

MaaT Pharma

Enhancing Survival through Microbiome Innovation

June 2024



## Disclaimer

This document has been prepared by MaaT Pharma (the "Company") and is for information and background purposes only.

While the information contained herein has been prepared in good faith, neither the Company, nor its shareholders, directors, officers, agents, employees, or advisors give, have given or have authority to give, any representations or warranties (express or implied) as to, or in relation to, the fairness, accuracy, reliability or completeness of the information in this document, or any revision thereof, or of any other written or oral information made or to be made available to any interested party or its advisers, including financial information (all such information being referred to as "Information"), and liability therefor is expressly disclaimed. Accordingly, neither the Company nor any of its shareholders, directors, officers, agents, employees, affiliates, representatives or advisers take any responsibility for, or will accept any liability whether direct or indirect express or implied, contractual, tortuous, statutory or otherwise, in respect of the accuracy or completeness of the Information or for any of the opinions contained herein or for any errors, omissions or misstatements or for any loss, howsoever arising from this document.

The information and opinions contained in this document are provided as of the date of this document only and may be updated, supplemented, revised, verified or amended, and thus such information may be subject to significant changes. The Company is not under any obligation to update the information or opinions contained herein which are subject to change without prior notice.

The information contained in this document has not been subject to independent verification and are qualified in their entirety by the business, financial and other information that the Company is required to publish in accordance with the rules, regulations and practices applicable to companies listed on the regulated market of Euronext in Paris, including in particular the risk factors and other information in the Company's Document d'enregistrement (Registration Document) registered by the French Autorité des marches financiers (Financial Markets Authority) (the "AMF") on October 1st, 2021 under no. I.21-0057 and its supplement on October 14, 2021 under no. I.21-0061 and in any other periodic report, which are available free of charge on the websites of the Company (https://www.maatpharma.com/) and the AMF (www.amf-france.org).

No representation, warranty or undertaking, express or implied, is made as to the accuracy, completeness or appropriateness of the information and opinions contained in this document. The Company, its subsidiaries, its advisors and representatives accept no responsibility for and shall not be held liable for any loss or damage that may arise from the use of this document or the information or opinions contained herein.

This document contains information on the Company's markets and competitive position, and more specifically, on the size of its markets. This information has been drawn from various sources or from the Company's own estimates which may not be accurate and thus no reliance should be placed on such information. Any prospective investors must make their own investigation and assessments and consult with their own advisers concerning any evaluation of the Company and its prospects, and this document, or any part of it, may not form the basis of or be relied on in connection with any investment decision.

This document contains certain forward-looking statements. These statements are not guarantees of the Company's future performance. These forward-looking statements relate to the Company's future prospects, developments and marketing strategy and are based on analyses of earnings forecasts and estimates of amounts not yet determinable.

Forward-looking statements are subject to a variety of risks and uncertainties as they relate to future events and are dependent on circumstances that may or may not materialize in the future. Forwardlooking statements cannot, under any circumstance, be construed as a guarantee of the Company's future performance and the Company's actual financial position, results and cash flow, as well as the trends in the sector in which the Company operates, may differ materially from those proposed or reflected in the forward-looking statements contained in this document. Even if the Company's financial position, results, cash-flows and developments in the sector in which the Company operates were to conform to the forward-looking statements contained in this document, such results or developments cannot be construed as a reliable indication of the Company's future results or developments. The Company does not undertake any obligation to update or to confirm projections or estimates made by analysts or to make public any correction to any prospective information in order to reflect an event or circumstance that may occur after the date of this document.

All persons accessing this document are deemed to agree to all the limitations and restrictions set out above.

# 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
- Our gutPrint® AI, linked to our coculture platform, is poised to deliver, potentially, clinical-ready candidates by 2025
- Largest European cGMP production facilities for Microbiome Ecosystem Therapies





## **Finance**

- Revenues of MaaT013 in aGvHD of 2.2m€ for 2023 from Early Access Program
- Cash position of 18.2m€ as of March 31, 2024. Post follow-on in May 2024, (approx. €17.3m€) cash runway extends into early Q1/2025
- Exploring options to extend cash runway, including non-dilutive and dilutive sources

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

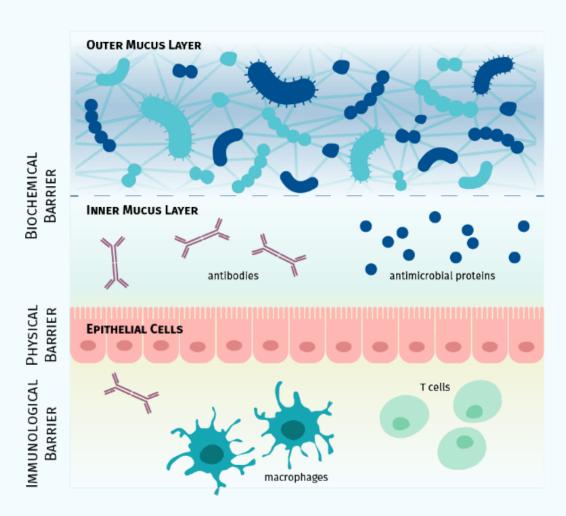
A rich and diversified gut ecosystem actively modulates the immune system functionality



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



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

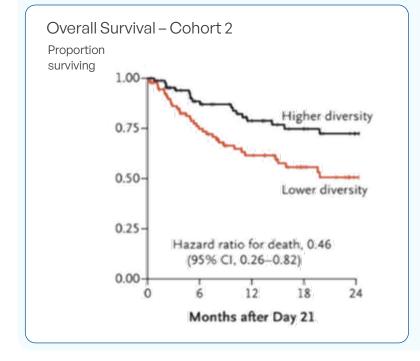


Cross-section of a healthy gut

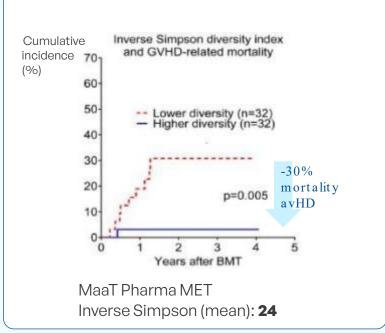
# In Oncology, a Higher Gut Microbiome Diversity is Associated with Increased Survival

Liquid Tumors Solid Tumors

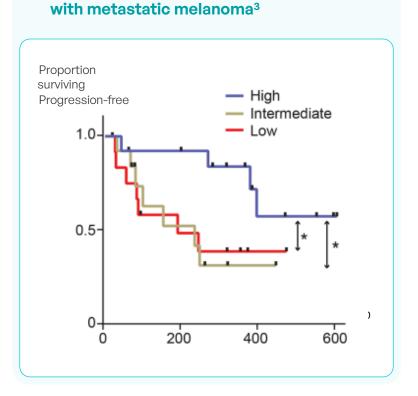
## Higher survival rate in patients receiving allo-HSCT\*1



#### Lower incidence and lower mortality from aGvHD\*2



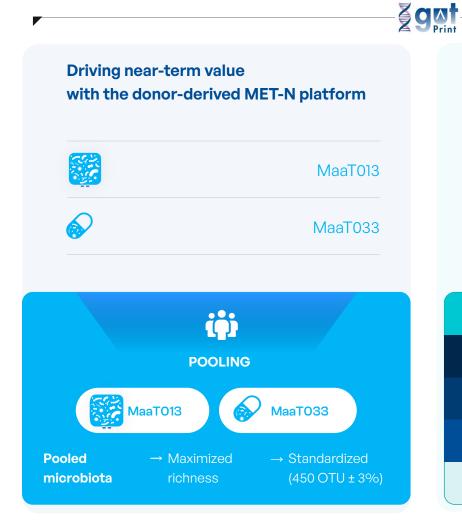
# **Higher response rate to ICI\* in patients**

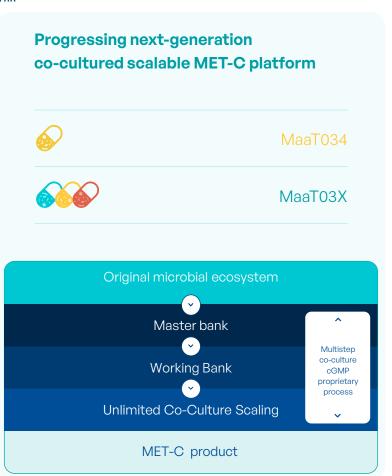


<sup>\*</sup> allo-HSCT: allogeneic hematopoietic stem cell transplantation; aGvHD: acute Graft-vs-host-Disease; ICI: Immune Checkpoint Inhibitors

Peled, J.U. & al N Engl J Med 2020;382;822-34; Ghani, 2021; Jeng RR, et al, Biol Blood Marrow Transplant 21 (2015) 1373e1383; Pamer, Blood, 2014; 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





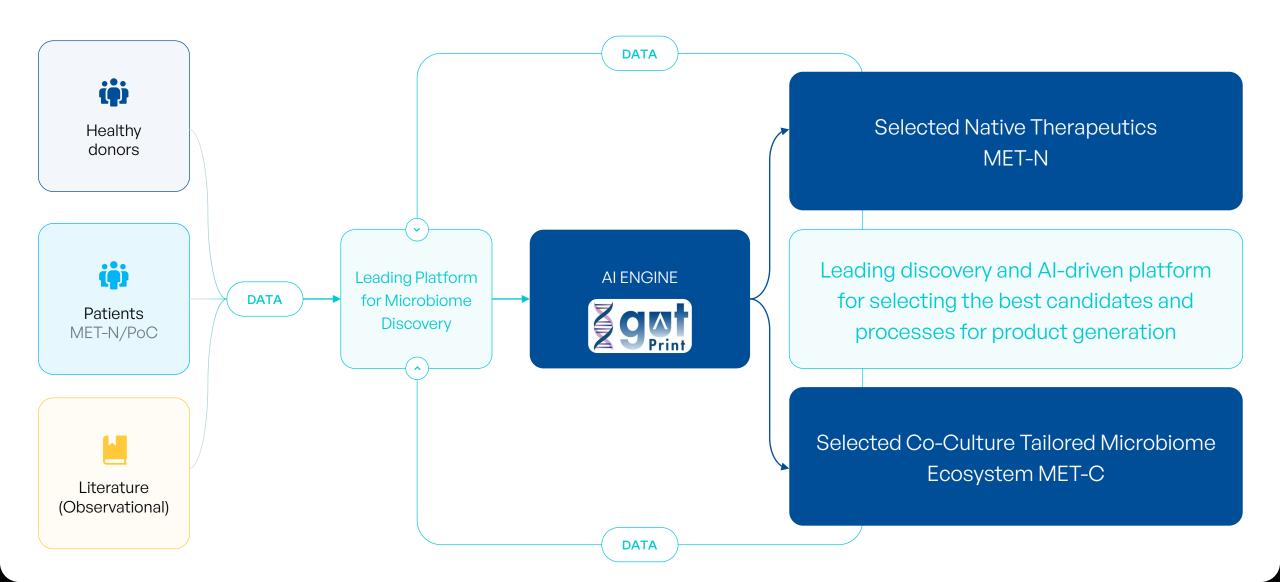
Leading capabilities in microbiome drug production



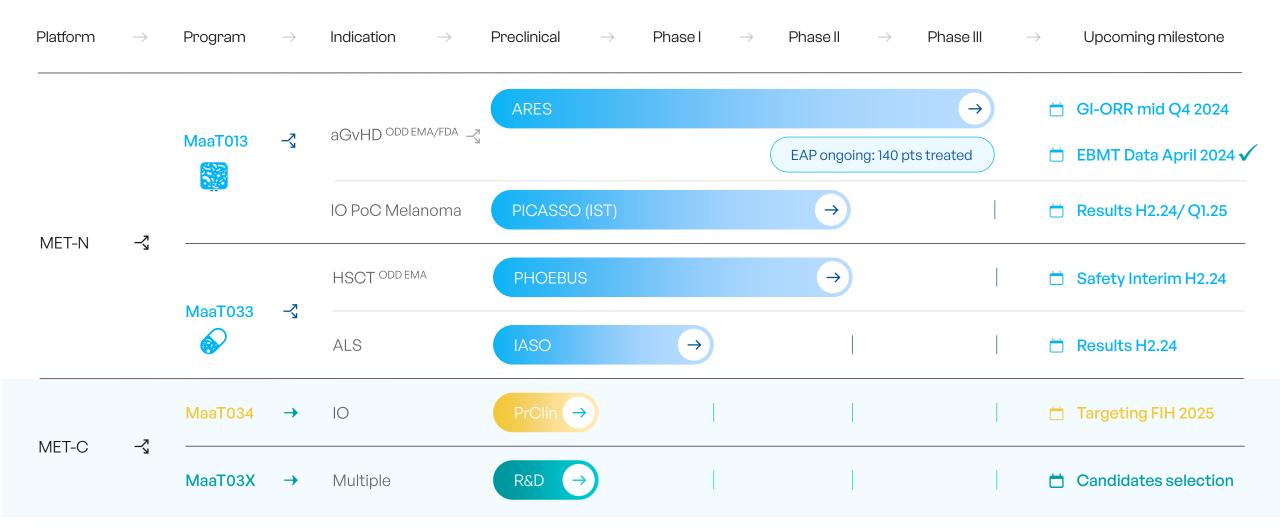
~10 000 treatable patients per year



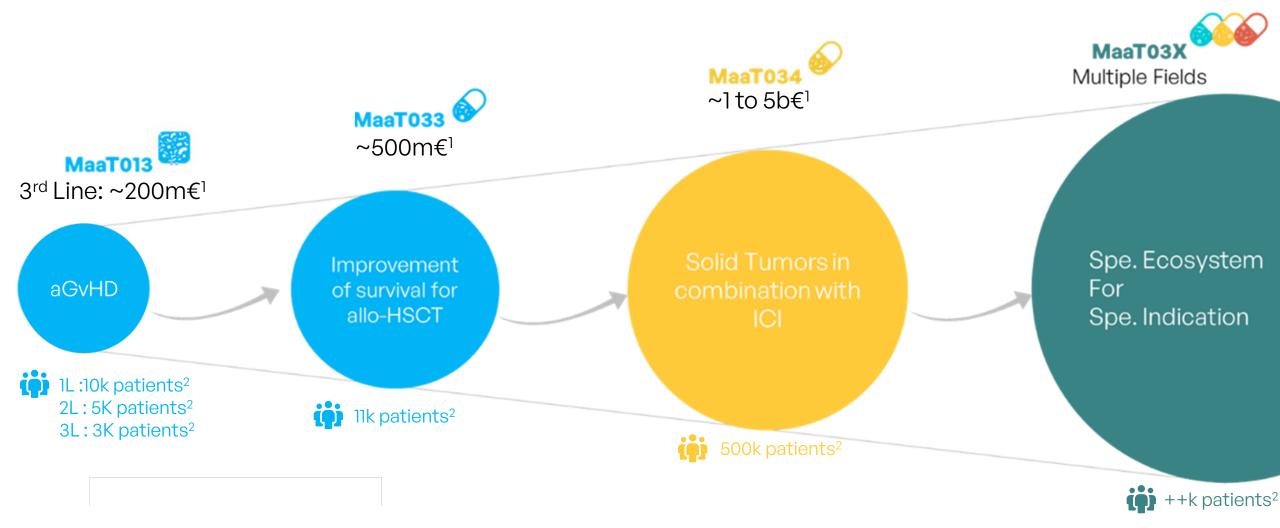
# Al-driven Research Engine Powered by Metagenomics Enabling Candidate Selection



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



# Targeting Multiple Attractive Markets with Unmet Medical Need





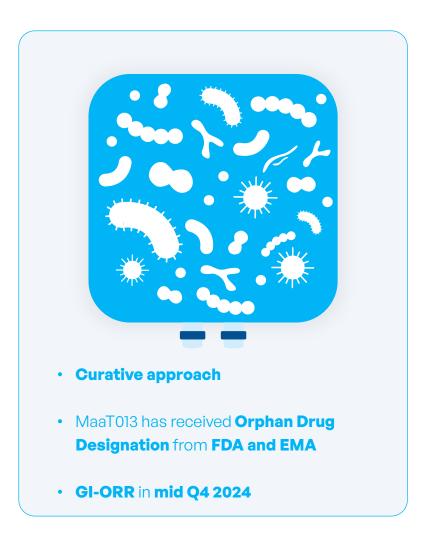


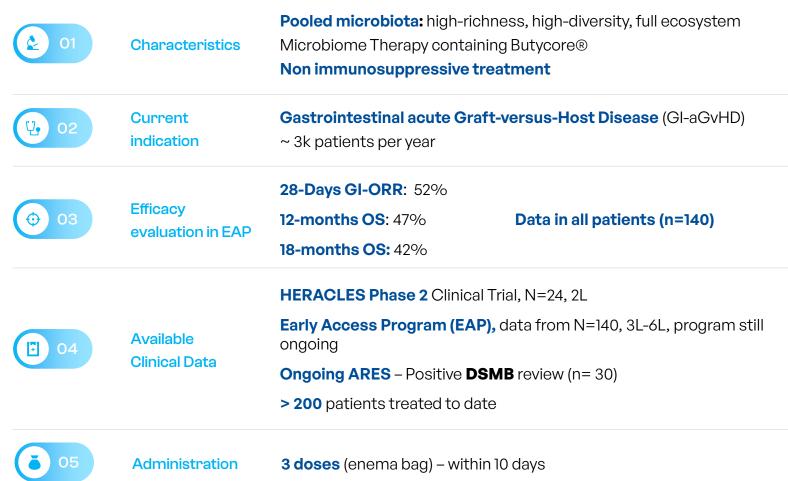
# Driving Near-Term Value with the Donor-Derived MET-N Platform

MET-N

CORPORATE PRESENTATION

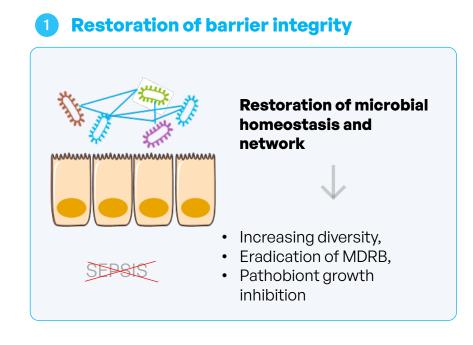
# Microbiome Restoration with MaaTO13: A Maximum-Density Product for Fast Engraftment in Acute Situations

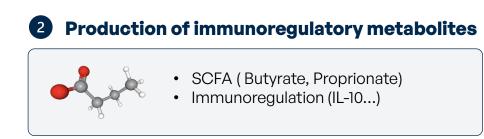


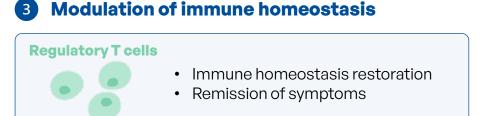


# MaaTO13, a Novel Agent to Treat aGvHD Acting by Restoring Immune Homeostasis and Gut Barrier Integrity









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

## **Treatment Paradigm**

- Ocrticosteroids are the 1st 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)

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

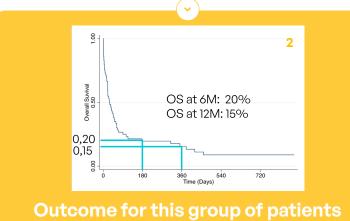
- There is **no** approved drug in 3L: lack of effective therapy
- Off label options have shown limited benefit, showing the critical need for a new treatment



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



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

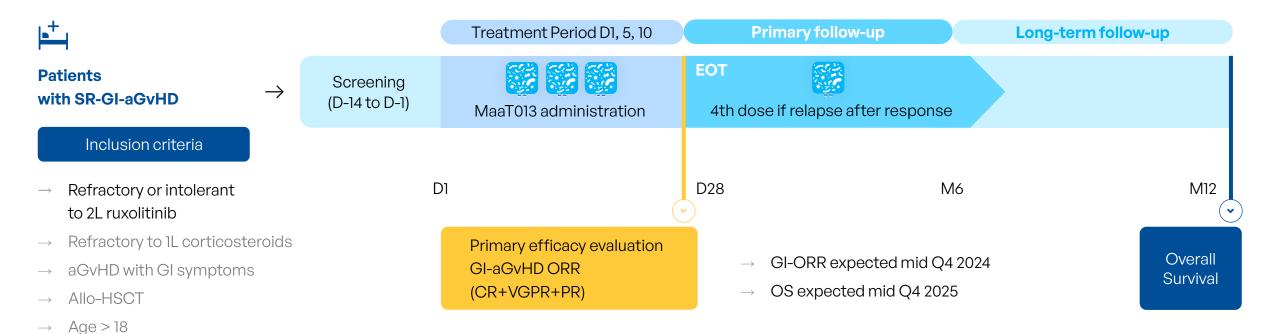


Outcome for this group of patients is dismal with a median survival of 28 days and a 15% OS at 1 year<sup>2</sup>



# 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





D: Day, M: Month, EOT: End of treatment; SR-Gl-aGvHD: Steroid-refractory gastro-intestinal acute Graft-versus-Host Disease; Gl-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

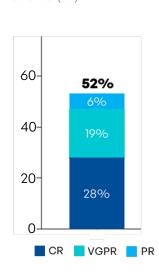


## **Global EAP populations**

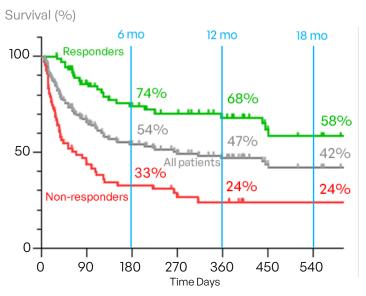
→ N=140, GI-aGVHD: 49% grade III, 40% grade IV, up to 6 lines of prior treatments (median 3) 121/140 received ruxolitinib

MaaT013 aGvHD





#### **Overall Survival Rate**



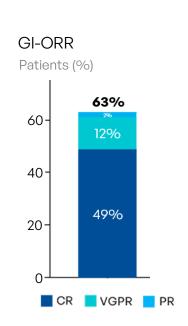
- High predominance of VGPR and CR responses in the EAP, suggesting a significant reduction in the disease burden
- Clinical response to MaaT013 translates to increased overall survival

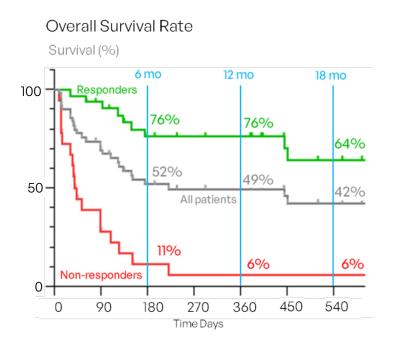
# The EAP Data Confirms Significant Improvement of Survival with High Level of Response

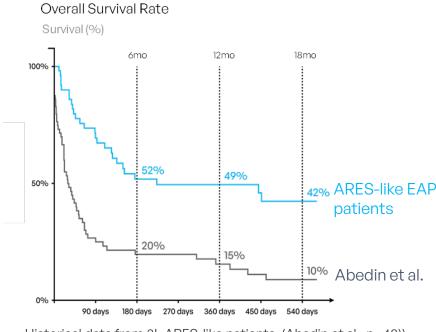


## **EAP: ARES like populations**

→ N=49. GI-aGVHD: 3L







- Historical data from 3L ARES-like patients (Abedin et al., n=48))
- No effective treatment in 3L with very low expected OS 2mo: 22%; 6mo: 20%; 12mo: 15% 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



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

Serves as PoC for MaaT034 in combination with ICI



# Checkpoint Inhibitors have revolutionized treatment of Solid Tumors but a large proportion of patients can't benefit from it due to primary resistance

# Primary Resistance Rate to Immune Checkpoint Inhibitors



Lung Cancer (NSCLC)

35 - 40 %



Skin Cancer (Melanoma)

**Up to 65 %** 



Around 19 million people diagnosed with cancer each year globally



Immune Checkpoint Inhibitors (ICI) significantly improved outcomes of patients with solid tumors and strategies to enhance responses remains a strong unmet medical need.



**Combination strategies** tested so far to improve responses to ICI remains mostly unsuccessful and/or associated with a **higher toxicity.** 

→ Urgent need to bring new combination therapies with ICI to safely increase the response rates and overall survival

# Growing clinical evidence that a Full-Ecosystem Gut Microbiome influences efficacy of ICI

#### 2021

FMT from ICI-responders could overcome resistance to ICI in non-responders
with metastatic melanoma

**⊘** 6/15

Non-responders

→ Responders
(Davar et al, 2021)

ers Non-responders

→ Responders
(Baruch et al, 2021)

**⊘** 3/10

2023

FMT from <u>healthy donors</u> increases
response of aPD1 in
<u>ICI-naive patients</u>
with metastatic melanoma

**3/20** 

**ICI-naive** → Responders (ORR=65 %, Routy *et al.* 2023)

2024

Microbiotherapy from healthy donors increases response of aPD1+aCTLA4 in ICI-naive patients

with metastatic melanoma

**35/20** 

**ICI-naïve** → Responders (ORR=75 %, Routy,. 2024)

.../35
First RCT
70 pts rand 1:1

(MaaT Pharma)

aPD1 historical response close to 33 %

aPD1+aCTLA4 historical response close to 59 %

FMT = (Hospital) Fecal Microbiota Transplantation

→ Leveraging the complete gut microbiome properties may be a game-changer in immuno-oncology

# MaaT013 Evaluated in Phase 2 Randomized Clinical Trial in Melanoma

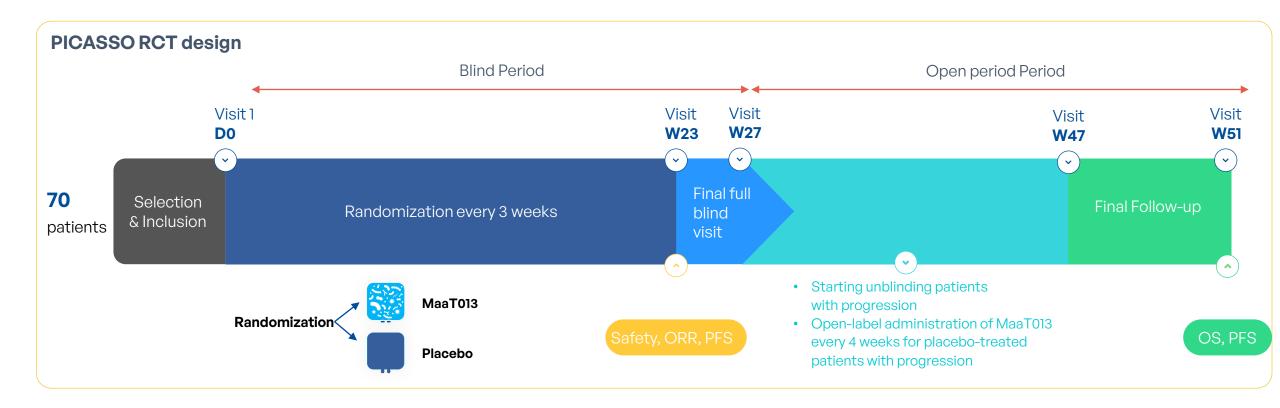
#### Recruitment completed Ph. 2a PICASSO trial

**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

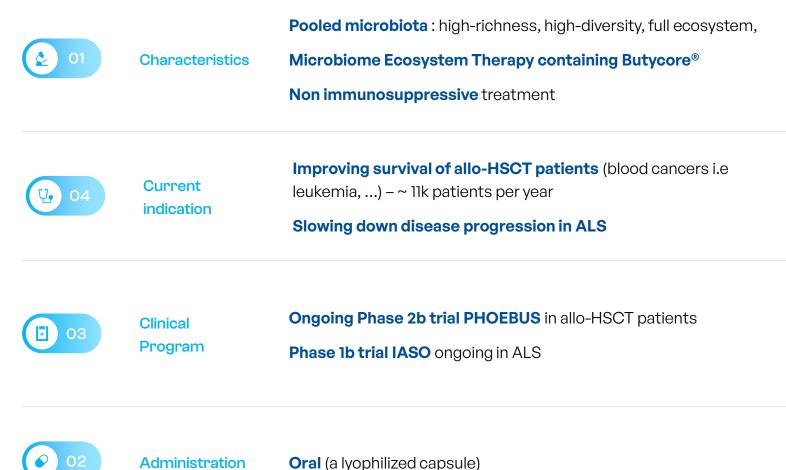
#### Key study endpoints after 23 weeks of treatment:

- MaaT013 safety profile vs placebo as add-on treatment to Ipilimumab + Nivolumab
- MaaT013 best-overall response rate vs placebo as add-on treatment to Ipilimumab + Nivolumab



# Ensuring Optimal Microbiota Function: MaaTO33 - The Oral Ecosystem Microbiome Capsule for Adjunctive and Maintenance Therapy



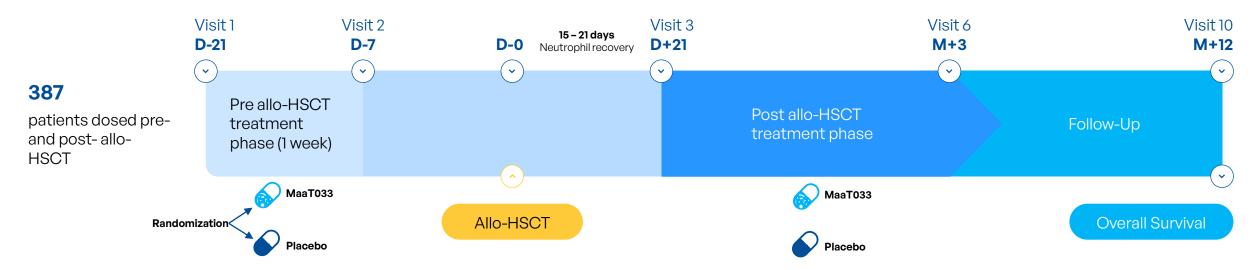


# MaaTO33: a Potential Adjunctive Treatment for Patients Receiving allo-HSCT



- → 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



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**

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

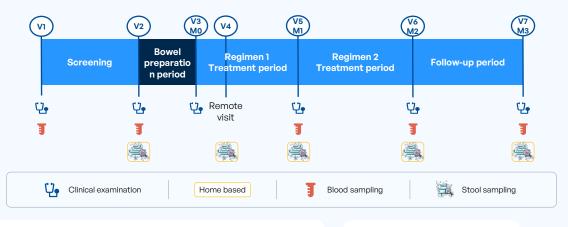
## Rationale for Exploratory Utilization of MaaT033 in ALS

- 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



- → 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:











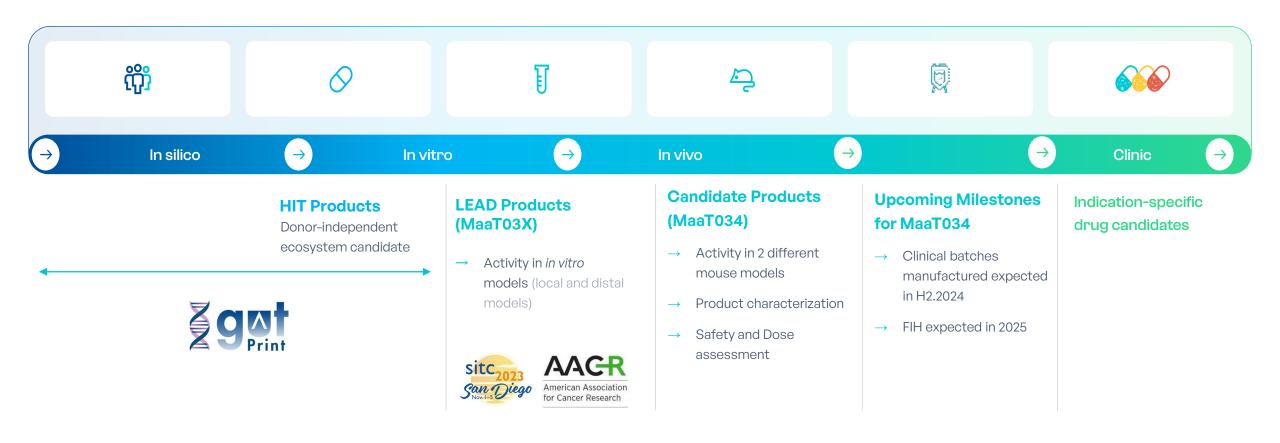


# Progressing the Next-Generation, Co-Cultured, Donor Independent MET-C Platform



CORPORATE PRESENTATION

# MET-C Product Generation is Driven by MaaT Pharma's Proprietary Predictive Al, 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 1,600m2 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



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



#### **Ongoing CSR global strategy:**

reforestation program in France (GoGreen) and "Cap Vert pour la forêt" program, etc.



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



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

## MaaTO33 (pooled capsule)

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

## MaaT034 (co-cultured capsule)

Candidate Selection

1st Clinical Batch Manufactured

2025

### MaaTO13 (pooled enema)

GvHD | Final Results (OS)

### MaaT033 (pooled capsule)

HSCT | PHOEBUS P2b Safety Interim 2

#### MaaTO34 (co-cultured capsule)

Solid Tumors IO | Target FIH 25

## MaaTO3X (co-cult. ind.-spec. caps)

Undisclosed | Next Steps

#### Finance

- Revenues of MaaT013 in aGvHD of 2.2m€ for 2023 from Early Access Program
- Cash position of 18.2m€ as of March 31, 2024. Post follow-on in May 2024, (approx. €17.3m€) cash runway extends into early Q1/2025
- Exploring options to extend cash runway, including nondilutive and dilutive sources



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

## MET-N

 $( \mathbf{\wedge} )$ 

#### Adressable **Patients**

#### **Creation Value**

Time:

Event:

1st Ind:

Market size:



#### MaaT013



Pooled enema

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

# MET-C

#### 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
- → 1to 5b€

#### MaaT03X



Co-cultured capsule Indication specific

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



MaaT Pharma has the largest Microbiome Ecosystem Therapies<sup>TM</sup> 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

3 GOOD HEALTH

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
<b>75</b> %	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
259	expenses  public interventions to increase awareness on microbiome

Governance		
38%	of women in the Board of directors	
<b>72</b> %	of women in the Executive team	

