



Corporate Presentation

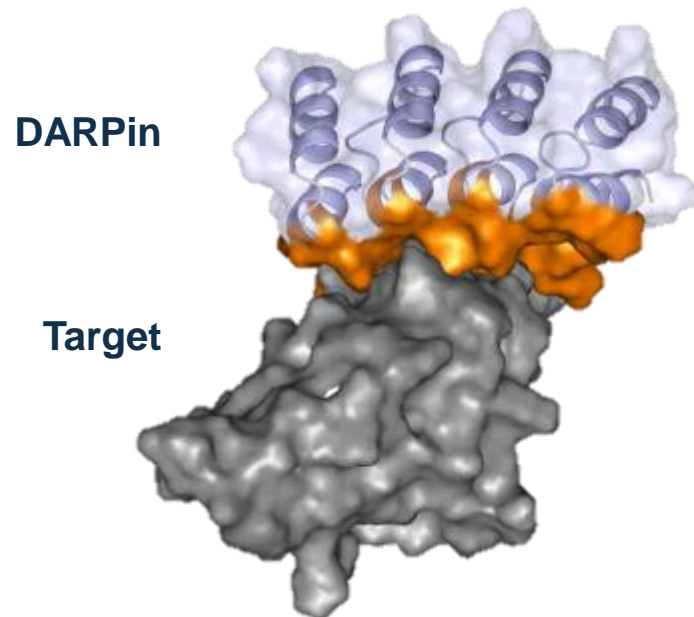
October 31, 2024

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The 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 solutions** for a defined medical problems not addressable by antibody designs
- Demonstrate **true patient value** with **early clinical readouts**
- Combine our **capabilities with world-class partners** to deliver innovative therapeutics

Corporate Highlights – Q3 2024

Radio-DARPin Therapy (RDT) & MP0712

- Successful RDT platform optimization to **reduce kidney accumulation** and **increase tumor uptake**
- **Strengthened collaboration with Orano Med** to co-develop four ^{212}Pb -labeled RDTs, incl. MP0712
- **MP0712, lead RDT targeting DLL3**, demonstrates attractive BioD, efficacy & safety profile (EANM 2024)

MP0533

- Novel **tetra-specific T-cell engager** for AML patients with high unmet need
- **Encouraging initial clinical data** (safety & efficacy) despite suboptimal exposure
- Phase 1/2a study on track (**DR 8** enrolling), dose intensification planned to explore the full potential of MP0533

Switch-DARPin & MP0621

- Demonstrated logic-gated immune activation for **Switch-DARPin platform**
- Proof-of-mechanism *in vivo* shown for **MP0621**, a **cKit x CD16a x CD47 Switch-DARPin** for next-gen HSCT
- **CD3 Switch-DARPin** as new platform for T cell engagers with enhanced function for solid tumors

MP0317

- Bi-specific CD40 agonist targeting FAP: **Favorable safety profile** and confirmed tumor-localized CD40 activation leading to **remodeling of tumor microenvironment** in patients

Operations

- Strong financial position with CHF 143.6 M in cash as of September 30, 2024, and proforma CHF 158 M following an offering on October 25, 2024, allowing **runway into 2027**

Pipeline

MODALITY	CANDIDATE	RESEARCH	PRE-CLINICAL	PHASE 1	PHASE 2	RIGHTS
Radio-DARPin Therapy (RDT)	MP0712	SCLC & NETs DLL3	Co-development*			MOLECULAR partners
	Undisclosed Programs	Solid Tumors	3 programs*			oranomed
	Undisclosed Programs	Solid Tumors	In-house programs			MOLECULAR partners
	Undisclosed Programs	Solid Tumors	2 partnered programs			NOVARTIS
Tetra-specific T-cell Engager	MP0533	r/r AML and AML/MDS CD33 x CD123 x CD70 x CD3				MOLECULAR partners
Switch-DARPin	MP0621	HSCT cKit x CD16a x CD47				MOLECULAR partners
	Undisclosed Programs	Immune Cell Engager				
Localized Agonist	MP0317	Advanced Solid Tumors FAP x CD40				MOLECULAR partners



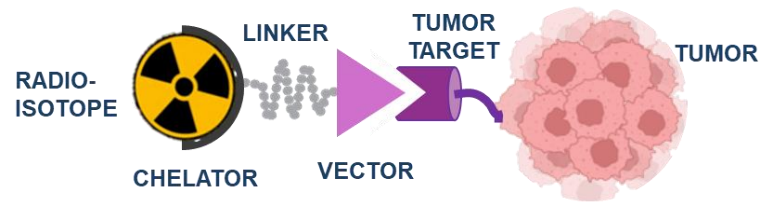
Radio-DARPin Therapy & MP0712 as first program

Platform & Pipeline



Targeted Radiotherapy: “Old” Modality Turned Hot Through Precision

A TARGETED RADIOTHERAPEUTIC:



- Potential to “**see what you treat**” and “**treat what you see**” in a powerful and targeted manner
- Proven clinical **benefit for oncology patients**;
 - Therapies with beta emitters established, data with alpha emitters on the rise
- **Supply chain** challenges remain
- **Opportunity**: Broaden the target & indication space with **vectors** amenable to selective tumor uptake

Example of a prostate cancer patient with extensive bone metastasis treated with ^{225}Ac -PSMA-617:

IMAGE → THERAPY → IMAGE



July 2017, PSA = 782 ng/ml
PET/CT, ^{68}Ga -PSMA-11

^{225}Ac -PSMA-617

8 MBq →

7 MBq →

8 MBq →



May 2018, PSA = 0.04 ng/ml
PET/CT, ^{68}Ga -PSMA-11

DARPinS Have the Potential to Broaden the Target Space

LMW Molecules

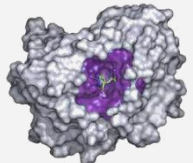


Generally good affinity and tumor uptake, low accumulation in kidneys

Limited number of targets with cavity where a LMW targeting moiety can be identified

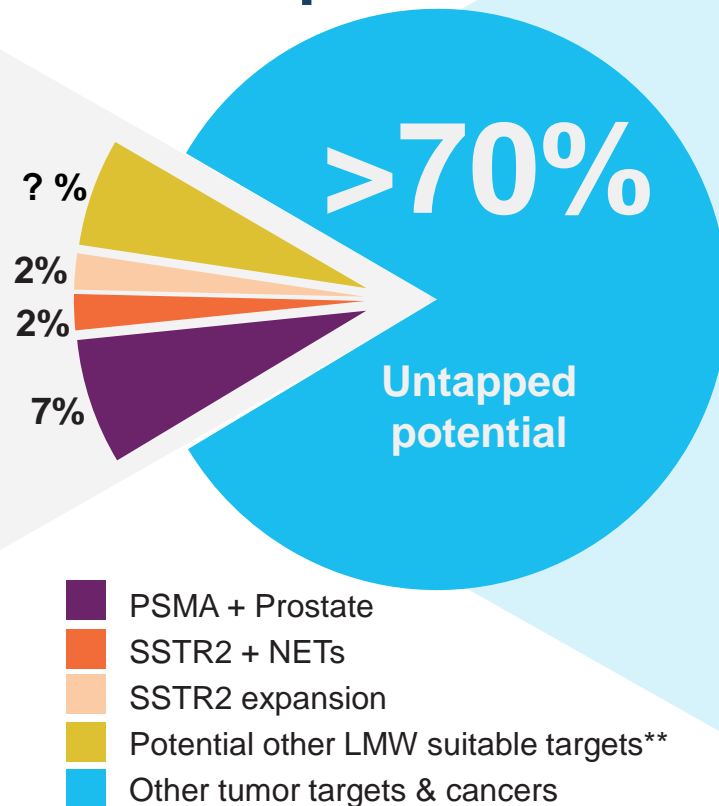
Target Examples:

PSMA
SSTR2



PSMA (1Z8L)

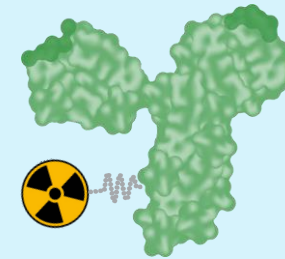
Target Space*



- PSMA + Prostate
- SSTR2 + NETs
- SSTR2 expansion
- Potential other LMW suitable targets**
- Other tumor targets & cancers

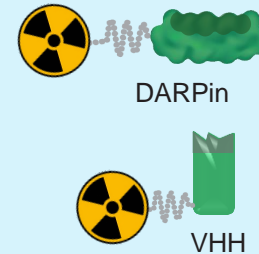
Expanding with other targeting moieties

High affinity & specificity binding of protein surfaces of broad range of tumor targets



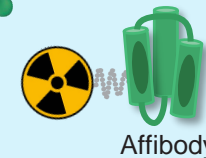
Antibodies

- Long circulation → risk of bone marrow toxicity
- Low tumor penetration



DARPin

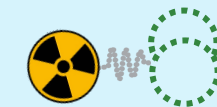
VHH



Affibody

Small proteins

- High kidney accumulation → risk of renal toxicity
- Suboptimal tumor uptake



Cyclic-peptides

- High kidney accumulation → risk of renal toxicity
- High affinity & selectivity can be challenging → target space?

Opportunity to Evolve DARPins to Radio-DARPins

Enabled by the robust architecture of the DARPin scaffold

Proteins < 60 kDa are reabsorbed by kidneys

Breast cancer patient imaged after treatment with a Her2 DARPin:

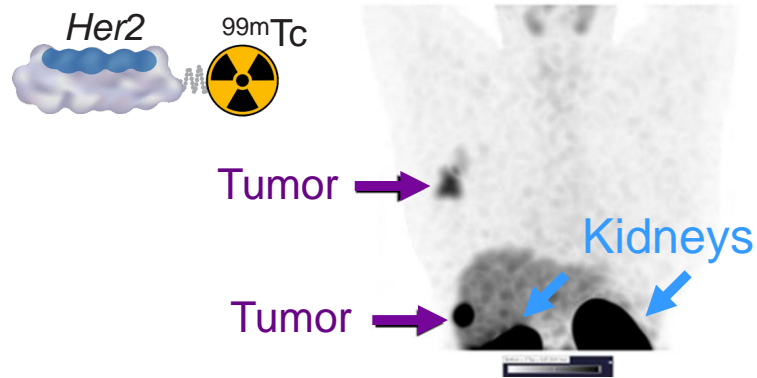
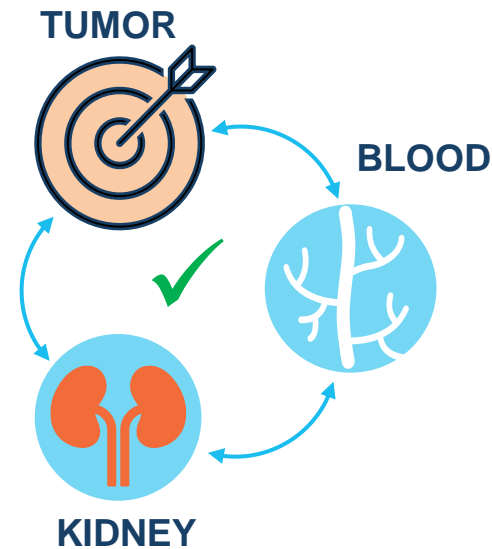


Image kindly provided by Dr. Bragina
Research Centrum for Oncotheranostics, Tomsk



Intrinsic DARPin properties



- ✓ **Small size** (~15 kDa)
→ Deep tumor penetration
→ Short systemic half-life
- ✓ **High affinity** (pM range)
→ Long tumor retention
- ✓ **High selectivity**
→ Low accumulation in other tissues
- ✓ **High stability**
→ Surface engineering

Unlocking DARPins for radiotherapeutic applications

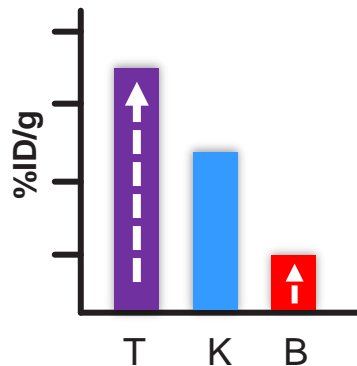
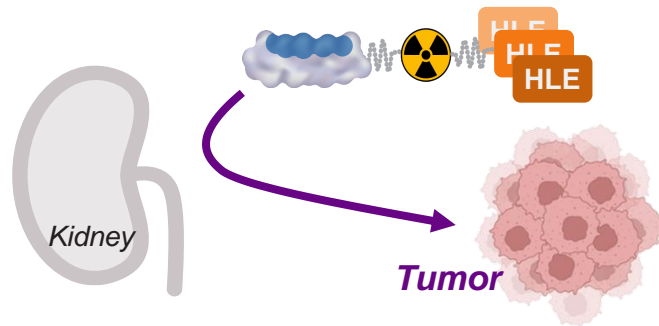
- Increase selective but moderate tumor uptake
- Reduce strong kidney accumulation

Radio-DARPin Platform Ready to Deliver Product Candidates

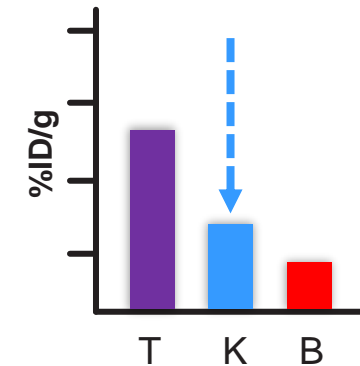
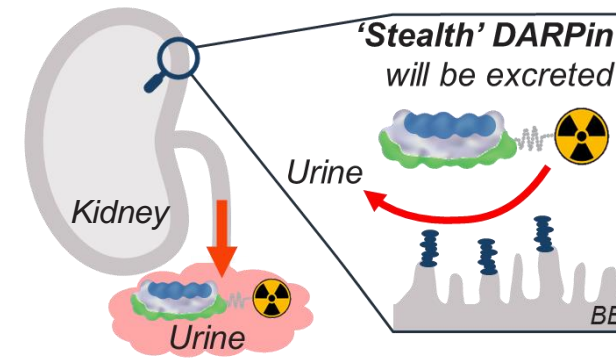
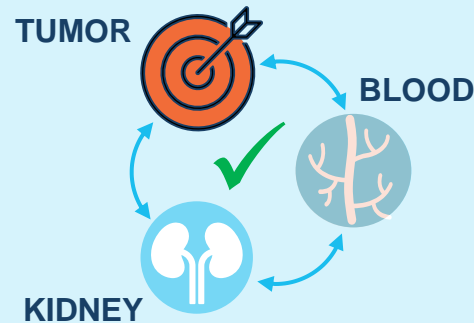
Increased tumor uptake
by half-life extension (HLE)*



Reduced kidney accumulation
by surface engineering (Stealth-DARPin)*



Optimized biodistribution
properties



MP0712: the First ^{212}Pb -DLL3 Targeted Radiotherapy

Combining distinctive DARPin features with the power of ^{212}Pb for efficacious cancer therapy

SCLC as indication

- Aggressive cancer with high unmet medical need
 - 2L: mPFS ~3m; 5y OS ~3%^{1,2}
- DLL3 is expressed in >85% of patients³

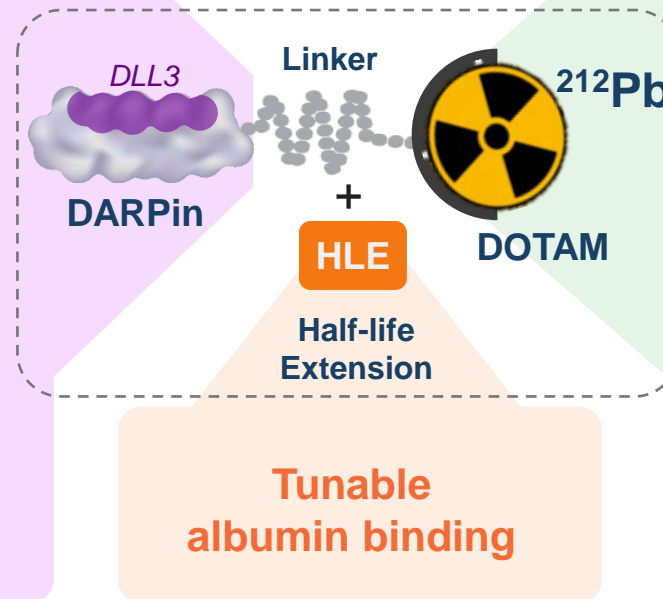
DLL3: a promising target

- Homogeneous tumor expression, but low expression level in patients
- No expression in healthy tissues
- New treatments with room for improvement: Tarlatamab (AMGEN) for 2L+; ORR ~40%

Diverse set of DARPins against DLL3

- Good developability
- Specific binding with high affinity

Product composition



^{212}Pb for targeted alpha therapy

- Strong cytotoxicity (dsDNA breaks)
- Single alpha decay (limited free daughters)
 - Limited irradiation of healthy tissues
- Relatively short half-life (10.6 h)
 - Fast energy deposition (efficacy)
 - Easier waste management

Co-Development with Orano Med

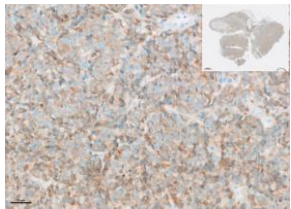
- The leader for ^{212}Pb & a committed partner
- Reliable & scalable ^{212}Pb production
- Independent production capacities (substantial inventory of purified ^{232}Th)

ASCO: Ph2 clinical data ^{212}Pb -DOTAMTATE (AlphaMedixTM) showed an ORR of 55.6%⁴

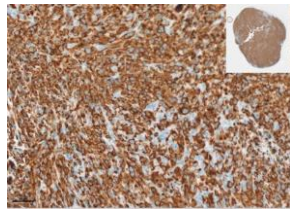
MP0712: Attractive BioD Profile and Tumor Specificity

DLL3 expression & distribution by IHC

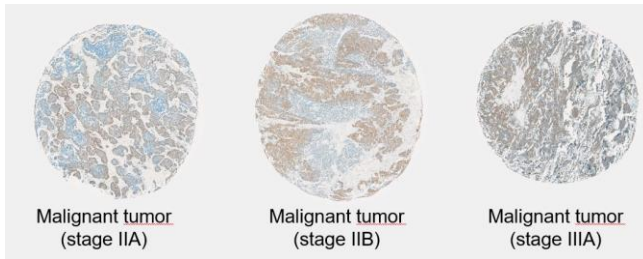
NCI-H82 tumors



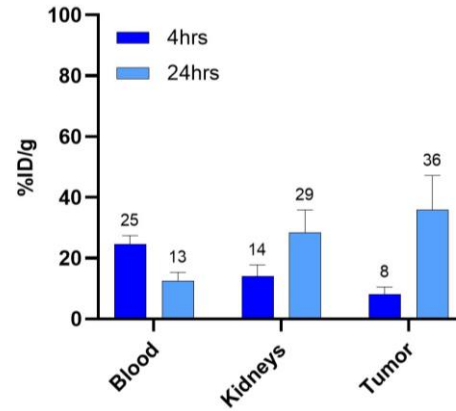
hDLL3-MC38 tumors



Human SCLC tumors



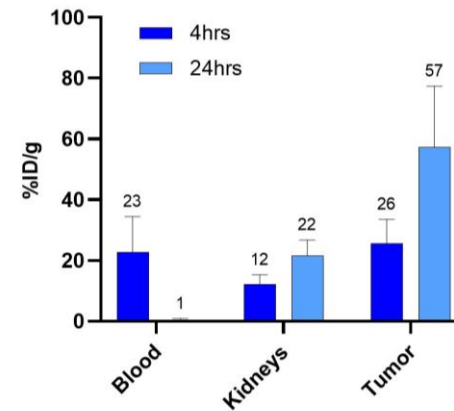
R2G2 mice xenografted *s.c.* with
NCI-H82



T:K at 4h = 0.6 / at 24h = 1.2



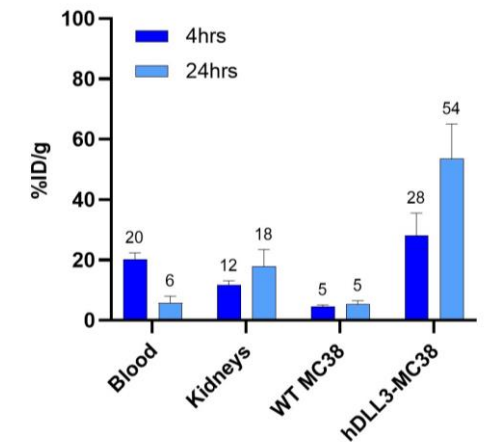
R2G2 mice xenografted *i.v.* with
NCI-H82



T:K at 4h = 2.1 / at 24h = 2.6



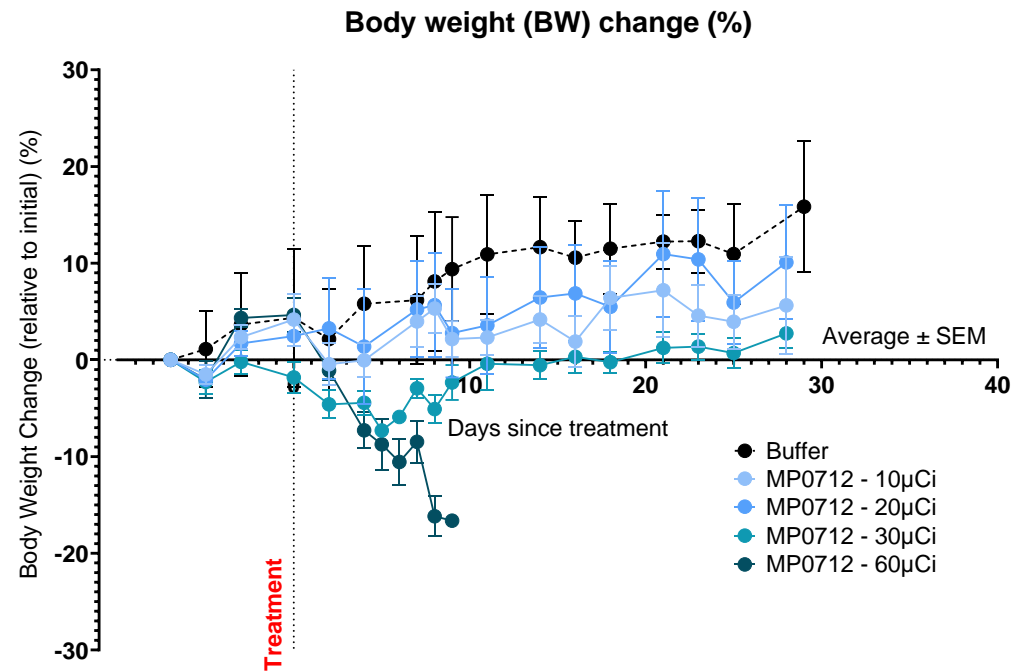
ATH nude mice double
xenografted *s.c.*
with **hDLL3-MC38** +
WT-MC38



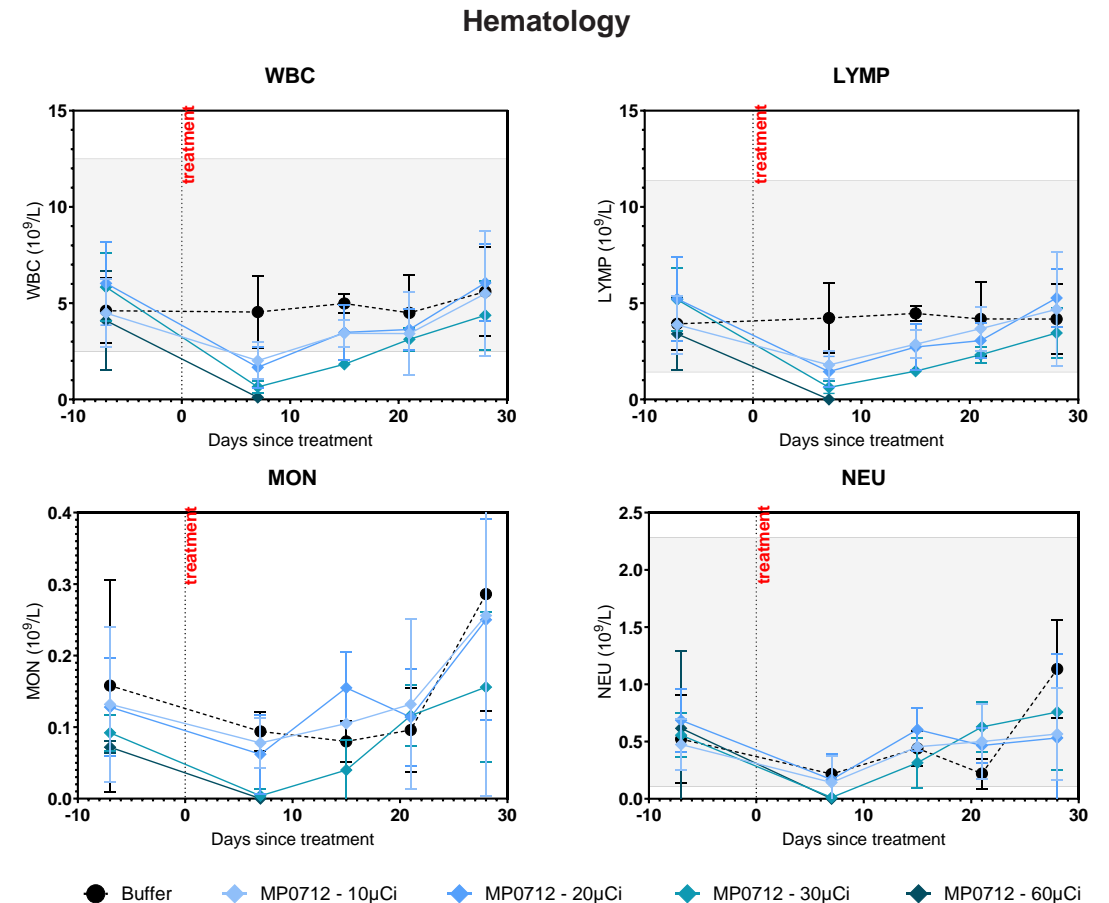
T:K at 4h = 2.4 / at 24h = 3

- MP0712 reached T:K ratios > 2 in mouse model matching clinically relevant DLL3 expression levels
- Selective uptake in DLL3-expressing tumors confirmed high target specificity of MP0712

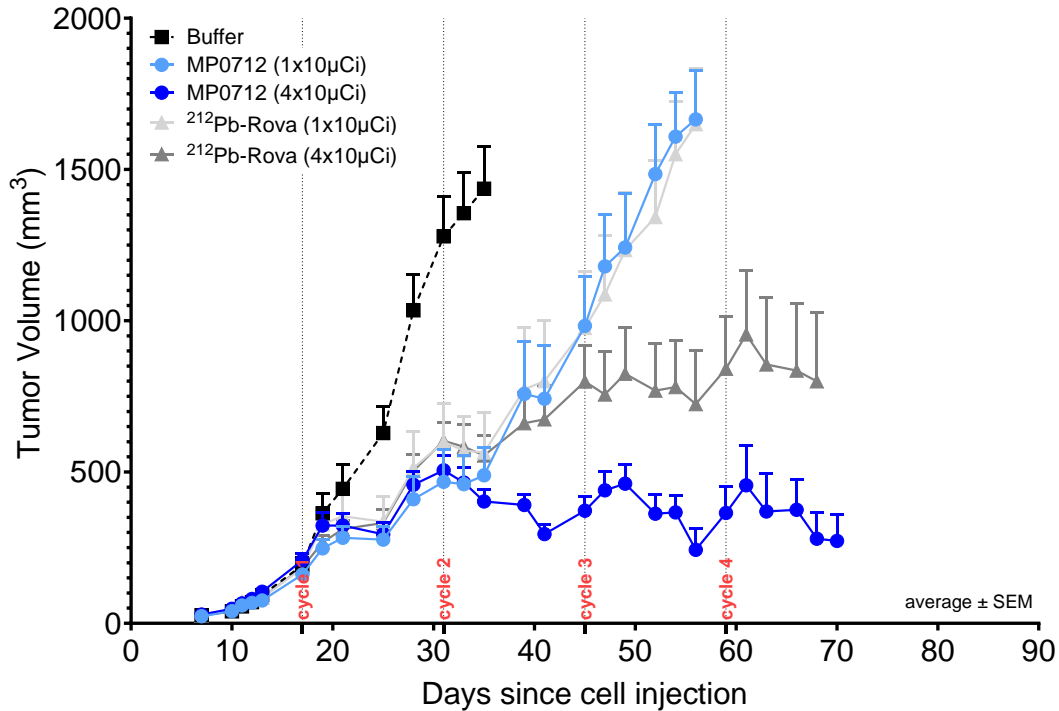
MP0712: Favorable Safety Profile



- Complete recovery of body weight loss after 10 days
- Complete recovery of hematologic profile after 28 days
- MP0712 treatment up to 30 μ Ci well tolerated



MP0712: Potent Efficacy at Clinically-Relevant Dose



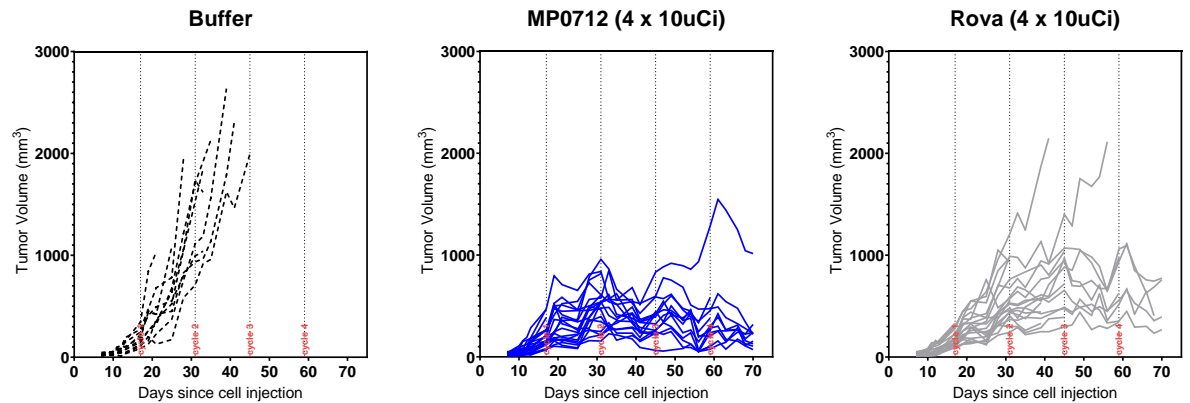
Median survival

Buffer	MP0712 1x10 μ Ci	MP0712 4x10 μ Ci	Rova 1x10 μ Ci	Rova 4x10 μ Ci
4.7 wks	7.9 wks	15.7 wks	7.9 wks	8.9 wks



R2G2 mice xenografted s.c. with
NCI-H82

Tumor growth curve for each animal



- **MP0712 induced tumor stabilization in NCI-H82 tumor model**

^{212}Pb has Key Advantages as Radioisotope Amenable to Radiotherapy

Selectivity

Localized and limited exposure of healthy cells with alpha particles

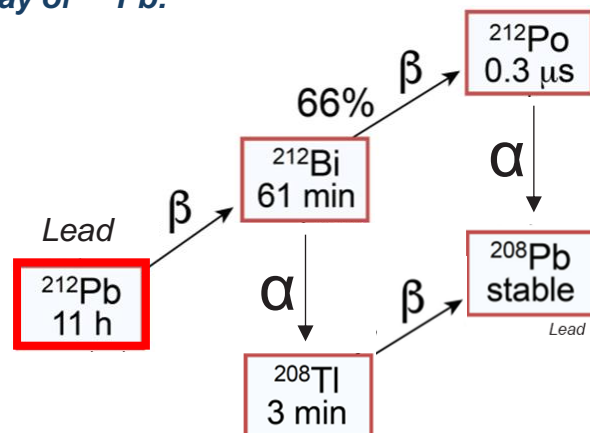
Safety

Clean decay profile: ^{212}Pb is an alpha precursor with low risk for long-lived free daughter radionuclides

Waste management

Less problematic thanks to short half-life

Decay of ^{212}Pb :



Adapted from Li et al., *Current Medicinal Chemistry*, 2020

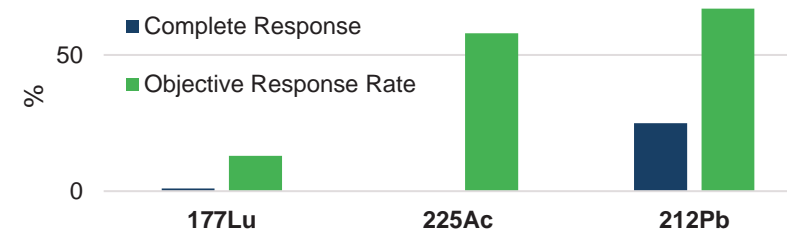
Efficacy

Short decay half-life leads to high energy deposition on tumor in short time frame

^{212}Pb demonstrated efficacy and good tolerability in GEP-NET patients treated with AlphaMedix™: 57% ORR in ph 1+2 combined (Strosberg et al, ASCO 2024)

^{212}Pb bears best-in-class potential for certain indications

	Beta	Alpha	
	177Lu (1)	225Ac (2)	212Pb (3)
Therapy	177Lu-DOTATATE (=Luthatera®)	225Ac-DOTATATE	212Pb-DOTAMTATE
Phase	Phase 3 NETTER-1	Comp. use	Phase 1
Patients (n)	111	26	12



Clinical data comparing ^{212}Pb with other radioisotopes in treatment-naïve NET patients treated with SSTR-targeting RLTs

Orano Med – Partner to Co-develop Radio-DARPin Therapies



“Endless” starting material as basis for ^{212}Pb supply

Leader in targeted alpha therapies

Large-scale, reliable, independent production and supply capabilities of ^{212}Pb

- Proprietary stockpile
- Achieve high purity of ^{212}Pb
- 4 GMP sites available or in construction across US and EU (incl. 2 AT Labs)
- Excellent logistics

Clinical capabilities demonstrated with ^{212}Pb and AlphaMedix™ in Phase 2 study in collaboration with RadioMedix

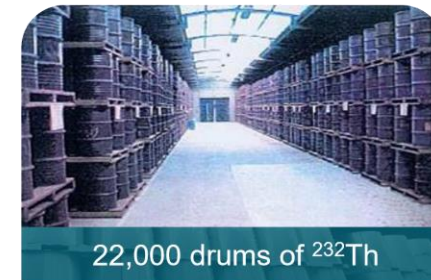
Strong partner for RDTs


Proven collaboration track record over past 2 years


- Trust, complementary and deep expertise

Co-development agreement signed in 2024:

- 50:50 cost and profit share
- Four RDT programs, including MP0712 (DLL3)
- Molecular Partners commercialization rights for DLL3

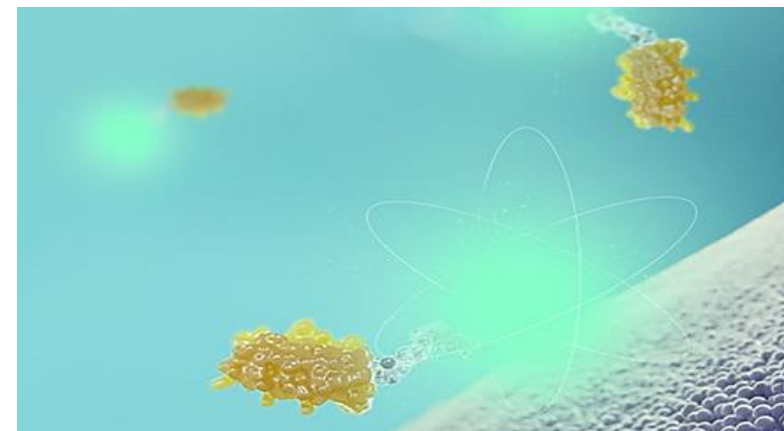


 Lead-212 is obtained **chemically** by **successive extractions and purifications** of the descendants of thorium-232

 Orano Med owns more than 20,000 drums of highly purified thorium-232 offering virtually unlimited supply

Summary – Radio-DARPin Therapy (RDT) & MP0712

- Successful RDT platform optimization **with attractive biodistribution profile** (tumor, kidney, blood)
- **MP0712 selected as Lead Candidate for targeted ^{212}Pb -DLL3 Radio-DARPin Therapy**: encouraging safety & efficacy *in vivo*
- IND-enabling package working towards completion; **initial clinical data expected in 2025**

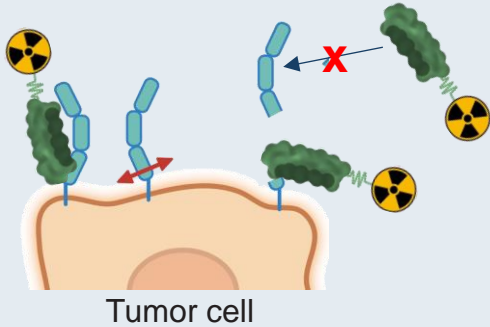


TARGET	RESEARCH	DEV.	RIGHTS
DLL3	MP0712		
Target 2			MOLECULAR partners
Target 3			oranomed
Target 4			
Target X			NOVARTIS
Target Y			
Several targets in evaluation			MOLECULAR partners

RDT Outlook:

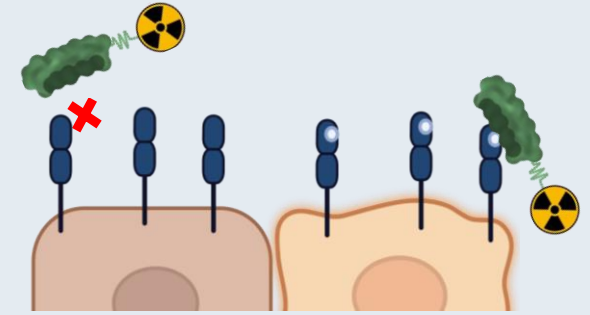
- **Advance MP0712 and additional pipeline candidates**
- **Continue to evolve RDT platform for next differentiated RDT programs**
- **Progress collaboration projects with Orano Med and Novartis**

Outlook: Leverage DARPin Differentiation to build RDT portfolio



Selectivity for **membrane-bound antigen** vs **shed antigen** for high tumor uptake

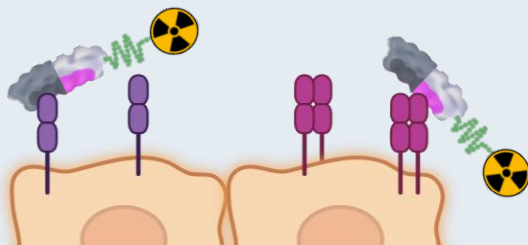
Selectivity to antigens with **high surface homology** to other targets



Healthy cell

Tumor cell

2in1 DARPin



Tumor cells

Bi-specific DARPins to achieve **broader distribution in tumors & overcome heterogeneity**, especially for targeted alpha therapy

Created in part with [BioRender.com](https://www.biorender.com)



MP0533

Tetra-specific T-cell Engager for AML

Patients with AML Have a High Unmet Medical Need

69 YEARS
OLD

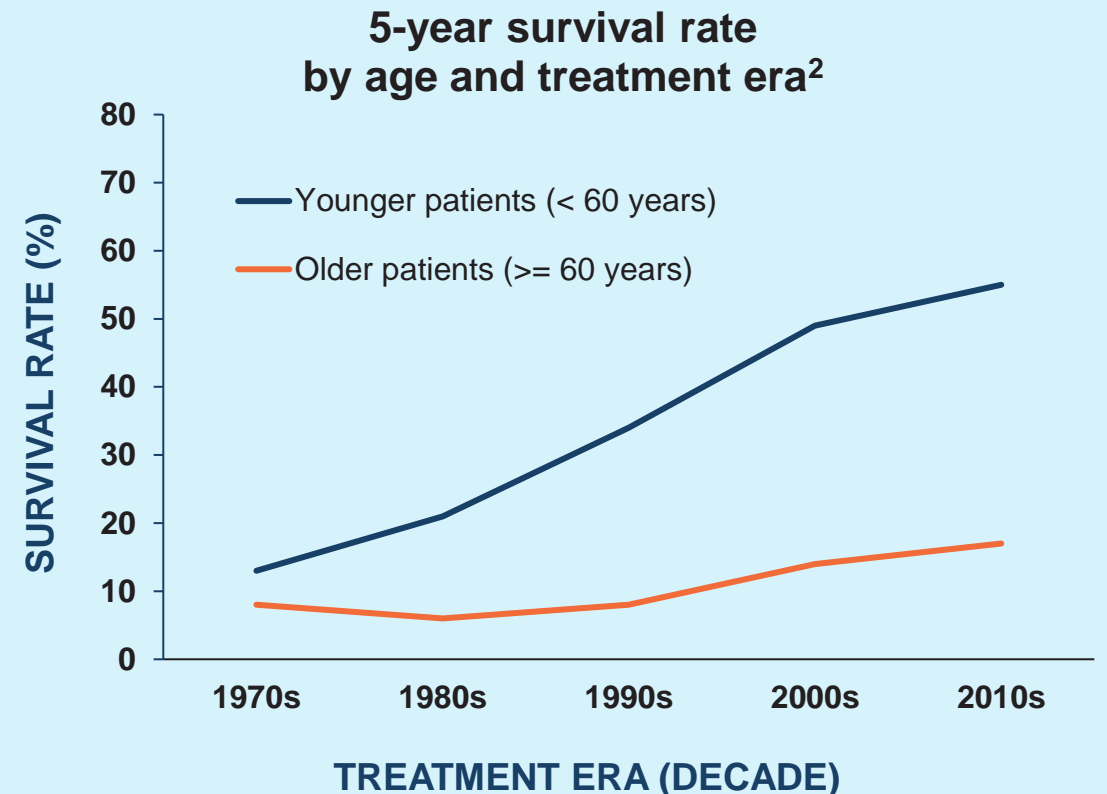
Median age of AML
patients at diagnosis¹

31.7%

Overall 5-year
survival rate¹

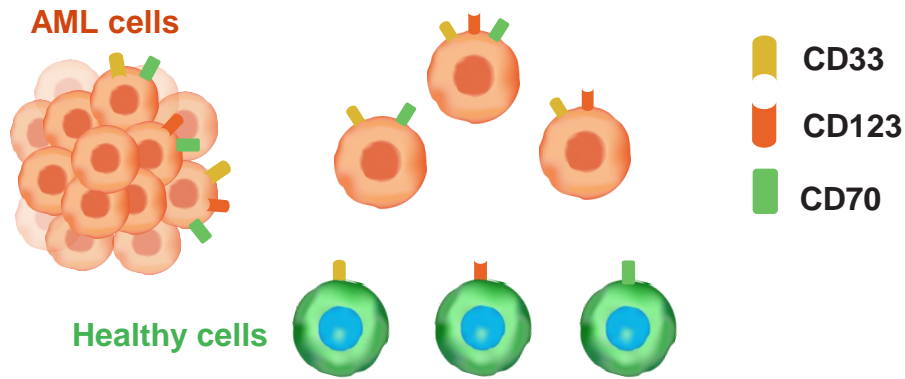
Despite 50 years of progress, elderly and frail patients are often not eligible for high-intensity conditioning and HSCT, and thus have limited treatment options and poor survival outcomes²

- Lack of broad and clean AML surface targets
- Risk of clonal escape even after high-intensity conditioning/HSCT



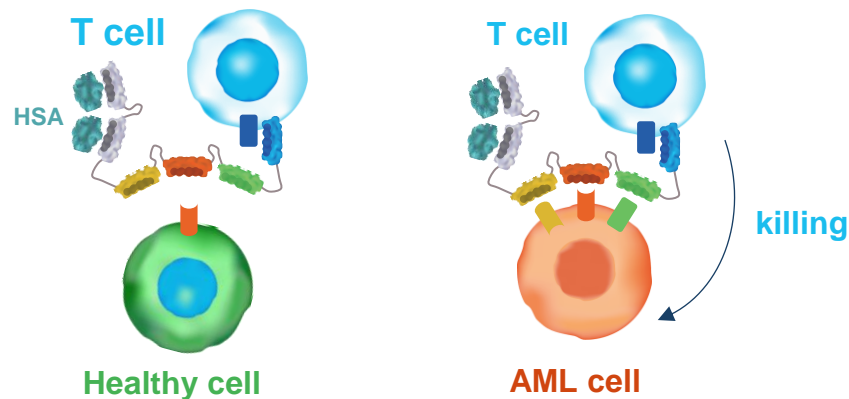
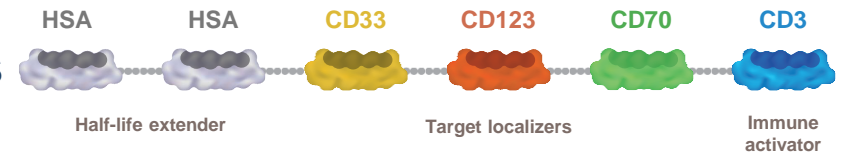
MP0533: Avidity-Driven Selectivity for Cancer Cells in AML

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 (e.g. HSCs) by their **co-expression of specific targets** (e.g. CD33, CD123, CD70)

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



- MP0533 is designed to induce **T cell-mediated killing preferentially when two or three 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, ensuring **long term disease control**

MP0533 Phase 1 Dose Escalation in R/R AML Patients

Rapid progress up to cohort 7 with need to explore higher doses

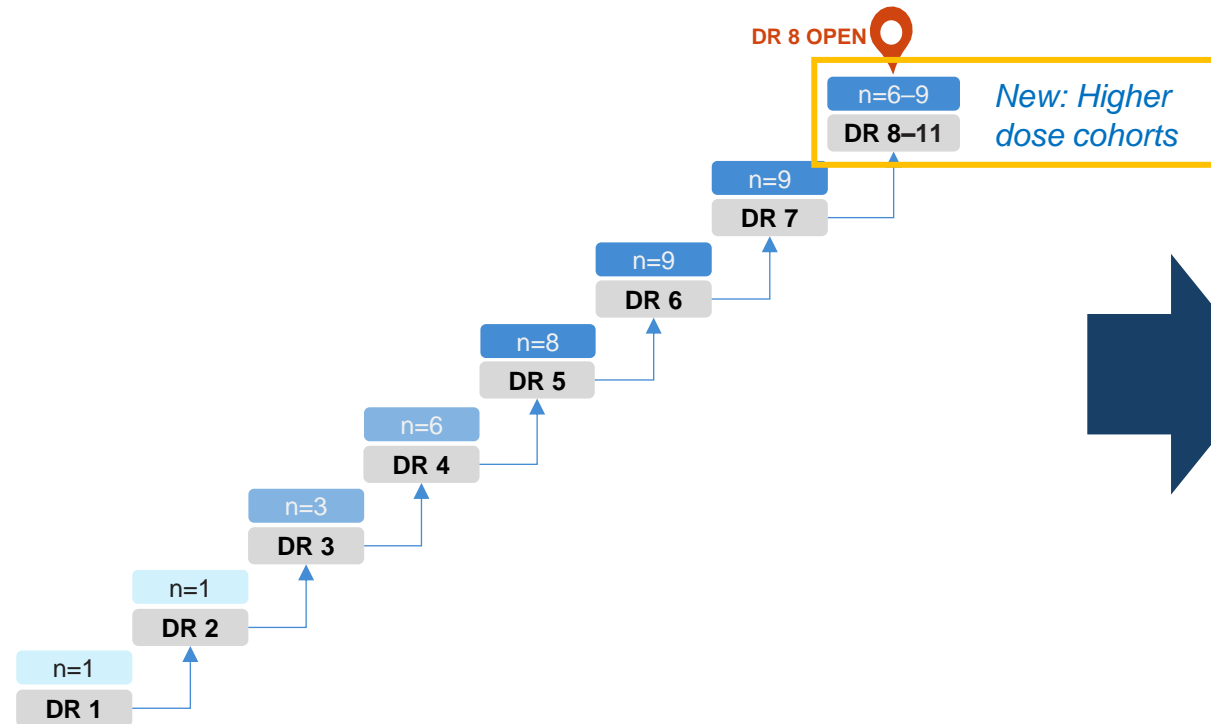
STUDY DESIGN

- FIH, single-arm, open-label, Phase 1/2a study of MP0533 monotherapy (NCT05673057)

STUDY OBJECTIVES

- Safety / tolerability
- PK / exposure
- Preliminary activity / PD
 - Clinical response as per ELN (incl. MRD status)
 - Blasts and LSCs counts
 - T-cell activity
 - MP0533 presence in BM
 - Target (co-)expression
 - Evolution of disease clonality

PHASE 1 DR ESCALATION OF MP0533 MONOTHERAPY



Study on-going across 9 sites in EU, DR 8 enrolling

PHASE 2A PoC OF MP0533 MONO- / COMBINATION THERAPY

MP0533 Phase 1 Patient Characteristics and Safety Profile

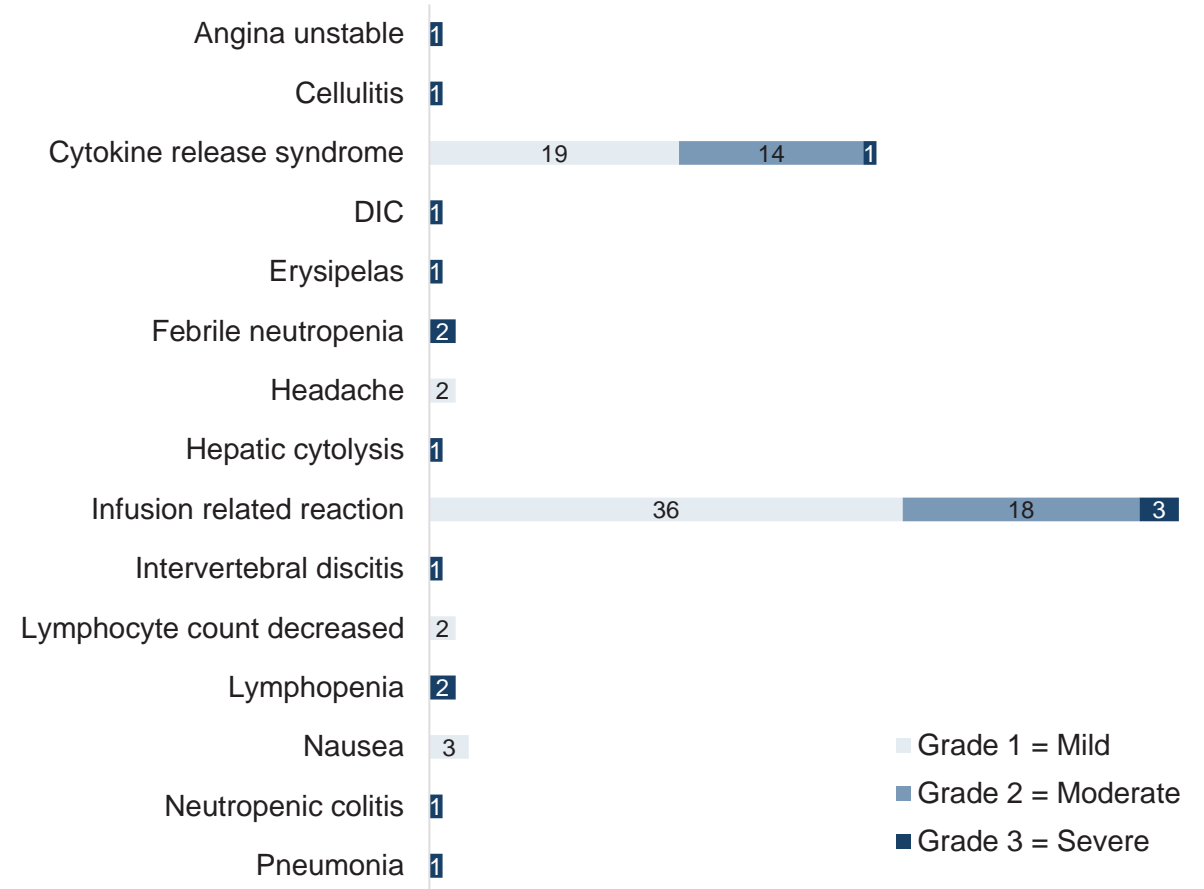
BASILINE CHARACTERISTICS	DR COHORTS 1-6 (n=28)
Sex, n (%)	
Female / male	14 (50) / 14 (50)
Age	
Mean / Median (range)	68 / 74 (22-82)
ECOG PS, n (%)	
0 / 1 / 2	11 (39) / 15 (54) / 2 (7)
Hematologic malignancy, n (%)	
AML / MDS/AML	19 (68) / 9 (32)
ELN risk category, n (%)	
Intermediate / adverse	4 (14) / 24 (86)*
No. of prior systemic treatment lines, n (%)	
1 / 2 / ≥3	12 (43) / 10 (36) / 6 (21)

*TP53 mutated: 7 (25%)

Acceptable safety profile for MP0533 reported for DR 1-6‡:

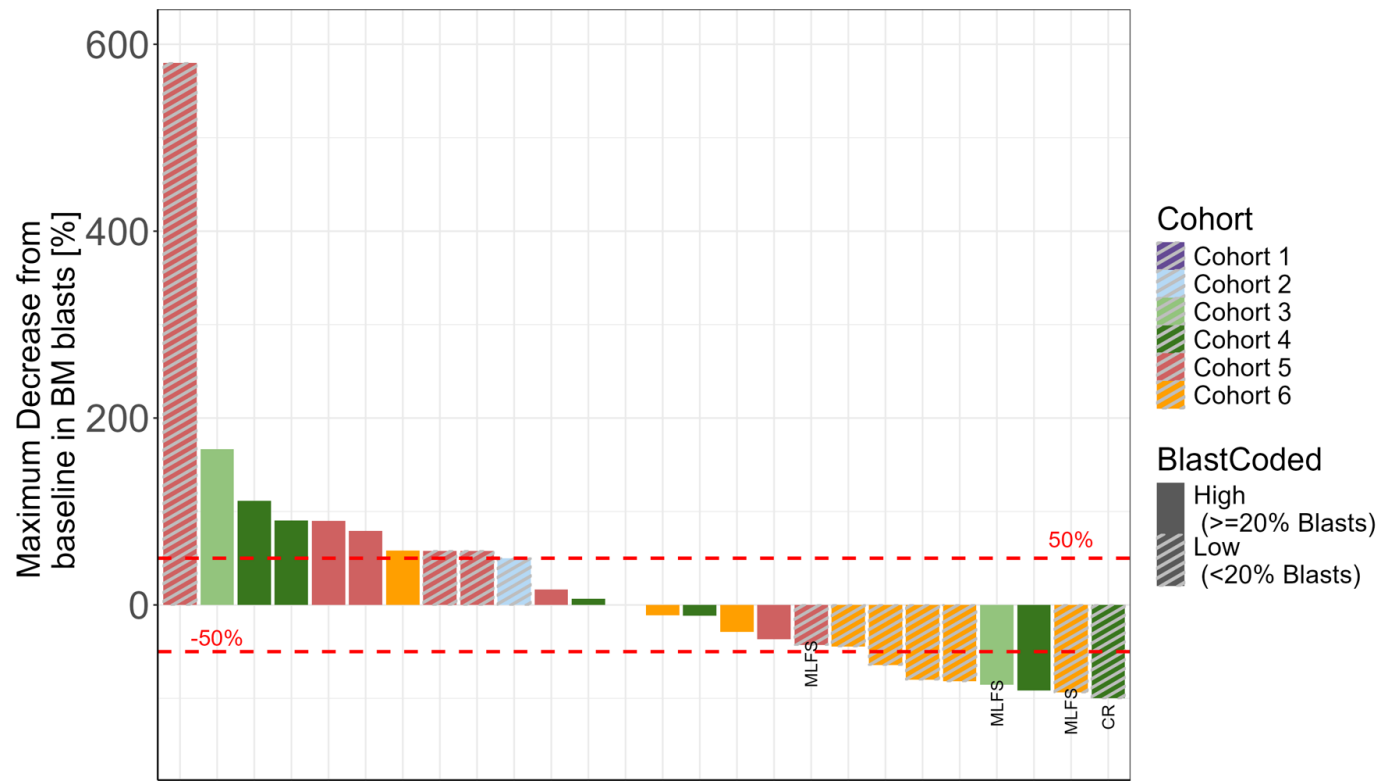
- IRR and CRS are the most frequent MP0533-related TEAEs (mostly Grade 1-2, occasional Grade 3)
- No DLTs up to DR 6

MP0533-RELATED TEAEs‡

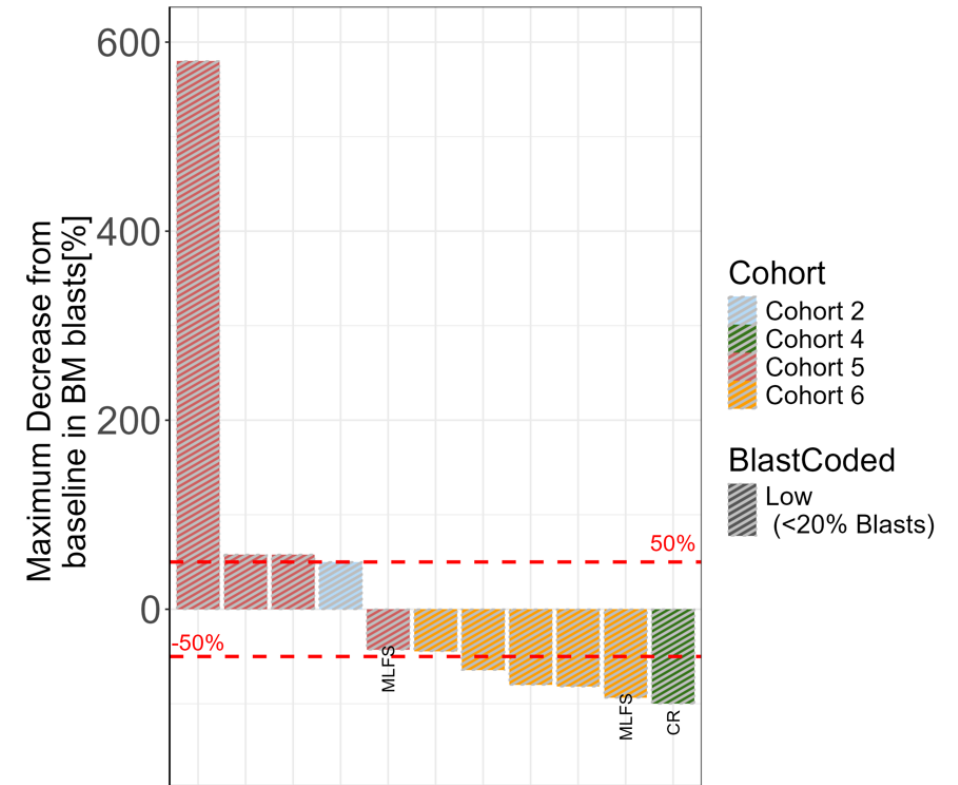


Encouraging Blast Reduction Observed, Particularly in Patients with Lower Disease Burden*

7 of 26 evaluable patients displayed >50% blast reduction in the bone marrow



5 of 11 patients with **lower disease burden*** displayed blast reduction >50 %



MP0533 Treatment & Clinical Response

Four responders reported in DR 3-6:

- CR in 1 patient at DR 4
- MLFS in 3 patients, 1 each at DR 3, DR 5 and DR 6

DR 8 enrolling patients



LEGEND

- ★ CR
- ★ CRi
- ★ MLFS
- No ELN response

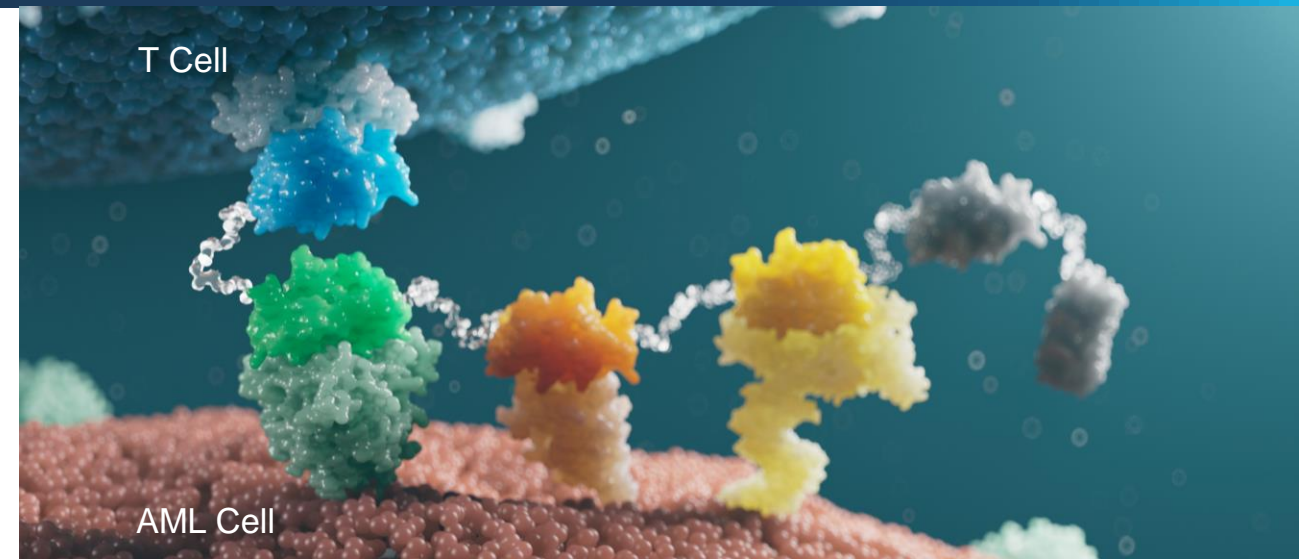
Response (2022 ELN¹) was assessed every 4 weeks until disease progression and results are presented as indicated

Arrows at the top indicate MP0533 administration at D1, D5, D8, D15 and weekly thereafter

- ↓ Step-up dosing is presented in green arrows
- ⇓ Color changes in blue arrows indicate start of a new 28-day cycle

MP0533 Summary

- **Rapid progress of MP0533 phase 1** with engaged clinical experts & sites
 - DR 8 enrolling, 28 patients treated in DR 1–6
- **Acceptable safety profile** supports higher dosing
 - IRRs & CRS as most frequent MP0533-related TEAEs
- **Initial antitumor activity** in highly heterogeneous r/r AML population
 - 4 responders reported (1 responder per cohort, DR 3–6)
 - Encouraging reduction in BM blasts observed
- Need to **improve suboptimal exposure** to **unleash the full potential of MP0533**
 - Increase response rate, depth and durability



Outlook

- Protocol being amended for **both higher & more frequent dosing** (in first weeks)
- Clinical update on the program at ASH 2024 and on the **amended dosing scheme in 2025**
- **Results** from these activities will **gate future development**



Switch-DARPin Platform & MP0621

Targeted and conditional activation of immune cells

Logic-gated Switch-DARPin(s) for Conditional Immune Activation

Swiss knives for enhanced immune engagers

1st Antigen Binder

- Anchoring to target cells
- Adding avidity for selectivity and address tumor heterogeneity

✓ CLINICALLY VALIDATED

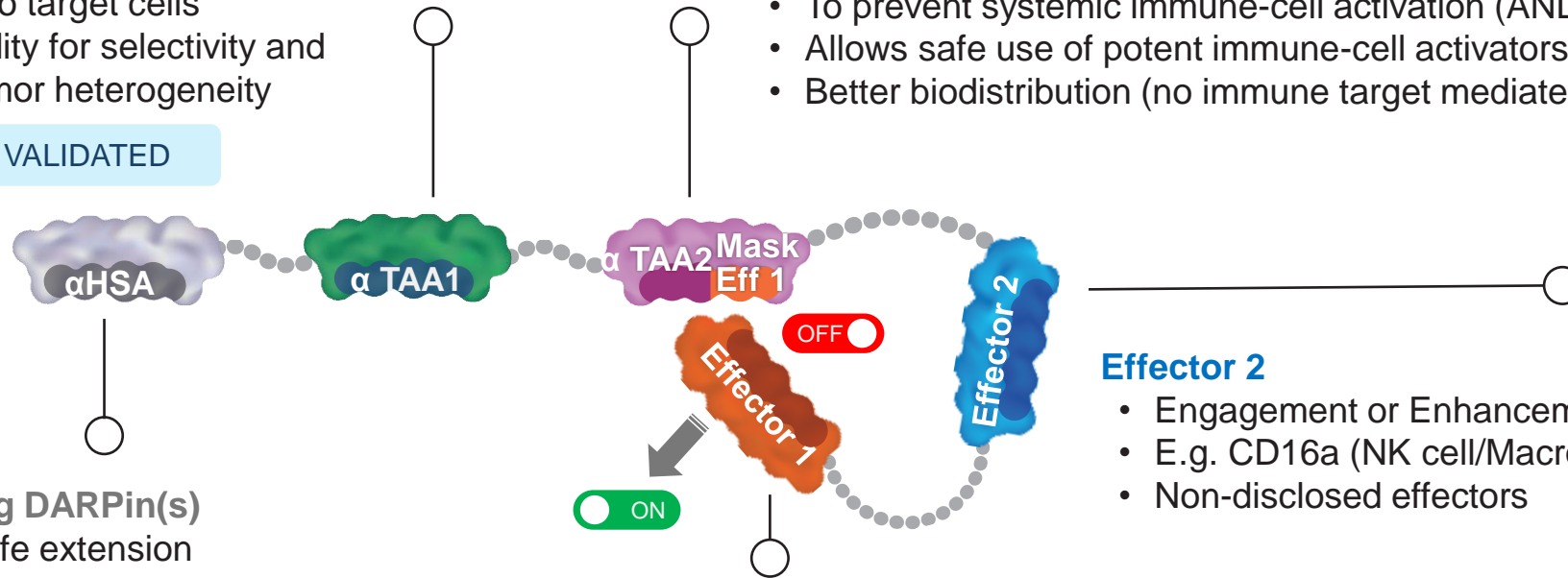
2-in-1 DARPin: Exclusive Binding to 2nd Antigen or Masking Effector 1

- To prevent systemic immune-cell activation (AND gate)
- Allows safe use of potent immune-cell activators
- Better biodistribution (no immune target mediated sink)

HSA Binding DARPin(s)

- For half-life extension

✓ CLINICALLY VALIDATED



Effector 2

- Engagement or Enhancement of immune response
- E.g. CD16a (NK cell/Macrophage engagement)
- Non-disclosed effectors

Effector 1 (Switched on/off by Masking DARPin)

- Engagement or enhancement of immune responses
- E.g. CD47 (block don't-eat-me signal)
- E.g. CD3 ("Signal 1" T-cell engagement)

✓ CD3 TCE CLINICALLY VALIDATED

MP0621: cKit x CD16a x CD47 Switch-DARPin

Next-Generation Conditioning Regimen for HSCT

Target cKIT to eliminate HSCs/LSCs

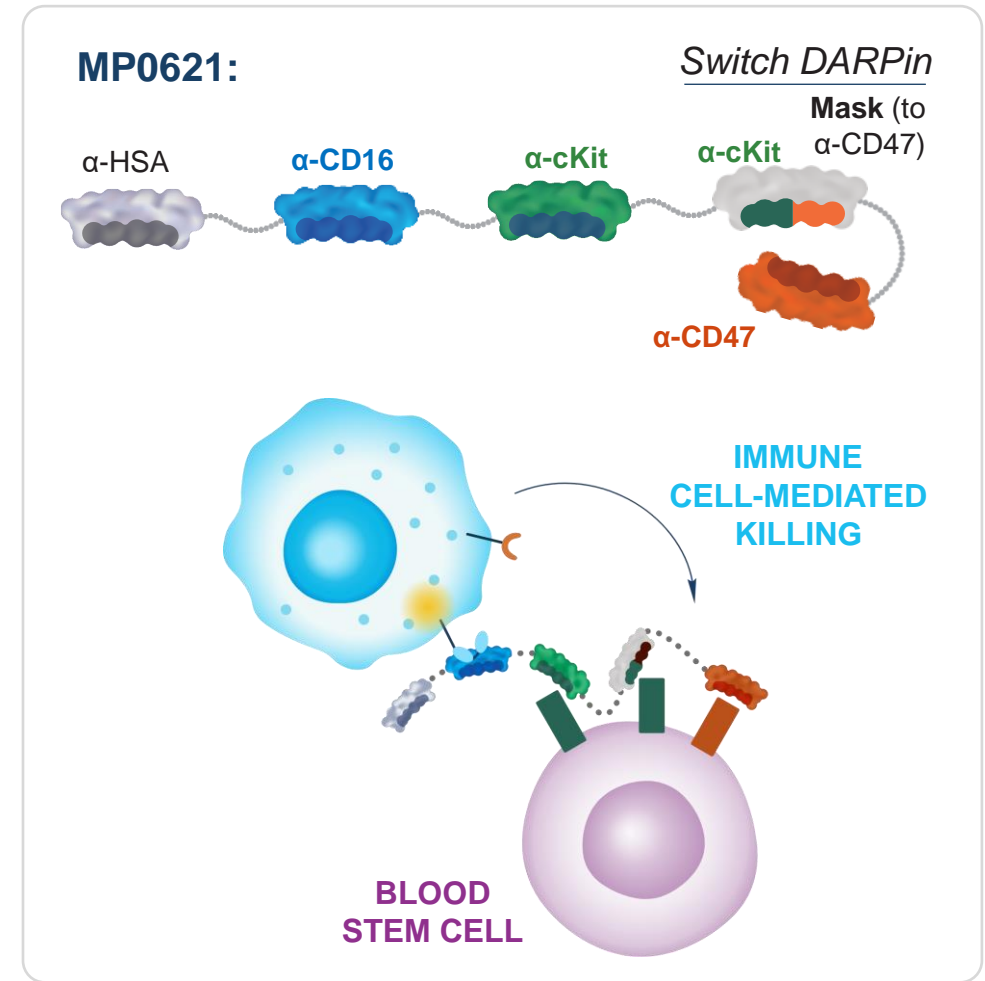
- cKIT is critical for stem cell maintenance and renewal^{1, 2}
- Simple antagonists (mAbs) to cKIT are not potent enough

Engage NK cells and macrophages (MΦ) via CD16a to kill HSCs/LSCs

- Effective and safe approach
- NK and MΦ activity is limited by CD47 expression on HSC/LSCs³

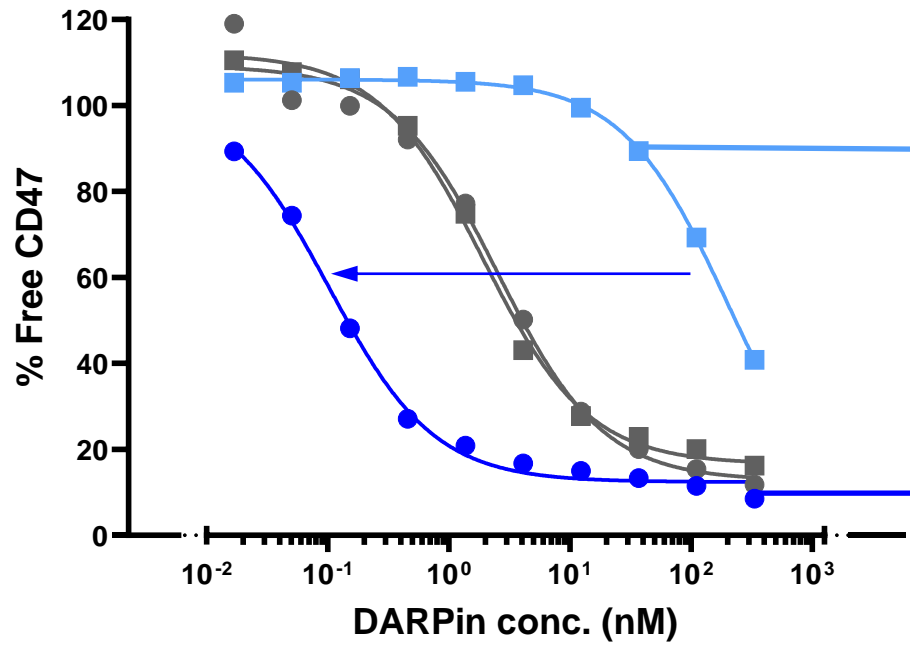
Conditionally block CD47 on LSCs/HSCs to boost NK cell and MΦ killing activity

- CD47 is expressed as “do-not-eat-me signal” and prevents killing of HSCs/LSCs^{1,3}
- Switch DARPin allows conditional local blocking of CD47 on HSCs/LSCs, prevents peripheral CD47 blockade

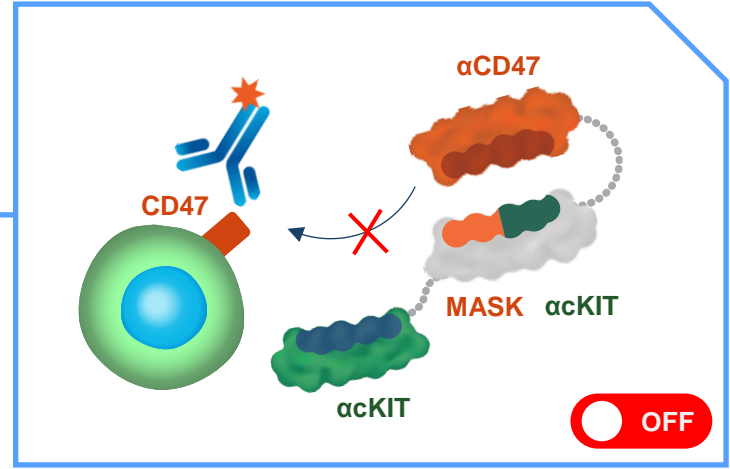


Switch-DARPin POC – CD47 is Blocked Only on cKit Positive Cells

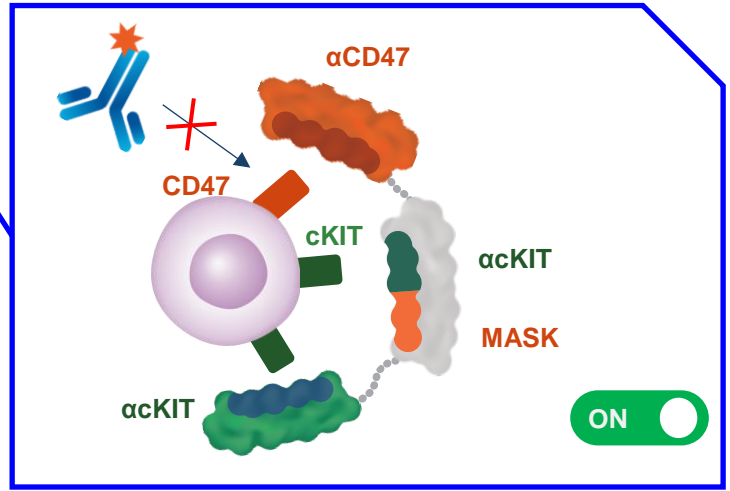
c-Kit-dependent CD47 blockade




- MP0621 on cKit⁺ cells
- MP0621 on cKit⁻ cells
- α-CD47 on cKit⁺ cells
- α-CD47 on cKit⁻ cells



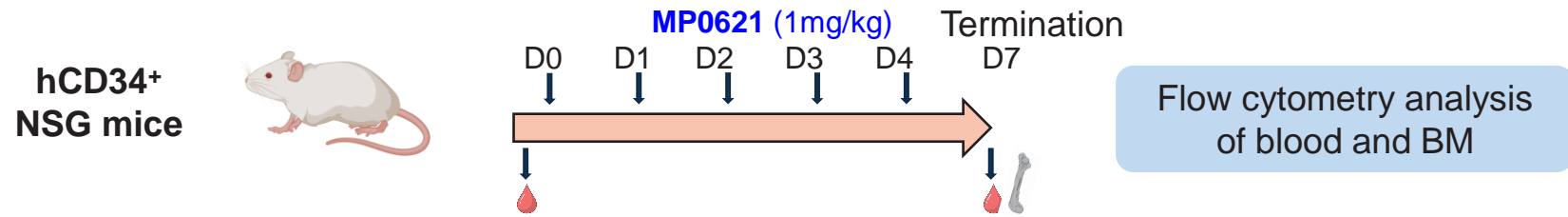
cKit Negative cells
Switch is OFF
CD47 is NOT blocked



cKit Positive cells
Switch is ON
CD47 is Blocked

 anti-CD47 detection agent

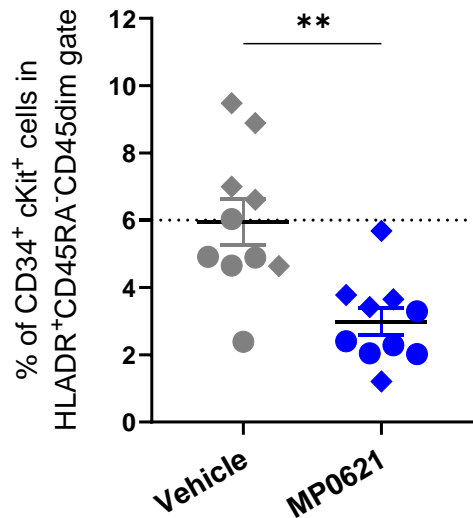
MP0621 Depletes cKit+ Cells in Bone Marrow Without Affecting Circulating Immune Cells in Humanized Mice



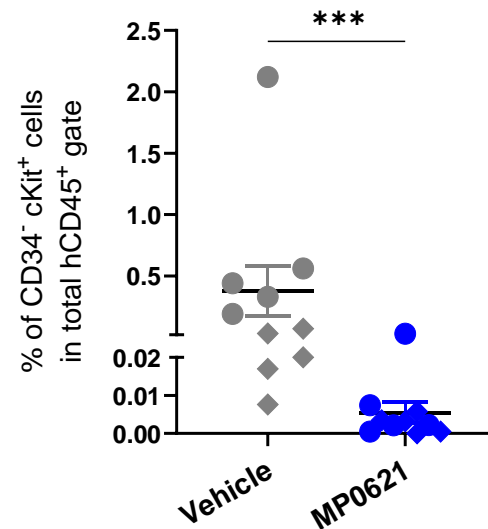
Targeted cKit⁺ cells depleted in bone marrow

Immune cells in blood

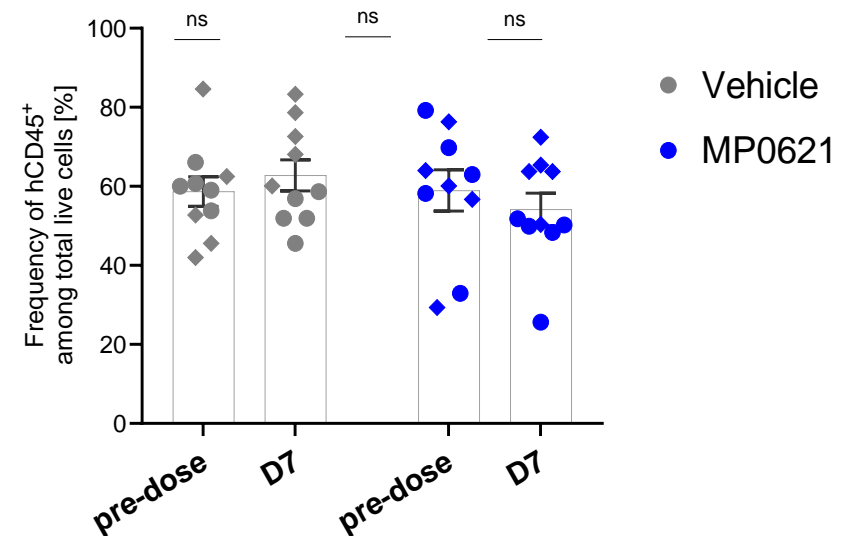
hcKit⁺ hCD34⁺ cells, incl. HSCs



hcKit⁺ hCD34⁻ cells

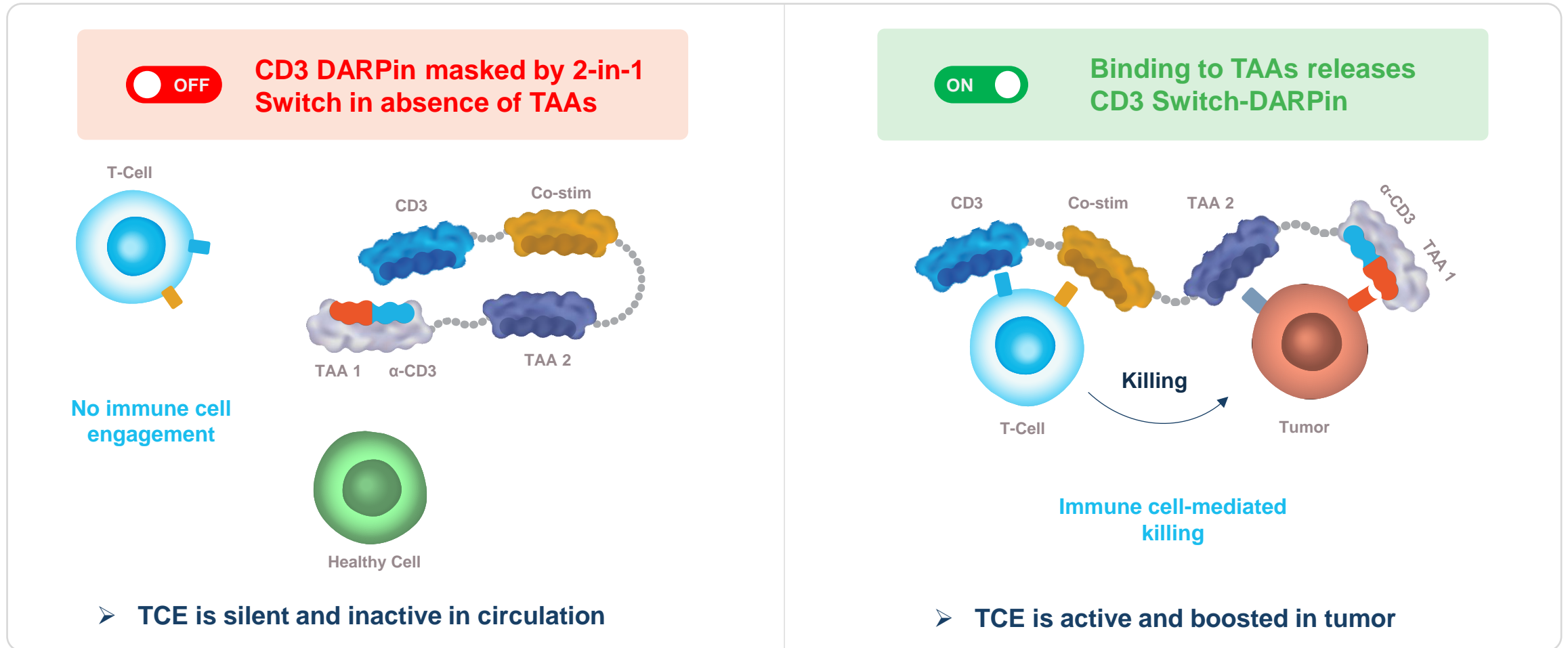


hCD45⁺ immune cells



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

Tackling current limitations of TCEs in solid tumors

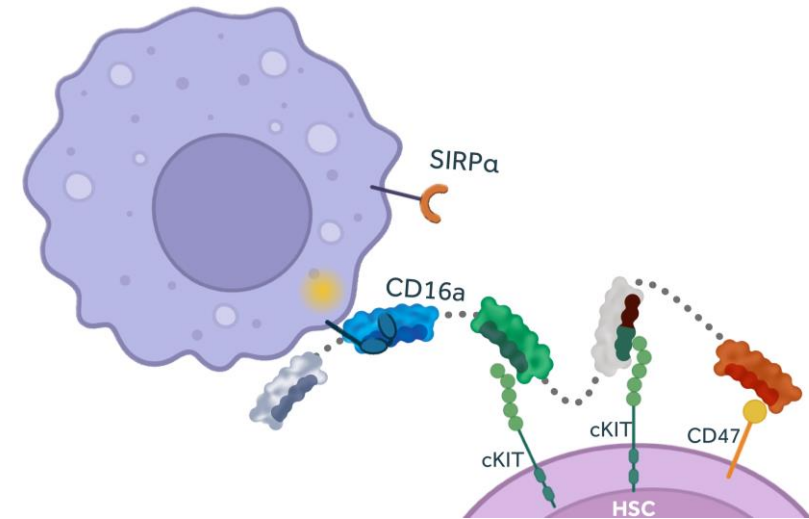


Outlook: Preclinical proof-of-concept to be presented at SITC 2024

Switch-DARPin & MP0621 – Summary

Summary

- ✓ Dual-binding DARPin (the “Switch”) provides a **logic-gated “on/off” function** to a multi-specific DARPin
- ✓ Conditional, target-specific immune activation demonstrated for **Switch-DARPin platform** *in vitro*
- ✓ MP0621: a **cKit x CD16a x CD47 Switch-DARPin** as next-gen conditioning for HSCT
- ✓ MP0621 effectively depletes targeted cells *in vivo* with a safe profile (EHA 2024)
- Introducing **CD3 Switch-DARPin as next-gen T cell engagers with enhanced function** to tackle current limitations in solid tumors



Outlook

- Update on MP0621 preclinical studies at ASH 2024
- Preclinical proof-of-concept on CD3 Switch-DARPin platform to be presented at SITC 2024



Outlook

2024 Outlook and Upcoming Milestones

Radio-DARPin Therapy (RDT) & MP0712

- Advance MP0712 into IND-enabling studies with **initial clinical data expected in 2025**
- Expand portfolio with additional **differentiated RDT programs**, update in H1 2025
- Continue to progress RDT collaborations with Orano Med and Novartis

MP0533

- Protocol being amended for both **higher & more frequent dosing** (in first weeks)
- Clinical update at ASH 2024, data on **amended dosing scheme expected in 2025**

Switch-DARPin & MP0621

- Update on MP0621 preclinical studies at **ASH 2024**
- Preclinical proof-of-concept on **CD3 Switch-DARPin platform** to be presented at SITC 2024

MP0317

- Final data from the FIH dose-escalation Phase 1 study to be presented at SITC 2024
- Clinical exploration of combinations possibly via **investigator-initiated trials**

CHF ~158 million cash* (incl. short-term time deposits) ensures **funding into 2027**



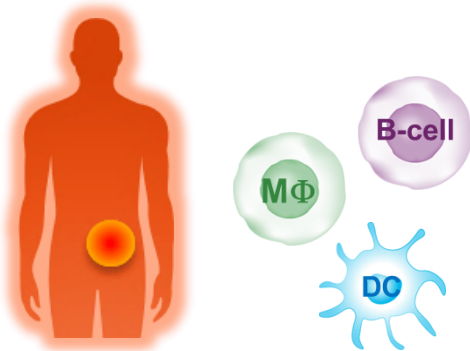
Thank You

MP0317

Tumor-localized Immunotherapy

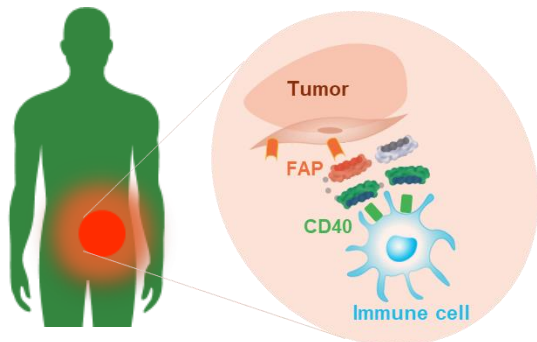
MP0317: Unlocking CD40 Activity Through Local Activation

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

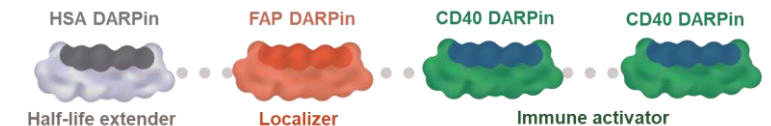


- **CD40 agonists** can activate **B cells, DCs and MΦ** to enhance the efficacy of IO drugs, especially in “cold tumors”
- **Systemic activation of CD40 via mAbs** has been hampered by **significant toxicities**, therefore **limiting their potential of reaching a therapeutically active dose**

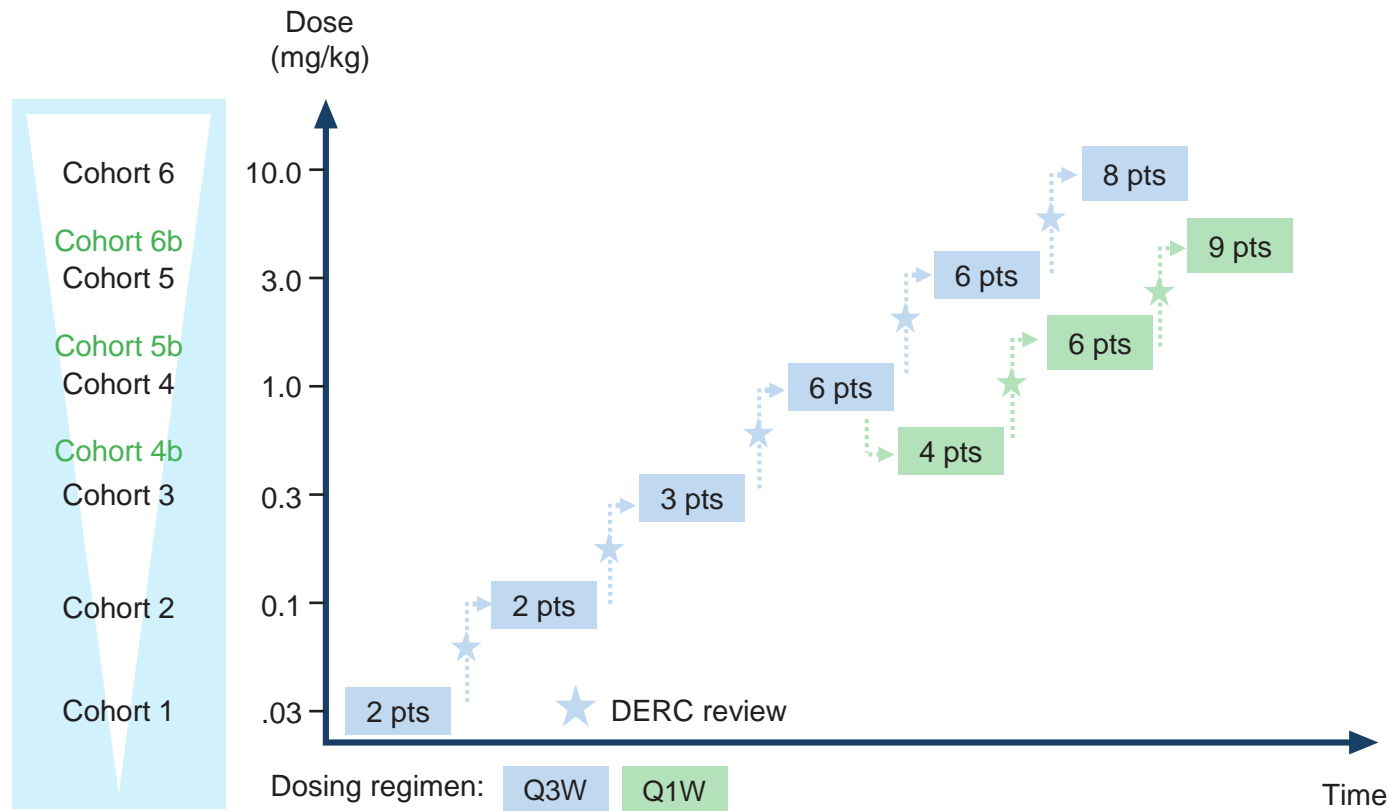
Solution: MP0317 – FAP-dependent tumor-localized CD40 activation



- **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 and Status



STUDY DESIGN

- **FIH, multi-center, dose-escalation study of MP0317 monotherapy** (9 dose cohorts; Q1W and Q3W dosing; NCT05098405)
- **Eligible patients:** adults with advanced solid tumors
- **Primary objectives:** safety/tolerability, recommended dose for expansion & combination
- **Secondary objectives:** PK, PD, and preliminary antitumor activity
- **Centers:** 4 sites in France and The Netherlands

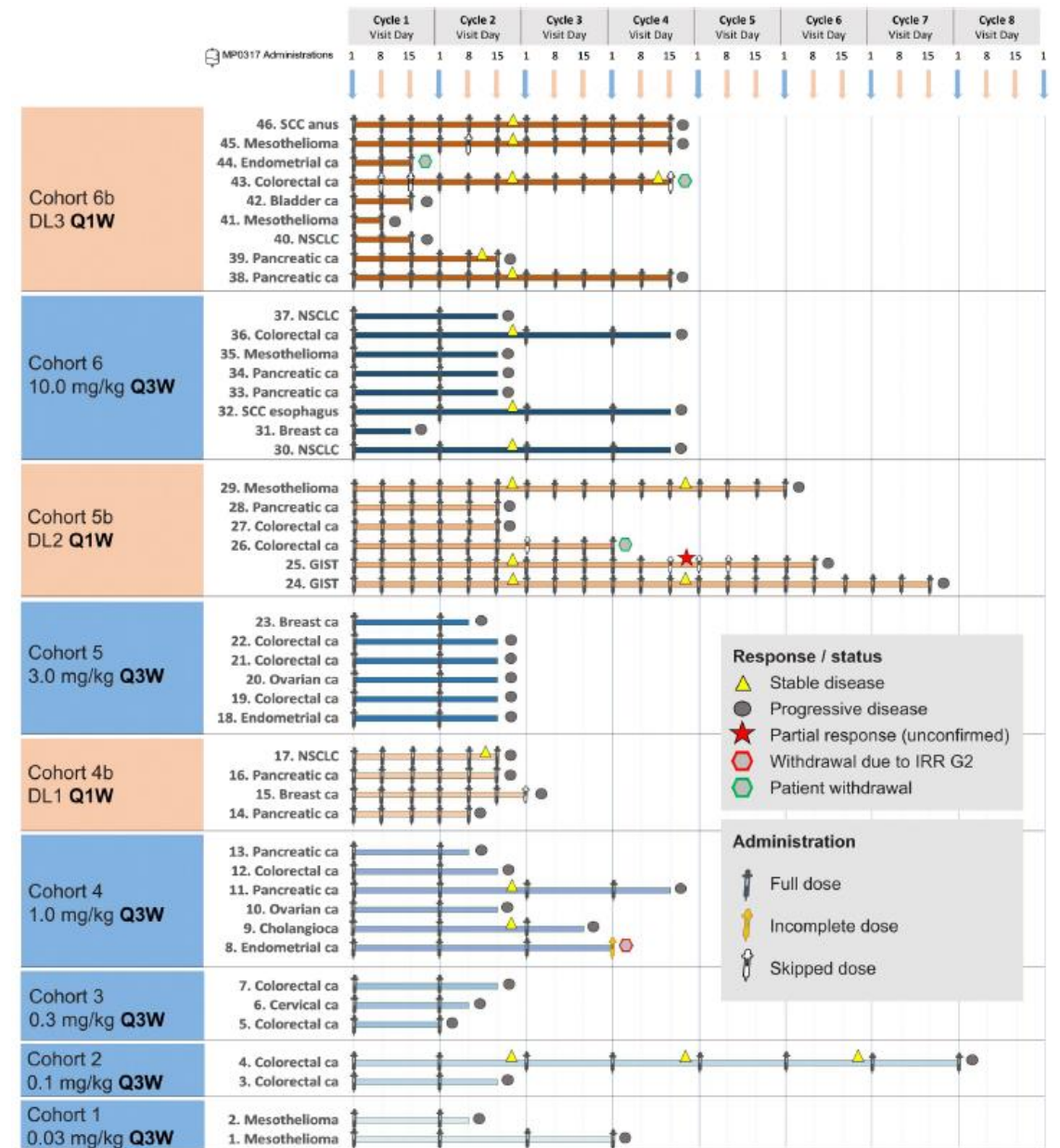
MP0317 Phase 1 Study

Summary:

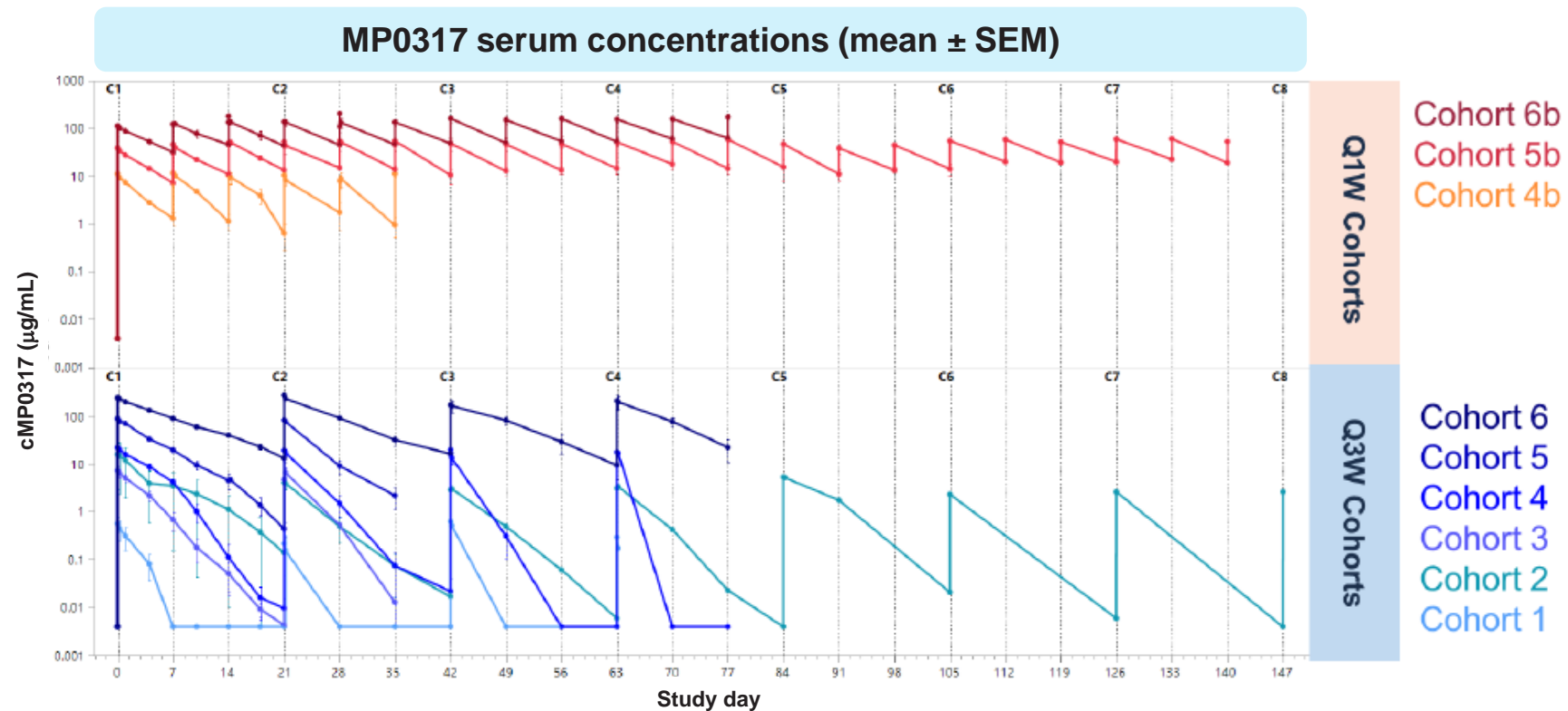
- A total of **46 patients treated** in 9 cohorts
 - Median age (range): 63 years (35–79)
 - Medial prior regimen (range): 4 (1–13)
- **Favorable safety profile** across all tested dose cohorts up to highest planned dose (10 mg/kg)
 - Only 1 patient with a DLT (cohort 6; Grade 3 AST and ALT increase)
 - Most frequent Ars: fatigue and Grade 1–2 IRRs
- **Clinical evidence** of tumor-localized CD40 pathway and immune cell activation, leading to **TME remodeling**

Outlook:

- Final data to be presented at **SITC 2024**
- Clinical exploration of combinations possibly via **investigator-initiated trials**



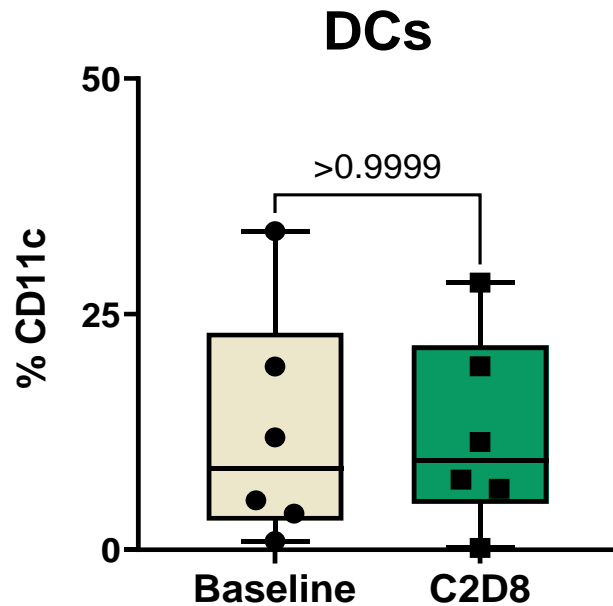
MP0317 Serum PK is Suitable for Q3 and Q1 dosing



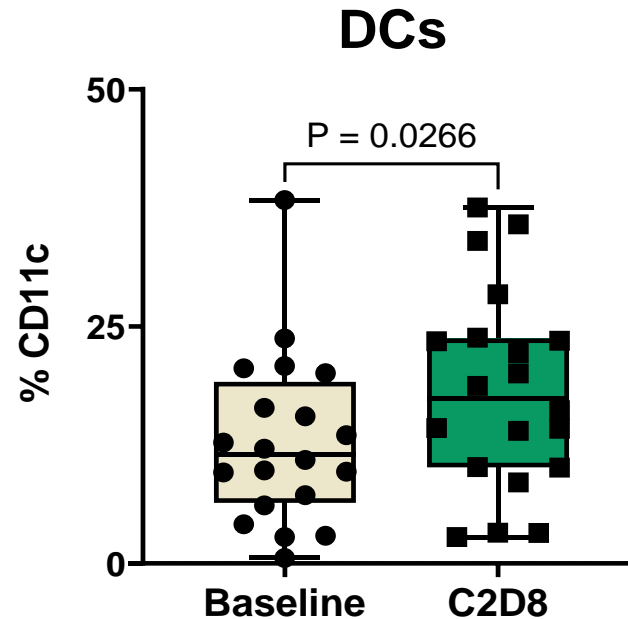
- PK profile is consistent with **half-life extended properties of DARPins**
- **MP0317 exposure shows dose-proportionality** throughout the treatment period analyzed
- **Sustained exposure** is observed at higher doses with both regimens overcoming TMDD and the impact of ADAs

MP0317 Tumor-localized CD40 Activation and TME Modulation

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



MP0317 higher doses and detected in tumor (n=20)



Evaluable paired tumor biopsies from treated patients were analyzed with mIF. 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.

- Bulk RNA sequencing in paired tumor biopsies (n=19) shows that MP0317 presence tends to be associated with:
 - Increase in abundance of plasma and T follicular helper cells
 - DC maturation gene signature
 - IFN γ downstream activation gene signature scores
- Increases observed in CXCL10 serum levels corroborate these findings