

MaaT Pharma

Enhancing Survival through Microbiome Innovation

October 2024



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MaaT Pharma

Late-Stage Biotech leading Microbiome Immunomodulator Innovation in Oncology



MaaT013 in Phase 3 in aGvHD

- **Recruitment completed for Phase 3** in aGvHD in Europe, expecting primary endpoint readout in **January 2025**
- **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, clinically-ready candidates by 2025
- **Largest European cGMP** production facilities for Microbiome Ecosystem Therapies







Revenues of MaaT013 in aGvHD of 1.7m€ for H1 2024 from Early

Access Program

- Cash position of 31.2m€ as of June 30, 2024. Post follow-on in May 2024, (approx. €17.3m€) cash runway extends into Q2/2025
- **Exploring options to extend cash** runway, including non-dilutive and dilutive sources

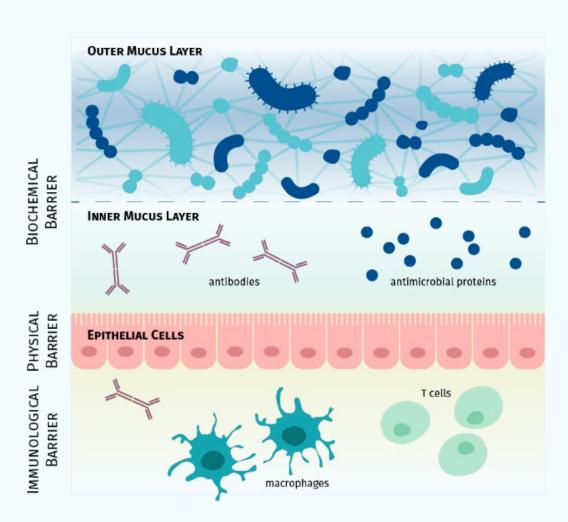
Host – Microbiota Interactions are Critical for a Functional Immune System



A rich & diversified gut ecosystem actively modulates the immune system functionality.



Diversity in microbial species leads to the production of a wide range of metabolites, such as short-chain fatty acids, which have anti-inflammatory properties and contribute to the regulation of immune responses.



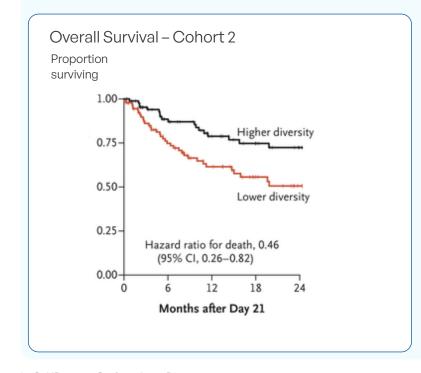
Cross-section of a healthy gut

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

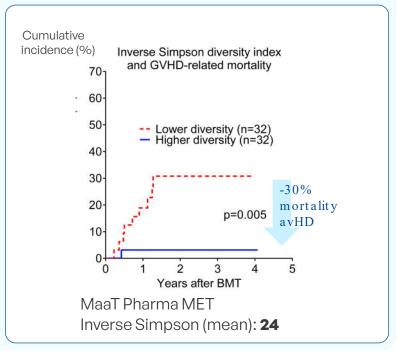
Hematopoietic Stem Celle Transplation (HSCT)

Immune Checkpoint Inhibitor (ICI)

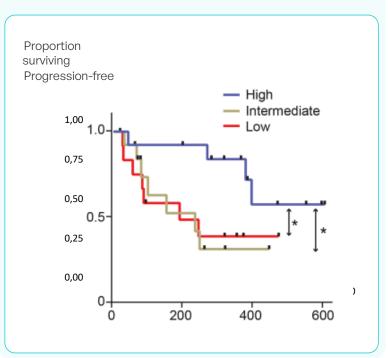
Higher survival rate in patients receiving allo-HSCT*1



Lower incidence and lower mortality from aGvHD*2



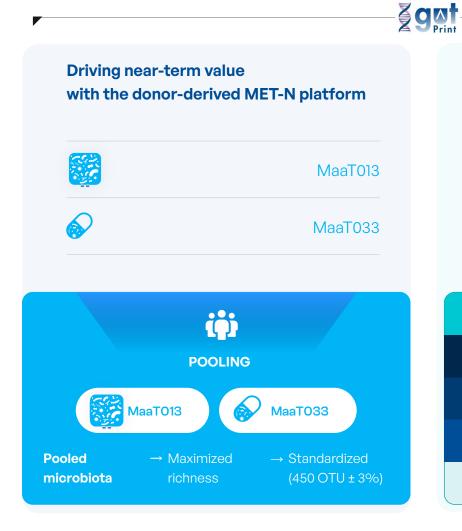
Higher response rate to ICI* in patients with metastatic melanoma³



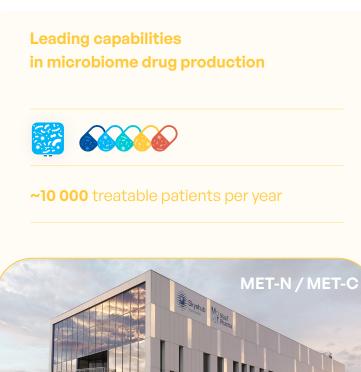
^{*} aGvHD: acute Graft-vs-host-Disease

^{&#}x27;; Peled, J.U. & al N Engl J Med 2020382:822-34; ² Ghani, 2021; Jenq RR. et al, Biol Blood Marrow Transplant 21 (2015) 1373e1383; Pamer, Blood, 2014; ³ Gopalakrishnan et al., Science, 2017, see also Routy et al, Science, 2018; Vetizou et al Science 2015;

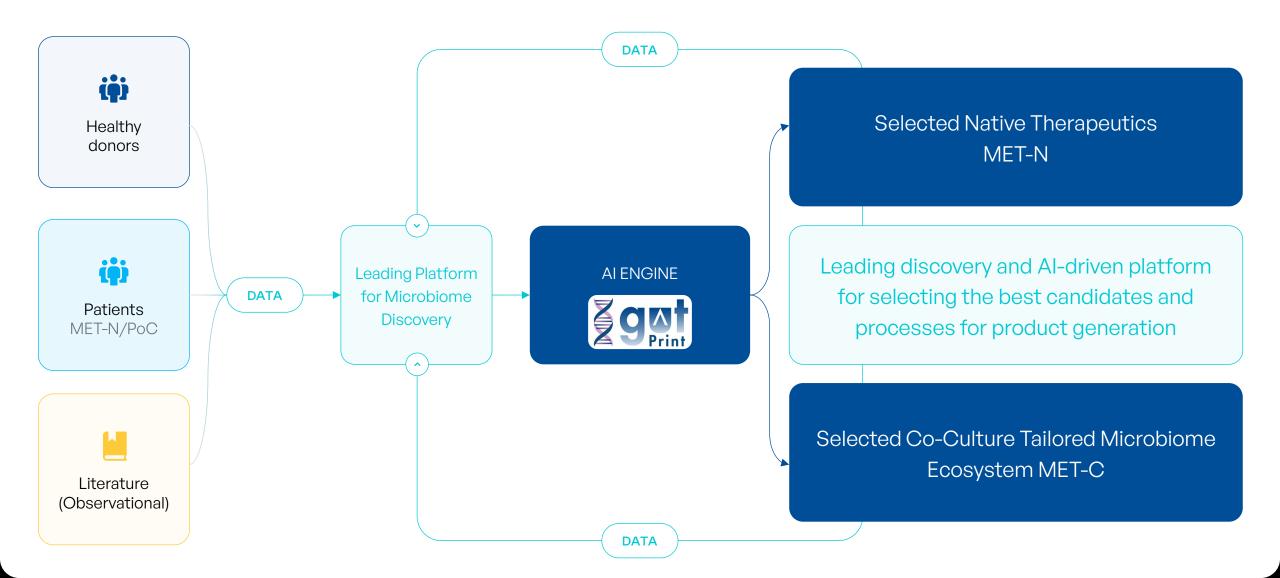
An Oncology-Focused Platform Fueling a Deep Pipeline of Drug Candidates







Al-driven Research Engine Powered by Metagenomics Enabling Candidate Selection

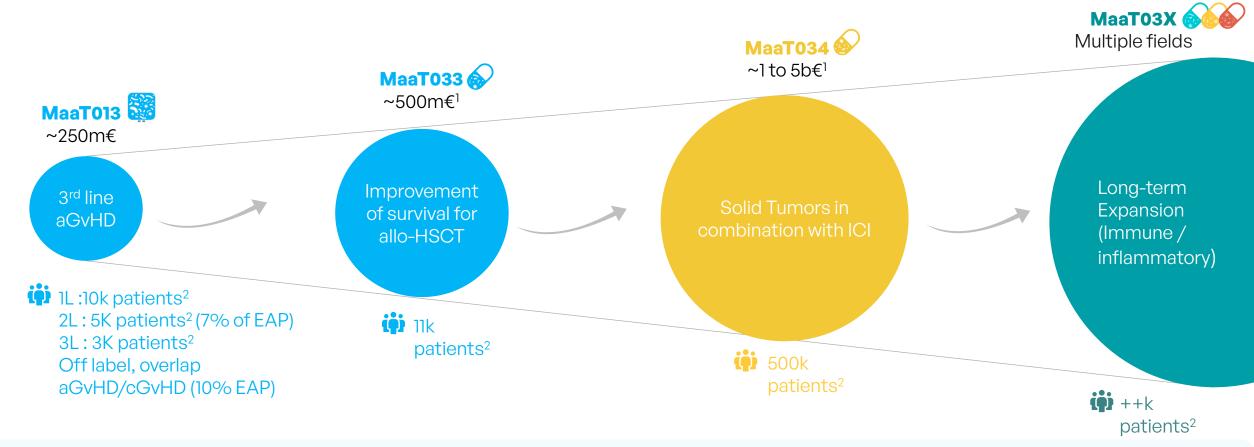


A Strong Pipeline With Multiple Near-Term Value Inflection Milestones



MaaT Pharma

Targeting Multiple Attractive Markets with Unmet Medical Need



MaaT013:

- Strong adoption from physicians with year-on-year growth and repeated requests from transplant centers
- In the EAP, we observe physicians use MaaT013 outside the third line setting: 7% in second line and 10% on overlapping a/cGvHD
- Potential expanded market estimated at ~440m€ EU/US





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

MET-N

CORPORATE PRESENTATION

Immuno-Modulation with MaaT013: A Maximum-Density Product for Fast Engraftment in Acute Situations



2 01	Characteristics	Pooled microbiota: high-richness, high-diversity, full ecosystem Microbiome Therapy containing Butycore® Non immunosuppressive treatment
<u>U</u> 02	Current indication	Acute Graft-vs-Host Disease with Gastrointestinal Involvement ~ 3k patients per year
Ф 03	Efficacy evaluation in EAP	28-Days GI-ORR: 52% 12-months OS: 47% 18-months OS: 42%
04	Available Clinical Data	HERACLES Phase 2 Clinical Trial, N=24, 2L ARES Phase 3 – Recruitment completed - Positive DSMB review (n= 30) – 3L
		Ongoing Early Access Program (EAP), N > 140, prior treatment median 2 (range 1-6)
3 05	Administration	> 250 patients treated to date 3 doses (enema bag) – within 10 days

Understanding and Addressing Acute Graft versus Host Disease

- → Acute Graft-versus-Host Disease, a severe complication following allo-HSCT
- → 50% of Allo-HSCT Patients at Risk

In aGvHD, donor immune cells attack the recipient's tissues primarily affecting the skin, liver and GI tract.

Skin GvHD Skin: Rash, itching





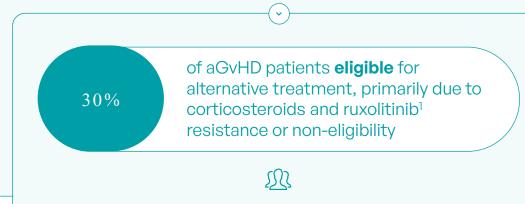




Unmet Medical Need: Acute Graft-versus-Host Disease (aGvHD) Resistant to Steroids and Ruxolitinib (3rd line of treatment)

Treatment Paradigm

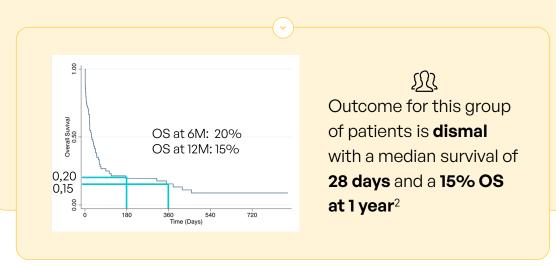
- Orticosteroids are the 1st line of treatment, but approximately 50% of patients do not achieve a sustained response
- Ruxolitinib is approved as a 2nd line of treatment for SR-aGvHD (FDA, 2019 & EMA, 2022)



Around 3,000 per year EU/US

Lack of effective therapy

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



→ GvHD is characterized by intestinal dysbiosis which is associated with higher mortality in hemato-oncology³

MaaT013 Restores Immune Homeostasis and Gut Barrier Integrity



Treatment of patients with hematological malignancies often results in microbiome dysbiosis, leading to aGvHD

Chemotherapy

Antibiotics

Irradiation

Immunosuppressants



latrogenic Dysbiosis

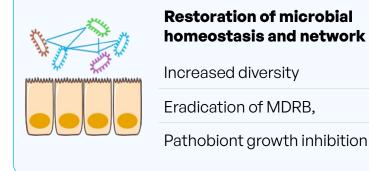
Profound imbalance of intestinal microbiota caused by medical treatments



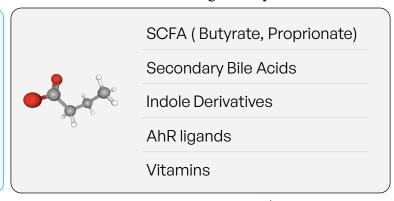
Gastrointestinal disorders (e.g., diarrhea, *C. difficile* infection)

Exacerbation and increased mortality of aGvHD

Restoration of barrier integrity



Production of immunoregulatory metabolites



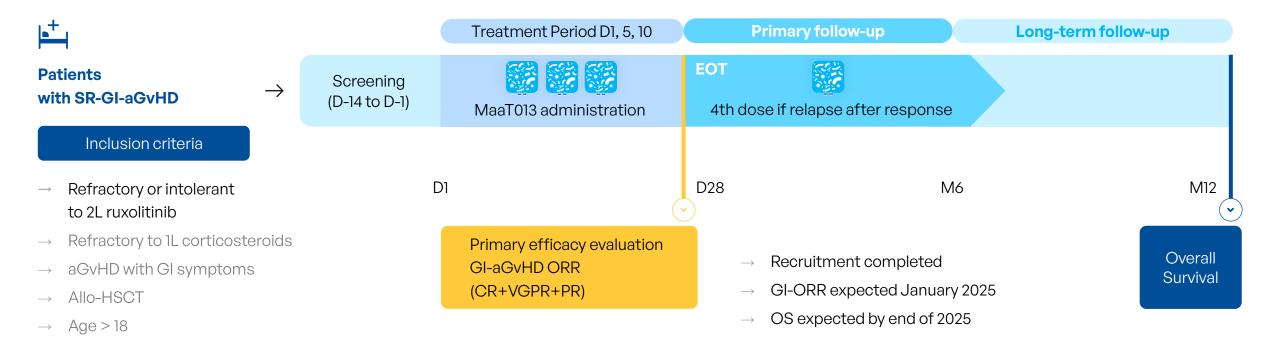
Modulation of immune homeostasis

Immune System	Immune homeostasis restoration	Treg sequestration to the gut
	 Better balance between Th17 and Treg → Anti-inflammatory cytokines (IL-10) → Pro-inflammatory cytokines (IL-6, TNF-α) 	ILCs modulation



ARES, a Pivotal Phase 3 Trial to Treat aGvHD in 3rd 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: ~ 250 m€

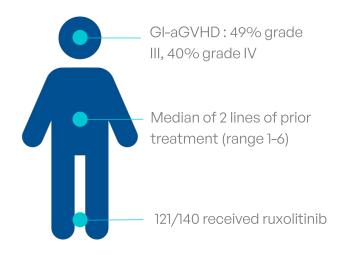
No Competitor in 3L

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



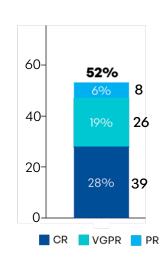
Global EAP Cohort - N=140

Patients' profile

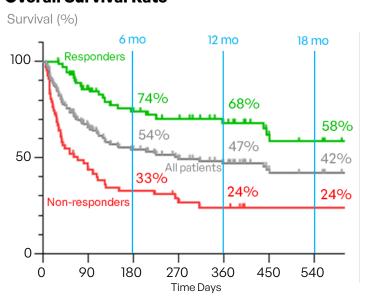


GI-ORR

Patients (%)



Overall Survival Rate



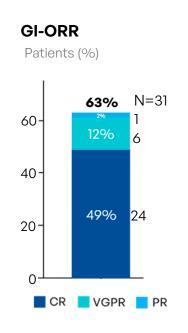
- High response rates (52%) in previously treated aGvHD patients
- High proportion of CR (39/73, 28%) and VGPR (26/73, 19%) among responders
- Effective aGvHD treatment with MaaT013 leads to prolonged patient survival

17

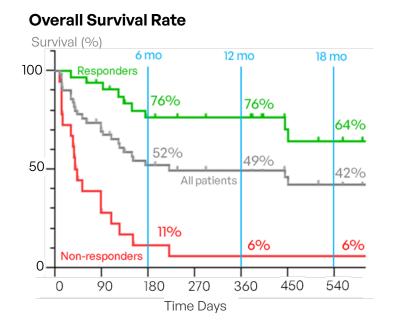
In 3L, the EAP Data Confirms Frequent Responses to MaaT013 Leading to Prolonged Survival

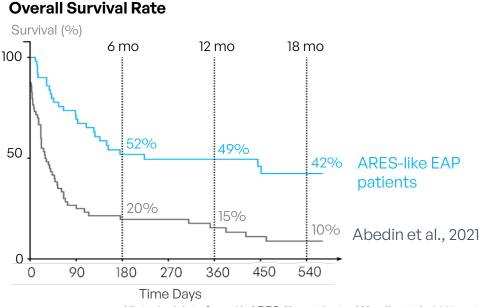


EAP: ARES like cohort - N=49, GI-aGvHD: 3L



MaaT013 aGvHD





- Historical data from 3L ARES-like patients (Abedin et al., 2021 n=48)
- No effective treatment in 3L with very low expected OS 6mo: 20%; 12mo: 15%¹ confirming strong unmet medical need
- Observed responses are almost invariably VGPR (6/31) and CR (24/31) at D28, indicating prompt and significant aGvHD control
- Remarkable improvement in overall survival (18-mo OS 42% vs 10% historical data) compared to REACH1 and Abedin et al. data 2021

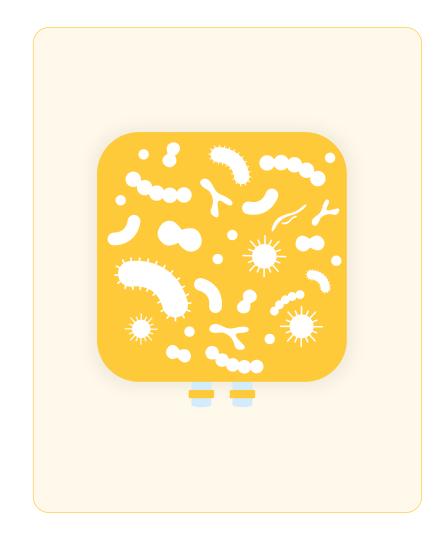
Remarkable Safety Profile of MaaT013 in Early Access Program



Very good safety compared to historical data in heavily pre-treated and fragile population

- ✓ Protective immunorestoration, not immunosupression
- ✓ Bi-annual ANSM* safety report required on Expanded Access Program (EAP)
- ✓ ARES (Ph 3) 1st DSMB review in October 26, 2023
- ✓ HERACLES (Ph 2)

*: ANSM: French Competent Authority



Proof-of-Concept with MaaTO13 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

(Davar et al, 2021)

→ Responders

→ Responders (Baruch et al, 2021)

Non-responders

⊘ 3/10

2023

FMT from healthy donors increases response of aPD1 in **ICI-naive patients** with metastatic melanoma

⊘ 13/20

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

⊘ 15/20

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

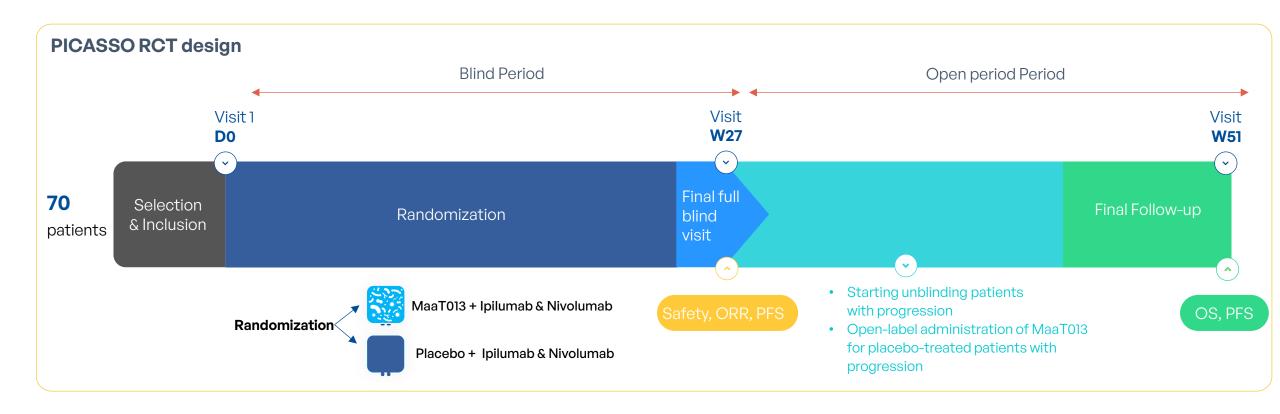
Recruitment completed Ph. 2a PICASSO trial

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

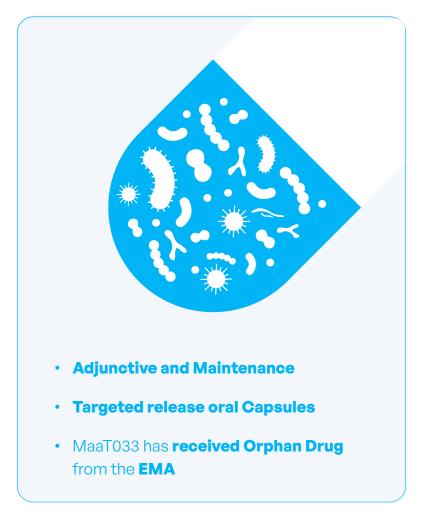
- → RCT [MaaT013 + ICI] vs. [Placebo + ICI] in 70 metastatic melanoma patients
- → Data expected Q4.24/Q1.25

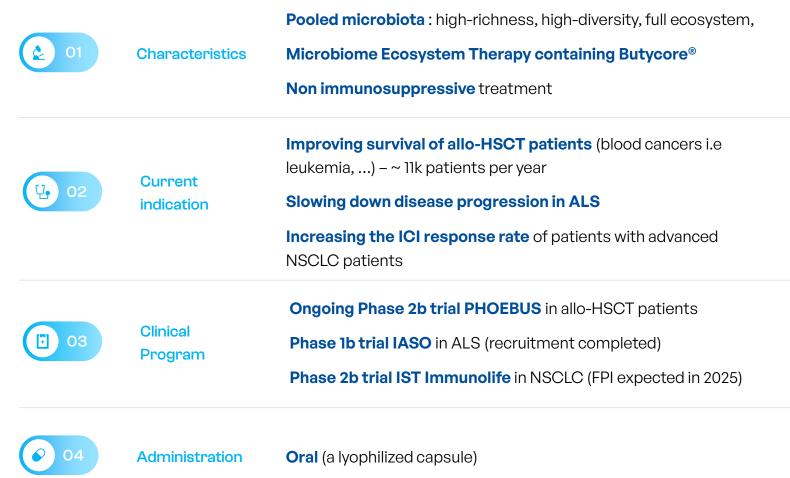
Key study endpoints after 23 weeks of treatment:

- 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 Capsule for Adjunctive and Maintenance Therapy



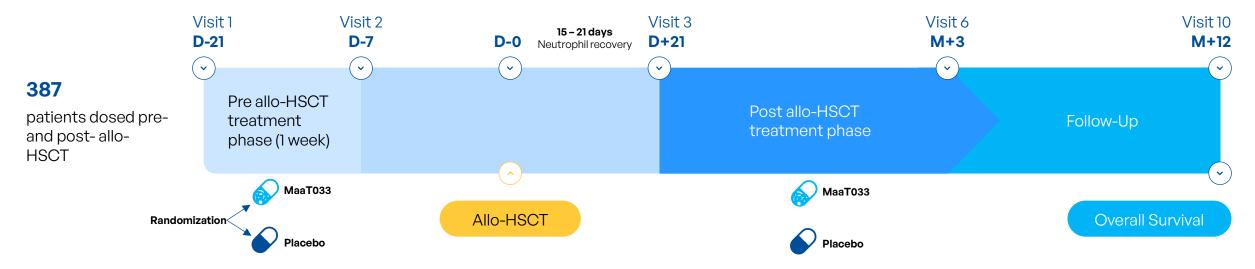


MaaT033: a Potential Adjunctive Treatment for Patients Receiving allo-HSCT



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

- → First positive DSMB (n=20*) in July 2024 safety DSMB are planned every
 6 months throughout the study
- Primary endpoint: efficacy of MaaT033 in improving overall survival at 12 months
- → Study started in November 2023, results are expected in 2027



Expansion to US sites subject to discussion with the FDA

*cutoff date: April 2024



Ongoing Phase 2b PHOEBUS



Safety Interim analysis on 60 patients in H1 2025



Based on expected duration of recruitment, OS primary endpoint expected in 2027



~ 11k patients per year

MaaTO33 Aims to Slow Down Amyotrophic Lateral Sclerosis Progression



Amyotrophic Lateral Sclerosis

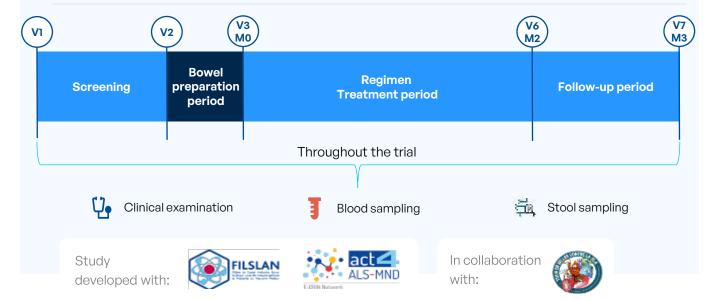
- \rightarrow Could affect up to 60,000 patients in US & EU by 2040 1
- \rightarrow Paralysis and death 3 to 5 years after diagnostic ²
- → Currently no curative treatment and few symptomatic treatments

Rationale for Exploratory Utilization of MaaTO33 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

- Study
- → Up to 15 patients in a pilot, open-label, Phase 1b study in France
- → **Key study endpoints**: safety and tolerability of MaaT033 | gut microbiota composition evolution | marker showing potential impact on disease progression
- → Study completed in **H1 2024**
- Results expected in H2 2024
- → Positive DSMB in Feb. 2024:

Trial to proceed as planned without modifications Good safety profile and generally well tolerated



¹ 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

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





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

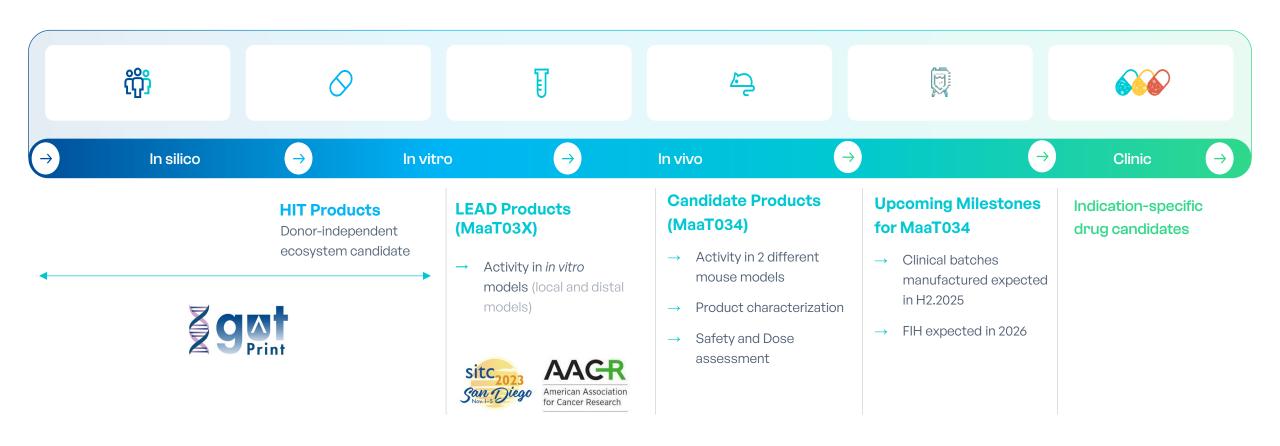
MET-C

ORPORATE RESENTATION

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October 2024

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

CORPORATE PRESENTATION

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/Q12025

MaaT013 (pooled enema)

GvHD | EAP long term follow-up EBMT24 ✓ IO Mela. | PICASSO P2a Results **Q4.24/Q1.25** GvHD | ARES P3 GI-ORR **January 2025**

MaaT033 (pooled capsule)

HSCT | PHOEBUS P2b 6-mo DSMB Study **Q2 24** ALS | IASO P1b Results **H2 2024**HSCT | PHOEBUS P2b Safety 6-mo DSMB **Q1 25**

MaaT034 (co-cultured capsule)

Selection of candidate

2025+

MaaTO13 (pooled enema)

GvHD | Final Results (OS)

MaaT033 (pooled capsule)

HSCT | PHOEBUS P2b Interim DSMB H1 25 NSCLC | IMMUNOLIFE P2a FPI H1 25

MaaT034 (co-cultured capsule)

1st Clinical Batch Manufactured H2 25 Solid Tumors IO | Target FIH 26

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

Undisclosed | Next Steps

Finance

- Revenues of MaaT013 in aGvHD of 1.7m€ for H1 2024 from Early Access Program
- Cash position of 31.2m€ as of June 30, 2024. Post follow-on in May 2024, (approx. €17.3m€) cash runway extends into Q2.2025
- 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

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Adressable Patients

Creation Value

Time:

Event:

1st Ind:

Market size:



MaaT013



Pooled enema

- → January 2025
- → P3 GI-ORR
- → aGvHD
- → 250m€

MaaT033



Pooled capsule

- → H1.2025
- → P2b DSMB
- → allo-HSCT
- → 500m€

MaaT034



Co-cultured caps.
Synthetic eubiotic microbiota

→ 2024

MET-C

- → Selection of candidate & 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 TherapiesTM 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.







Patients are our priority. We are committed to our patients and to the protection of human health by respecting environmental protection, valuing our employees and maintaining strong governance practices.

Our daily work is shaped by the following four core guidelines:

- 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	,

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sumption ees on

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	

