

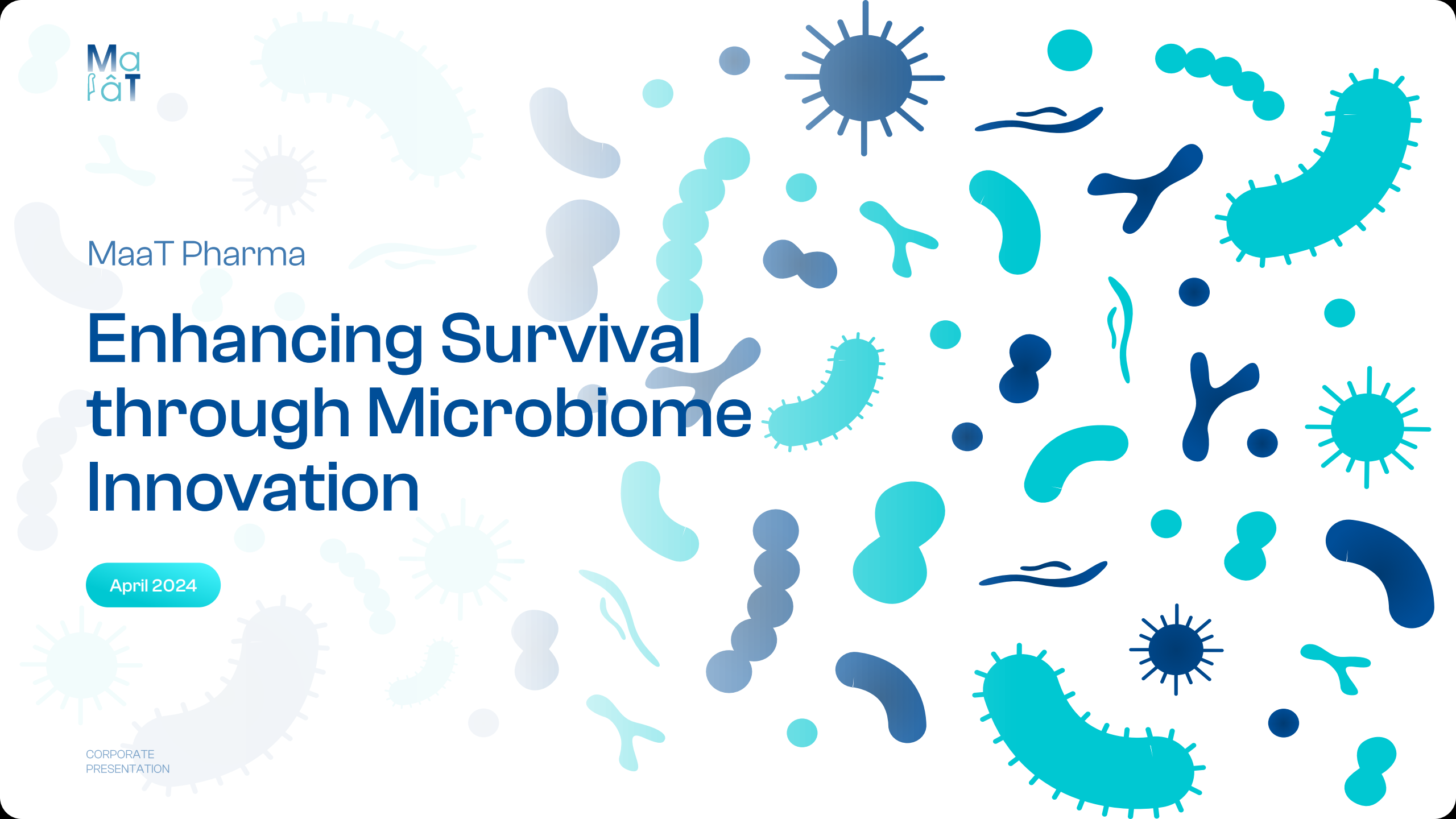
MaaT

MaaT Pharma

# Enhancing Survival through Microbiome Innovation

April 2024

CORPORATE  
PRESENTATION



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# Late-Stage Clinical Biotech, Leading the Way in Microbiome Therapies in Oncology



## MaaT013 in phase 3 in aGvHD

- Lead asset MaaT013 in **Phase 3 in aGvHD in Europe, expecting primary endpoint readout in mid Q4**
- Strong data from Early Access Program** published in April (1y OS 49% vs 15% historical data)
- US IND Open** – Readiness Phase before launch ongoing



## Deep oncology pipeline

- Donor-derived** and **co-culture** platforms **driving candidate development** with **2 clinical** and 1 preclinical assets
- Largest European cGMP** production facilities for microbiome ecosystem therapies
- Predictive AI-engine **gutPrint®**



## Finance



- Revenues** from **of MaaT013 in aGVHD of 2.2m€ for 2023 from Early Access Program**
- Cash position of 24.3m€** as of 23/12/31 & **Cash runway** to the **end of Q3 2024**
- Strong institutional shareholder base**



# Host – Microbiota Interactions are Critical for a Functional Immune System

A rich and diversified gut ecosystem actively modulates the immune system functionality



01

A diversified microbiome contributes to the **education and modulation of our immune system** throughout life

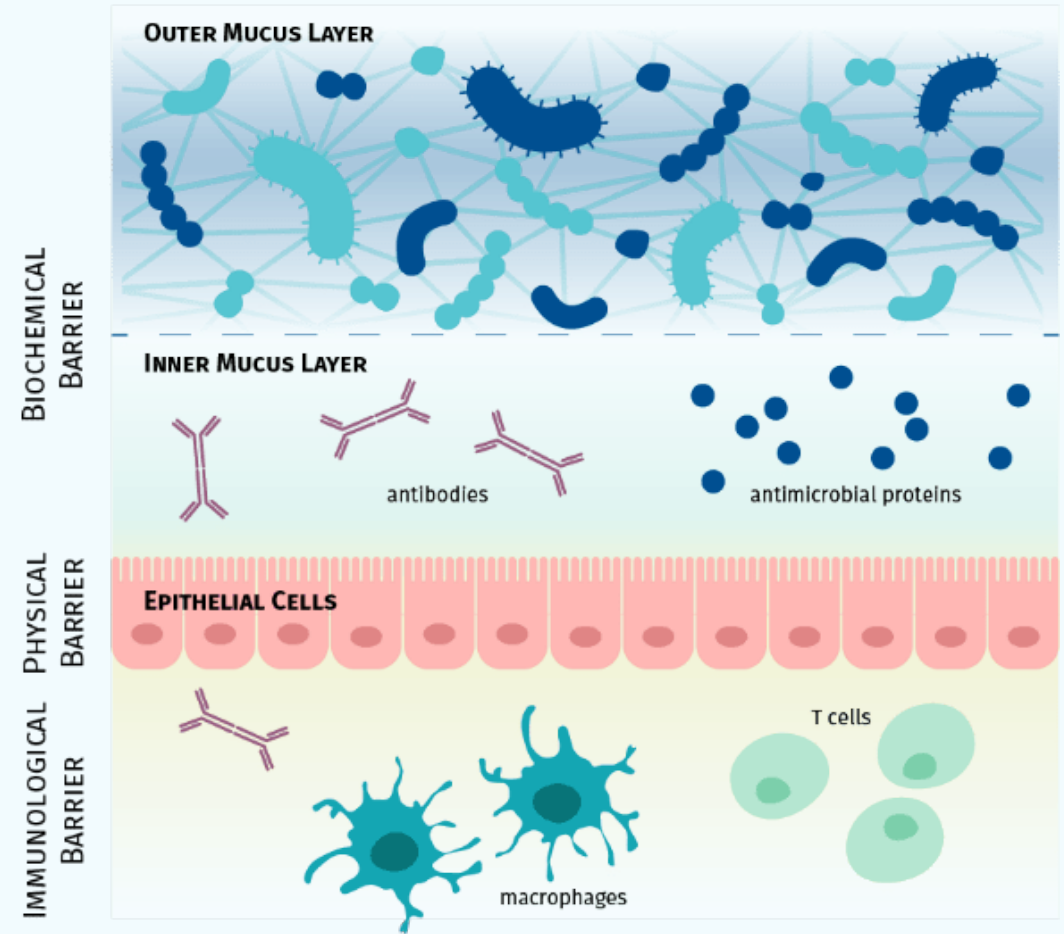


02

Bacterial **richness** and mucus layer prevent colonization by pathogens and improve gut barrier

80%

Cellular host defense localized in the gut



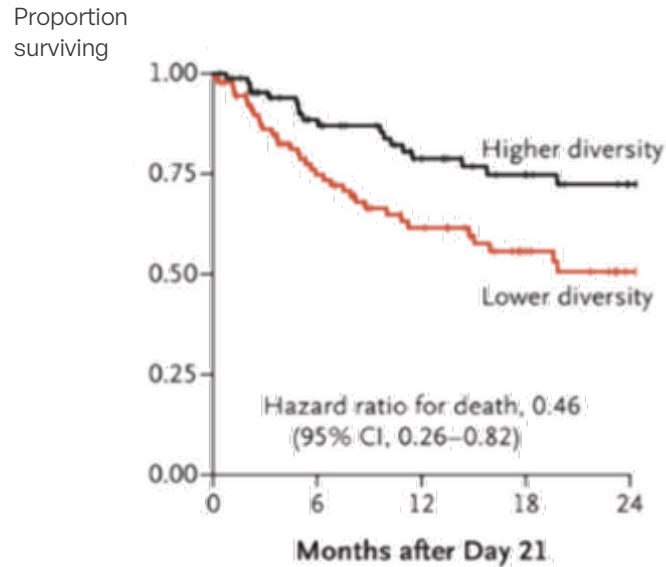
Cross-section of a healthy gut

# In Oncology, a Higher Gut Microbiome Diversity is Associated with Increased Survival<sup>1</sup>

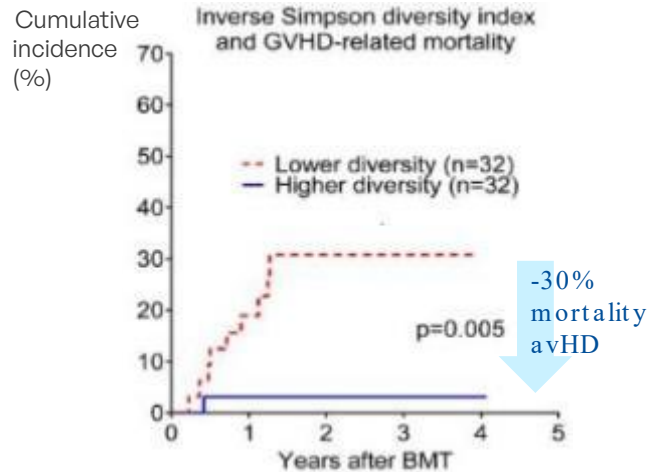
## Liquid Tumors

### Higher survival rate in patients receiving allo-HSCT\*<sup>1</sup>

Overall Survival – Cohort 2



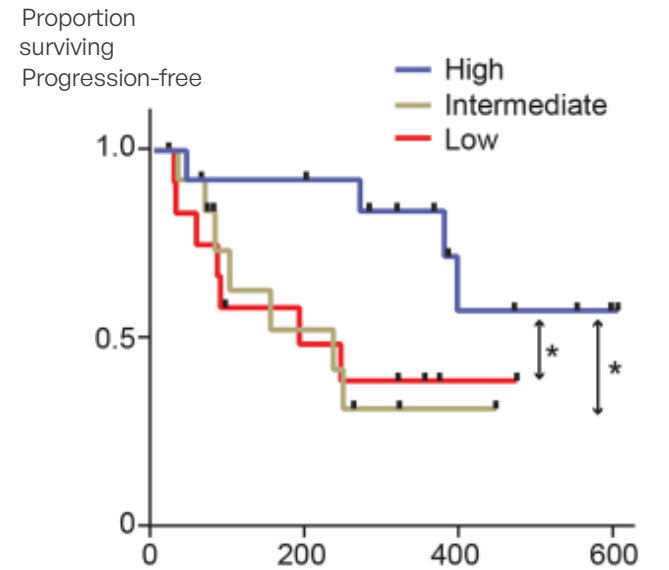
### Lower incidence and lower mortality from aGvHD\*<sup>2</sup>



MaaT Pharma MET  
Inverse Simpson (mean): **24**

## Solid Tumors

### Higher response rate to ICI\* in patients with metastatic melanoma<sup>3</sup>



\* allo-HSCT: allogeneic hematopoietic stem cell transplantation; aGvHD: acute Graft-vs-host-Disease; ICI: Immune Checkpoint Inhibitors

<sup>1</sup> Peled, J.U. & al N Engl J Med 2020;382:822-34; <sup>2</sup> Ghani, 2021; Jenq RR. et al, Biol Blood Marrow Transplant 21 (2015) 1373e1383; Pamer, Blood, 2014; <sup>3</sup> Gopalakrishnan et al., Science, 2017, see also Routy et al, Science, 2018; Vetizou et al Science 2015;

# An Oncology Microbiome Platform Fueling a Deep Pipeline of Drug Candidates



Driving near-term value with the donor-derived MET-N platform



MaaT013



MaaT033



POOLING



MaaT013



MaaT033

Pooled microbiota

→ Maximized richness

→ Standardized richness (450 OTU ± 3%)

Progressing next-generation co-cultured scalable MET-C platform



MaaT034



MaaT03X

Original microbial ecosystem



Master bank



Working Bank



Unlimited Co-Culture Scaling

MET-C product

Multistep co-culture cGMP proprietary process

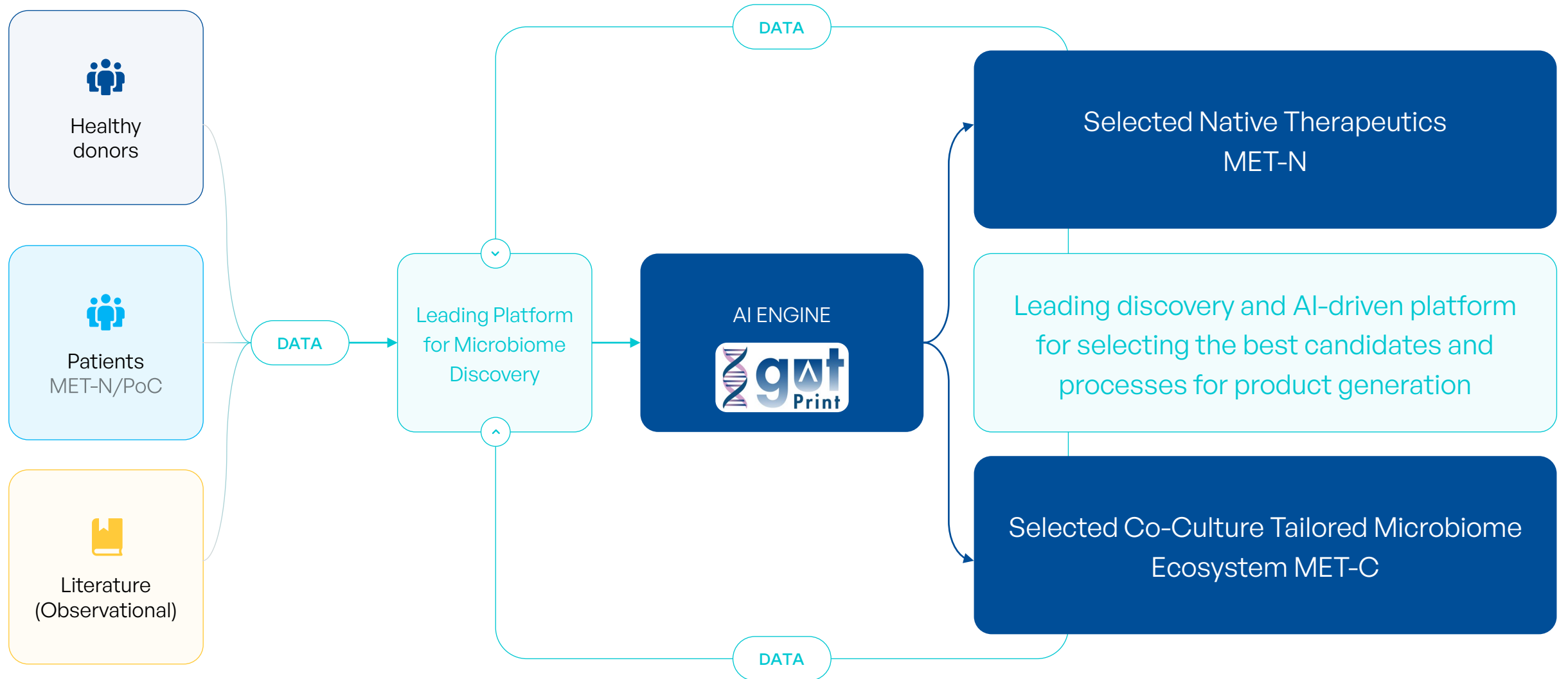
Leading capabilities in microbiome drug production



~10 000 treatable patients per year



# AI-driven Research Engine Powered by Metagenomics Enabling Candidate Selection



# A Strong Pipeline With Multiple Near-Term Value Inflection Milestones

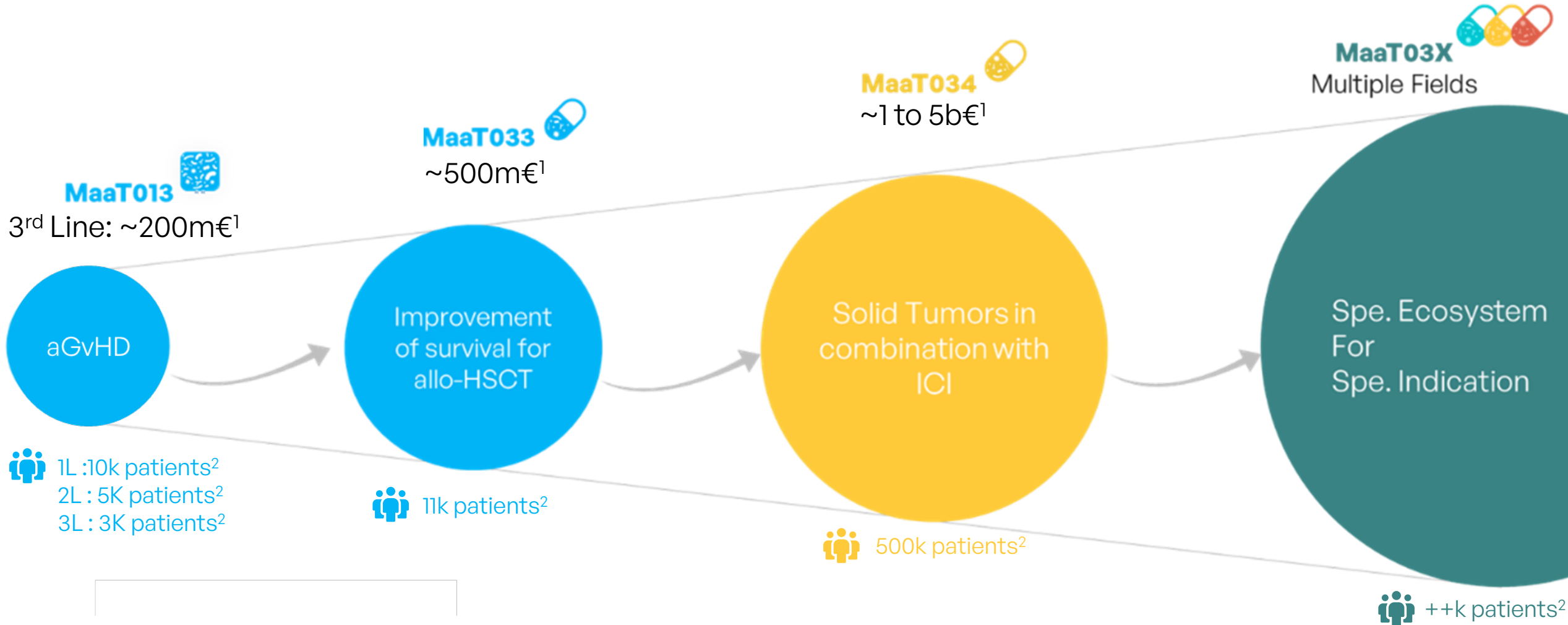
Platform → Program → Indication → Preclinical → Phase I → Phase II → Phase III → Upcoming milestone

Platform	Program	Indication	Preclinical	Phase I	Phase II	Phase III	Upcoming milestone
MET-N	MaaT013	aGvHD <sup>ODD EMA/FDA</sup>	ARES	EAP ongoing: 140 pts treated			<ul style="list-style-type: none"> <li>GI-ORR mid Q4 2024</li> <li>EBMT Data April 2024</li> </ul>
		IO PoC Melanoma	PICASSO (IST)				Results H2.24/ Q1.25
		HSCT <sup>ODD EMA</sup>	PHOEBUS				Safety Interim H2.24
MET-C	MaaT033	ALS	IASO				Results H2.24
	MaaT034	IO	PrClin				Targeting FIH 2025
	MaaT03X	Multiple	R&D				Candidates selection

aGvHD: acute Graft versus Host Disease ; IO: Immuno-Oncology ; PoC: Proof of Concept ; HSCT: Hematopoietic Stem Cell Transplantation ; ALS: Amyotrophic Lateral Sclerosis ; IST: Investigator Sponsored Trial



# Targeting Multiple Attractive Markets with Unmet Medical Need



<sup>1</sup>PYS EU5, US; <sup>2</sup>Per year

# Driving Near-Term Value with the Donor- Derived **MET-N** Platform

MET-N

# Microbiome Restoration with MaaT013: A Maximum-Density Product for Fast Engraftment in Acute Situations



- **Curative approach**
- MaaT013 has received **Orphan Drug Designation** from **FDA and EMA**
- **GI-ORR in mid Q4 2024**



01

## Characteristics

**Pooled microbiota:** high-richness, high-diversity, full ecosystem  
Microbiome Therapy containing Butycore®  
**Non immunosuppressive treatment**



02

## Administration

**3 doses** (enema bag) – within 10 days



03

## Available Clinical Data

**HERACLES Phase 2** Clinical Trial, N=24, 2L

**Early Access Program (EAP)**, data from N=140, 3L-6L, program still ongoing

**Ongoing ARES** – Positive **DSMB** review (n= 30)

> **200** patients treated to date



04

## Efficacy evaluation in EAP

**28-Days GI-ORR:** 52%

**12-months OS:** 47%

**18-months OS:** 42%

**Data in all patients (n=140)**

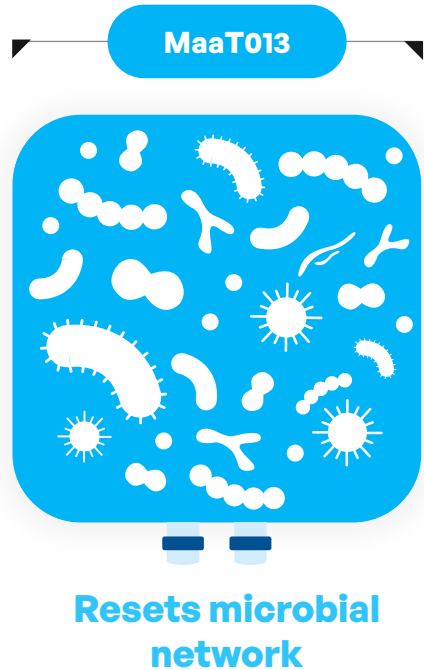


05

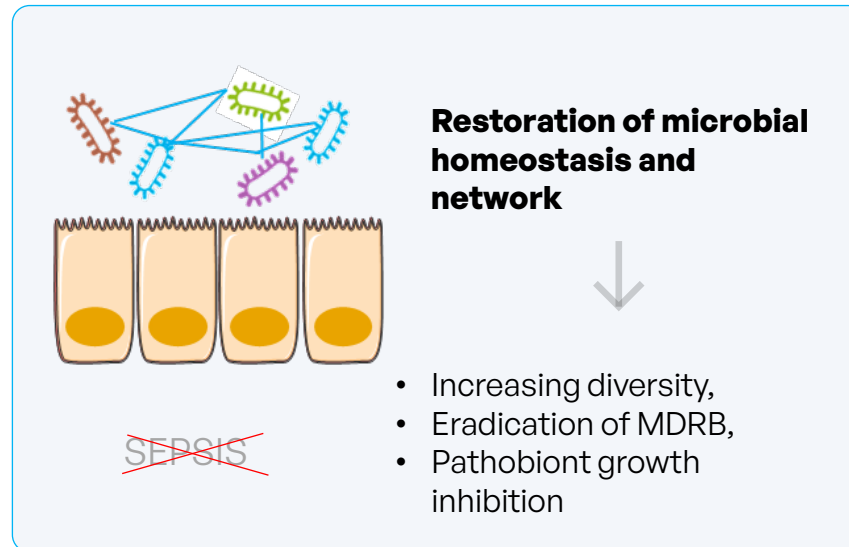
## Current indication

**Gastrointestinal acute Graft-versus-Host Disease (GI-aGvHD)**  
~ 3k patients per year

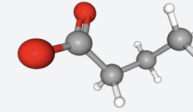
# MaaT013, a Novel Agent to Treat aGvHD Acting by Restoring Immune Homeostasis and Gut Barrier Integrity



## 1 Restoration of barrier integrity



## 2 Production of immunoregulatory metabolites



- SCFA ( Butyrate, Propionate)
- Immunoregulation (IL-10...)

## 3 Modulation of immune homeostasis

### Regulatory T cells



- Immune homeostasis restoration
- Remission of symptoms

Based on preclinical and ongoing clinical studies: MaaT013 could restore microbiome diversity, regenerates gut barrier's protective effect, and significantly curbs inflammation.

# Unmet Medical Need: Acute Graft-versus-Host Disease (aGvHD) Resistant to Steroids and Ruxolitinib (3<sup>rd</sup> line of treatment)

## Acute Graft-versus-Host Disease

- > aGvHD is a condition where transplanted cells attack the recipient's body
- > Is life-threatening when not controlled by a treatment, and induces long-term complications for those who do survive



Affects 50% of stem cell transplanted patients, 10,000 people a year EU/US

## Treatment Paradigm

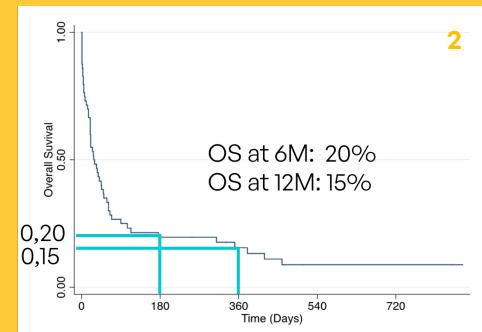
- > Corticosteroids are the 1<sup>st</sup> line of treatment, but 50% of patients do not achieve a sustained response
- > Ruxolitinib is approved as a 2<sup>nd</sup> line of treatment for SR-aGvHD (FDA, 2019 & EMA, 2022)



30% of aGvHD patients eligible for alternative treatment, primarily due to corticosteroids and ruxolitinib<sup>1</sup> resistance or non-eligibility  
Around 3,000 per year EU/US

## c. 30% of patients have no effective treatment option

- > There is **no** approved drug in 3L: lack of effective therapy
- > Off label options have shown limited benefit, showing the critical need for a new treatment



Outcome for this group of patients is dismal with a median survival of 28 days and a 15% OS at 1 year<sup>2</sup>

> Intestinal dysbiosis is associated with higher mortality in hemato-oncology<sup>3</sup>

# ARES, a Pivotal Phase 3 Trial to Treat aGvHD in 3<sup>rd</sup> Line Showing *“high efficacy and low toxicity”* as Concluded by the DSMB

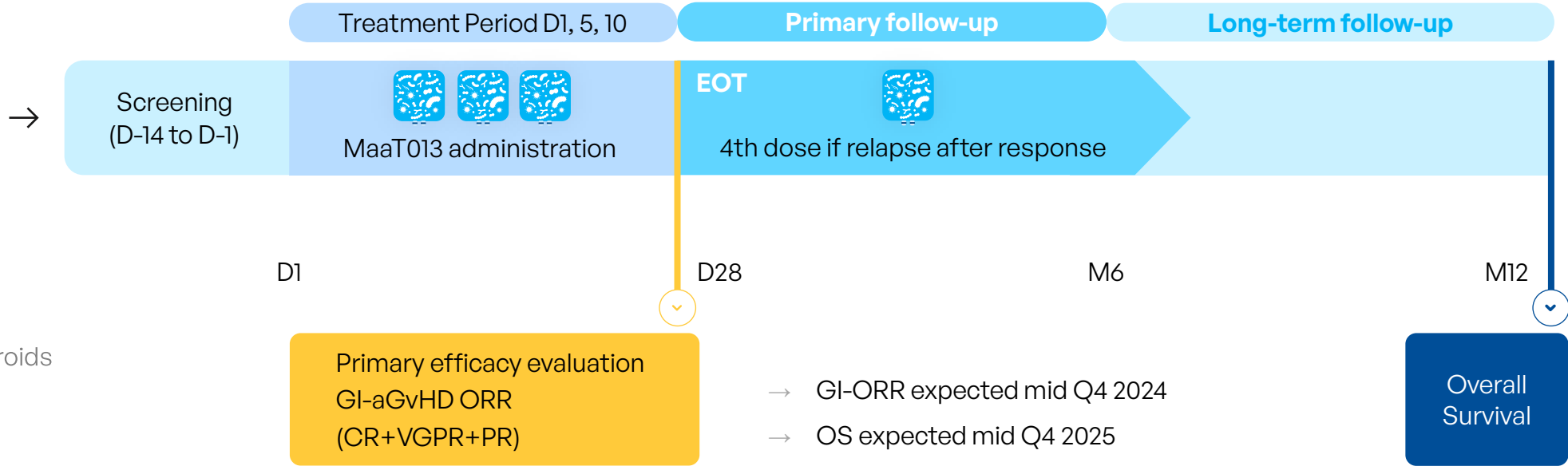
ARES



Patients with SR-GI-aGvHD

### Inclusion criteria

- Refractory or intolerant to 2L ruxolitinib
- Refractory to 1L corticosteroids
- aGvHD with GI symptoms
- Allo-HSCT
- Age > 18



D: Day, M: Month, EOT: End of treatment ; SR-GI-aGvHD: Steroid-refractory gastro-intestinal acute Graft-versus-Host Disease ; GI-ORR: Gastrointestinal Overall Response Rate; CR: Complete Response; VGPR: Very Good Partial Response; PR: Partial Response  
 \* DSMB review on 30 patients on October 2023

### DSMB\* main conclusions:

- Good safety profile
- ORR higher than pre-defined protocol



Commercial launch date anticipated in 2026



Market potential:  
~ 200 m€  
No Competitor in 3L

# The EAP Data Confirms Significant Improvement of Survival with High Level of Response

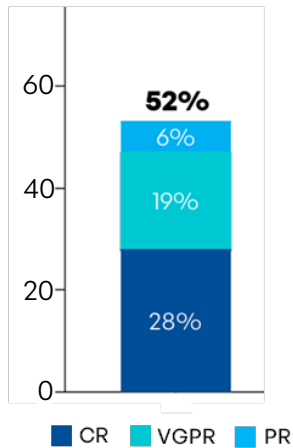


Data presented at EBMT 2024

## Global EAP populations

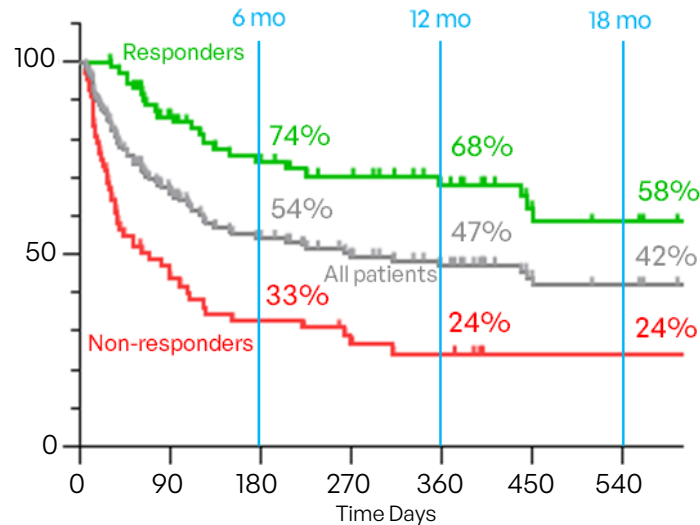
GI-ORR

Patients (%)



Overall Survival Rate

Survival (%)

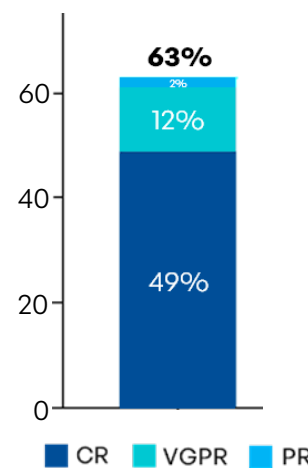


→ N=140, GI-aGVHD : 49% grade III, 40% grade IV, up to 6 lines of prior treatments (median 3) 121/140 received ruxolitinib

## ARES-like populations from EAP

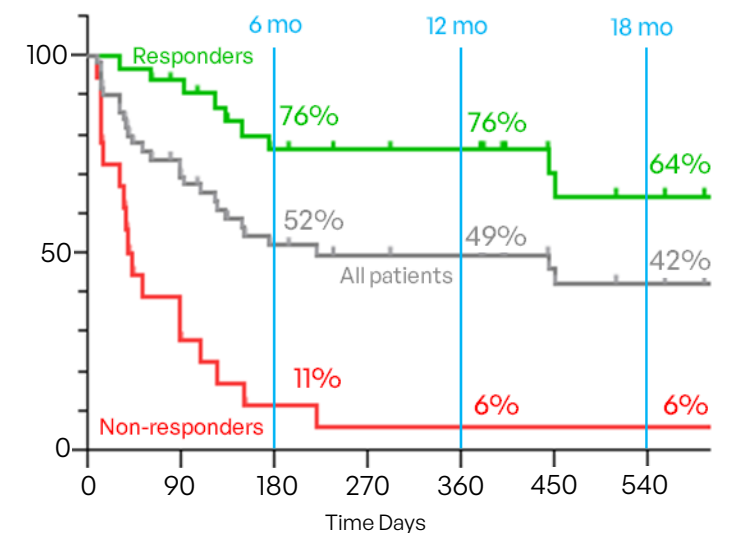
GI-ORR

Patients (%)



Overall Survival Rate

Survival (%)



→ N=49 ARES-like population ruxolitinib-refractory in 2nd line, MaaT013 given in 3rd line

- No effective treatment approved in 3L with very low expected OS 2mo: 22% ; 6mo: 20% ; 12mo: 15%<sup>1</sup> confirming strong unmet medical need
- High predominance of VGPR and CR responses in the EAP, suggesting a significant reduction in the disease burden
- Good safety and high efficacy translating in a significant increase in overall survival compared to REACH1 and Abedin et al. data - 2021<sup>1</sup>

<sup>1</sup>Expected OS of Steroid and Ruxolitinib resistant aGvHD patient at : 2 mo: 22% (REACH1 trial), 6mo: 20% and 12mo: 15% (Abedin et al.)



## Proof-of-Concept with MaaT013 in Combination with ICI In Metastatic Melanoma

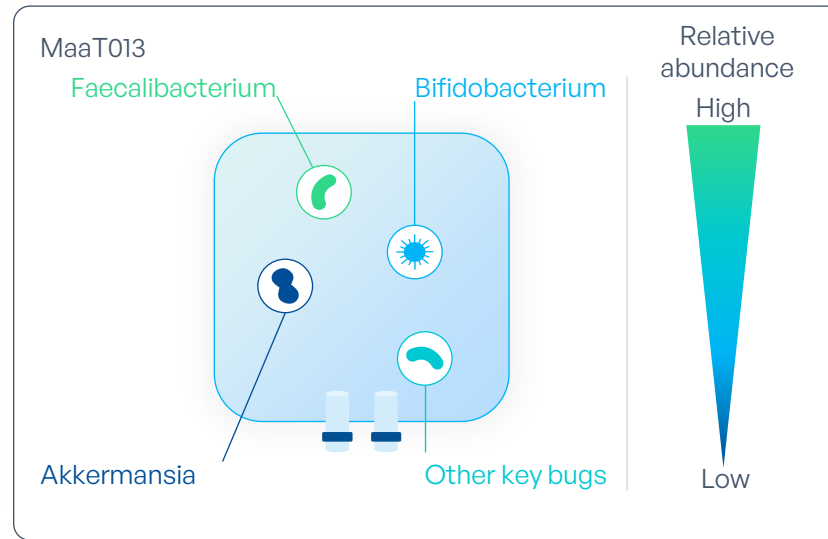
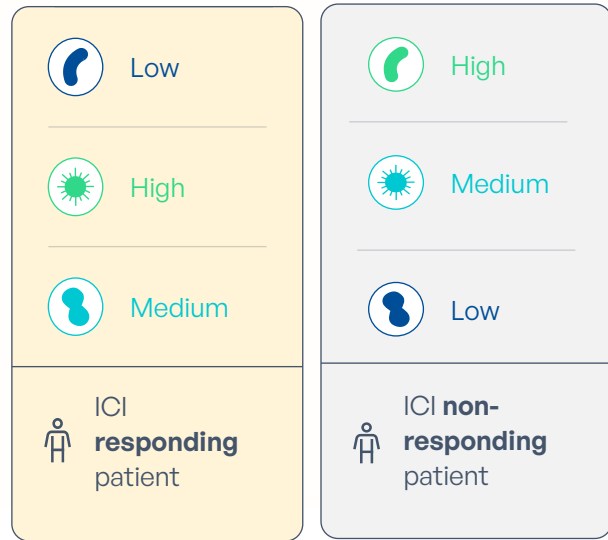


Serves as PoC for MaaT034 in  
combination with ICI

MET-N



# MaaT013 Evaluated in Phase 2 Randomized Clinical Trial in Melanoma

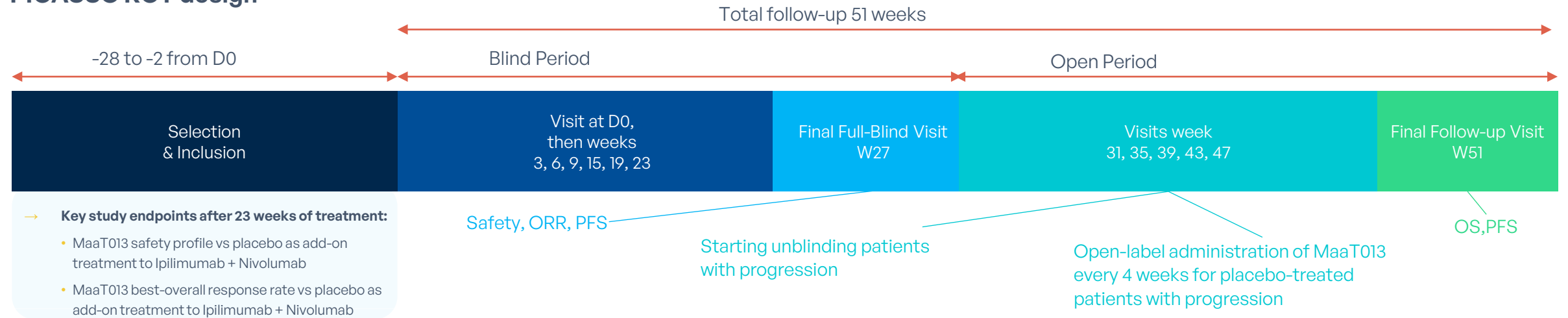


**Recruitment completed Ph. 2a PICASSO trial**

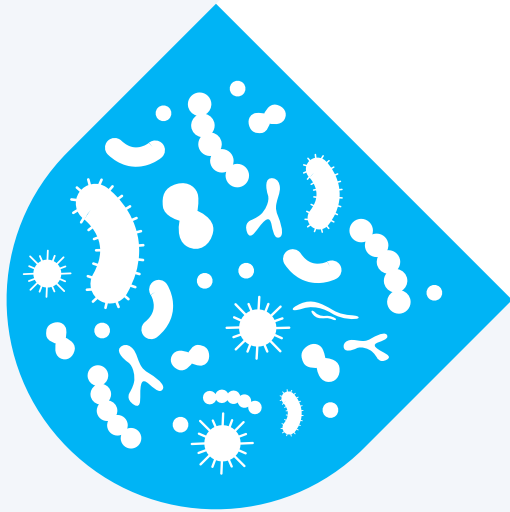
**Investigator led trial** (Assistance Publique - Hôpitaux de Paris – sponsor) and in collaboration with Institut Gustave Roussy

- RCT [MaaT013 + ICI] vs. [Placebo + ICI] in 70 metastatic melanoma patients
- **Data expected H2.24/Q1.25**

## PICASSO RCT design



# Ensuring Optimal Microbiota Function: MaaT033 – The Oral Ecosystem Microbiome Capsule for Adjunctive and Maintenance Therapy



- **Adjunctive and Maintenance**
- **Targeted release oral Capsules**
- MaaT033 has **received Orphan Drug** from the **EMA**



01

## Characteristics

**Pooled microbiota** : high-richness, high-diversity, full ecosystem,  
**Microbiome Ecosystem Therapy containing Butycore®**  
**Non immunosuppressive** treatment



02

## Administration

**Oral** (a lyophilized capsule)



03

## Clinical Program

**Ongoing Phase 2b trial PHOEBUS** in allo-HSCT patients  
**Phase 1b trial IASO** ongoing in ALS



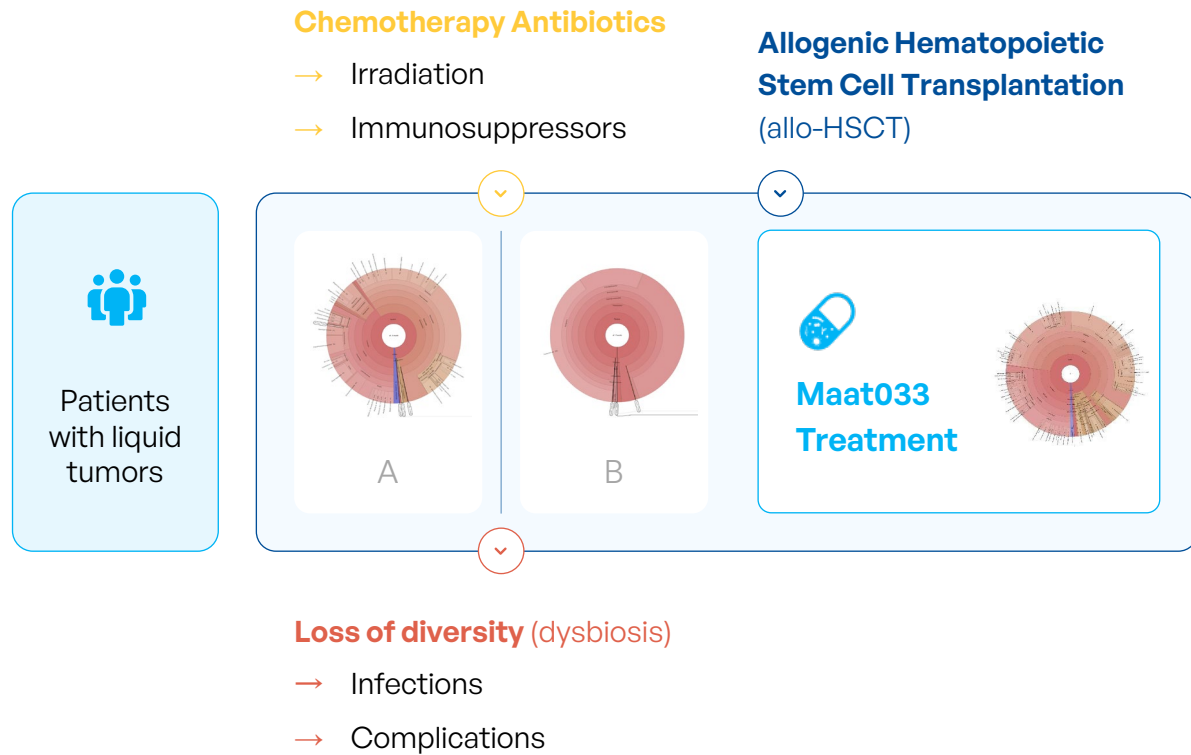
04

## Current indication

**Improving survival of allo-HSCT patients** (blood cancers i.e leukemia, ...) – ~ 11k patients per year  
**Slowing down disease progression in ALS**

# MaaT033 to Ensure Optimal Gut Microbiota to Improve Survival in Patients Receiving Allogeneic HSCT

Intestinal dysbiosis is associated with higher mortality in hemato-oncology<sup>1</sup>



<sup>1</sup> Peled et al., NEJM 2020



United States

**C. 7,800**

Primary procedures



EU 5

**C. 9,600**

Primary procedures



Japan

**C. 3,000**

Primary procedures

Additional

**7% - 10%**

Recurrent procedures

Approximately

**22,500**

procedures / year

EBMT aHSCT Survey, 2017 (published in Bone Marrow Transplantation (2019) 54:1575 – 1584), Global Data 2020

# MaaT033: a Potential Adjunctive Treatment for Patients Receiving allo-HSCT



- **387** patients in a **randomized, double-blind, placebo-controlled international** study
- **56 sites** targeted globally

- Primary endpoint: **efficacy** of MaaT033 in **improving overall survival at 12 months**
- Study started in **November 2023**, results are expected in **2026**

**387** patients dosed pre- and post- allo-HSCT



<sup>1</sup> Expansion to US sites subject to discussion with the FDA

**Ongoing Phase 2b PHOEBUS** | **Safety Interim analysis on 60 patients in H2 2024** | **Based on expected duration of recruitment, OS primary endpoint expected in 2026** | **~ 11k patients per year**

# MaaT033 Aims to Slow Down Amyotrophic Lateral Sclerosis Progression



## Amyotrophic Lateral Sclerosis

- Could affect up to 60,000 patients in US & EU by 2040<sup>1</sup>
- Paralysis and death 3 to 5 years after diagnostic<sup>2</sup>
- Currently no curative treatment and few symptomatic treatments

## Rationale for Exploratory Utilization of MaaT033 in ALS

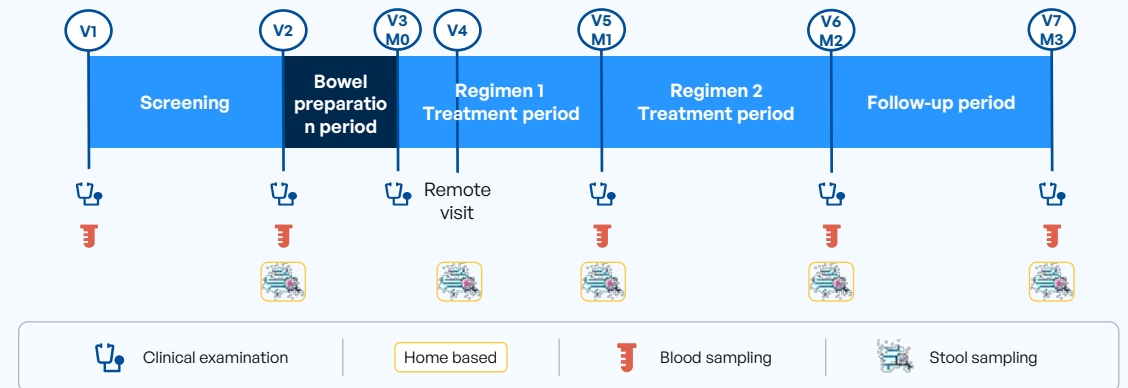
- Microbiota-Gut-Brain axis has the potential to become the new standard to treat neurodegenerative diseases, including ALS
- MaaT033 safety profile and oral administration is suitable for ALS
- Strong support from medical community & patients
- A cost-effective way of testing neurodegenerative field in an indication with high medical need

<sup>1</sup> Arthur, K., Calvo, A., Price, T. et al. Projected increase in amyotrophic lateral sclerosis from 2015 to 2040. Nat Commun 7, 12408 (2016). <https://doi.org/10.1038/ncomms12408>

<sup>2</sup> <https://tousensellescontrelasla.fr/la-sla-cest-quoi/>

## Study

- Up to **15 patients** in a **pilot, open-label, Phase 1b** study **in France**
- **Key study endpoints:** assess safety and tolerability of MaaT033 and gut microbiota composition evolution
- Study started in **2023** → **Results** expected in **H2 2024**
- **Positive DSMB** in **Feb. 2024:**  
Trial to proceed as planned without modifications  
Good safety profile and generally well tolerated



Study developed with:



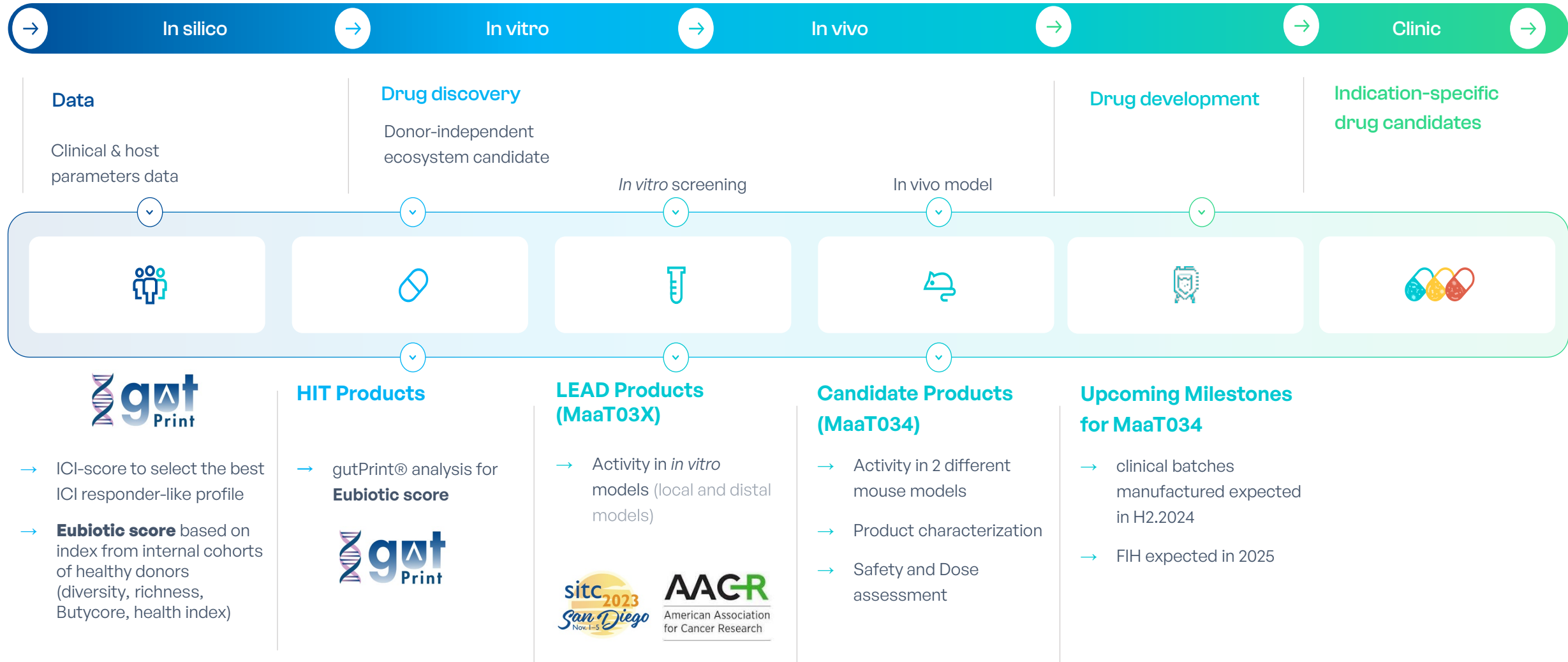
In collaboration with:



# Progressing the Next- Generation, Co-Cultured, Donor Independent **MET-C** Platform

MET-C

# MET-C Product Generation is Driven by MaaT Pharma's Proprietary Predictive AI, Eubiotic Score and *in vitro* and *in vivo* validation processes



# End-to-End In-house cGMP Manufacturing

All MET



# Europe's Largest Specialized cGMP Manufacturing Facility for Microbiome Ecosystem Therapies

A dedicated 1,600m<sup>2</sup> site (expandable) to support demands until 2034 for **MET-N clinical and future commercial production, R&D, and clinical batches of MET-C products (MaaT034 & MaaT3X family)** (est. first step):

~10 000 treatable patients per year

MaaT013

9.000  
pouches / year

MaaT033

1.300.000  
capsules / year

MaaT03X

Up to 300.000  
capsules / year



01

**Fully integrated Manufacturing and development platform** for a streamlined product development, scaleup and GMP process.



02

**Ongoing CSR global strategy:** reforestation program in France (GoGreen) and “Cap Vert pour la forêt” program, etc.



03

**Option to expand manufacturing facilities** to double manufacturing capabilities.



Status

**Production started in September 2023**



Partnership  
with



# Key Takeaways

# Multiple Near-Term Value Inflection Milestones

2024

## MaaT013 (pooled enema)

GvHD | EAP long term follow-up EBMT24 ✓  
 GvHD | ARES P3 GI-ORR mid Q4 24  
 IO Mela. | PICASSO P2a Results H2.24/Q1.25

## MaaT033 (pooled capsule)

HSCT | PHOEBUS P2b Safety Interim H2  
 ALS | IASO P1b Results H2

## MaaT034 (co-cultured capsule)

Candidate Selection  
 1<sup>st</sup> Clinical Batch Manufactured

2025

## MaaT013 (pooled enema)

GvHD | Final Results (OS)

## MaaT033 (pooled capsule)

HSCT | PHOEBUS P2b Safety Interim 2

## MaaT034 (co-cultured capsule)

Solid Tumors IO | Target FIH 25

## MaaT03X (co-cult. ind.-spec. caps)

Undisclosed | Next Steps

Finance

- **Revenues from of MaaT013 in aGVHD of 2.2m€ for 2023 (record year)**
- **Cash position of 24.3m€** as of 23/12/31
- **Cash runway to the end of Q3 2024**

# A Robust Value Creation Strategy Driven by Leading Expertise in Microbiome-based Therapeutics

## MET-N

## MET-C

Addressable Patients



### Creation Value

Time:  
Event:  
1<sup>st</sup> Ind:  
Market size:



MaaT013



Pooled enema

- Mid Q4 2024
- P3 GI-ORR
- aGvHD
- 200m€

MaaT033



Pooled capsule

- H2.2024
- P2b DSMB
- allo-HSCT
- 500m€

MaaT034



Co-cultured caps.  
Synthetic eubiotic microbiota

- 2024
- Candidate selection & PICASSO PoC Results
- ICI combo in solid tumor
- 1 to 5b€

MaaT03X



Co-cultured capsule  
Indication specific

- 2025+
- New program reveal
- Multiple Indications
- Multiple Markets



MaaT Pharma has the largest Microbiome Ecosystem Therapies™ production facility in Europe, which is the foundation of the Company's ability to scale and produce drug candidates in a cGMP environment

# Corporate Social Responsibility



**MaaT Pharma aims to become the source of Microbiome excellence providing patients with safe and innovative medicines.** The Company develops products from sustainable biological matters, driving optimal impact of Microbiome.



**Patients are our priority.** We are committed to our patients and to the protection of human health by respecting environmental protection, respecting our employees and ensuring good governance practices. Our way of working every day is driven by the 4 guidelines below:

- Innovate and raise awareness to **deliver better care,**
- Contribute to employees-growth within a **people-oriented ecosystem,**
- Place **ethics and transparency** at the core of the Company’s strategy,
- Control and measure our **impact on the environment.**

## 2023 CSR indicators

### Social

- 34 y-o** is the average age of permanent employees
- 36%** Percentage of PhD, PharmD, MD among employees involved in research
- 75%** Training Plan Completion Rate

### Environment

- 2394 tCO2e** Carbon footprint
- 361 kWh /Employee** Energy consumption per employees on site

### Societal

- 85%** of operating expenses related to R&D as a proportion of total operating expenses
- 259** public interventions to increase awareness on microbiome

### Governance

- 38%** of women in the Board of directors
- 72%** of women in the Executive team

The background of the slide is filled with various stylized, colorful illustrations of microscopic organisms, including bacteria, viruses, and fungi, in shades of teal and blue. The organisms are scattered across the page, creating a dynamic and scientific atmosphere.

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# Thank you

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