



Custom Built Biology for Patients

Corporate Presentation

Kempen Conference

April 2022

Molecular Partners AG, Switzerland
(SIX: MOLN, NASDAQ: MOLN)



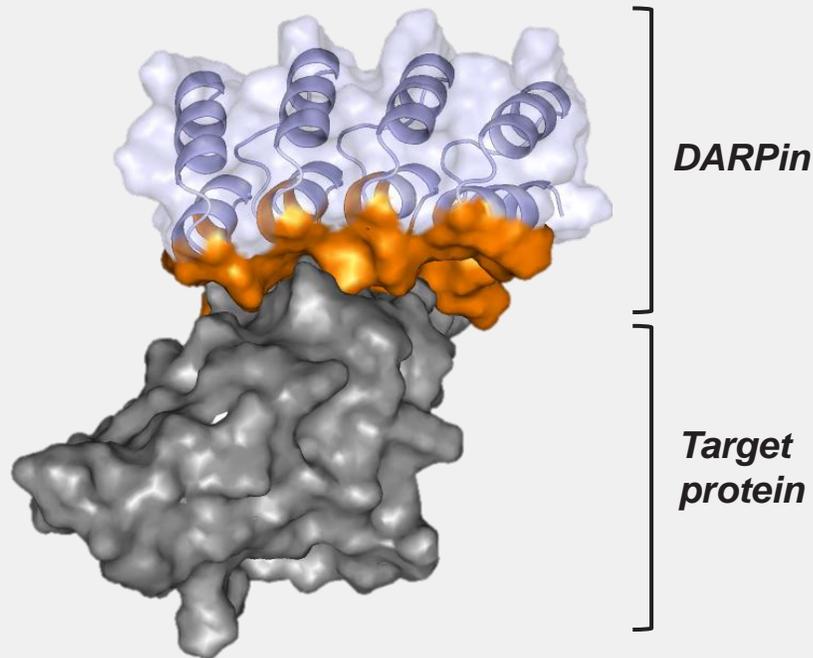
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DARPin: The Core of our Drug Engine

DARPin are binding proteins derived from natural ankyrin repeat proteins



DARPin KEY PROPERTIES

DARPin ADVANTAGE



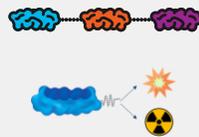
Small size
(15 kDa)

- Deep tissue penetration
- High molar concentration



Rigid protein
scaffold

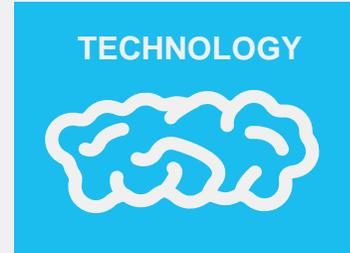
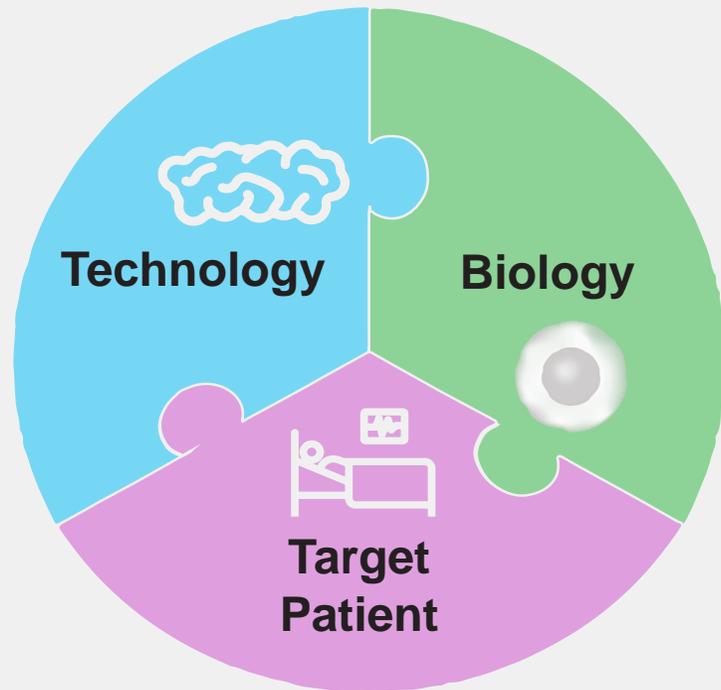
- Ultra-high binding affinity and selectivity



Simple & robust
architecture

- Turn-key multispecifics
- Easy coupling of payloads

MP Strategy – building on our Strengths



We leverage the advantages of the **DARPin technology** to provide unique solutions to impact biology and bring value to patients



Our candidates' design aims to **directly change the course of disease biology** and allow testing in a model with **high translatable value**



We aim to drive **true patient value** with **early clinical read-outs**



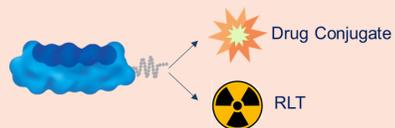
We strive to **collaborate** with the best scientists and clinicians in the field from ideation to clinical trials

Translating DARPin Properties into Differentiated Therapeutics

Delivery Vectors “Radical Simplicity”

RLT & DDC

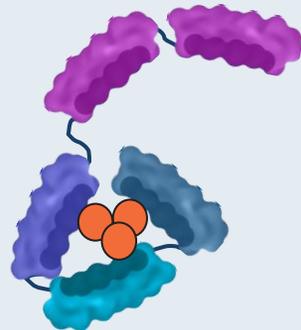
Small size: high affinity delivery, limited systemic exposure



Multispecificity enabled possibilities

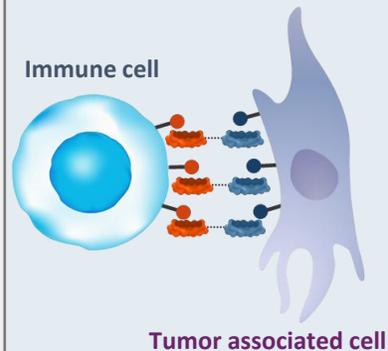
Ensovibep

Cooperative binding to inhibit SARS-Cov-2 and prevent escape



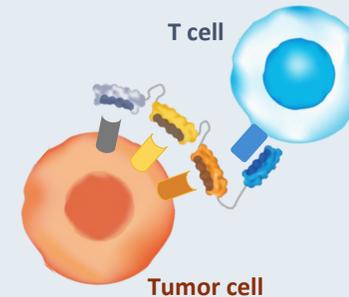
MP0310 & MP0317

Tumor localized clustering activates effector cells in tumor



MP0533

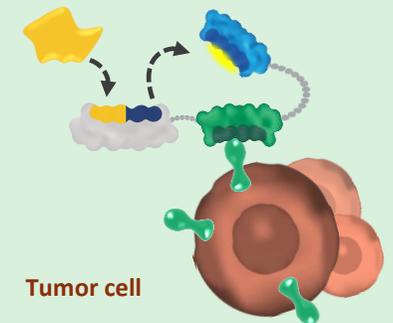
Avidity driven TCE for tumor specificity and heterogeneity



Conditional activation “Radical Complexity”

SWITCH

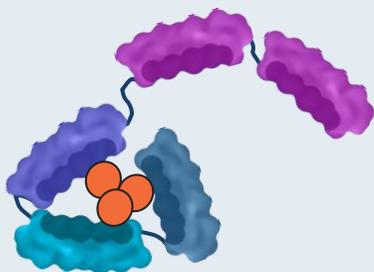
Programming highly potent effectors to omit off-tumor activity



Synergistic Partnerships Built on a Versatile Drug Class

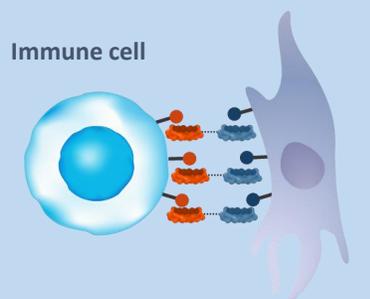
Ensovibep

- Leverage production, global development and distribution of Novartis for ensovibep
- CHF 60 million upfront
- CHF 150 million received upon option exercise
- 22% royalty on sales



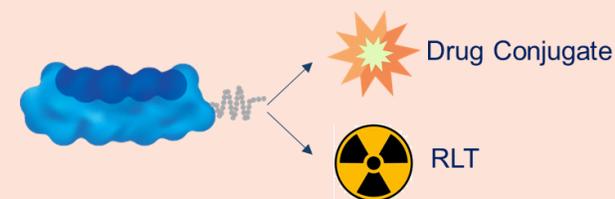
AMG 506 / MP0310

- Partnership with Amgen to combine AMG 506 / MP0310 with BiTE[®] molecules
- Phase 1 conducted by MP to develop for combination studies
- \$50 million upfront
- ~\$500m in milestones and mid teen royalties



Radioligand therapeutics

- Combining small size and high affinity and specificity of DARPins with Novartis' radioligand expertise
- 2 cancer antigen targets
- \$20 million upfront
- Up to \$560 million and low double-digit royalties



Pipeline



Pipeline						
CANDIDATE / FOCUS	RESEARCH	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	RIGHTS
Ensovibep – Covid	Covid ambulatory – EMPATHY					NOVARTIS MOLECULAR partners
Next Gen Covid	Future VoC*					
AMG506 / MP0310 FAP x 4-1BB	Solid tumors					AMGEN
MP0317 FAP x CD40	Solid tumors					MOLECULAR partners
MP0533 CD3 x CD33+CD70+CD123	AML					MOLECULAR partners
Abicipar VEGF	wet AMD – CEDAR & SEQUOIA					MOLECULAR partners
Radio Ligand Therapy	Solid tumors					NOVARTIS

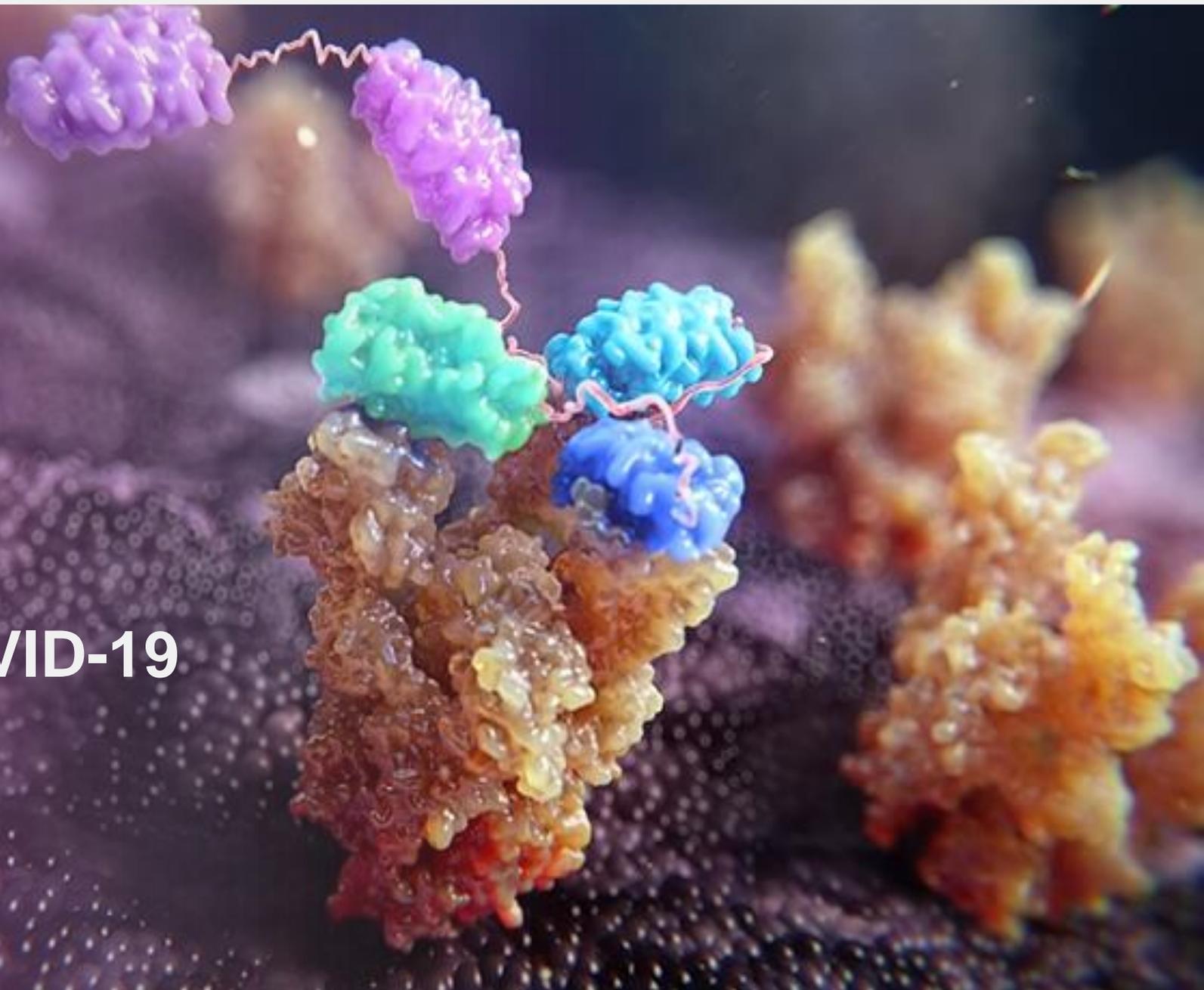
Platform Discovery

Radical simplicity & Conditional Activation	MOLECULAR partners
Additional Infectious Diseases	



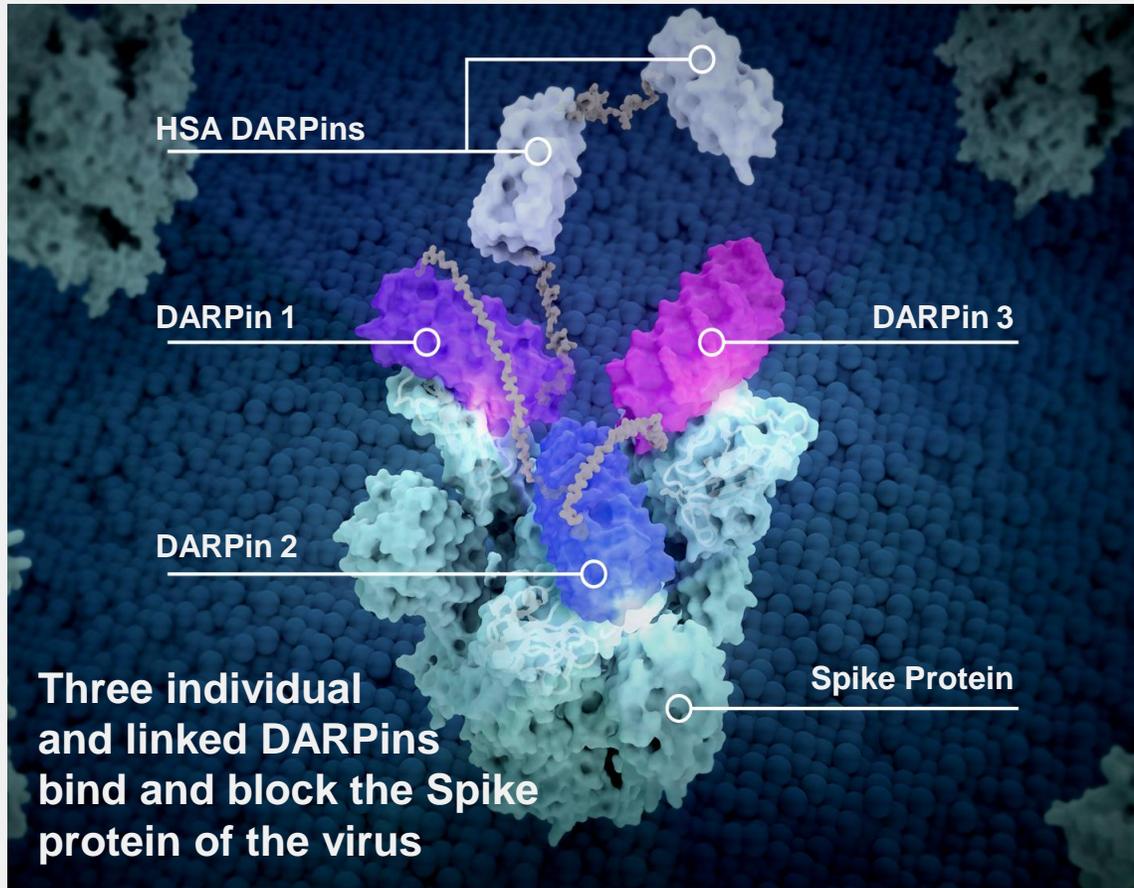
Ensovibep:

**Advancement of COVID-19
Clinical Program**



Structure and Features of Ensovibep Neutralizing the SARS-CoV-2 Spike Protein

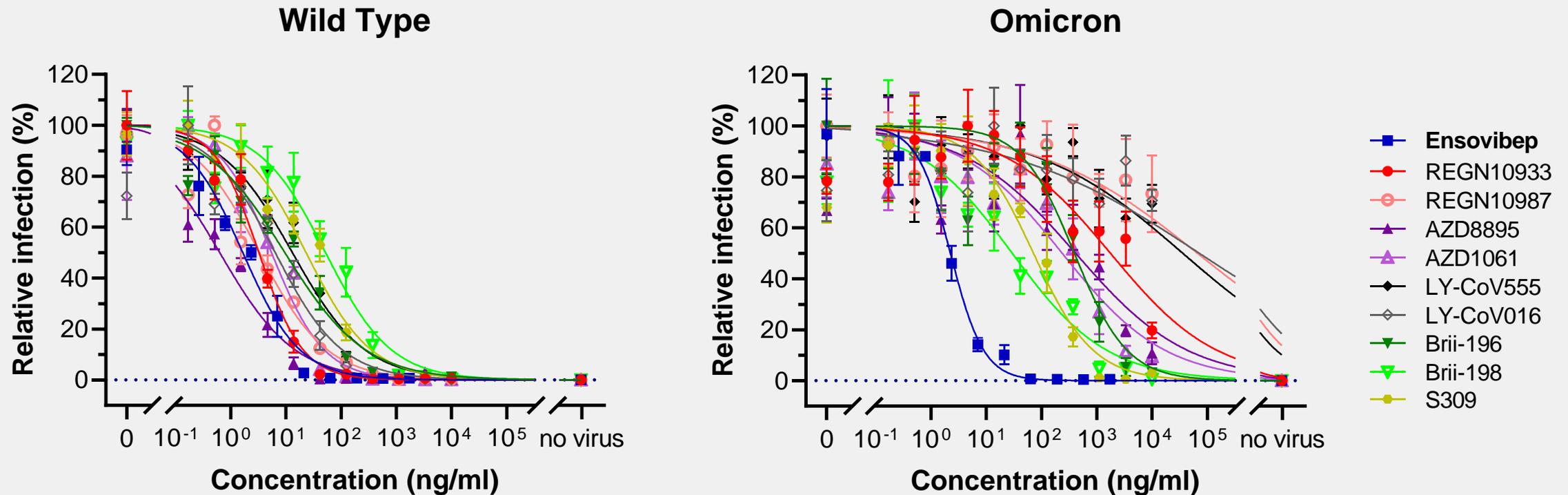
3D model of a DARPin molecule



Characteristics

- High potency: high binding affinity and avidity leads to one of the highest anti-viral potencies reported to date
- Pan variant activity: cooperative binding of different sites allows blocking of all described variants of concern, so far
- Simple administration: long-half life, high solubility and low dose activity can allow for single administration via i.v., i.v. bolus, or s.c. injection
- Supply: microbial manufacturing in *E.Coli*

Ensovibep Retains Full Activity Against Omicron



EMPATHY Part A (Phase 2) Clinical Design and Endpoints

Objective	Demonstrate superiority of ensovibep, compared to placebo, in reducing SARS-CoV-2 viral load through Day 8 and select a dose for Phase 3
Population	<ul style="list-style-type: none"> • Ambulatory symptomatic patients diagnosed with COVID-19 • Onset of symptoms within 7 days prior to dosing • Positive Rapid Antigen Test on the day of dosing • Vaccinated patients allowed
Primary Endpoint	• Time-weighted viral load reduction through through Day 8
Key Secondary Endpoints	<ul style="list-style-type: none"> • Reduction in ER visits and/or hospitalizations (≥ 24 hours) and/or death up to Day 29 • Time to sustained clinical recovery (resolution or improvement in clinical symptoms) up to Day 29

Cohorts

ensovibep 600mg i.v. ~ 100 pts

ensovibep 225mg i.v. ~ 100 pts

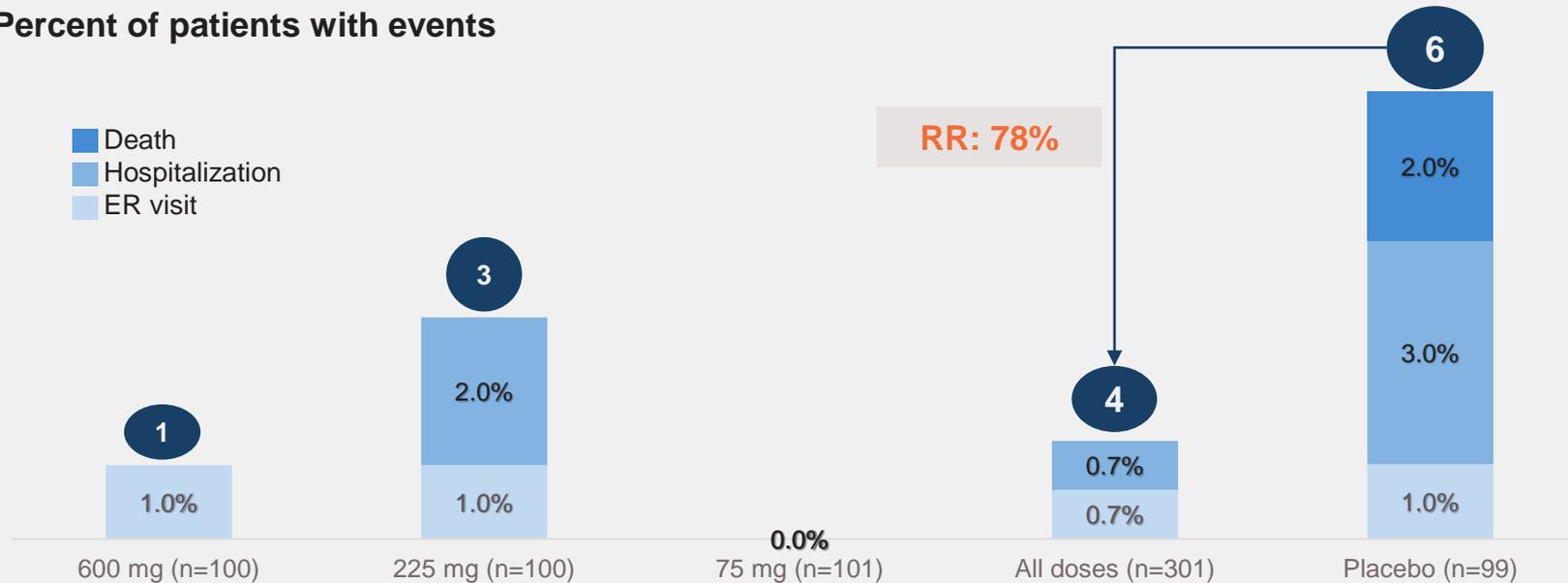
ensovibep 75mg i.v. ~ 100 pts

placebo i.v. ~ 100 pts

EUA Submission Supported by Secondary Endpoint in Reductions in Hospitalization and/or ER Visit, or Death

Patients with hospitalization and/or ER visit related to COVID-19 or death

Percent of patients with events



Numbers indicate absolute number of patients

Note:

In the hierarchy of ER-visit/hospitalization/death- patients are counted in the highest category

- ER visits exclude those resulting in hospitalization/ death
- Hospitalizations exclude those that resulted in death

Novartis Deal Terms and Next Steps



Deal Terms

- Novartis option exercise for in-licensing of ensovibep: CHF 150m
 - CHF 60m previously received at signing of option agreement (20m cash/40m MOLN shares)
- Royalty of 22% on sales in commercial countries
 - Molecular Partners has agreed to forgo royalties in lower income countries and is aligned with Novartis' plans to ensure affordability based on countries' needs and capabilities.

Next Steps

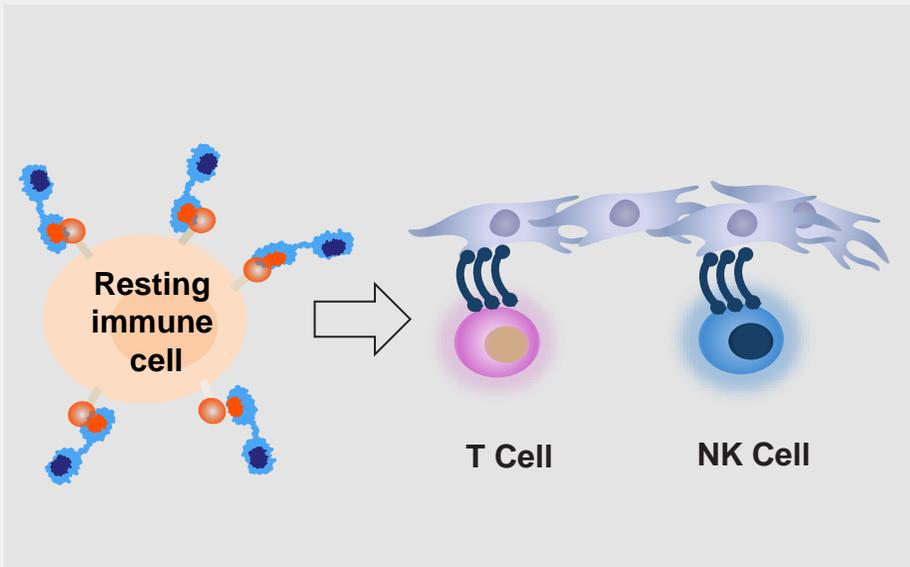
- EUA submitted and review ongoing
- Discussion with appropriate federal agencies regarding supply agreements of ensovibep
- Phase 3 initiation
- Planned initiation of subcutaneous Phase 2/3 study (led by Novartis)



MP0310 and MP0317

Multispecific Immune Activators

AMG 506/MP0310: Localized Activation of 4-1BB



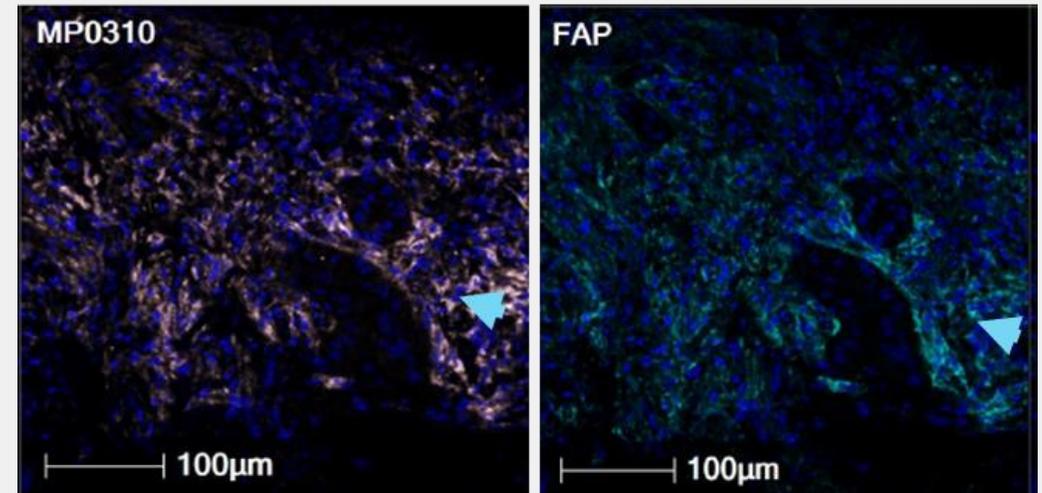
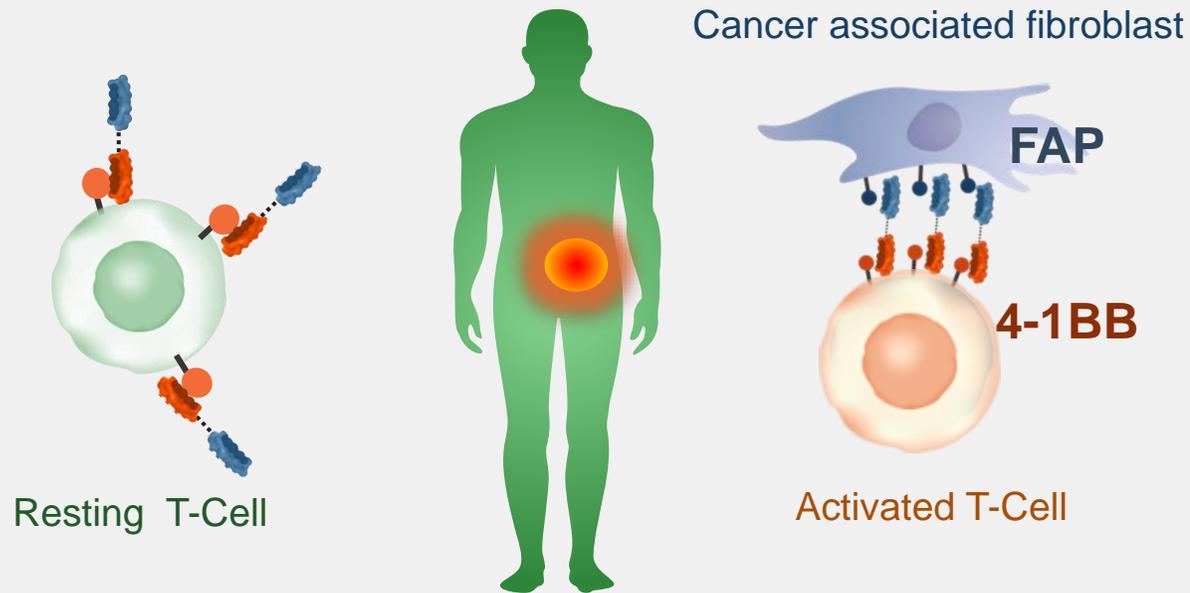
- **Good safety profile without major systemic toxicity**
 - No liver toxicity or systemic activation of immune cells
 - IRRs frequent but manageable
- MP0310 is observed in tumor tissue
- Tumor biopsies show tumor-localized immune response consistent with the MoA
- **Presently investigating appropriate dosing schedule for sustained activity**
- \$50m upfront, ~\$500m in milestones plus royalties

AMG 506 / MP0310: Localized Activation of 4-1BB FAP – an Ideal Target for Tumor-localized Activity



- Immune-cell activation via 4-1BB is associated with liver tox
- MP0310/AMG506 is designed to activate immune cells in the tumor only via FAP clustering

MP0310 & FAP staining in human biopsies from Phase 1 trial

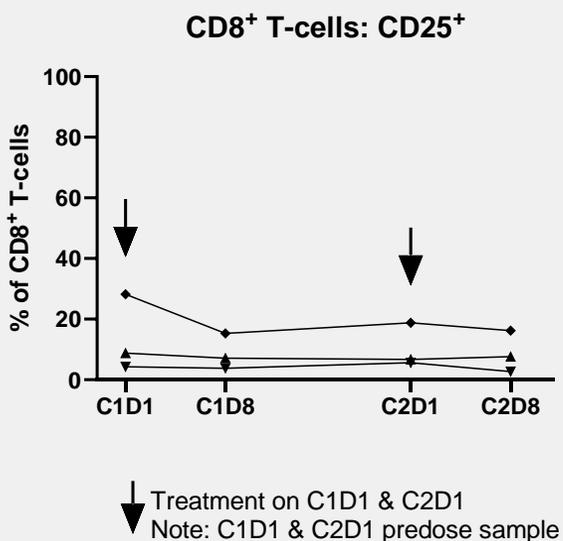


- When dosed systemically, MP0310 binds to and co-localizes with FAP

PD Activity in Paired Biopsies Supports AMG 506 / MP0310 MoA on 4-1BB Activation

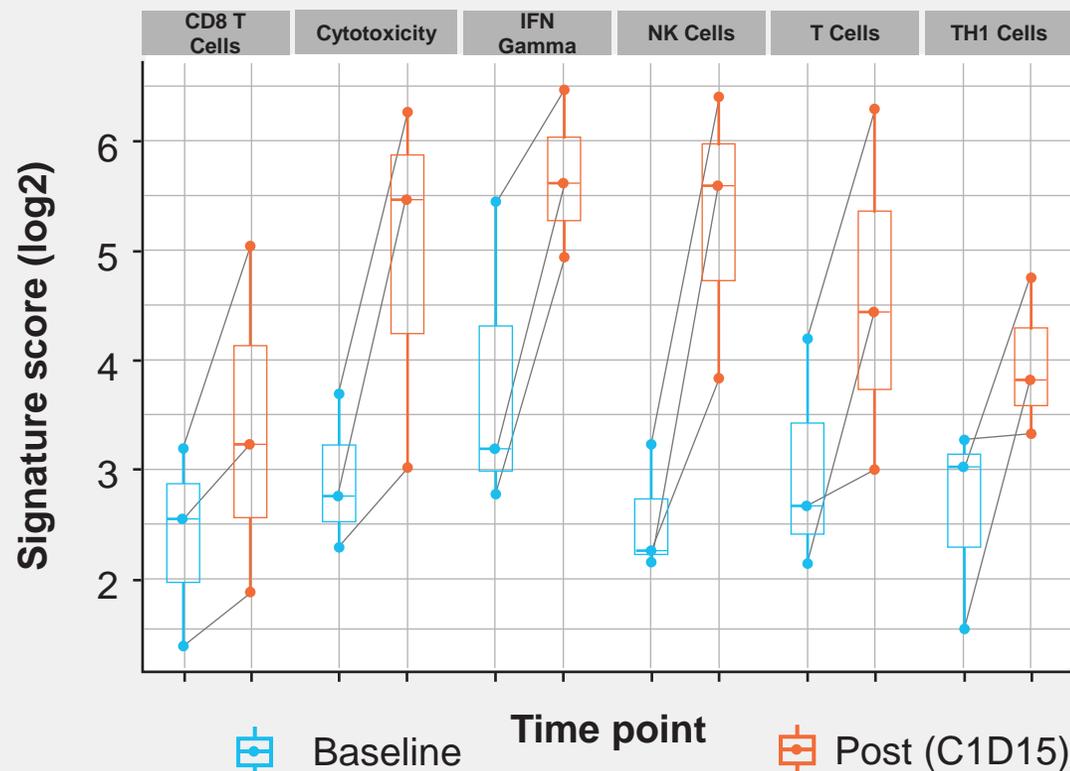


BLOOD



- In the blood, immune cells remain inactive (CD8⁺ & CD4⁺ T-cells, Treg, NKT, B-cells, NK)

TUMOR



- In the tumor, T-cells and NK cells are activated

MP0317: Localized Activation of CD40



Target Patient



- Solid tumor patients with positive FAP expression
- Many patients still fail to benefit from current immunotherapy options, or relapse

Disease Biology



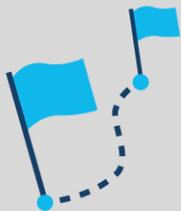
- CD40 is a potent activator of dendritic cells, macrophages, and B cells, and has long been considered an attractive immunotherapy target
- Prior attempts at targeting CD40 have shown anti-tumor activity but remain hampered by toxicity issues

DARPin Advantage



- MP0317 is designed to activate CD40 in a context dependent manner, by anchoring to FAP and activating via clustering
- Preclinical data show local activation of immune cells while limiting off target toxicity

Expected Milestones



- **FIH studies initiated in Q4 2021**
- **Initial data anticipated in H2 2022**
- Rapidly explore expansion arms in phase 1b

Summary: Multispecific Immune Activators



AMG 506 (MP0310)

- Phase 1 mechanistic POC established
 - Demonstrating localized activity of 4-1BB in the TME
 - No elevated liver enzymes or systemic tox identified
- Ideal program for T-cell engagers (CD3) and potentially PD-1 checkpoint inhibition
- Ongoing weekly dosing will determine optimal go-forward treatment regimen
- Data available for Amgen evaluation in Q3

MP0317

- Following the same concept of activation by clustering to FAP
- MP0317 evaluating if a safer administration of CD40 can be therapeutically beneficial to patients
- CD40 stimulation would allow for multiple combination treatments
- Phase 1 ongoing, initial FIH data available later 2022



**MP0533:
Tri-specific T-cell
Engager for AML**

MP0533: Tri-specific T-cell Engager for AML

Target Patient



- ~20,000 people are diagnosed with AML every year
- Over 50% of patients die in the first year
- High relapse rates

Disease Biology



- Persistence of LSCs* is the driver of relapse
- Tumor antigens in AML are also found on healthy cells
- On-target toxicity (not clean targets) limit current T-cell engager approaches

DARPin Advantage



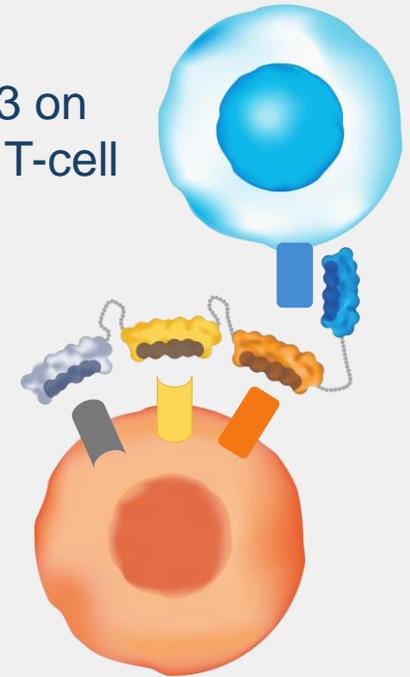
- Avidity driven multispecific DARPin to target LSCs
- T cell are activated only when 2 or more TAA's** are bound
- Open therapeutic window for “difficult” targets

Expected Milestones



- FIH clinical studies initiating in late 2022

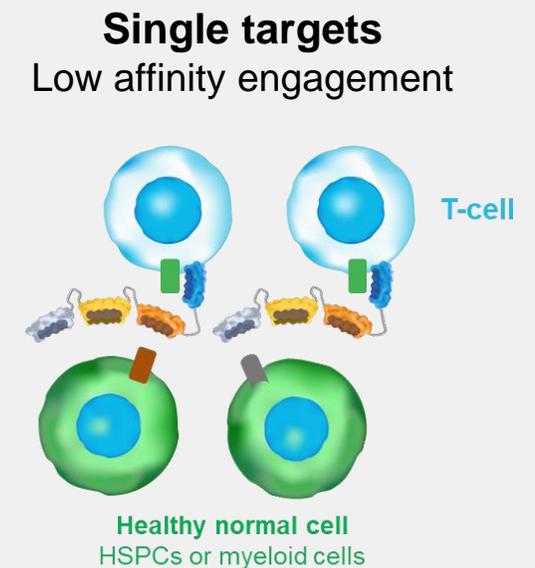
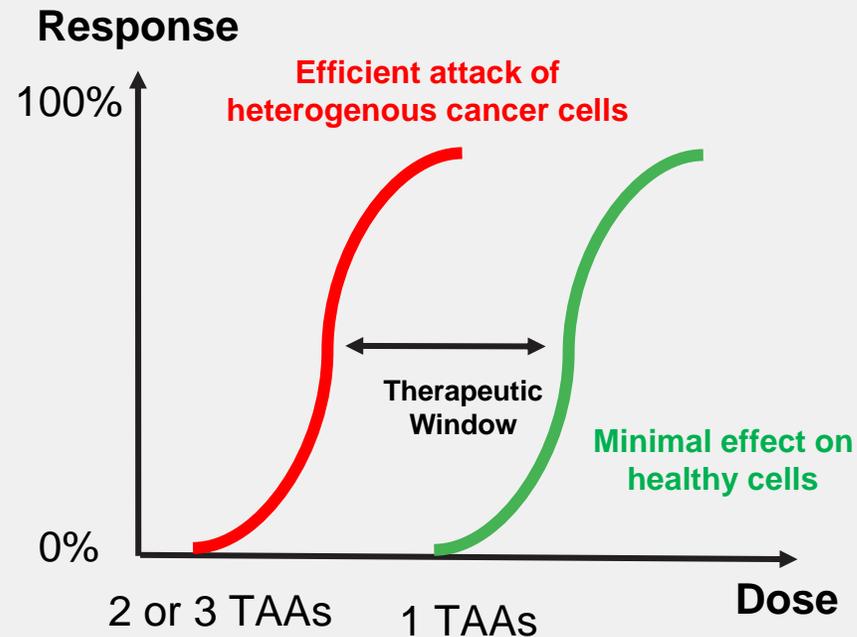
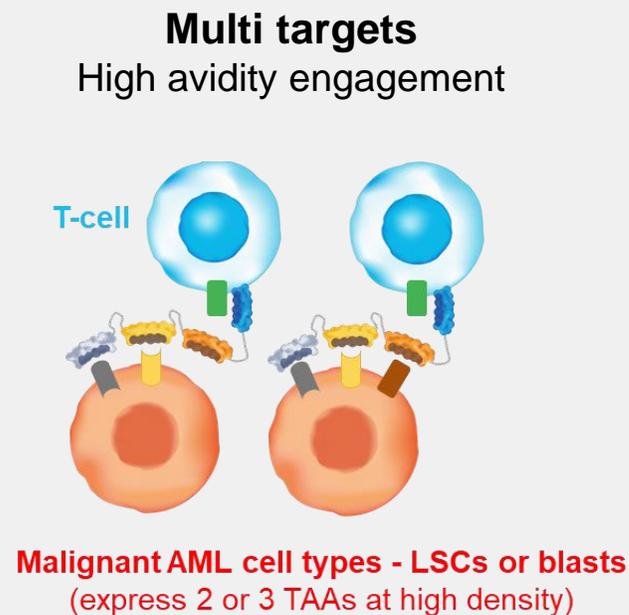
CD3 on
the T-cell



CD33, CD70, CD123
on the AML blast or LSC

Unlock the value of “not-clean” targets to kill Leukemic Stem Cells and blasts in AML

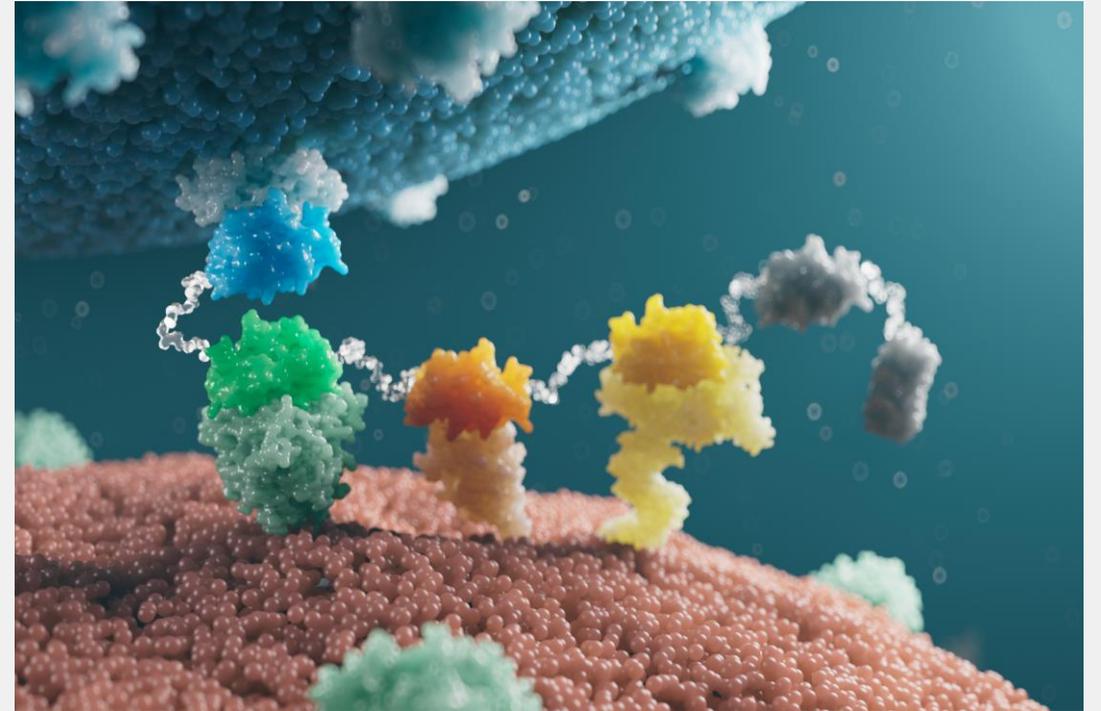
- Persistence of LSCs is the driver of relapse in AML
- Targets in AML are also on healthy cells, leading to on-target toxicity (not clean targets)
- Goal: avidity-driven killing of LSCs and blast, while less killing of HSPs



MP0533: a Unique DARPin Solution for AML Patients

- **Properties of an ideal AML drug:**
 - Ensure long term control of the disease by eliminating LSCs ✓
 - Control tumor heterogeneity by targeting multiple Ag ✓
 - Increase the therapeutic window ✓
 - Limited killing of healthy HSCs
 - Reduced CRS

➤ **Phase 1 clinical trial initiation H2 2022**





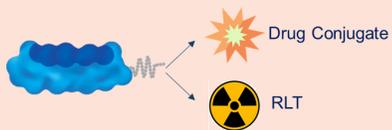
Research Activities

Translating DARPin Properties into Differentiated Therapeutics

Delivery Vectors “Radical Simplicity”

RLT & DDC

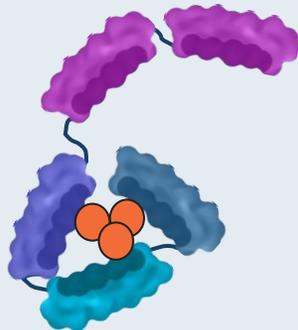
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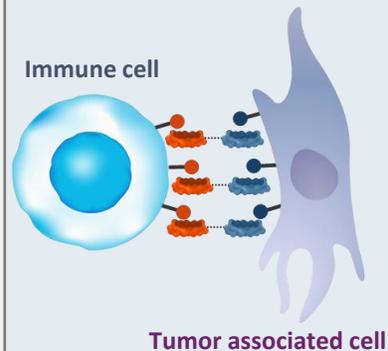
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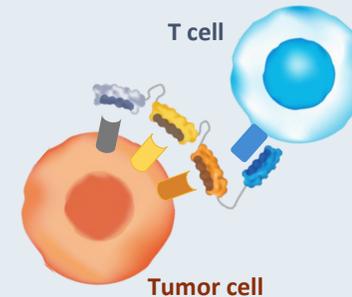
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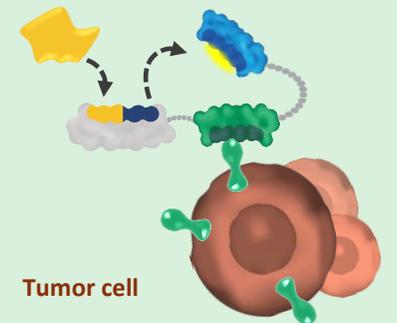
Avidity driven TCE for tumor specificity and heterogeneity



Conditional activation “Radical Complexity”

SWITCH

Programming highly potent effectors to omit off-tumor activity





Summary & Outlook

Financial Guidance for Full-Year 2022

- Total expenses of CHF 75-85 million, of which around CHF 8 million non-cash effective costs
- Total cash as of February 28, 2022 – CHF 291.3 million, which include funds received from Novartis in January 2022
- The Company is funded into 2025, excluding any potential payments from R&D partners. Guidance subject to progress and changes of pipeline

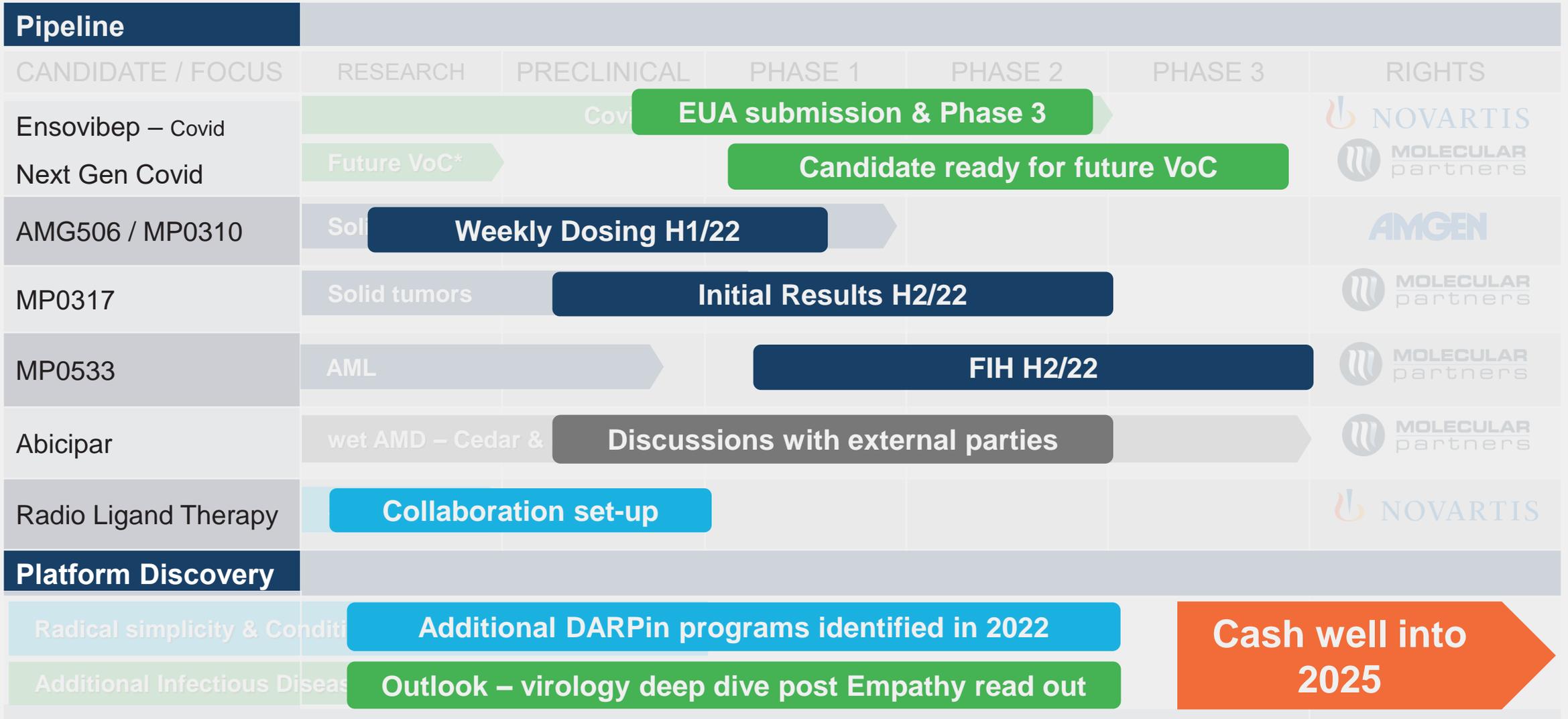
Pipeline Inflection Points



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Next Gen Covid	Future VoC*					
AMG506 / MP0310 FAP x 4-1BB	Solid tumors					AMGEN
MP0317 FAP x CD40	Solid tumors					MOLECULAR partners
MP0533 CD3 x CD33+CD70+CD123	AML					MOLECULAR partners
Abicipar VEGF	wet AMD – Cedar & Sequoia					MOLECULAR partners
Radio Ligand Therapy	Solid tumors					NOVARTIS

Platform Discovery						
Radical simplicity & Conditional Activation						MOLECULAR partners
Additional Infectious Diseases						

Pipeline Inflection Points





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