



Corporate Presentation

November 2023

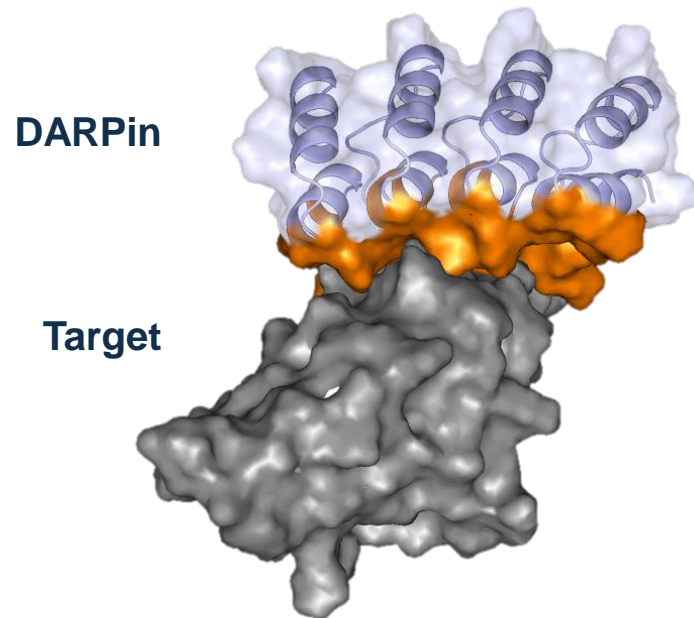


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DARPin Modality and Molecular Partners' Strategy



What we invented

- New class of therapeutics: Designed Ankyrin Repeat Proteins (**DARPins**)
- DARPins to **close the gap between small molecules and antibodies**
- 7 clinical-stage compounds, **>2500 patients treated**

How we apply it

- **Unique DARPin solution** for a defined medical problem not addressable by antibody designs
- Demonstrate **true patient value** with **early clinical read out**
- Combine our **capabilities with world-class partners** to deliver innovative therapeutics

Highlights Q3 2023

MP0533

- Novel tetra-specific T cell engager for R/R AML and high-risk MDS
- Phase 1 dose-escalation study well on track, **currently enrolling at dose regimen (DR) 5**
- **Favorable safety profile** in DR1-3, **first responder in DR3**: additional data (up to DR4) at **ASH 2023**

MP0317

- Bi-specific targeting FAP and CD40 for tumor-localized immune activation
- Phase 1 study in R/R solid tumors, dose escalation fully enrolled
- **Favorable safety profile up to highest dose, tumor-localized CD40 activation leading to remodeling of tumor microenvironment** in patients presented at **SITC 2023**

Radio-DARPin Therapy Platform

- RDT platform successfully optimized to **reduce kidney accumulation & increase tumor uptake**, progress presented at EANM 2023
- Selected tumor-associated protein DLL3 as a first in-house target
- Novartis collaboration further progressing

Operations

- Strong financial position with CHF ~207 M in cash (incl. short term deposits) as of Sept. 30, 2023
- **Capitalized well into 2026**

Pipeline

— Oncology

— Radio-DARPin Therapy

— Virology¹

— Ophthalmology²

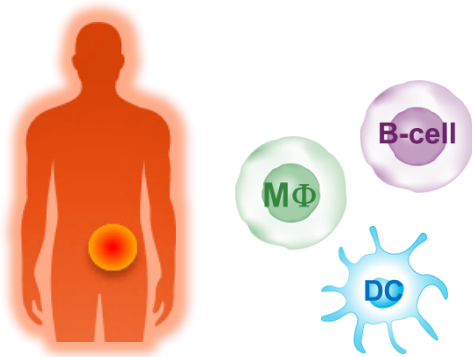
CANDIDATE	RESEARCH	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	RIGHTS
MP0317 FAP x CD40	Solid Tumors					MOLECULAR partners
MP0533 CD33+CD70+CD123 x CD3	AML					MOLECULAR partners
Immune Cell Engagers						MOLECULAR partners
Radio-DARPin Therapy Platform	DLL3 and 2 nd target ongoing	In-house programs				MOLECULAR partners
	Solid Tumors	Partnered programs				NOVARTIS
Virology						MOLECULAR partners
Abicipar VEGF	Wet AMD					MOLECULAR partners

MP0317

Tumor-localized Immunotherapy

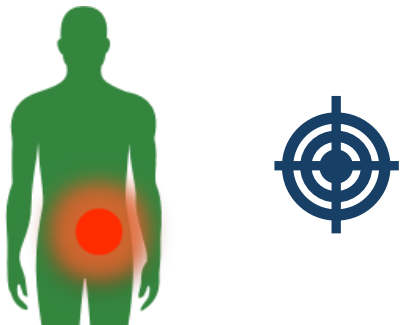
MP0317: Unlocking CD40 Activity by Local Activation

Problem: Toxicity of CD40 Antibodies Has So Far Limited Their Activity

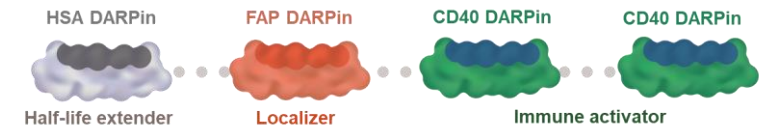


- **CD40 agonists** can activate **B cells, DCs and MΦ** to enhance the efficacy of IO drugs, especially in “cold tumors”
- **Systemic activation of CD40 via mAbs** has been hampered by **significant toxicities**, therefore **limiting their potential of reaching a therapeutically active dose**

Solution: MP0317 – FAP-dependent tumor-localized CD40 activation

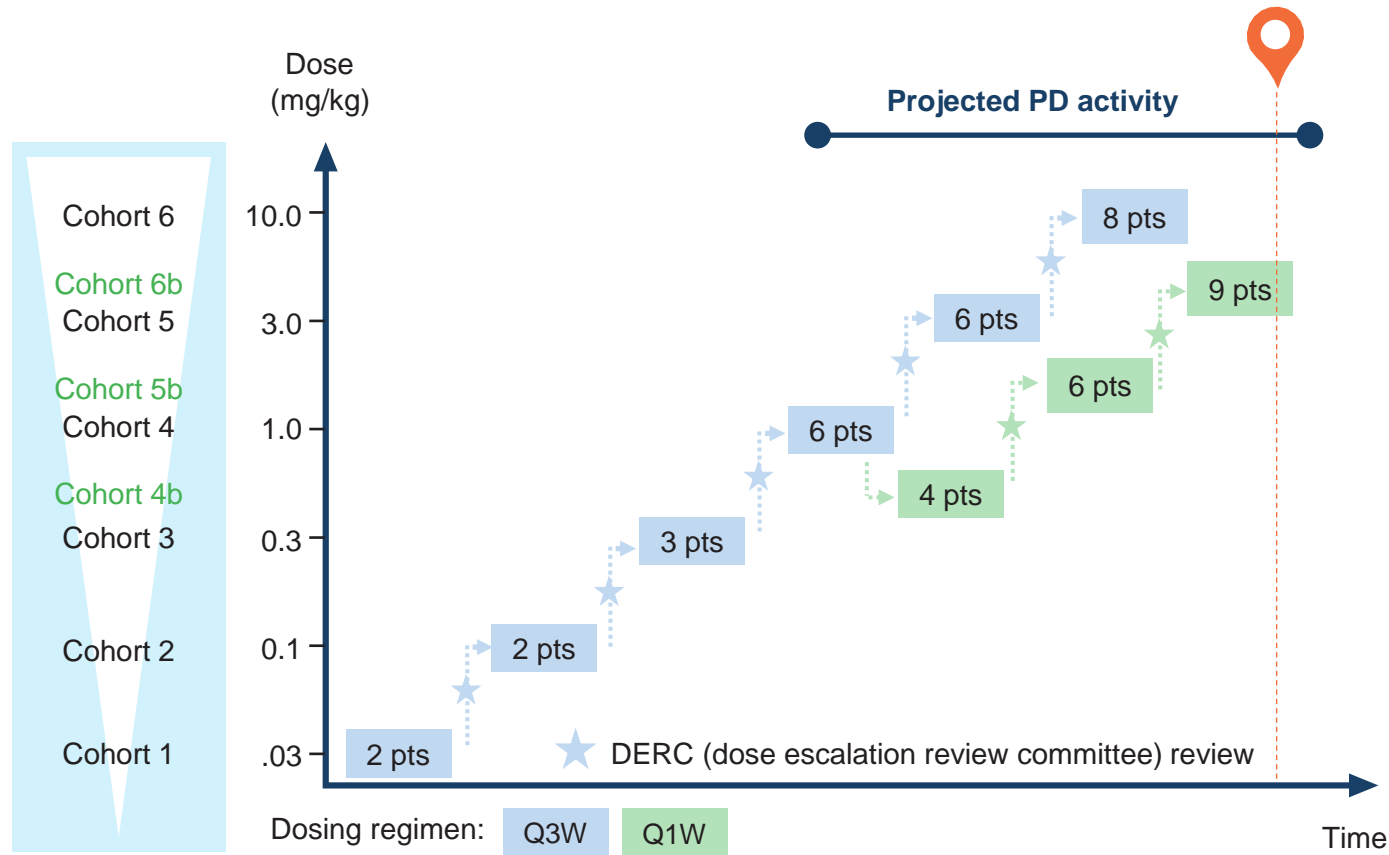


- **FAP is a validated tumor target** overexpressed in at least 28 different cancer types and its expression is not downregulated during disease progression
- **MP0317** is designed to bind tumor-localized FAP and induce CD40-mediated **activation of immune cells in the tumor**, thereby overcoming systemic toxicity and allowing a **wider therapeutic dosing range**



MP0317 Phase 1 Study Design & Status

First-in-human, multicenter, dose-escalation study in adults with advanced solid tumors



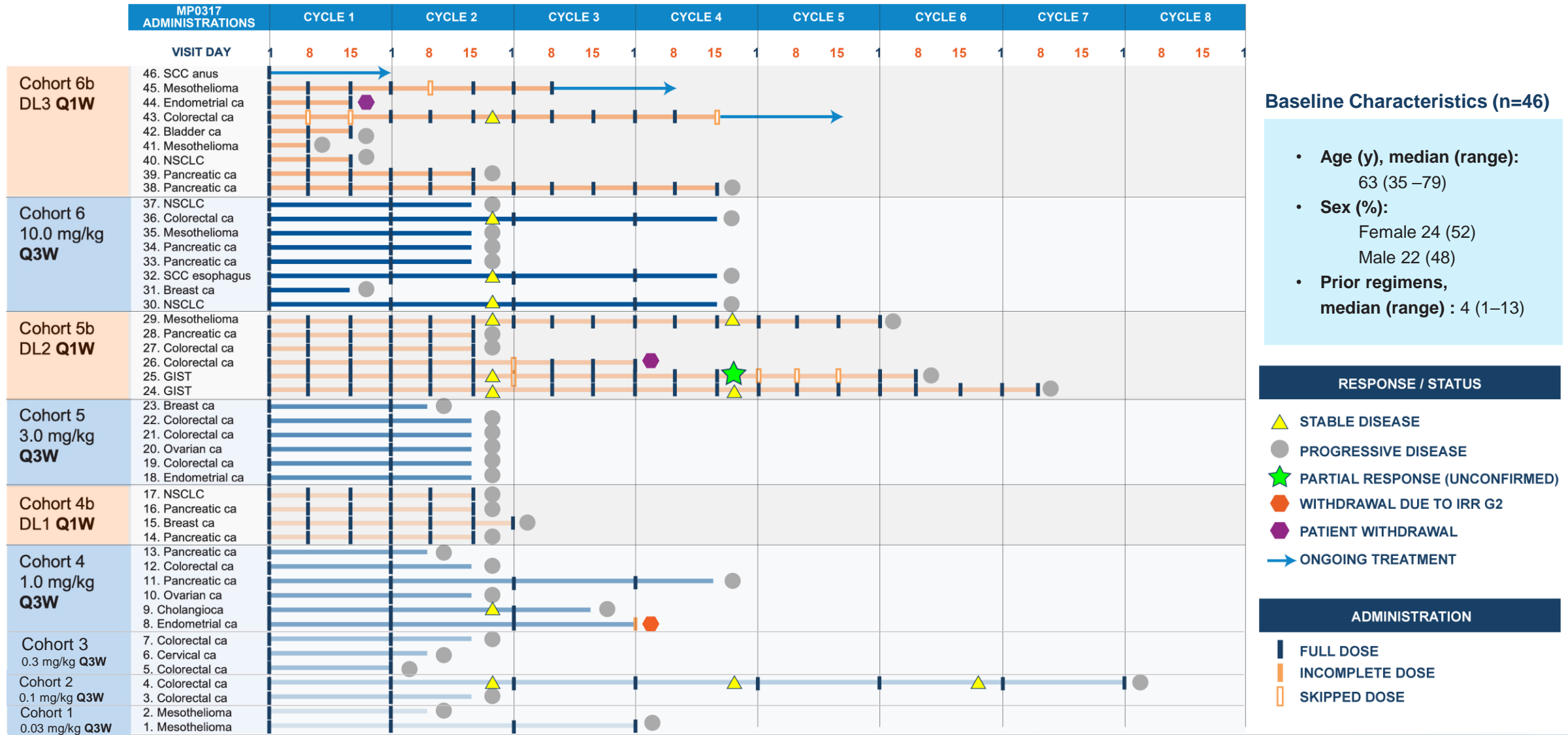
Primary Study Objectives

- MP0317 safety and tolerability
- Recommended dose for expansion and combination

Updated Data Presented at SITC 2023

- **Enrollment completed** at highest planned doses in dose escalation part
- **Favorable safety profile** up to highest planned dose; one DLT observed
- Tumor-localized CD40 pathway and immune cell activation, leading to **remodeling of TME**

MP0317 Study Status & Patient Characteristics



Baseline Characteristics (n=46)

- **Age (y), median (range):**
63 (35 –79)
- **Sex (%):**
Female 24 (52)
Male 22 (48)
- **Prior regimens, median (range) : 4 (1–13)**

RESPONSE / STATUS

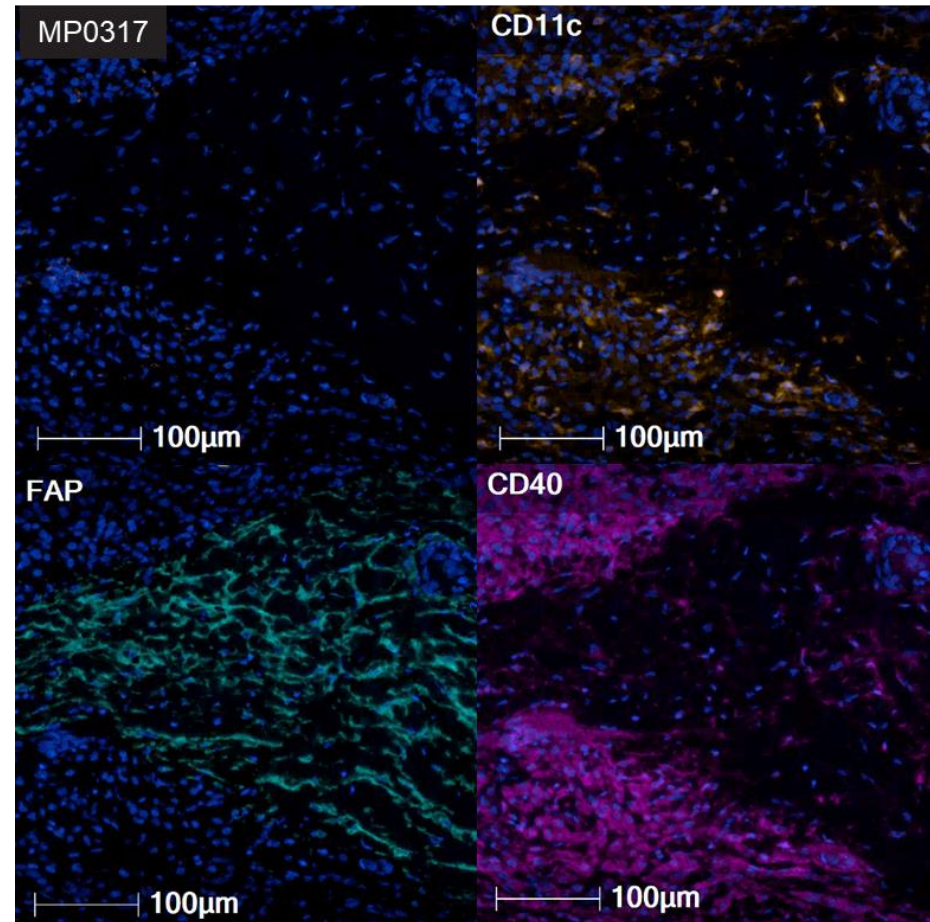
- ▲ STABLE DISEASE
- PROGRESSIVE DISEASE
- ★ PARTIAL RESPONSE (UNCONFIRMED)
- ⬢ WITHDRAWAL DUE TO IRR G2
- ◆ PATIENT WITHDRAWAL
- ONGOING TREATMENT

ADMINISTRATION

- ▮ FULL DOSE
- ▮ INCOMPLETE DOSE
- ▮ SKIPPED DOSE

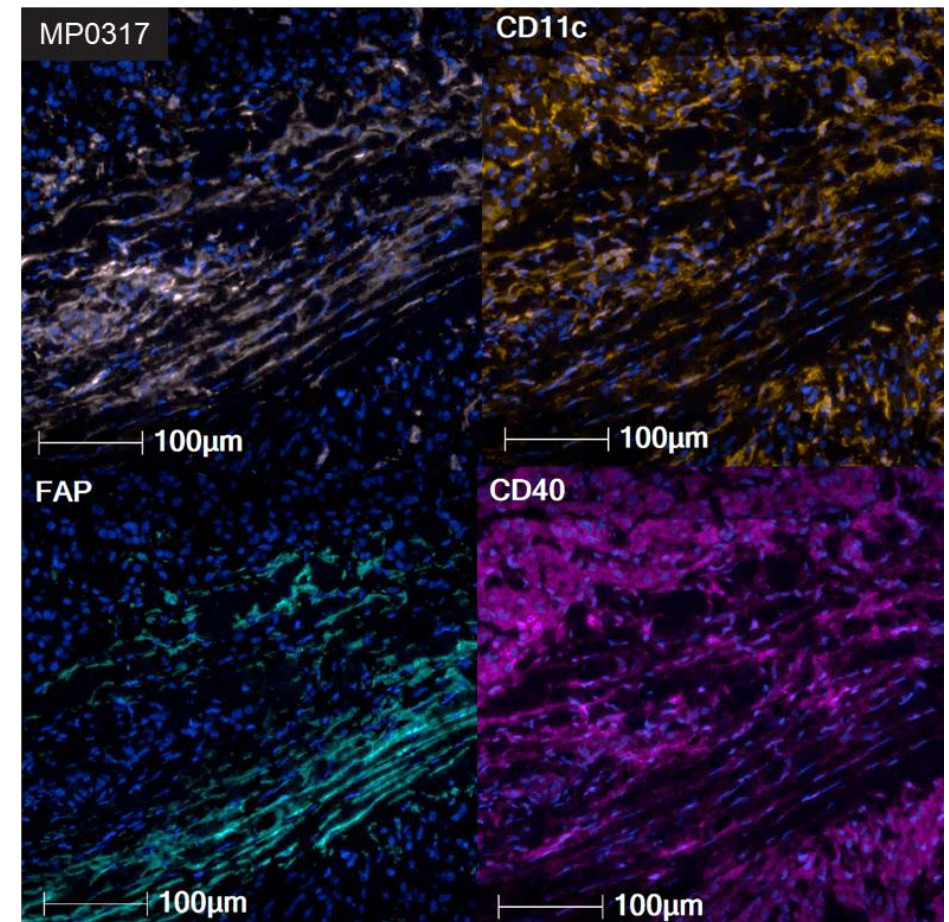
MP0317 co-localizes with FAP and CD40 in tumors – concomitant increase in intra-tumoral DCs observed

PRIOR TO TREATMENT



Minimal DC presence in FAP-positive tumor area

CYCLE 2 DAY 8



High DC infiltration in FAP-positive tumor area in MP0317 presence

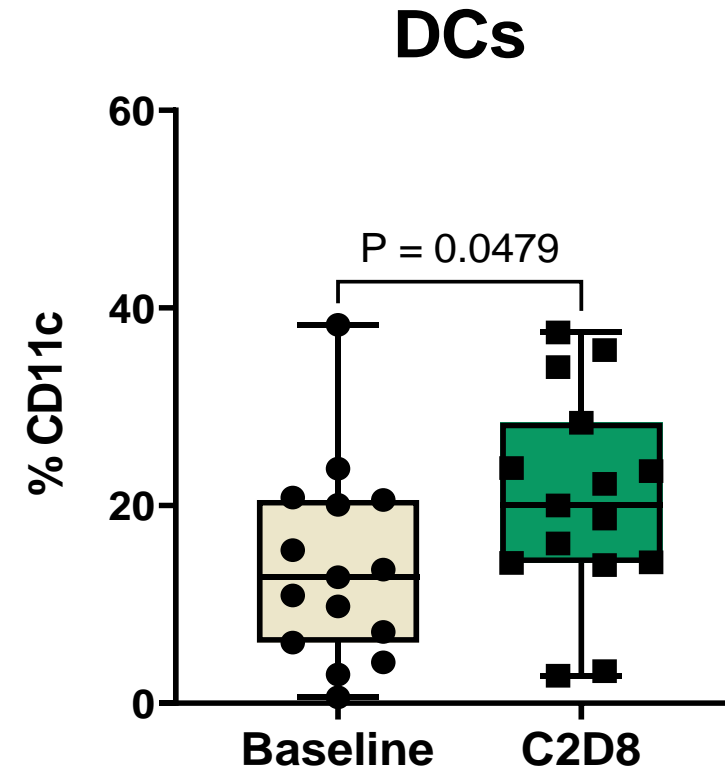
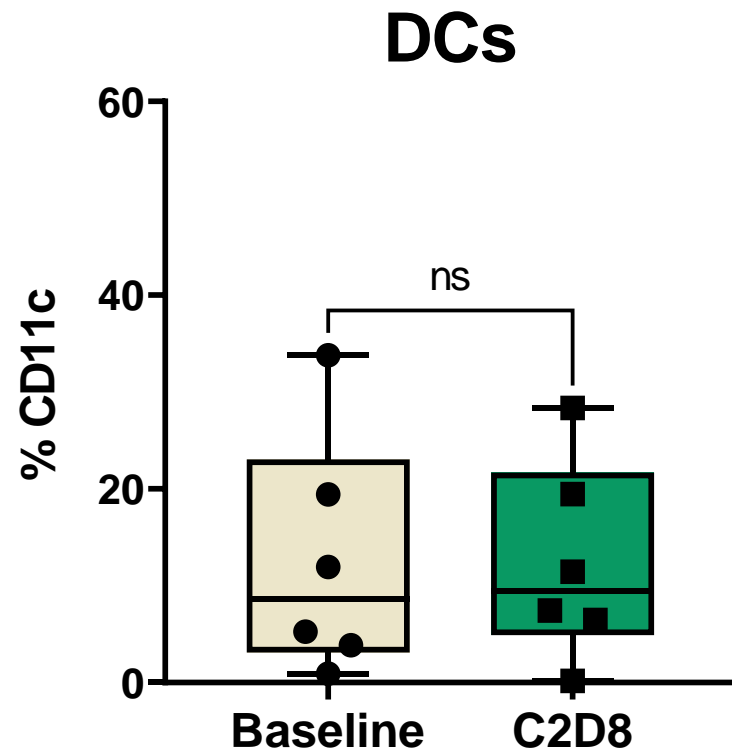
DC
infiltration

MP0317

Increase in intra-tumoral DCs observed post MP0317 treatment

MP0317 low* doses or not detected in tumor (n=6)

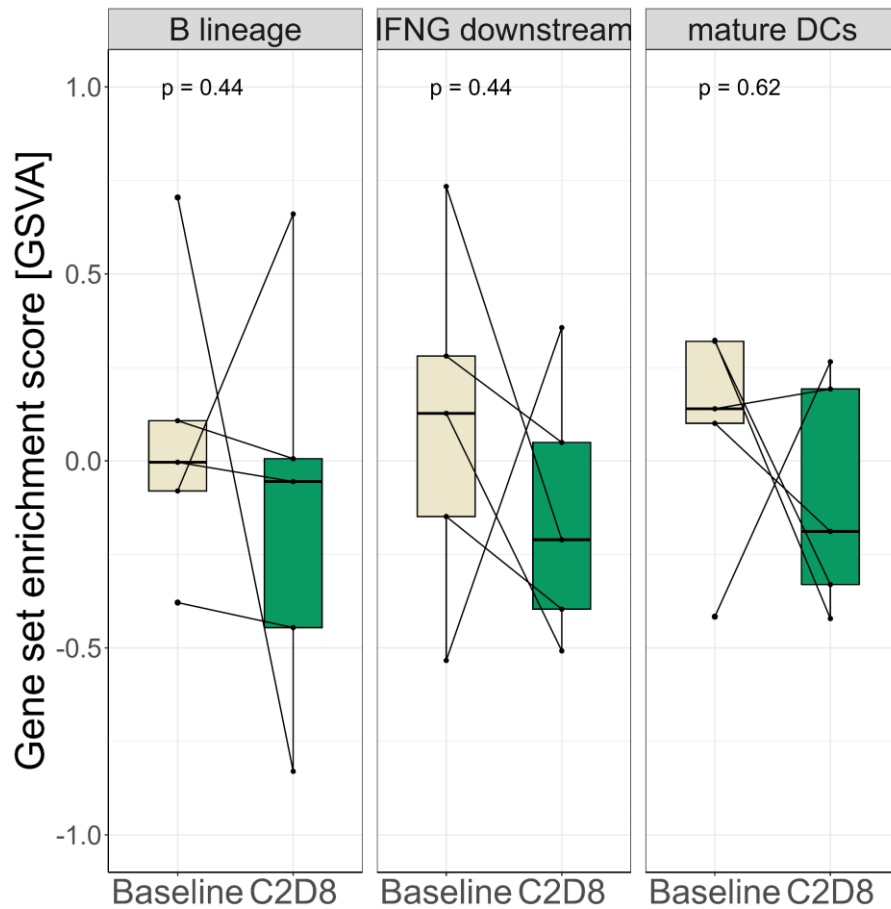
MP0317 higher** doses and detected in tumor (n=15)



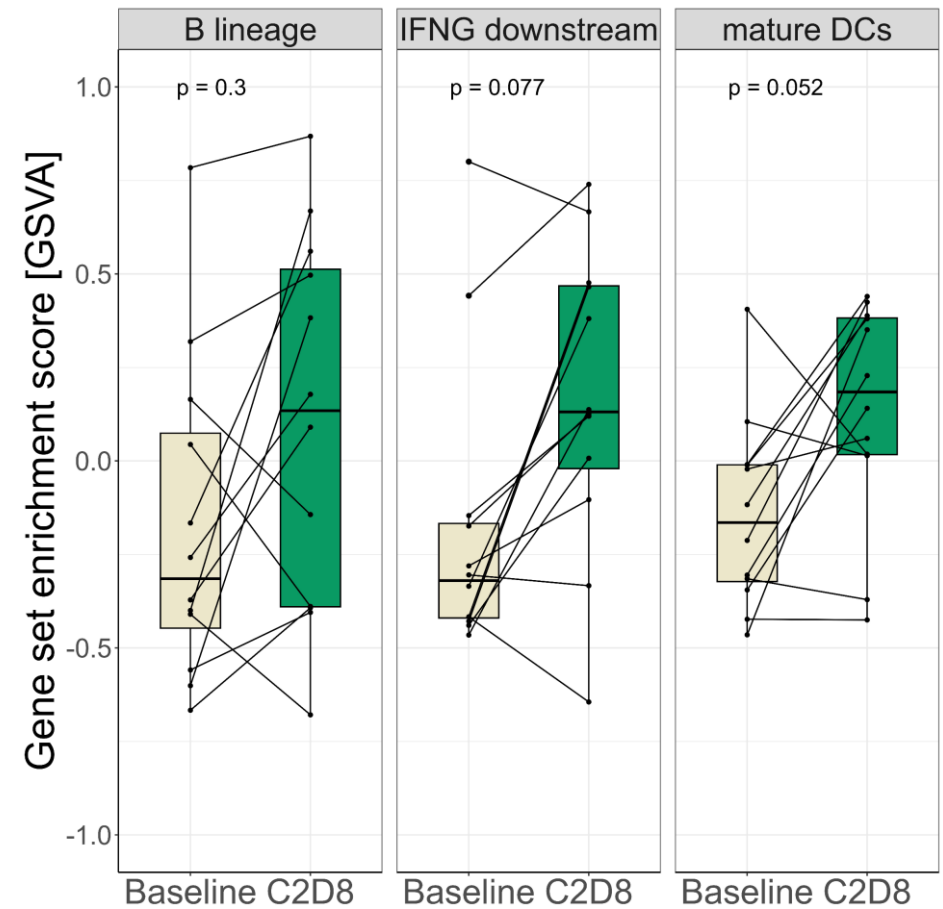
Treated patients up to Cohort 6 with evaluable paired biopsies for mIF (n=21). *Low doses = ≤ 0.1 mg/kg; **Higher doses = ≥ 0.3 mg/kg. Upper (75%), median, and lower (25%) percentiles are indicated. P-values are derived from paired ranked sum Wilcoxon test.

Increased immune cell infiltration, DC maturation and IFN γ production observed in tumors post MP0317 treatment

MP0317 low* doses or not detected in tumor (n=5)



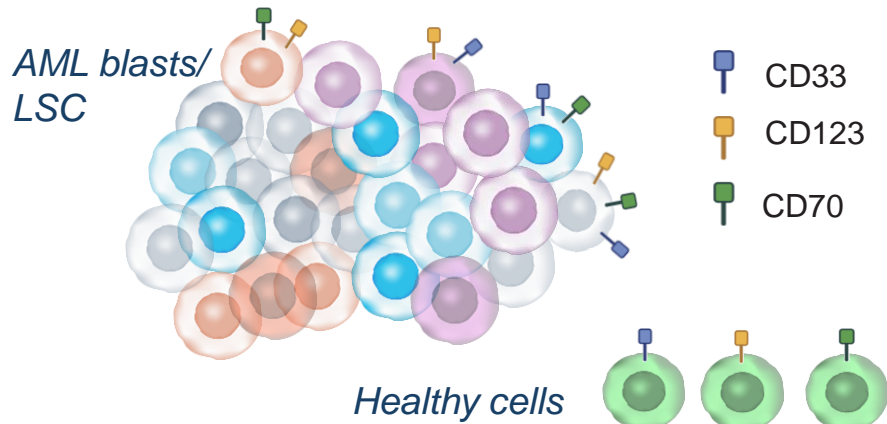
MP0317 higher** doses and detected in tumor (n=12)



MP0533 Tetra-specific T cell Engager for AML

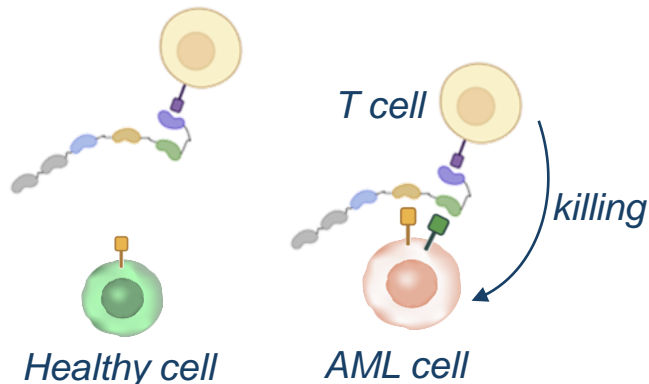
MP0533: Avidity-guided selectivity for cancer cells in AML

Problem: AML tumor-associated antigens are expressed on healthy cells

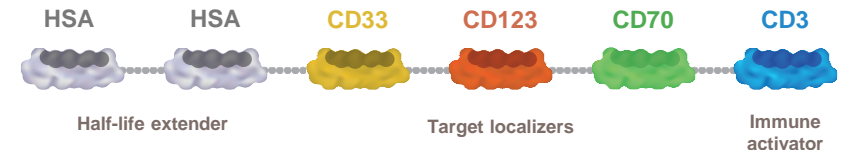


- **AML remains a deadly disease** and persistence of **leukemic stem cells (LSCs)** drives relapse
- **AML cell population is heterogeneous**: individual AML blasts and LSCs lack a clean target. AML cells can be differentiated from healthy cells by their **co-expression of specific targets** (e.g. CD33, CD123, CD70)

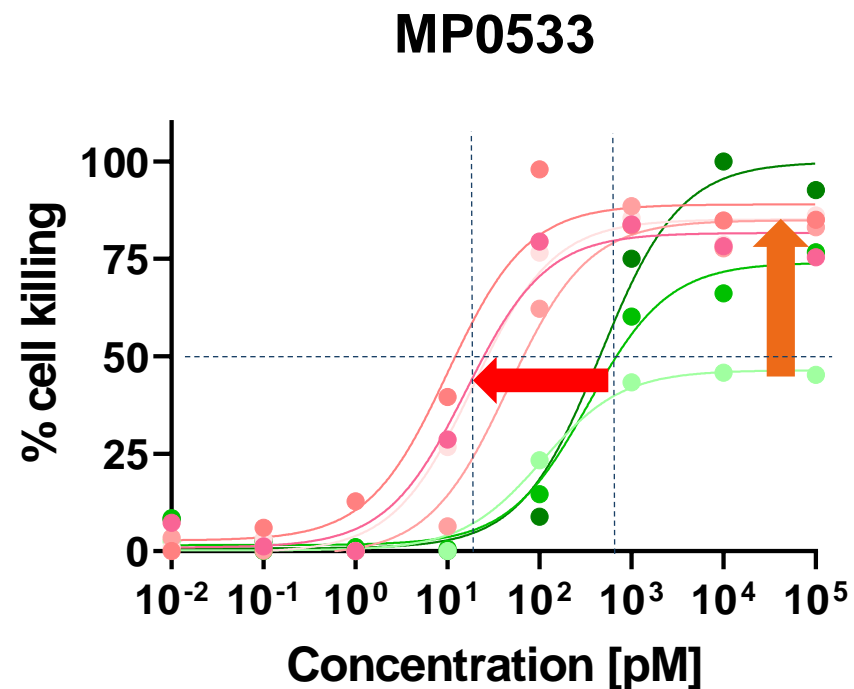
Solution: MP0533 – Avidity-driven selectivity & killing by T cells



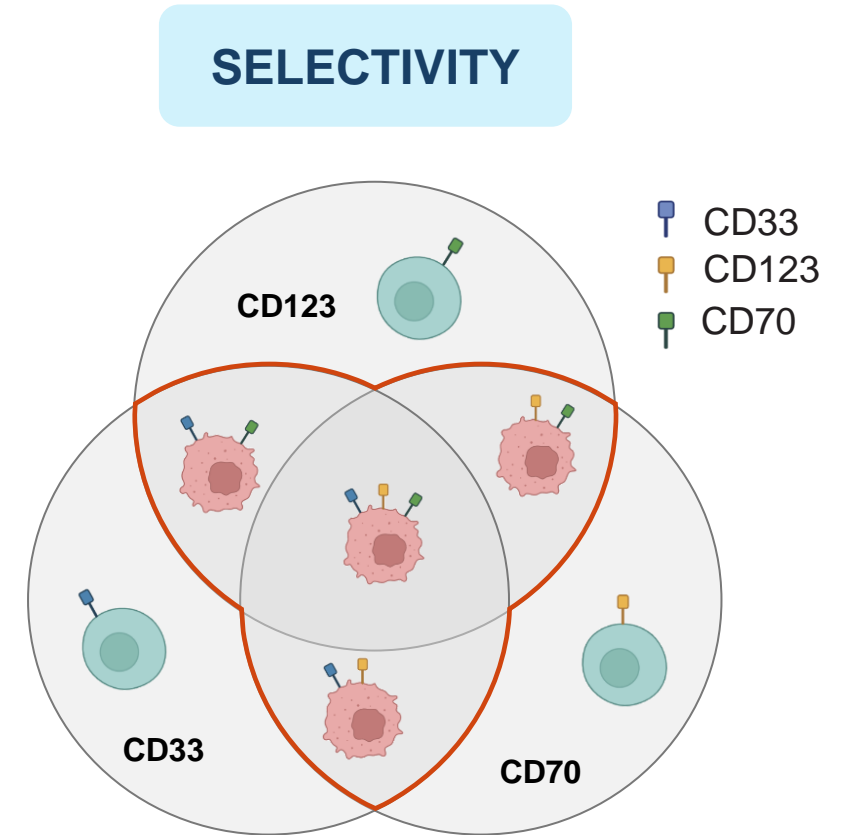
- MP0533 designed to induce **T cell-mediated killing preferentially when 2 or 3 target antigens (CD33, CD123, CD70) are co-expressed**
- MP0533 is hypothesized to preserve healthy cells hence **opening a therapeutic window**
- MP0533 has the potential to kill all AML cells (blasts and LSCs) despite heterogeneity hence ensuring **long term disease control**



MP0533 Induces Specific Killing of AML Cells Expressing 2 or 3 TAAs



- TAA's expressed on MOLM-13 cells
- CD33+CD123+CD70+
 - CD33+CD70+
 - CD123+CD70+
 - CD33+CD123+
 - CD33+
 - CD123+
 - CD70+



MP0533 Shows Preferential Killing of CD34+ LSCs over HSC

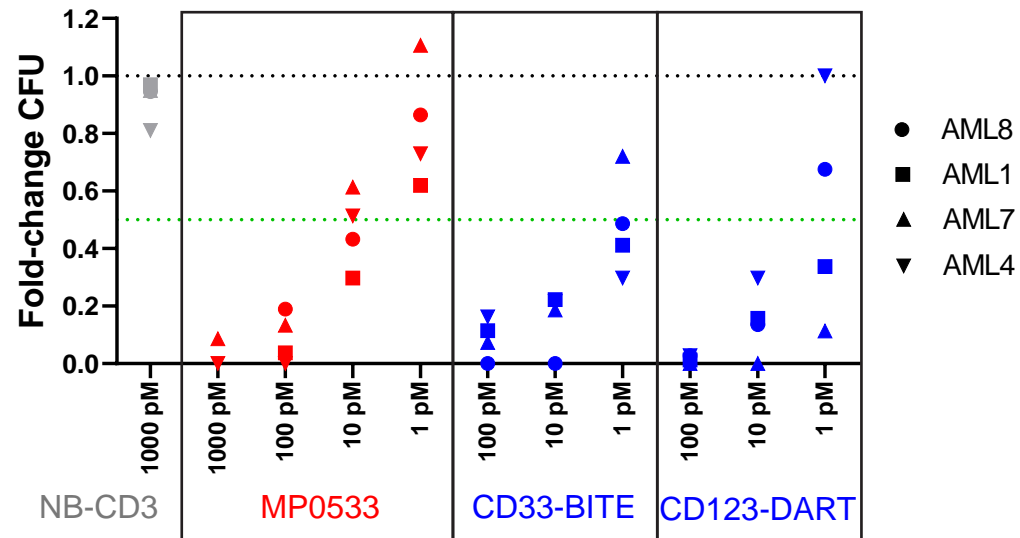
Larger therapeutic window as compared to CD123-CD3 DART and CD33-CD3 BiTE

Sorted CD34+ LSC or HSC
+ Healthy donor T cells (E:T = 1:1)

MP0533 or controls
14 days colony forming assay

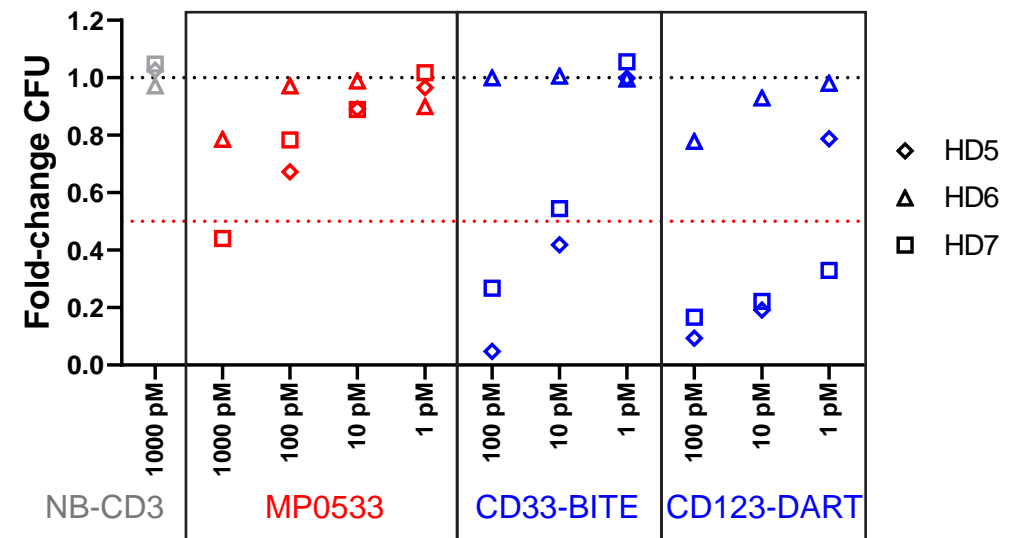
Counting of Colony
Forming Units (CFU)

Allogeneic killing of **AML CD34+ LSC**



Efficacy

Allogeneic killing of **healthy donor CD34+ HSC**



Safety

MP0533 Phase 1 Dose-escalation Trial in R/R AML patients

Patient population

AML or MDS/AML R/R to HMA, induction CT or allogenic HSCT
N=20–45 patients

Endpoints

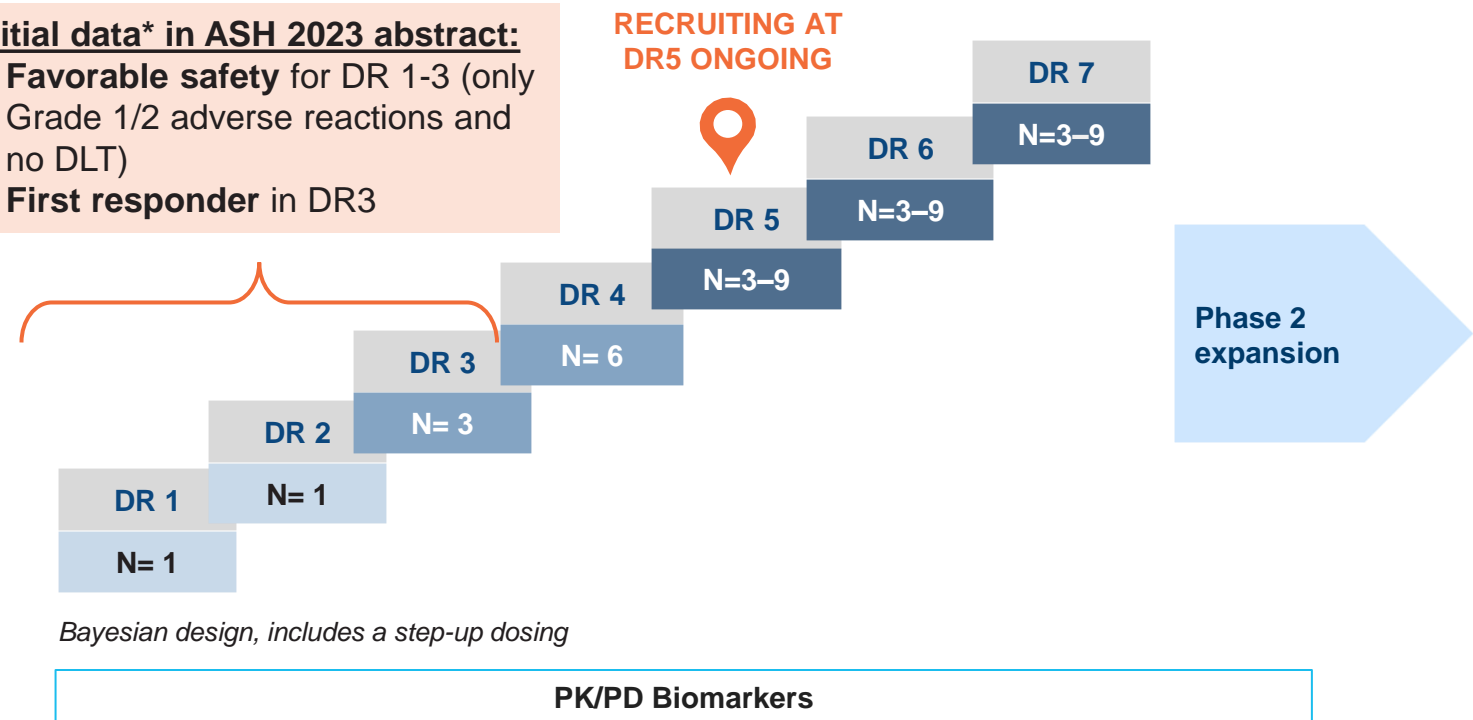
DLTs, safety, tolerability
antileukemic activity
PK, T-cell activation, cytokine release

Centers

7 sites open across Europe
(NCT05673057)

Initial data* in ASH 2023 abstract:

- **Favorable safety** for DR 1-3 (only Grade 1/2 adverse reactions and no DLT)
- **First responder** in DR3

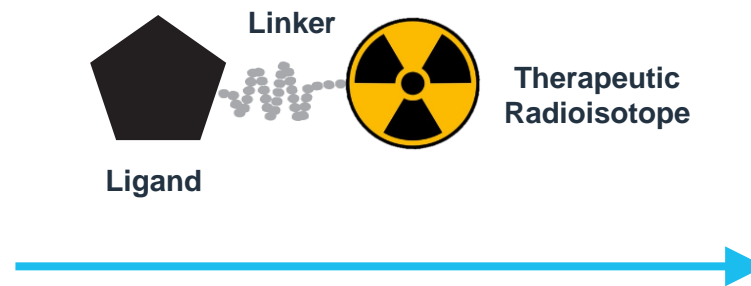
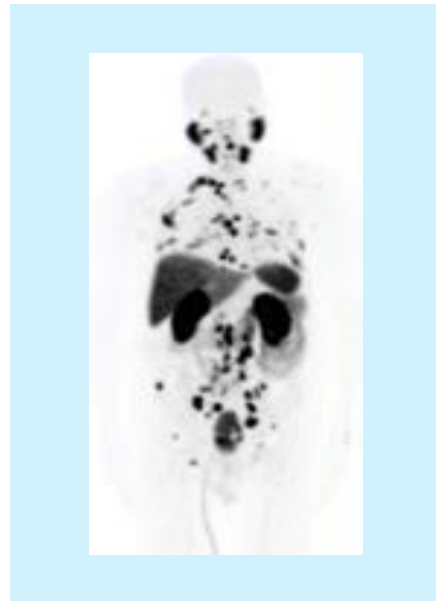


Study open and recruiting, initial results up to DR 4 to be presented at ASH 2023

Radio-DARPin Therapy Platform

Precision Oncology by Targeted Radioligand Therapy

Effective radioligands deliver a sufficiently large dose of radioactivity to the tumor for cell killing, while sparing healthy tissues



Ligand:
Specific targeting of tumor cells

Therapeutic radioisotope:
DNA damage to kill tumor cells

Radio-DARPin Therapeutics (RDTs): Platform to Expand the Targetable Space in Nuclear Oncology

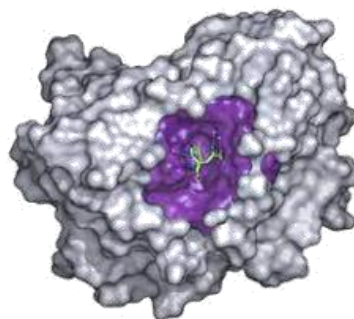
IDEAL RADIO PLATFORM PROPERTIES

- High affinity
- High specificity
- Short systemic half-life
- Low kidney uptake
- **Broad target range**



Most effective for

Targets where a small molecule ligand with high affinity & specificity can be generated or is available

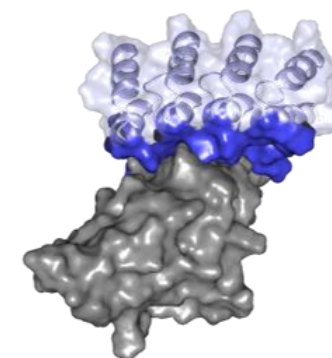


Example targets: PSMA...



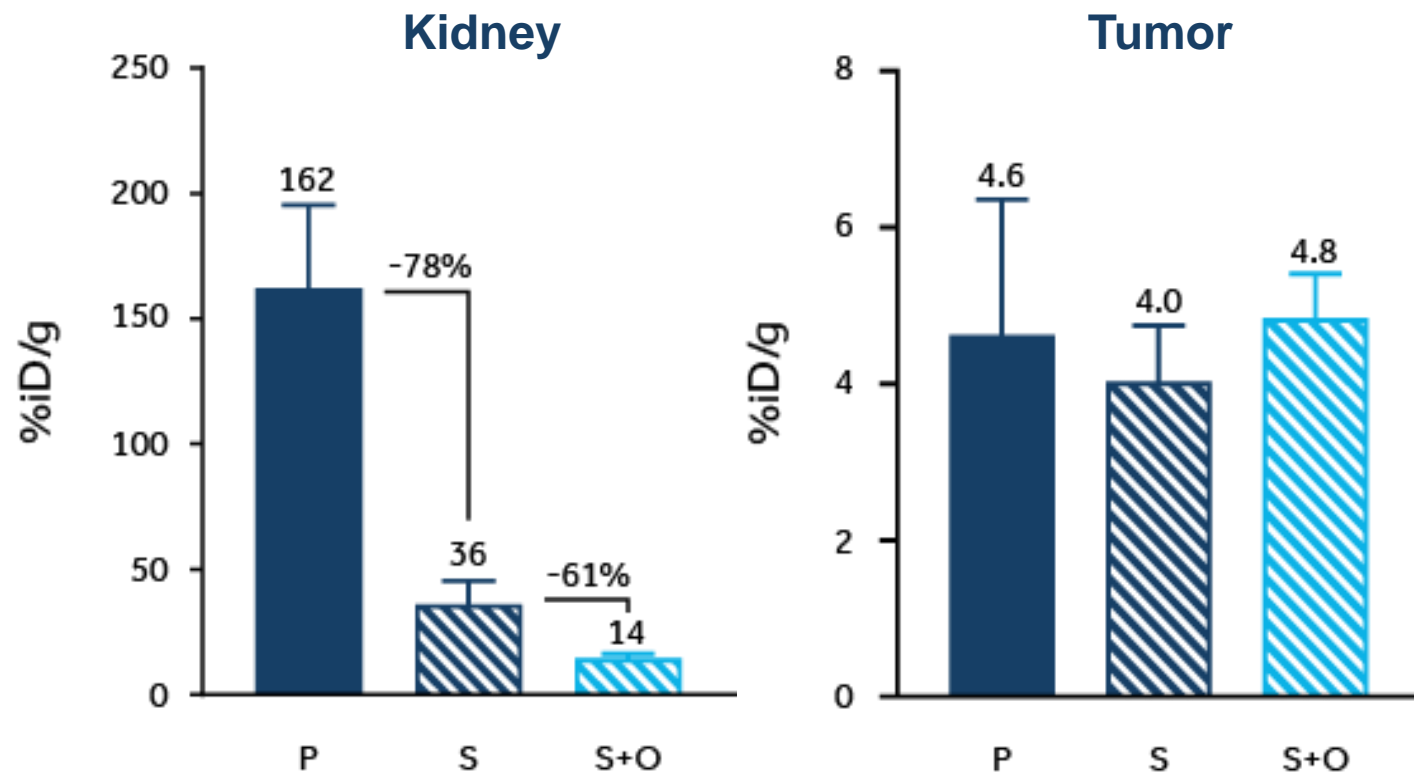
Most effective for







Targets that are challenging for peptides or small molecules (for desired specificity & affinity)



Example targets: Her2, DLL3, ...

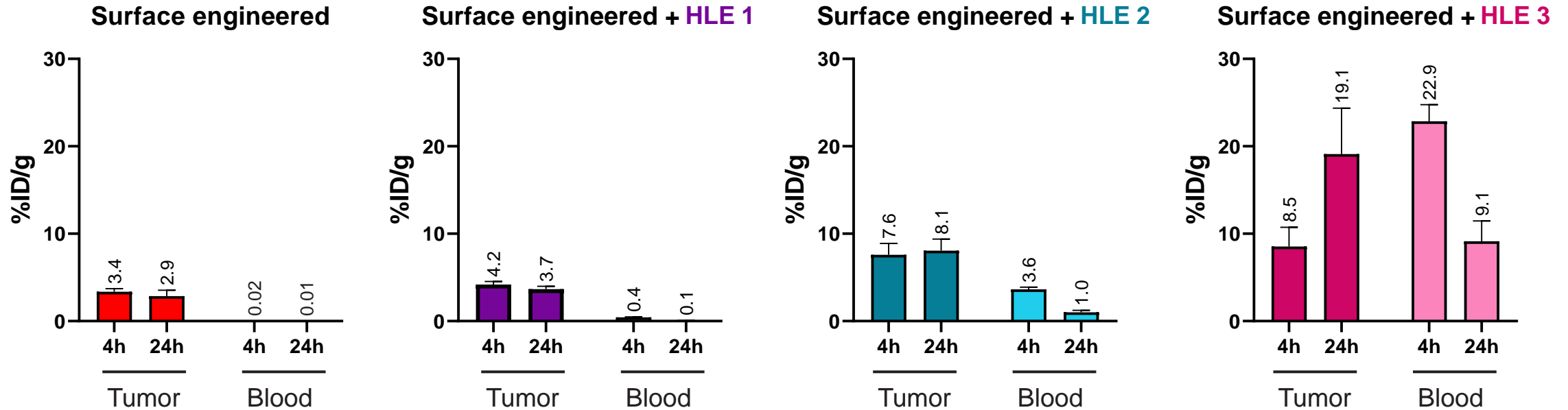
Surface Engineered Radio-DARPin Shows Strongly Reduced Kidney Uptake



After 4 hour timepoint		T/K*
	P: Parental 	1/35
	S: Surface Engineered 	1/9
	S+O: Surf. Eng. + Orthogonal 	1/3

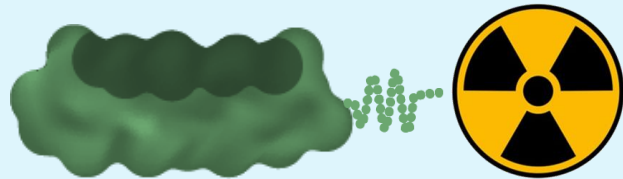
*tumor to kidney ratio
 **Orthogonal = MP proprietary kidney blocking or saturating agent

Systemic Half-life Extension (HLE) Increases Tumor Uptake



- Serum albumin binding results in increased blood levels that correlate with higher tumor uptake
- **HLE toolbox with different “strengths” allows RDT properties tailored to specific needs & payloads**

Collaborating with World Leader in Radio-Oncology



- \$20m up front*
- Up to \$560m in potential milestones
- Up to double-digit royalties
- Exclusive for two tumor antigens

Outlook

Outlook & Upcoming Milestones

MP0533	<ul style="list-style-type: none">• Initial encouraging Phase 1 results in R/R AML at ASH 2023• Additional data (response durability and depth) expected in H1 2024• Clinical expansion in Europe and preparation of potential US IND application
MP0317	<ul style="list-style-type: none">• Full Phase 1 proof-of-mechanism and safety data in H1 2024• Partnering for clinical development in combination settings
Radio DARPin Therapy Platform	<ul style="list-style-type: none">• Build on reduced kidney accumulation, focus on tumor accumulation• Evaluation and nomination of additional targets• Establish clinical and supply collaborations with radionuclide companies
Next Opportunities for DARPins	<ul style="list-style-type: none">• Presentation of SWITCH concept at PEGS Europe 2023• Leverage DARPin platform for next-generation immune cell engagers

CHF ~207 million cash (incl. short-term time deposits) ensures funding well into 2026*

