



# Extending the Boundaries of Targeted Cancer Therapies with Radio-DARPin and Next-Gen Immune Cell Engagers

Full-Year Earnings Call

March 7, 2025

Nasdaq, SIX Swiss Exchange: MOLN

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# Agenda & Speakers

**Welcome &  
Introduction**



**Seth Lewis**  
SVP IR & Strategy

**Highlights 2024  
& Outlook 2025**



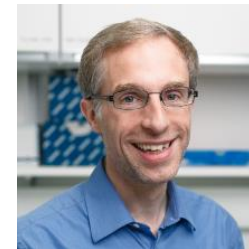
**Patrick Amstutz**  
CEO

**Financial  
Overview**



**Robert Hendriks**  
SVP Finance

**MP0712 &  
Radio-DARPin**



**Michael T Stumpp**  
EVP Projects

**MP0533 &  
Switch-DARPin**



**Philippe Legenne**  
CMO

# Corporate Highlights 2024



## MP0712 & Radio-DARPin Therapy

- **MP0712** targeting **DLL3** completed **IND-enabling studies**
- **Mesothelin** (MSLN) as second target in Radio-DARPin pipeline, leveraging DARPin uniqueness
- **Strategic collaboration** with **Orano Med expanded to ten** <sup>212</sup>Pb programs

## MP0533 T-Cell engager

- Phase 1/2a study (cohort 8 on-going) with **novel tetra-specific T-cell engager** for AML patients
- **Optimized dosing** schedule in cohort 8 **shows improved response rate and depth**

## Switch-DARPin

- Proof-of-mechanism for MP0621, a cKit x CD16a x CD47 Switch-DARPin for next-gen HSCT
- **Switch-DARPin** platform **expanded to CD3** for logic-gated **T cell engagers**
- **T-cell co-stimulating** DARPins validated (CD2 and others) for solid tumor application


## MP0317

- Phase 1 data for FAP-mediated CD40 activation demonstrates **remodeling of tumor microenvironment**, favorable safety profile in patients with advanced solid tumors; combination IIT in preparation

## Operations

- **Strong financial position** with CHF 149.4M in cash as of December 31, 2024
- Capital raise of \$20M in October 2024 allowing **runway well into 2027**

# Pipeline

MODALITY	CANDIDATE	RESEARCH	PRE-CLINICAL	PHASE 1	PHASE 2	PHASE 3
Radio-DARPin Therapy (RDT) 	<b>MP0712 (DLL3)</b>	SCLC & NETs <i><sup>212</sup>Pb - DLL3</i>		Co-development*		
	<b>RDT x MSLN</b>	Ovarian <i><sup>212</sup>Pb - MSLN</i>		Co-development*		
	Undisclosed Programs	Solid Tumors		Up to 8 programs*		
Next-Gen Immune Cell Engagers	<b>MP0533</b>	r/r AML and AML/MDS <i>CD33 x CD123 x CD70 x CD3</i>				
	<b>Switch-DARPin T Cell Engager</b>	<i>CD3 x costim x TAAs</i>				
	MP0621 (Switch-DARPin)	HSCT <i>cKit x CD16a x CD47</i>				
	MP0317	Advanced Solid Tumors <i>FAP x CD40</i>				



# Financial Overview

# Key Figures FY2024

<i>(CHF million, except per share and FTE data)</i>	<b>FY 2024</b>	<b>FY 2023</b>	<b>change</b>
<b>Revenues</b>	<b>5.0</b>	<b>7.0</b>	<b>(2.0)</b>
<b>Total operating expenses<sup>1</sup></b>	<b>(66.2)</b>	<b>(68.1)</b>	<b>1.9</b>
<b>Operating result</b>	<b>(61.2)</b>	<b>(61.1)</b>	<b>(0.1)</b>
<b>Net financial result</b>	<b>7.2</b>	<b>(0.9)</b>	<b>8.1</b>
<b>Net result</b>	<b>(54.0)</b>	<b>(62.0)</b>	<b>8.0</b>
<b>Basic net result per share (in CHF)</b>	<b>(1.59)</b>	<b>(1.89)</b>	<b>0.30</b>
<b>Net cash from / (used in) operations</b>	<b>(59.2)</b>	<b>(59.0)</b>	<b>(0.2)</b>
<b>Cash balance (incl. s.t. deposits) as of Dec 31<sup>2</sup></b>	<b>149.4</b>	<b>186.9</b>	<b>(37.5)</b>
<b>Number of FTE's as of Dec 31</b>	<b>158.5</b>	<b>167.5</b>	<b>(9.0)</b>

# Financial Guidance\* for 2025

- Total operating expenses of CHF 55-65 million, of which around CHF 7 million non-cash effective costs
- ~**CHF 149 million** cash & cash equivalents (incl. short-term time deposits) ensure comfortable **funding well into 2027** (excl. any potential payments from R&D partnerships)

\* Guidance subject to progress and changes of pipeline



# Radio-DARPin Therapy & MP0712

Custom-engineered to create  
vectors ideal for radiopharmaceuticals

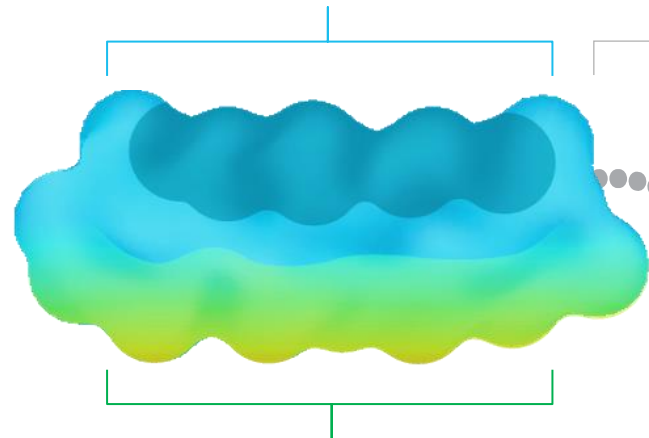


# Radio-DARPin as Versatile Therapeutic Candidates

Combining versatile DARPin features with the power of  $^{212}\text{Pb}$  for next-gen Targeted Alpha Therapy

## DARPin: IDEAL VECTOR FOR RADIOPHARMACEUTICALS

- Proven selective targeting
- High affinity, tumor retention
- Broad target space
- Small size

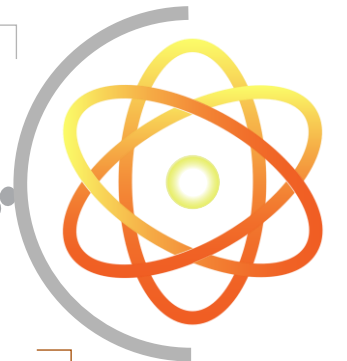


## SURFACE ENGINEERING

- High stability
- Reduce kidney accumulation

## LINKER & CHELATOR

- Established DOTAM



## $^{212}\text{Pb}$ : ALPHA-EMITTING THERAPEUTIC ISOTOPE

- Proven clinical efficacy
- Fast & high energy deposition
- Safe profile
- Ideal waste management

## HALF-LIFE EXTENDER

- Half-life tuning
- Promote tumor uptake

# Orano Med – Pioneer of Targeted Alpha Therapy

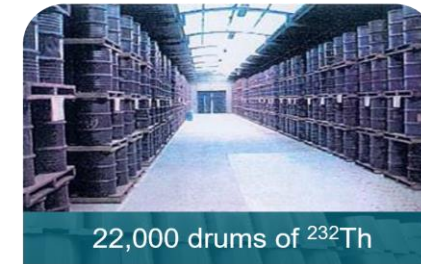


## Targeted Alpha Therapy with $^{212}\text{Pb}$ :

- **Alpha therapy:**
  - **High cytotoxicity:** DNA double-strand break, high energy deposition, no need for receptor internalization
  - **Targeted effect:** short range of action with potential to target microlesions, less impact on healthy tissues
- **Lead-212 ( $^{212}\text{Pb}$ ):**
  - **Short-half life** (~11h): out-patient administration, easy waste management, high-dose rate, flexible dosing
  - **Clean decay** with single alpha emission and effective chelating agent, limiting circulation of free daughter isotopes

## Orano Med as Pioneer of $^{212}\text{Pb}$ Targeted Alpha Therapy:

- Unique, independent **supply of  $^{212}\text{Pb}$**  as alpha emitting therapeutic isotope
- **Large scale GMP manufacturing** capabilities
- Fully integrated research and preclinical development platform
- **Clinical capabilities** demonstrated with  $^{212}\text{Pb}$  and lead program AlphaMedix™, outlicensed to Sanofi in Sept 2024, in Phase 2
- Strong collaboration partner with MP since early 2023

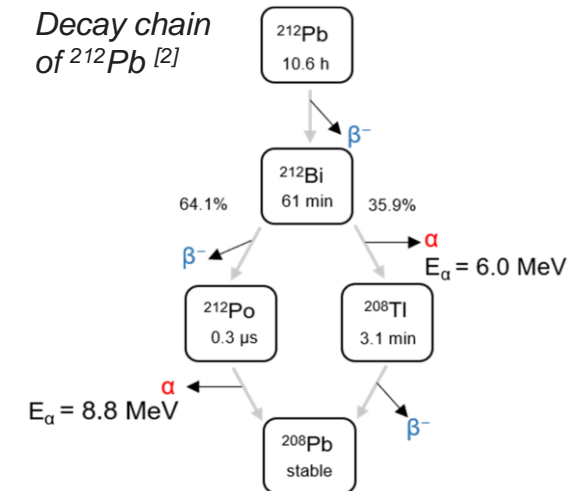


22,000 drums of  $^{232}\text{Th}$

Lead-212 is obtained **chemically** by **successive extractions and purifications** of the descendants of thorium-232



Decay chain of  $^{212}\text{Pb}$  [2]



# Global Partnership to Develop $^{212}\text{Pb}$ Radio-DARPin Therapeutics

Combining DARPin versatility with the power of  $^{212}\text{Pb}$  for next-gen Targeted Alpha Therapy



**MOLECULAR PARTNERS**  
PIONEERS of *DARPin THERAPEUTICS*



**ORANO MED**

PIONEERS of *TARGETED ALPHA THERAPY*



## FULL VALUE CHAIN PARTNERSHIP

### World class technologies & capabilities combined



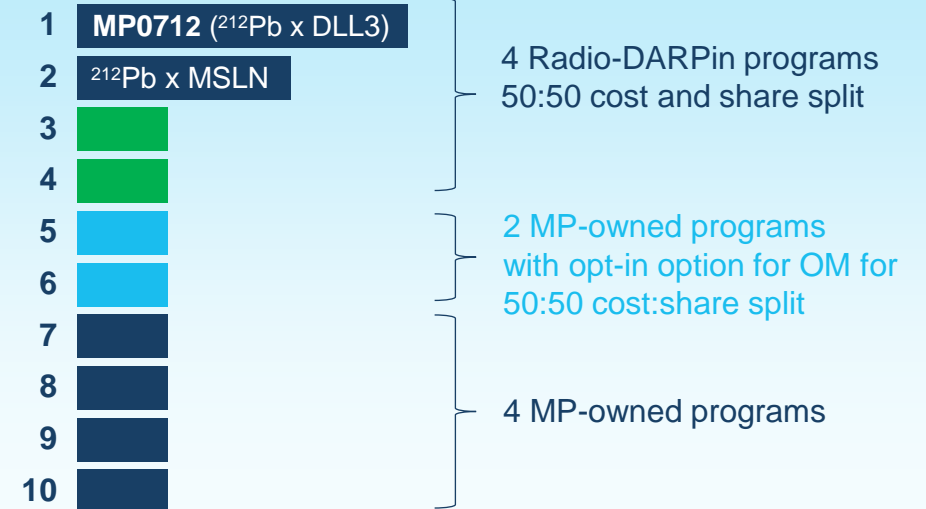
INDIANA, US:  
Industrial scale manufacturing  
Global shipping hub  
ATLab US

TEXAS, US:  
Preclinical development  
GMP supply for early  
clinical phases

SWITZERLAND:  
Preclinical assessment  
DARPin engine, fast &  
high throughput

FRANCE:  
 $^{212}\text{Pb}$  starting  
material  
ATLab Europe

### Pipeline of Ten $^{212}\text{Pb}$ Radiotherapy products



# MP0712, the first $^{212}\text{Pb}$ -DLL3 Targeted Radiotherapeutic for SCLC

## SCLC: critical unmet need, limited treatment options

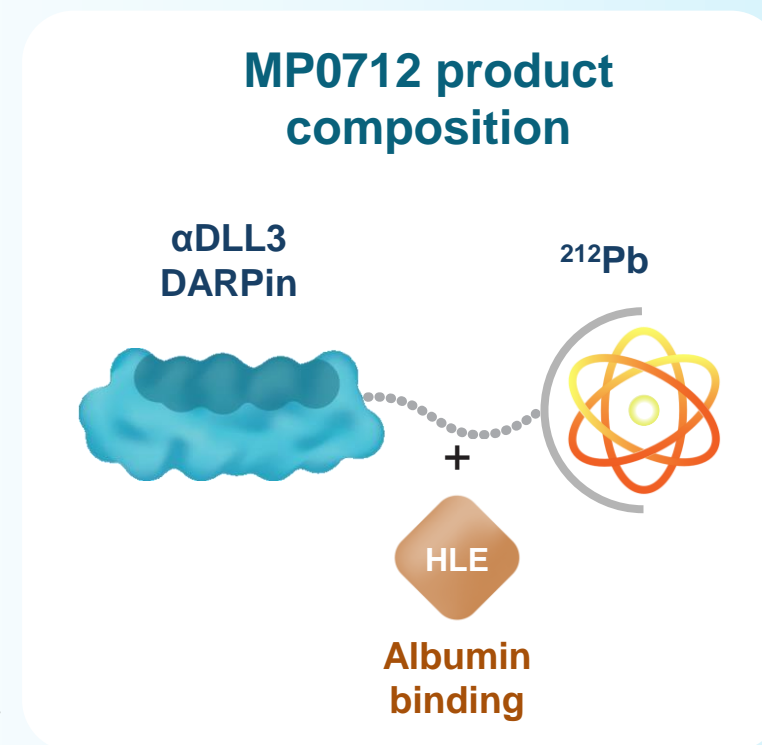
- Median progression free survival (mPFS) ~3 months<sup>1,2</sup>
- 5y overall survival (OS) ~3%<sup>1,2</sup>

## DLL3: a validated target for SCLC

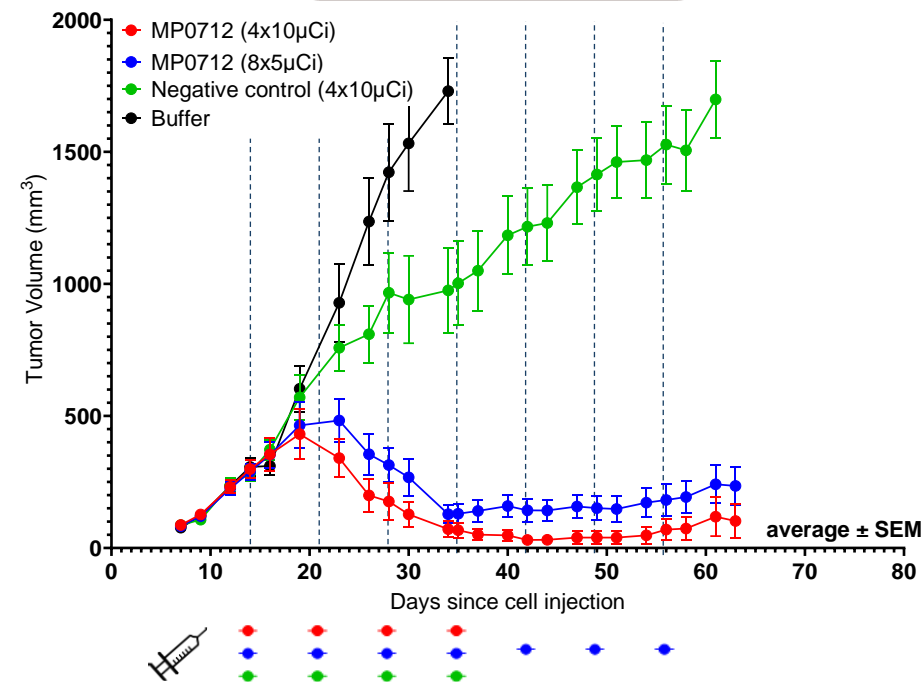
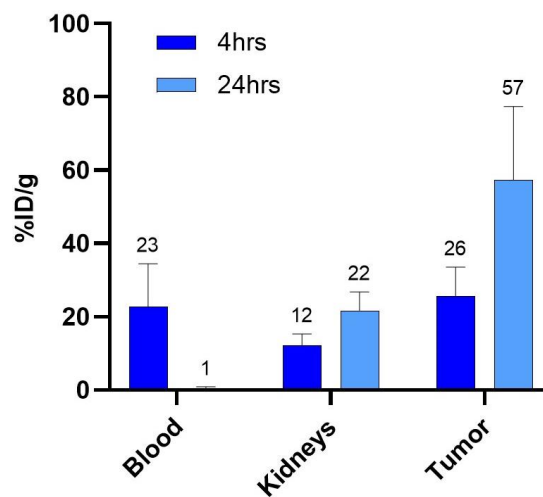
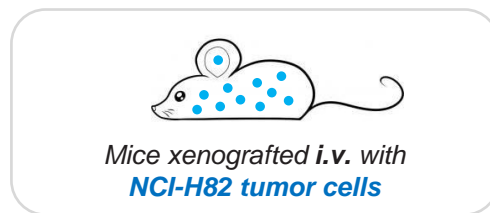
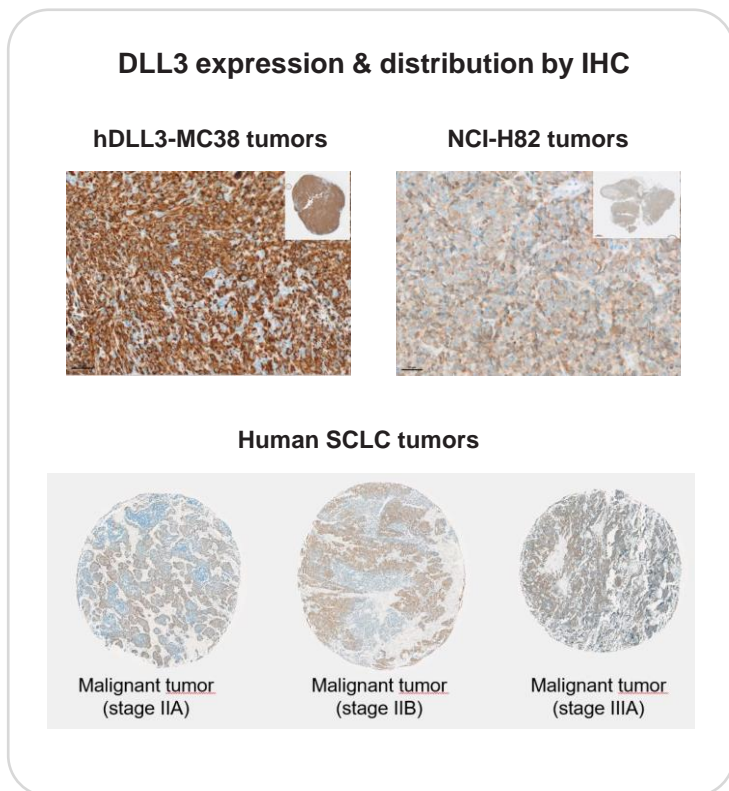
- Expressed in >85% of SCLC patients<sup>3</sup> and in neuroendocrine cancers
- No expression in healthy tissues
- Tarlatamab<sup>4</sup>, approved DLL3 targeting drug (T cell engager)
  - ORR ~40%, DOR 9.7 months, PFS 4.3 months

## MP0712: targeted delivery of alpha radiation with $^{212}\text{Pb}$

- DLL3 DARPin optimized for selective delivery of payload to tumor
- $^{212}\text{Pb}$  payload: high energy alpha emissions in short time frame, works with low target copy number (no need for internalization)
- Potential for combinations with immunotherapy



# MP0712: Potent Efficacy at Clinically-Relevant Dose



- MP0712 reached T:K ratios > 2 in mouse model matching clinically relevant DLL3 expression levels
- MP0712 induces complete and durable tumor regression in NCI-H82 tumor model at 10µCi injected every week

# Outline of MP0712 Clinical Development Strategy

- Patients: Focus on Small Cell Lung Cancer (SCLC), secondly on Neuro Endocrine Prostrate (NEPC)
- Biodistribution and dosimetry Phase 0 and Phase 1 studies to start in H2 2025, initial clinical data by YE

## IMAGING & DOSIMETRY



### Phase 0 – Imaging of MP0712 with $^{203}\text{Pb}$ (biodistribution/dosimetry)

Main objective: Imaging and Full Dosimetry to support dose strategy for  $^{212}\text{Pb}$

*N = 5–10 patients*

### Purpose:

→ Build confidence to reach relevant therapeutic level in tumor lesions

## THERAPY



### Phase 1/2a study (2 parts: part 1= phase 1; part 2=phase 2a)

**Part 1:  $^{212}\text{Pb}$  Dose Escalation**, Main objective: Safety, RP2D; *N = 15–20 patients*

**Part 2 – Dose Expansion and PoC – SCLC+NEC**; Main objective: Efficacy signals, confirm RP2D (*n=30*)

### Registration study

2L+ SCLC patients

### Phase 2s

- 1–2L combination with IO SCLC
- PoC in NEPC patients

# $^{212}\text{Pb}$ x MSLN Targeted Radio-DARPin for Ovarian Cancer

*Combining distinctive DARPin features with the power of  $^{212}\text{Pb}$  for next-gen targeted alpha therapy*

## Ovarian Cancer (OC): high medical need and marginal progress

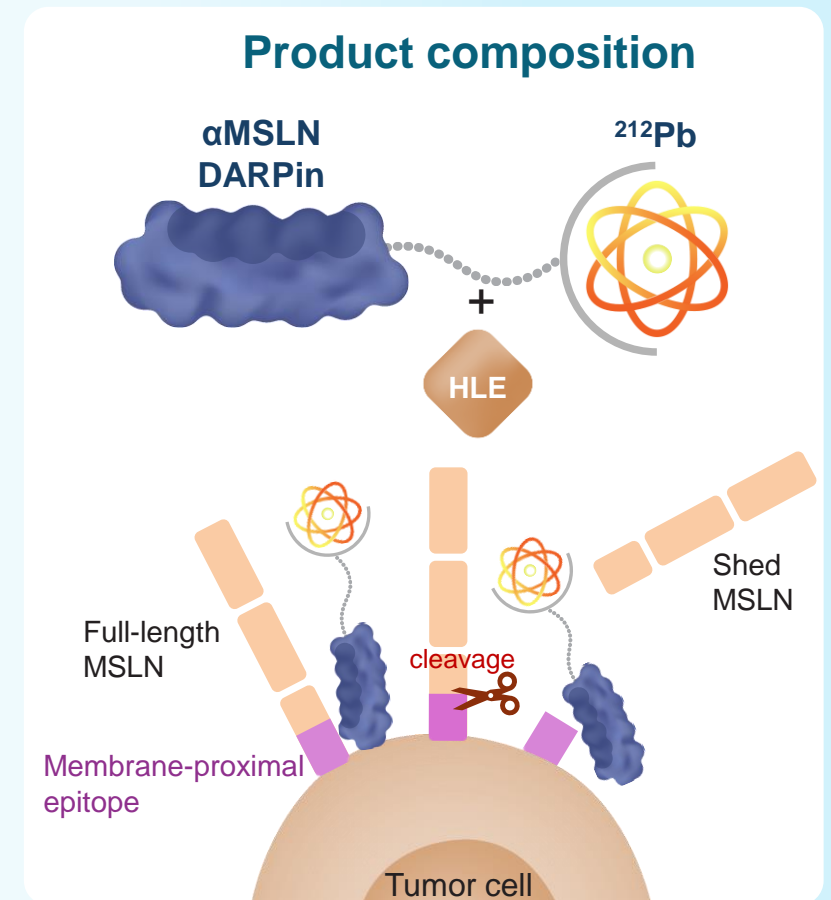
- > 50% patients die within 5y post-diagnosis (diagnosis often in late stage)
- Poor treatment options: ~80% recurrence rate post 1L chemo, limited 2L options (FR-alpha targeted Tx relevant for only 40% patients)

## Mesothelin (MSLN): a promising target for OC as 1<sup>st</sup> indication

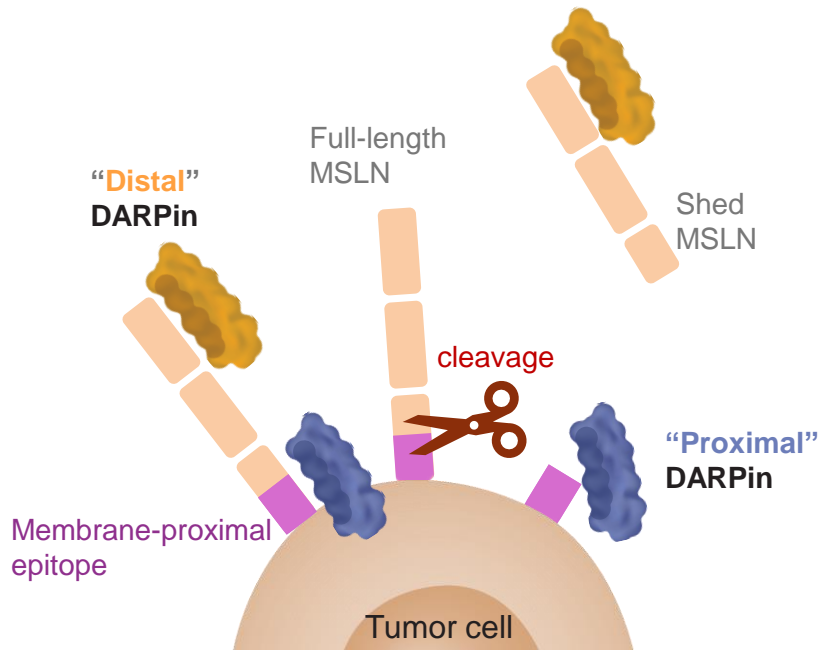
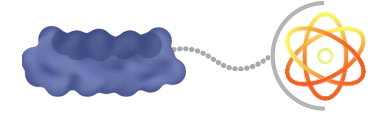
- Highly expressed in OC (>80% prevalence), expression maintained in metastases
- Shed MSLN detected in serum of OC patients, might limit efficacy of MSLN-targeted therapies<sup>1,2,3,4</sup> (e.g., CAR T/NK, ADC, TCE in development)

## RDT x MSLN: targeted delivery of alpha radiation with $^{212}\text{Pb}$

- MSLN DARPin targets **membrane-proximal epitope** (and not shed MSLN)
- $^{212}\text{Pb}$  payload: high energy alpha emissions in short time frame
- Potential for combinations with immunotherapy (incl. next-gen TCEs)

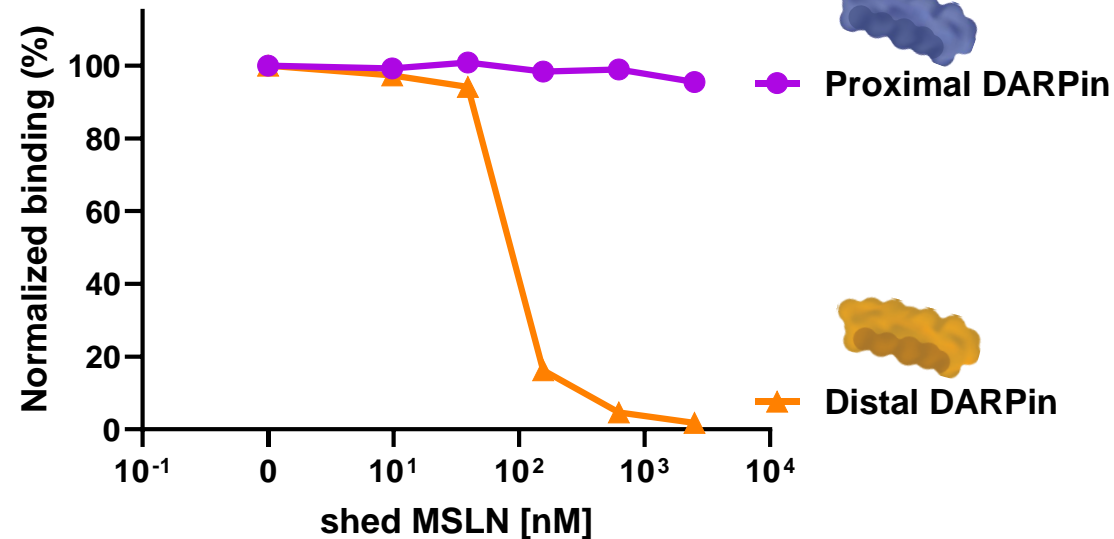


# DARPin activity is maintained despite shed MSLN



## OVCAR-8 Cell binding competition assay

100nM DARPin with increasing concentration of shed MSLN



**Binding maintained in presence of shed MSLN**

**Binding inhibited in presence of shed MSLN**

Preclinical update on MSLN at AACR 2025



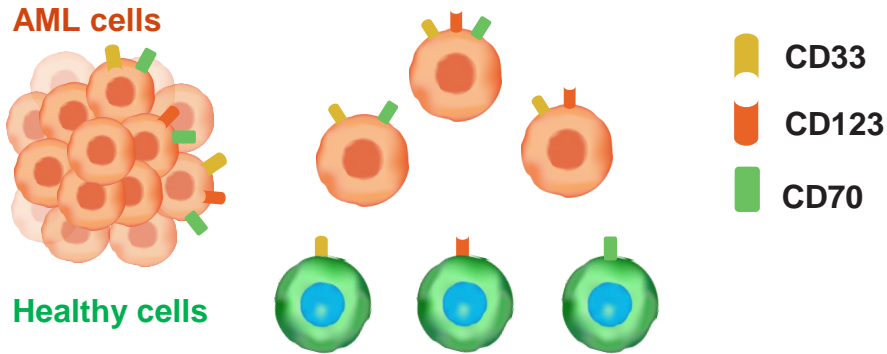
# MP0533

Tetra-specific T-cell Engager for AML



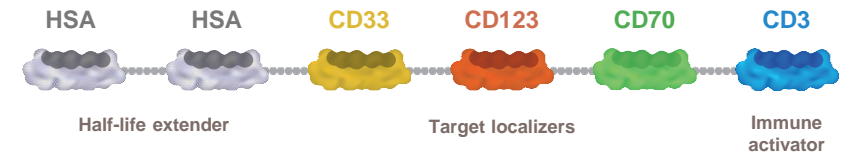
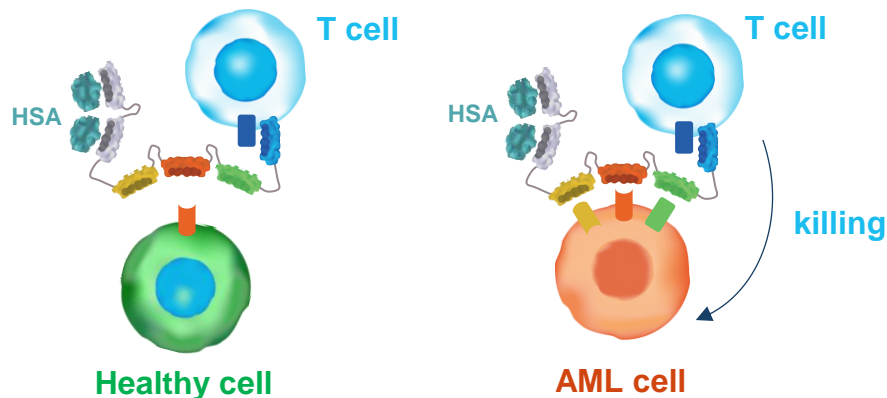
# MP0533 Enables Avidity-Driven Selective Killing of AML Cells

AML-associated antigens are also expressed on healthy cells



- AML bears a high risk of relapse due to persistent LSCs
- AML cell population is heterogeneous → differentiation from healthy cells (e.g., HSCs) feasible through their co-expression of CD33, CD123, CD70

MP0533: avidity-driven selectivity and T cell-mediated killing



- MP0533 designed to induce **T cell-mediated killing** preferentially when **2 or 3 AML-associated antigens** are co-expressed
- Potential to **kill all AML cells (blasts and LSCs)** despite heterogeneity, ensuring long-term disease control

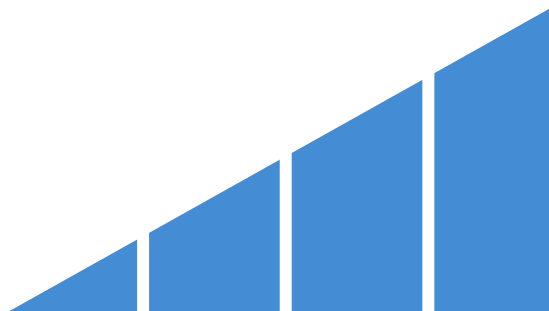
# MP0533 Phase 1/2a Study in Patients with R/R AML/MDS

*Protocol amendment to optimize MP0533 exposure*

## Initial Protocol

### Dose Escalation (DR 1–7)

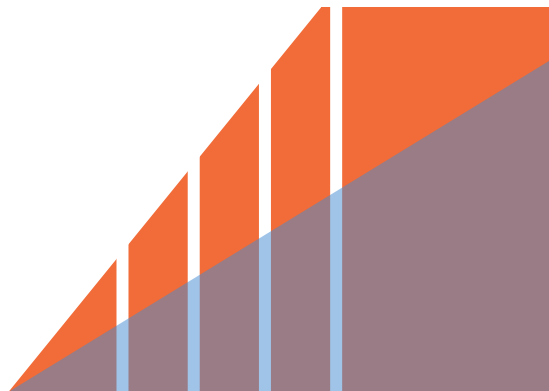
- Limited clinical activity
- Target-mediated drug disposition (TMDD) = **low exposure**
- Loss of exposure (LoE) in some patients (ADA)



## Amended Dosing Scheme

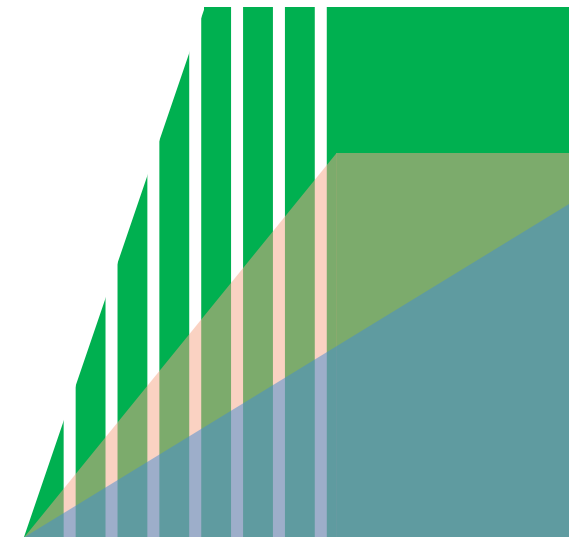
### Intermediate densification (DR 8)

- **Additional Day 12 dose** allows **steeper & faster dose escalation (step-up-dosing)**, addressing TMDD



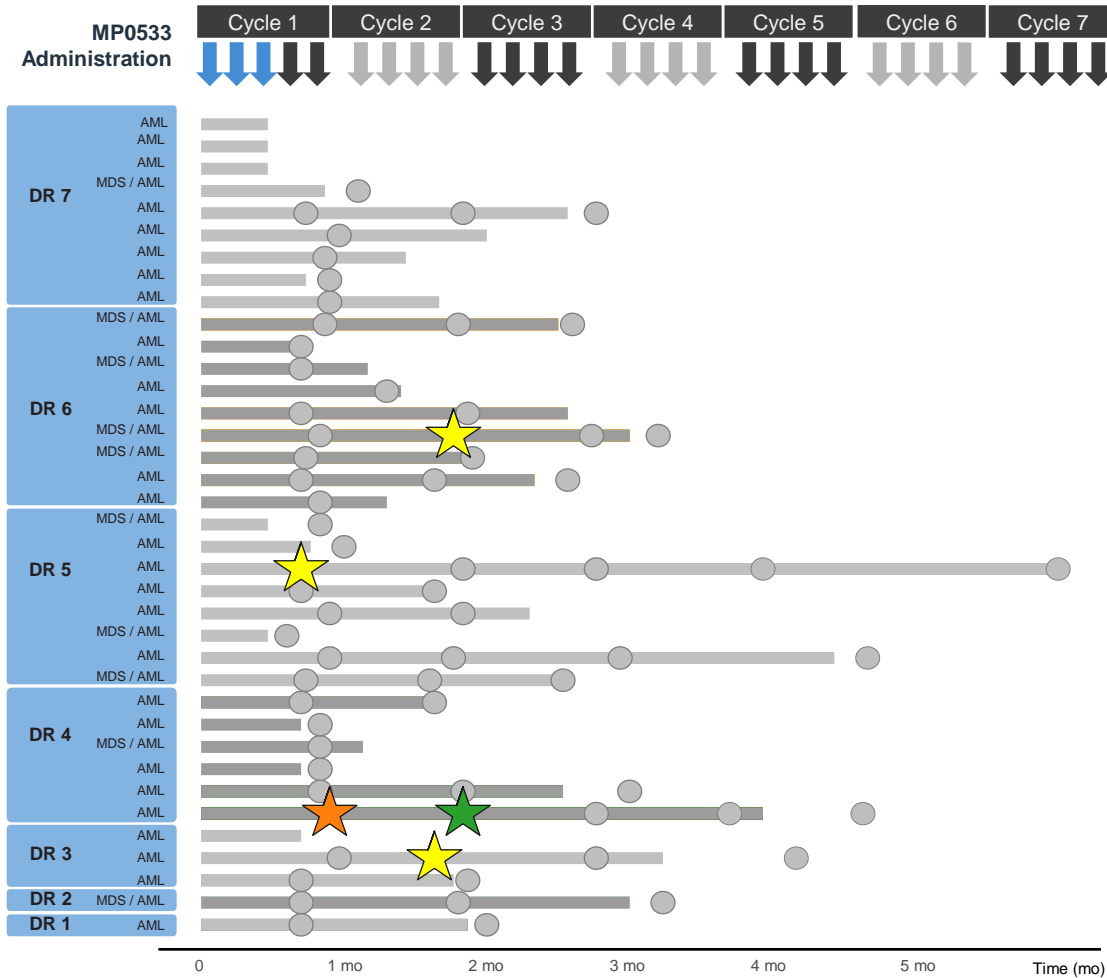
### Further dose densification (DR 9–10)

- **High dose frequency for 1<sup>st</sup> cycle**
- Premedication for **LoE mitigation**

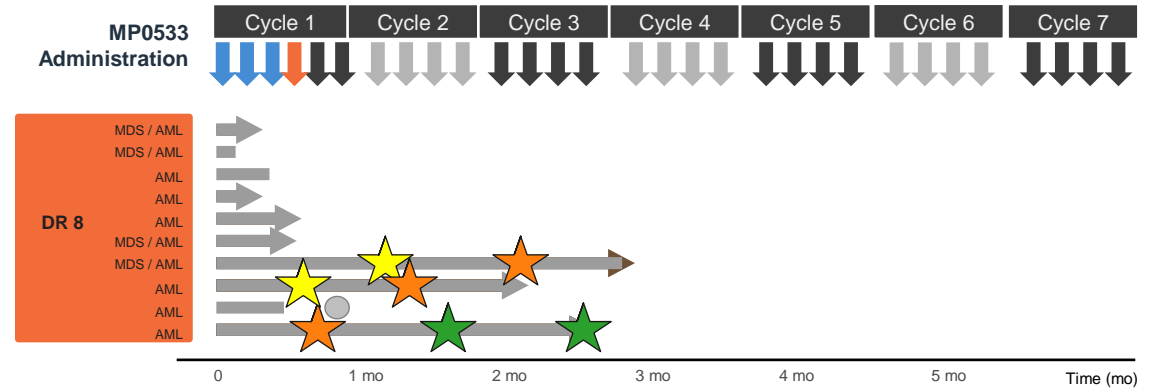


# MP0533 Treatment and Clinical Response

## Dose Escalation (DR 1–7)



## Intermediate densification (DR 8)



**DR 1–7:** 4 responders reported, manageable safety

**DR 8:** 3 responders and manageable safety reported to-date, evaluation on-going

**DR 9+** (further dose densification): update in 2025

### Legend

Response (2022 ELN<sup>1</sup>) was assessed every 4 weeks until disease progression and results are presented as:

★ CR    ★ CRi    ★ MLFS    ○ No ELN response

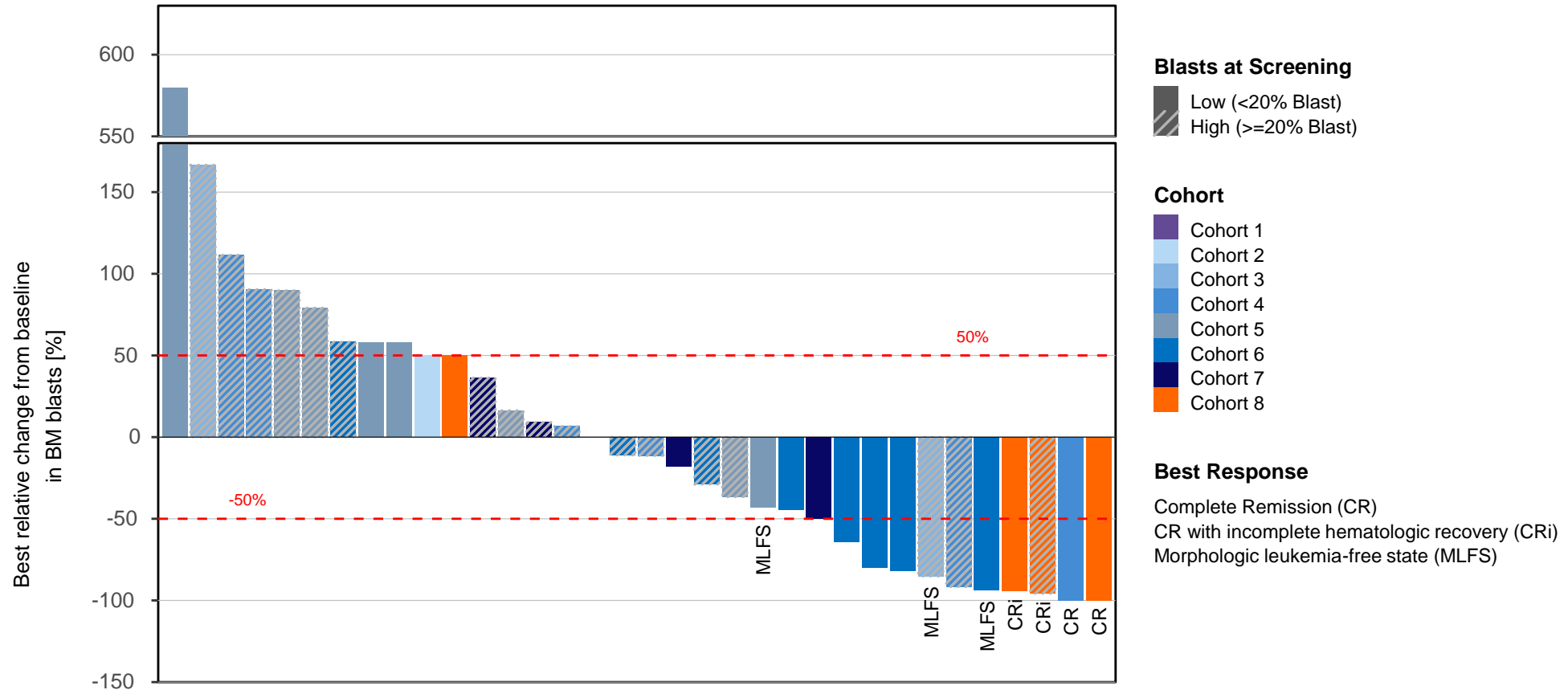
→ Patients with ongoing treatment at data cut-off    — Patients who discontinued treatment

Arrows at the top indicate MP0533 administration at D1, D5, D8, D12 (DR 8 only), D15 and weekly thereafter

↓ Step-up dosing at DR 1–7    ↓ D12 dose at DR 8

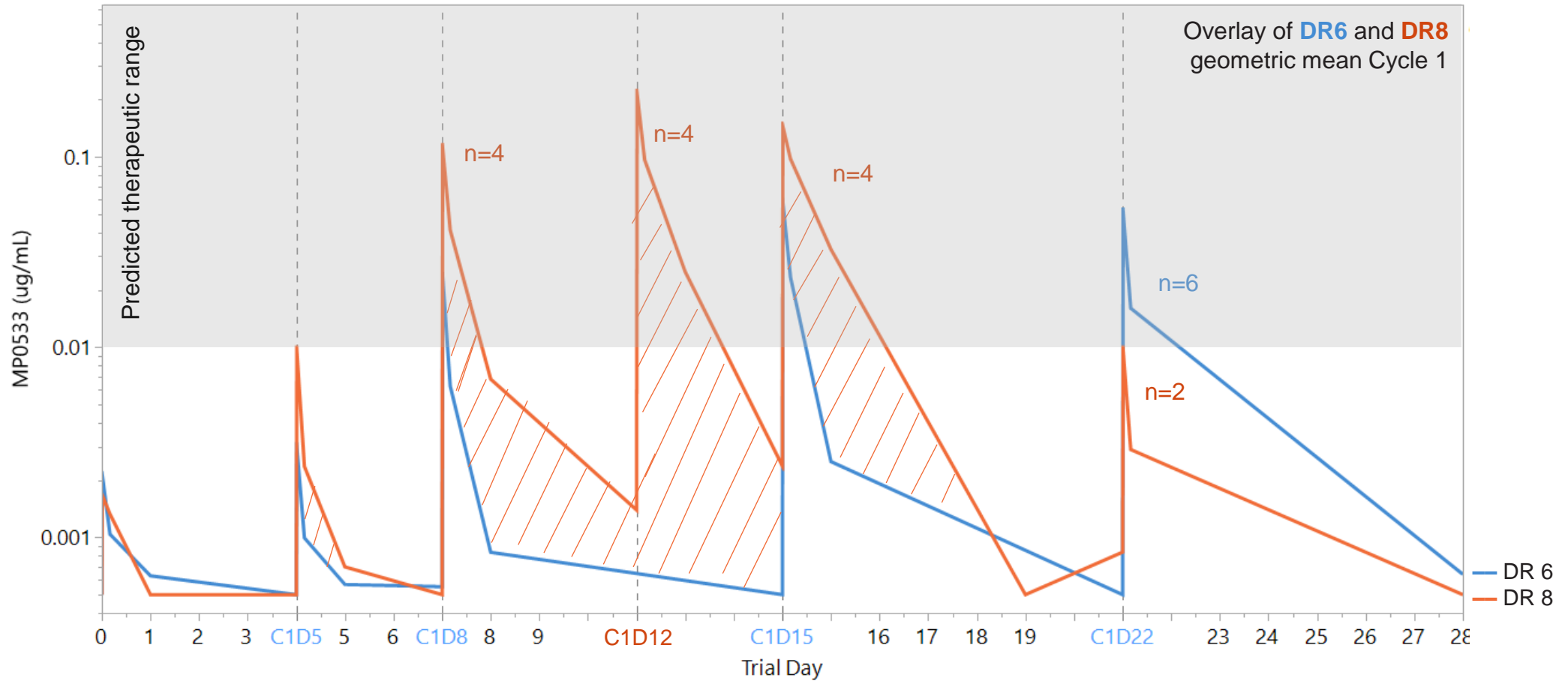
⇓ Color changes in grey: start of a new 28-day cycle

# Encouraging Blast Reduction Observed, Particularly in Patients with Lower Disease Burden at Baseline\*



➤ Improved response rate and depth observed to-date in Cohort 8

# Improved MP0533 exposure at DR 8 with steeper and denser step-up dosing regimen



# MP0533 Phase 1/2a Study in Patients with R/R AML/MDS

## *Protocol amendment to optimize MP0533 exposure*

### Objectives of protocol amendment:

- To improve the exposure profile of MP0533 in patients
- To increase the rate, depth and duration of clinical responses

### Hypotheses:

- Dose densification may overcome target-mediated drug disposition (TMDD), contributing to increasing response rate and depth
- Premedication with B cell depleting agent may mitigate loss of exposure (LoE), contributing to increasing duration and depth of responses

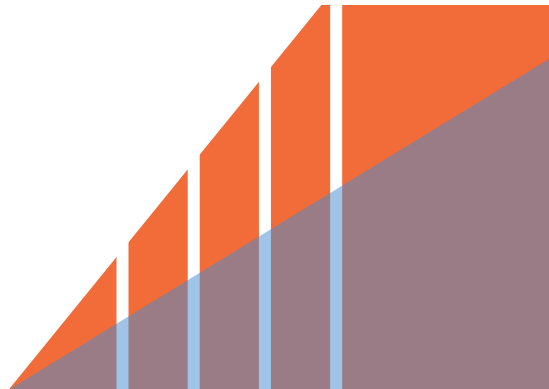
### Outlook:

- Full DR 8 clinical data in H1 2025
- Initial data from DR 9 in H2 2025

### Amended Dosing Scheme

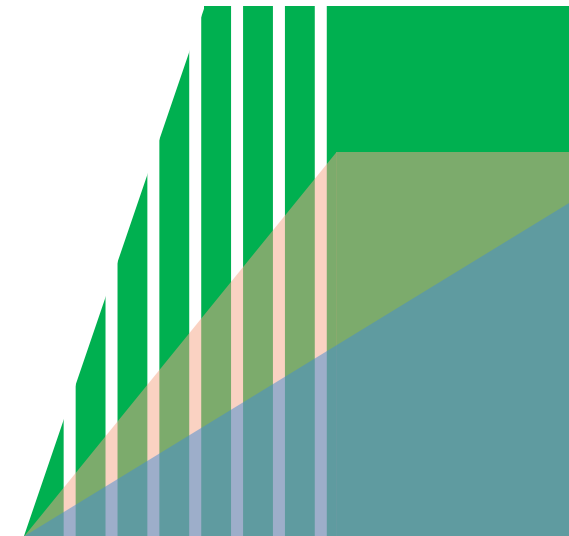
#### Intermediate densification (DR 8)

- **Additional Day 12 dose** allows **steeper & faster dose escalation (step-up-dosing)**, addressing TMDD



#### Further dose densification (DR 9–10)

- **High dose frequency for 1<sup>st</sup> cycle**
- Premedication for **LoE mitigation**



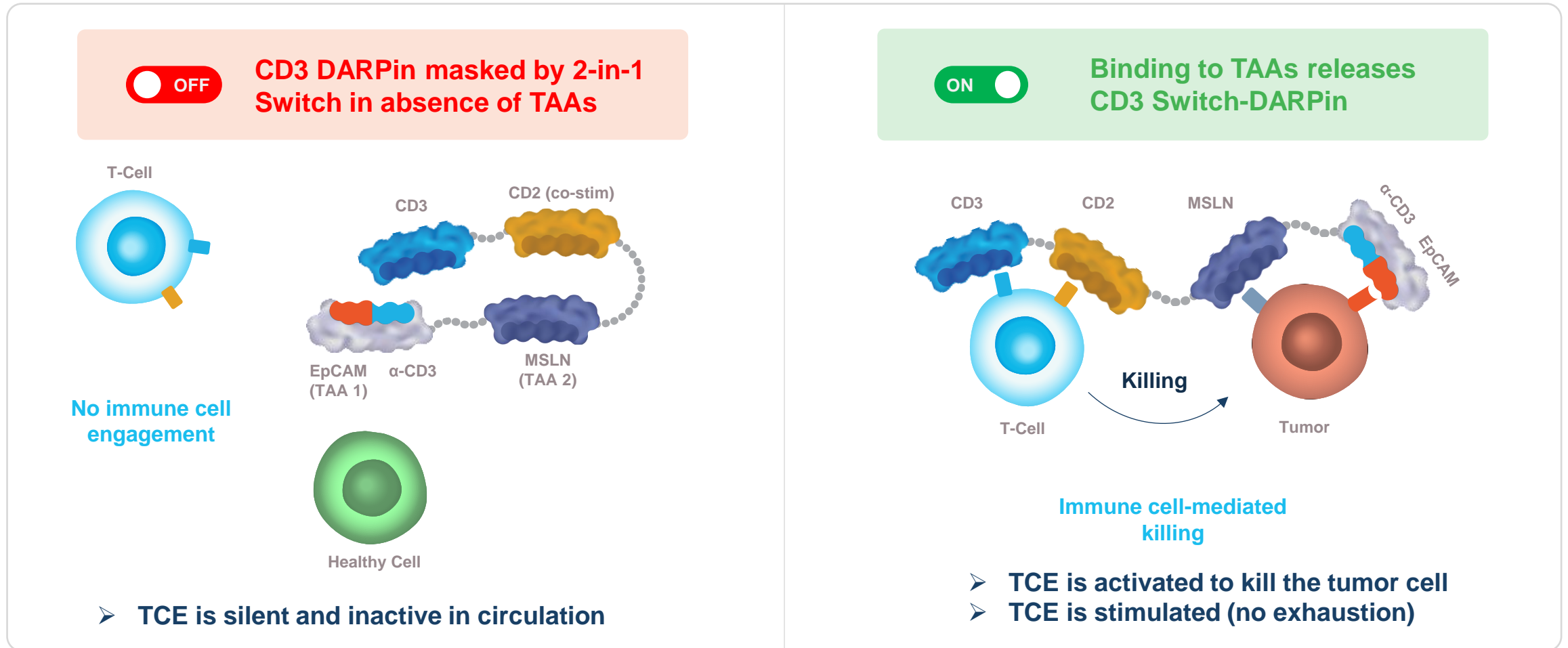


# Switch-DARPin Platform

Next-generation T cell engagers

# CD3 Switch-DARPin for Next-gen TCEs with Enhanced Function

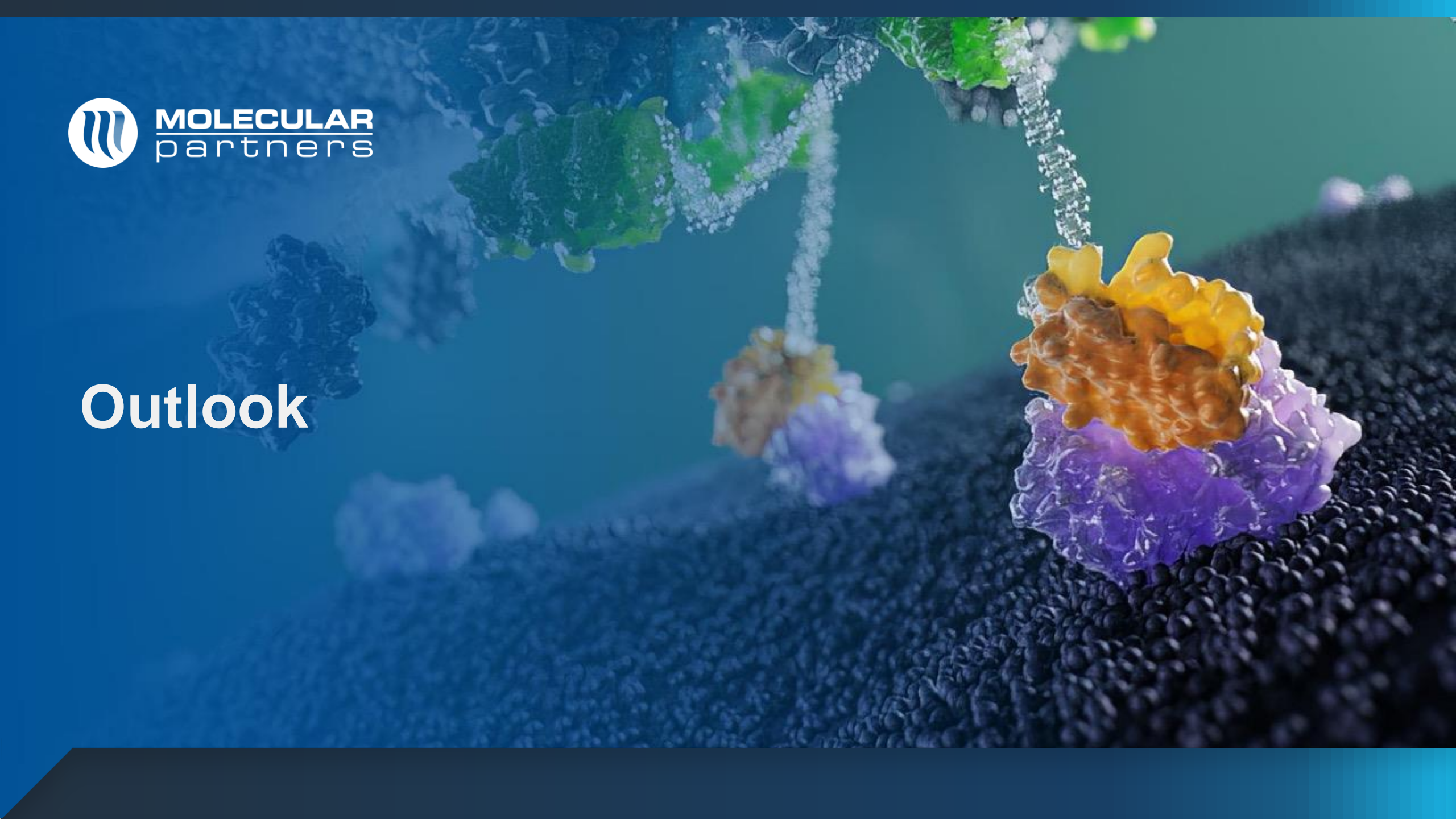
*Tackling current limitations of TCEs in solid tumors*



Preclinical update on CD3 Switch T cell engager at AACR 2025



# Outlook



# 2025 Outlook and Upcoming Milestones



## MP0712

- First-in-Human studies to start in 2025 (Phase 0 and Phase 1)
- Initial clinical data by end 2025 (imaging & dosimetry)
- Initial efficacy and safety data in H1 2026

## Radio-DARPin Therapy (RDT)

- MSLN preclinical update at AACR 2025, therapeutic candidate selection
- Additional  $^{212}\text{Pb}$  x RDT programs nominated, in collaboration with Orano Med

## MP0533

- Comprehensive clinical data from Phase 1 cohort 8 in H1 2025
- Protocol implementation of improved dosing regime, H1 2025
- Data from additional cohorts on amended dosing scheme in H2 2025

## Switch-DARPin

- Preclinical update on CD3 Switch T cell engager at AACR 2025
- Evaluation of partnering opportunities with Switch platform, including MP0621 (cKit) & T-cells

**CHF ~149 million cash\*** (incl. short-term time deposits) ensures **funding well into 2027**



*Twenty Years of Pioneering  
DARPin Therapeutics for Patients*

# Thank You





Back-up





# Financial Overview

# Balance Sheet

<i>(CHF million)</i>	FY 2024	FY 2023	FY 2022	FY 2021	FY 2020
<b>Non-current assets</b>	4.2	5.9	7.5	8.5	9.7
<b>Other current assets</b>	4.8	5.6	5.6	31.4	4.1
<b>Cash balance</b>	149.4 <sup>1</sup>	186.9	249.1	132.8	173.7
<b>Shareholders' equity</b>	141.6	176.4	235.2	107.3	107.2
<b>Non-current liabilities</b>	6.1	7.5	9.8	18.5	22.7
<b>Current liabilities</b>	10.8	14.4	17.3	46.9	57.7

<sup>1</sup> Includes CHF 85.6 million of short-term time deposits

Note: Rounding differences may occur

# Income Statement

<i>(CHF million)</i>	FY 2024	FY 2023	FY 2022	FY 2021	FY 2020
<b>Revenues / other income</b>	<b>5.0</b>	7.0	189.6	9.8	9.3
<b>R&amp;D expenses</b>	<b>(48.6)</b>	(48.8)	(50.7)	(55.7)	(56.1)
<b>SG&amp;A expenses</b>	<b>(17.6)</b>	(19.4)	(22.3)	(17.5)	(11.6)
<b>Operating result</b>	<b>(66.2)</b>	(61.1)	116.6	(63.4)	(58.3)
<b>Net financial result</b>	<b>7.2</b>	(0.9)	1.2	(0.4)	(4.4)
<b>Net result</b>	<b>(54.0)</b>	(62.0)	117.8	(63.8)	(62.8)

Note: Rounding differences may occur

# Cash Flow Statement

<i>(CHF million)</i>	FY 2024	FY 2023	FY 2022	FY 2021	FY 2020
Net cash from / (used in) operations	<b>(59.2)</b>	(59.0)	118.6	(91.0)	(29.0)
Net cash from / (used in) investing <sup>1</sup>	<b>40.5</b>	44.6	(101.1)	(22.2)	(21.7)
Net cash from / (used in) financing <sup>2</sup>	<b>14.4</b>	(1.2)	(1.6)	50.6	113.2
Exchange gain / (loss) on cash	<b>0.9</b>	(5.1)	0.3	0.7	(4.5)
Net cash increase / (decrease)	<b>(3.4)</b>	(20.6)	16.1	(61.9)	58.0
Cash balance at year end	<b>149.4</b>	186.9	249.1	132.8	173.7

<sup>1</sup> Includes movements in short-term time deposits

<sup>2</sup> For 2024 this includes the October 2024 capital raise, for 2021 the funds received from the NASDAQ listing; for 2020 this includes two capital raises

Note: Rounding differences may occur