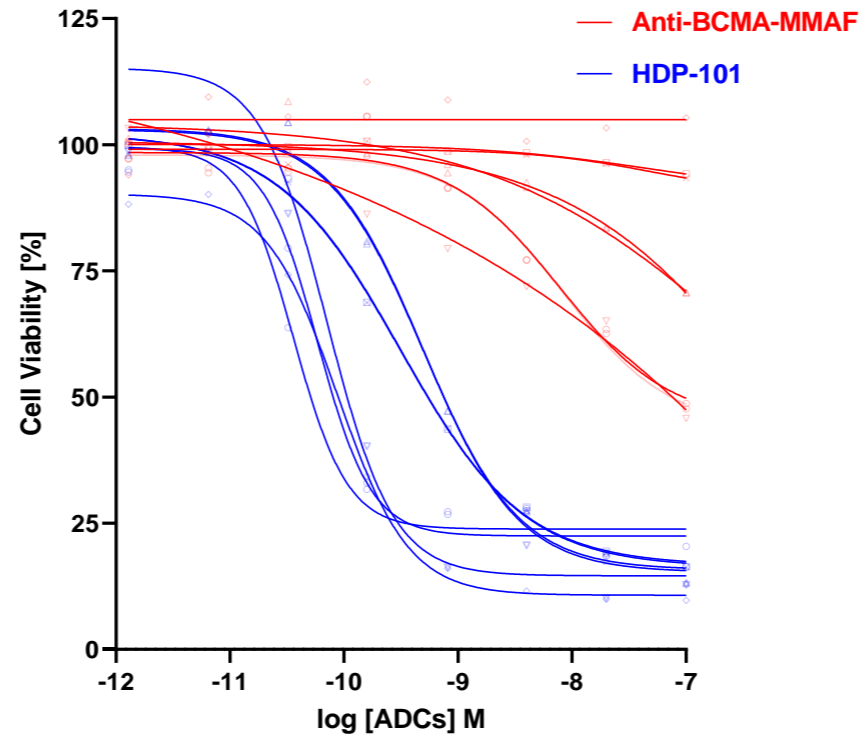
# We know ATACs work

HDP-101 is highly efficacious in primary myeloma cells from patients

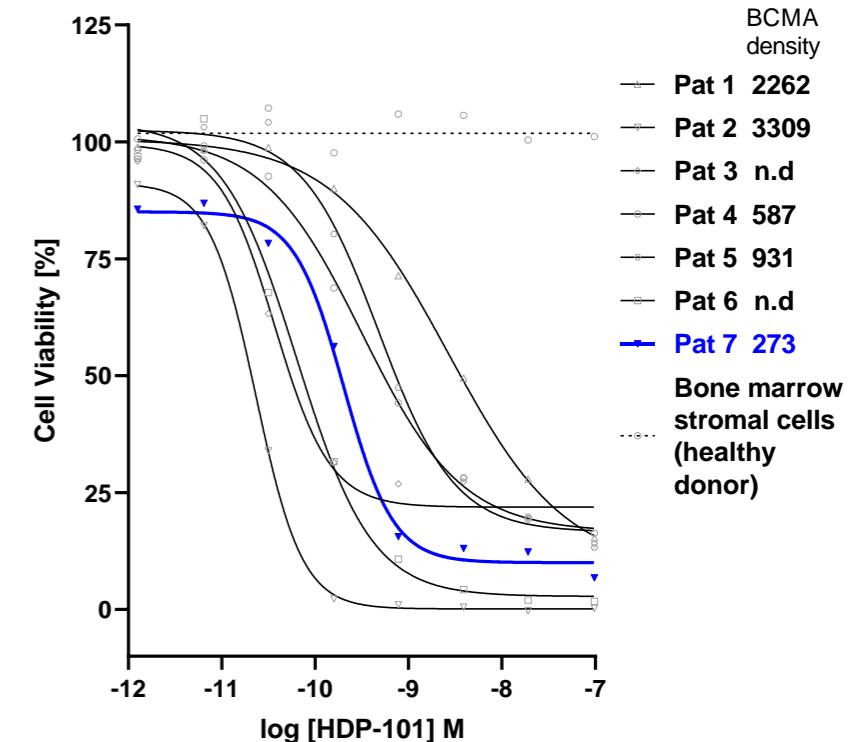
Overcomes resistance in patients refractory to SOC



More efficacious than other payloads by killing non-dividing tumor cells



Overcomes resistance through antigen escape by killing cells with ultra-low antigen expression

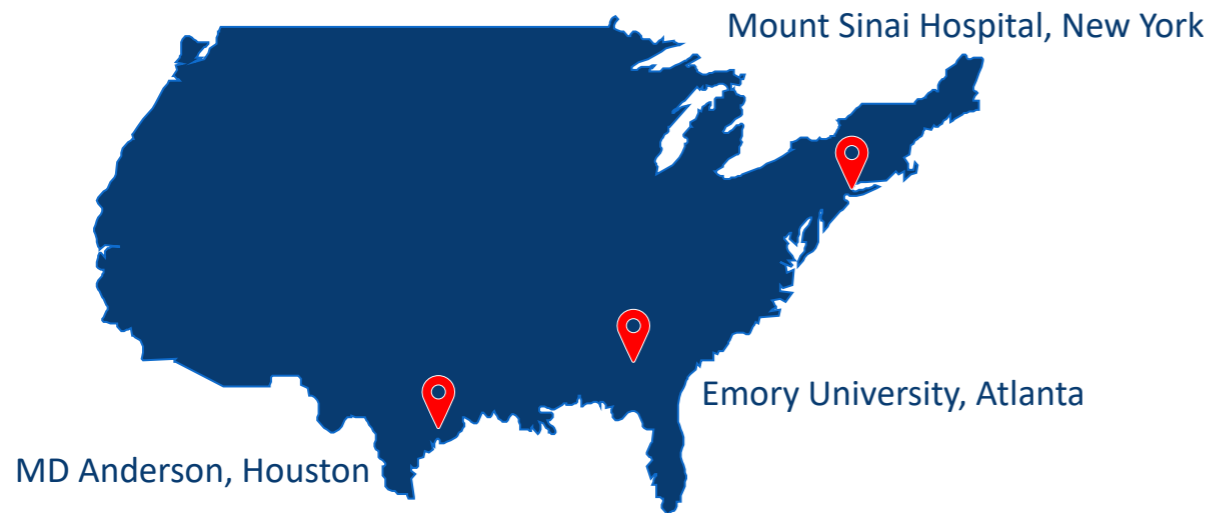


HDP-101 overcomes multiple types of resistance in patient cells

# HDP-101 Phase I/II study

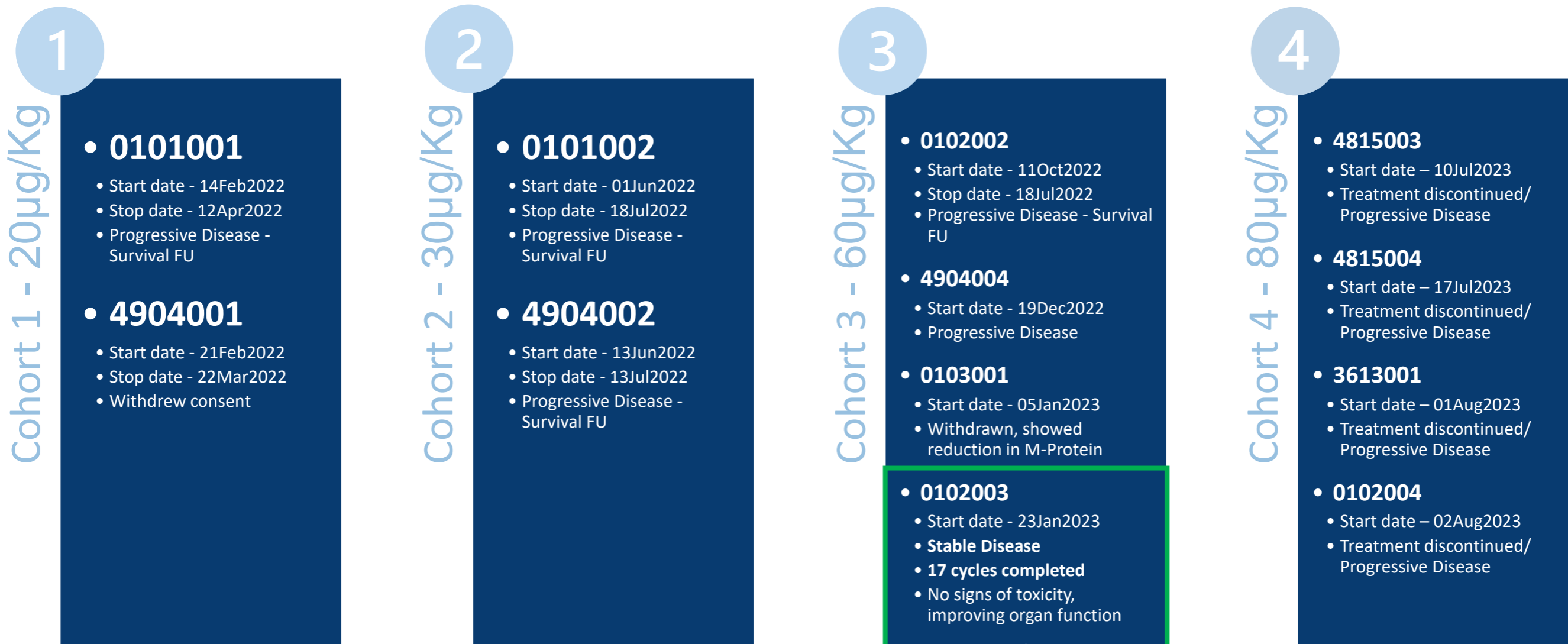
## Study status

- Four patient cohorts (20, 30, 60 and 80  $\mu\text{g}/\text{kg}$ ) completed, 12 patients in total
- Latest Safety Review Committee conclusions (September 2023):
  - Treatment is safe and well-tolerated in the four cohorts
  - Continue dose escalation
- Dose escalation continues with 100  $\mu\text{g}/\text{kg}$  in the fifth cohort



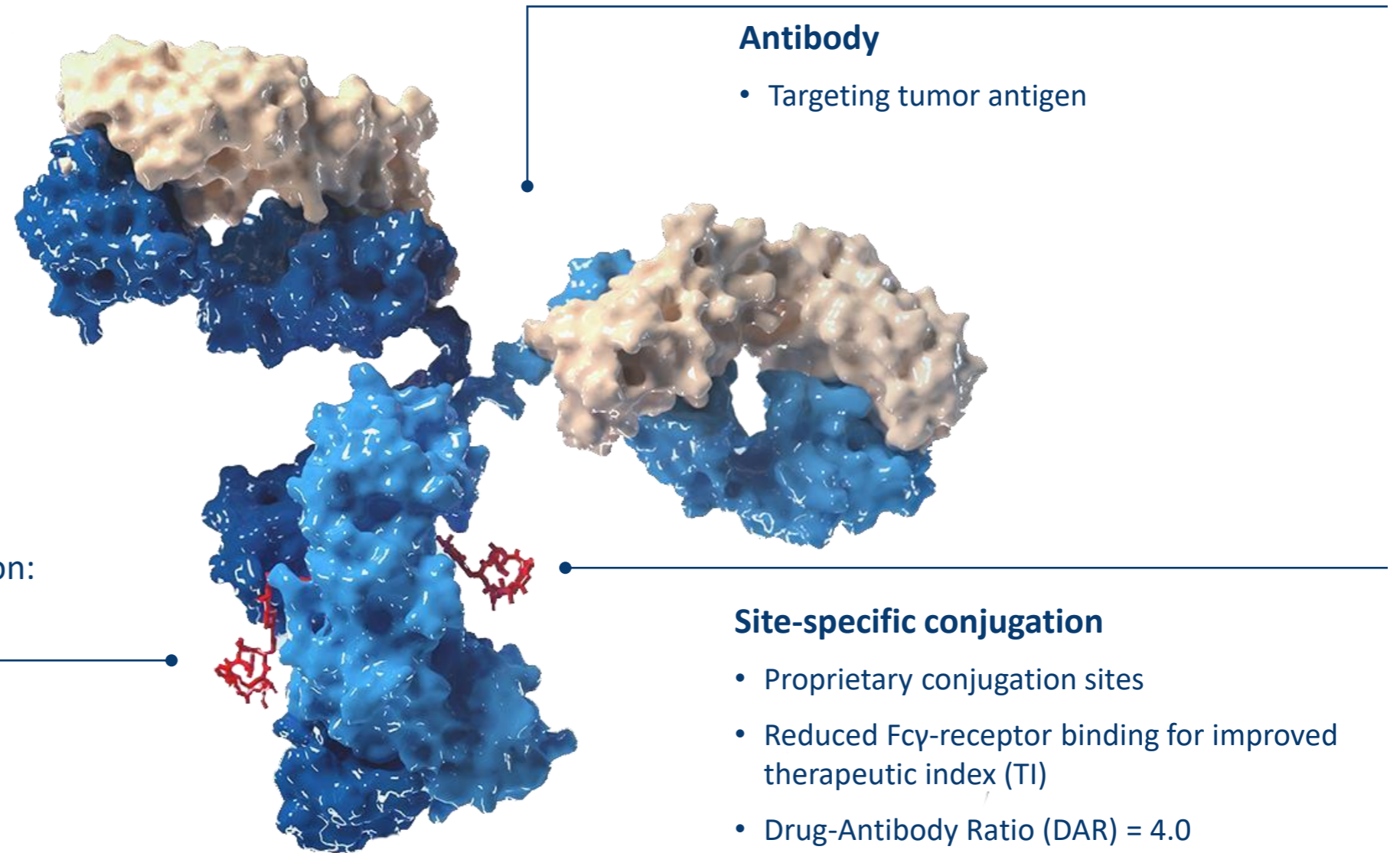
# HDP-101: It works in the clinic

- 1 patient from cohort 3 with SD for 14 cycles, on monotherapy for 11 months



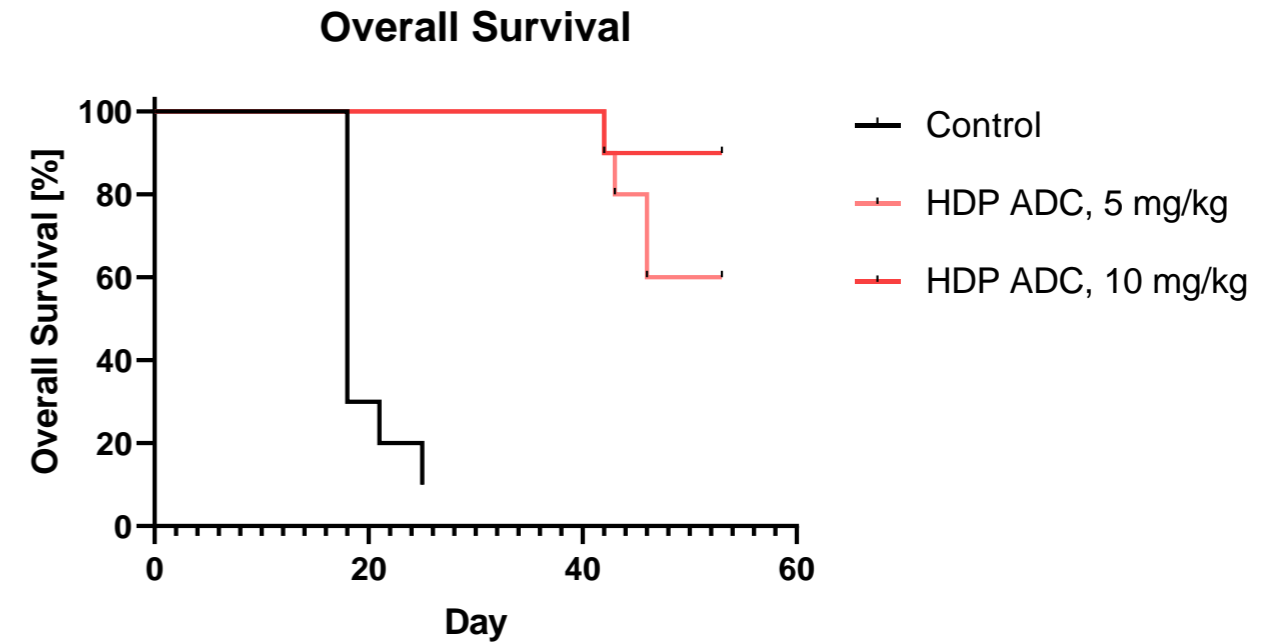
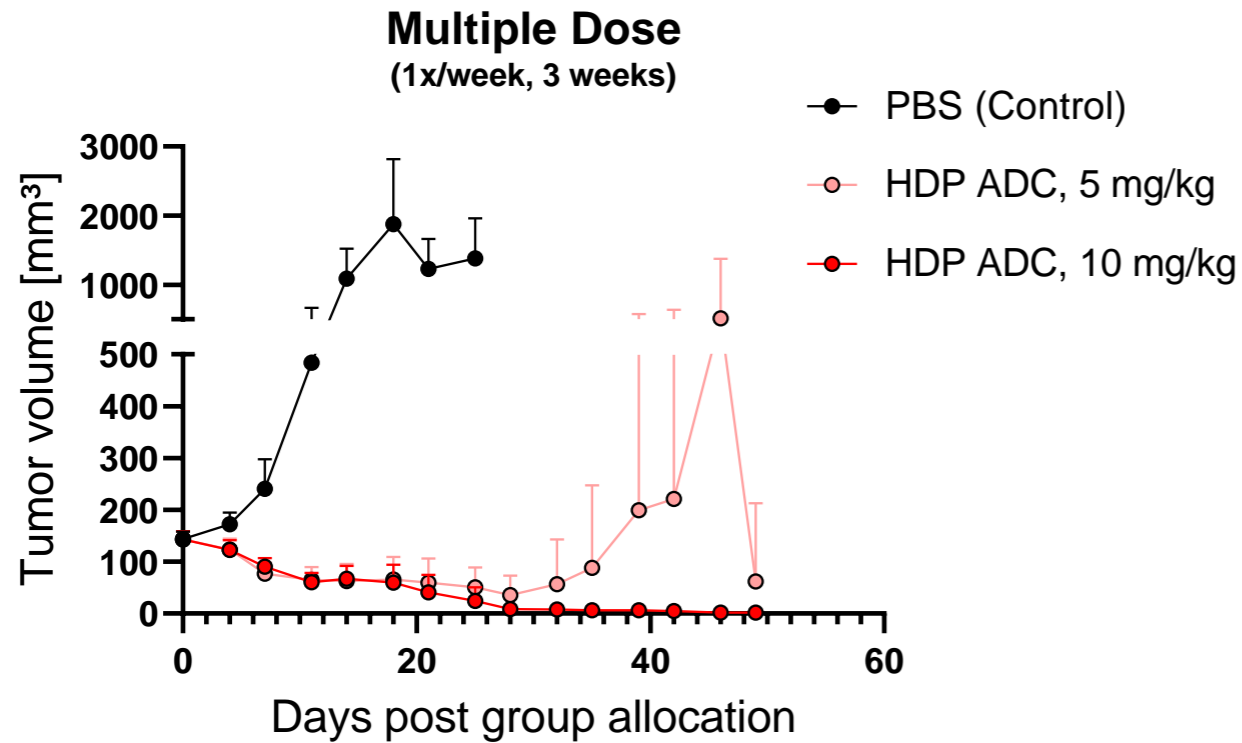
Dose elevated to 80 µg/kg

# ADCs with TOPOI inhibitor as a payload





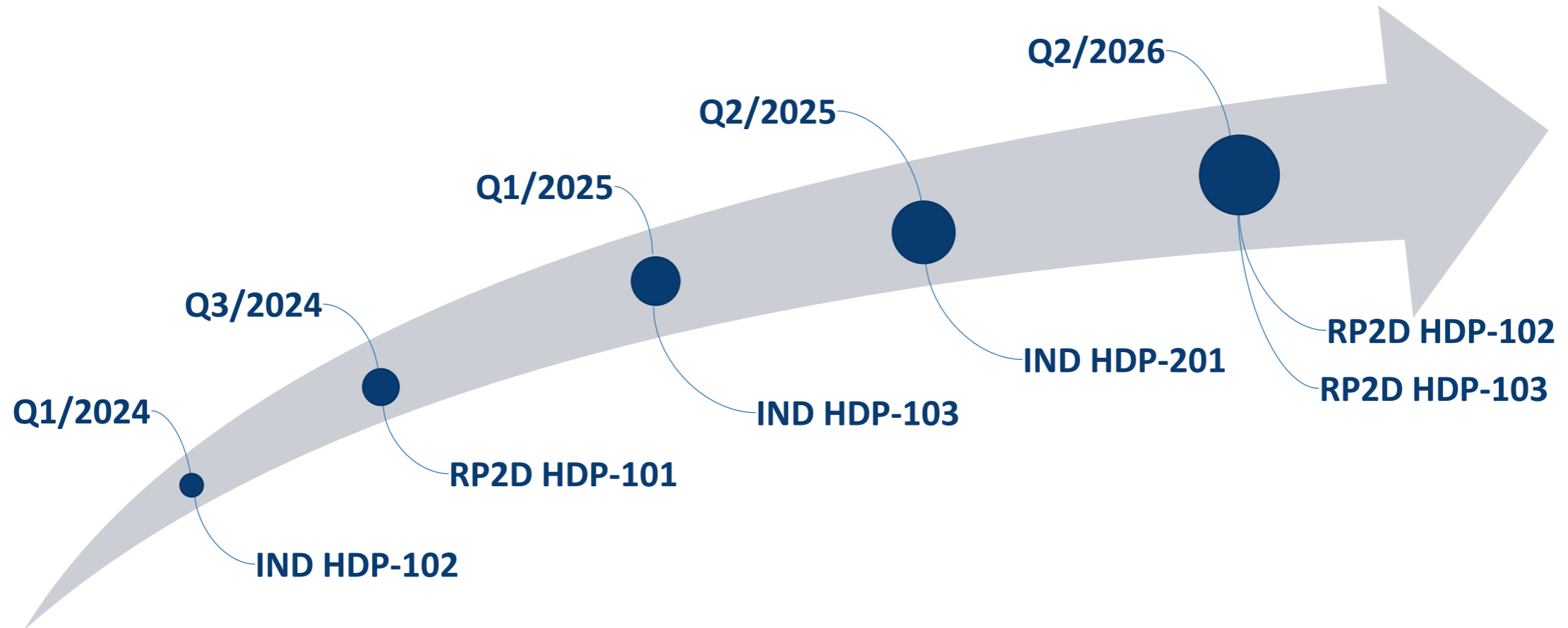
# Strong efficacy of HDP's Topo 1 ADC upon multiple dose treatment

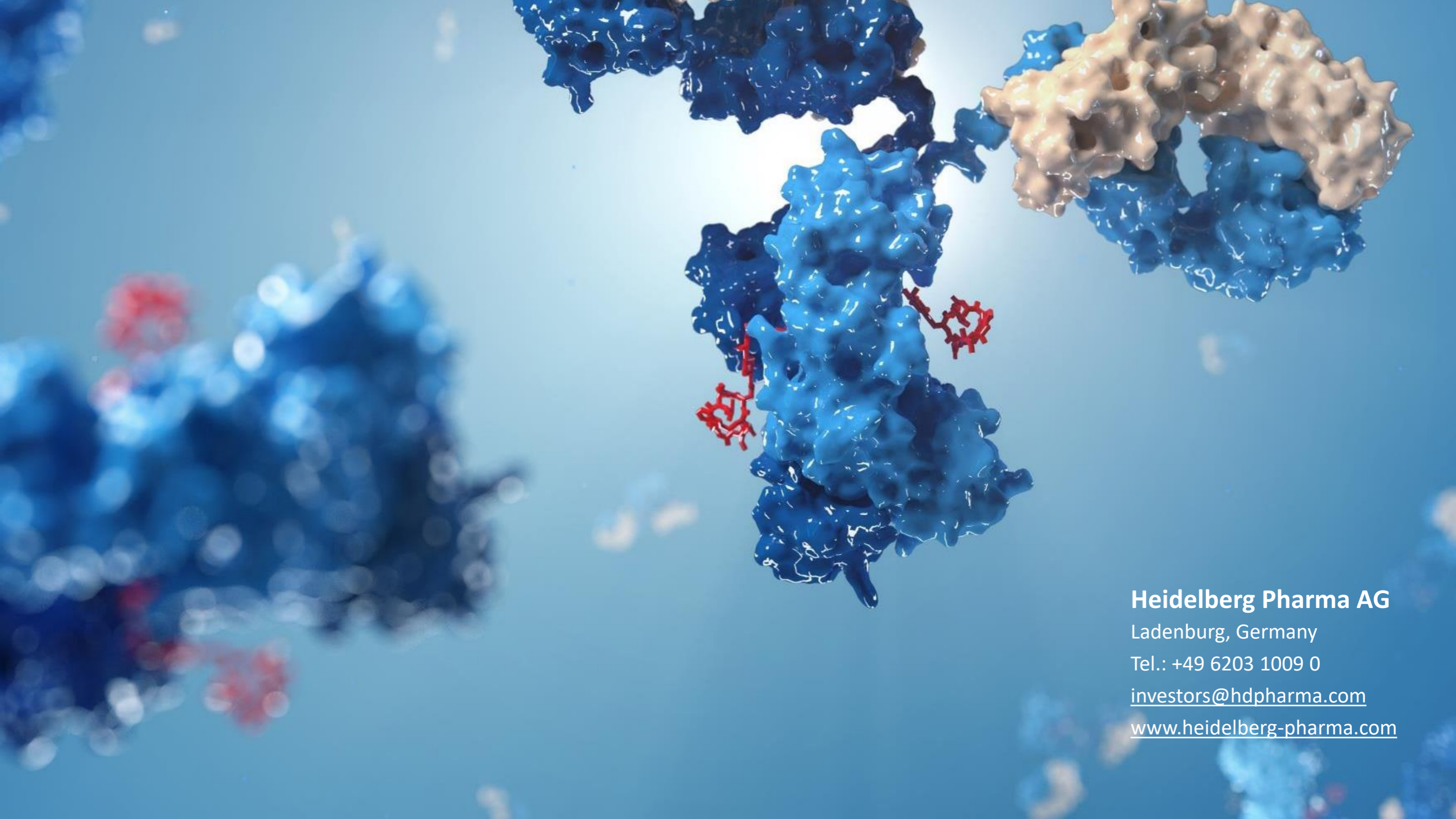


- Efficacy of HDP ADC similar to Deruxtecan ADC
- Only half the amount of toxin (DAR 4 vs DAR 8-10)

# Outlook

- We are a clinical-stage company with the goal of becoming a leading global ADC player
- Multiple inflection points over the next 36 months with potential to many-fold increase of company valuation





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