



# Topline Results Phase 3 MANIFEST-2 Study

*Pelabresib in First-Line Myelofibrosis*

November 2023

Gail, Living with Myelofibrosis  
since 2018

# Forward-Looking Statements

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This communication contains certain forward-looking statements concerning the MorphoSys group of companies, including the expectations regarding Monjuvi's ability to treat patients with relapsed or refractory diffuse large B-cell lymphoma ("DLBCL"), the further clinical development of tafasitamab, including ongoing confirmatory trials, additional interactions with regulatory authorities and expectations regarding future regulatory filings and possible additional approvals for tafasitamab as well as the commercial performance of Monjuvi. The words "anticipate", "believe", "estimate", "expect", "intend", "may", "plan", "predict", "project", "would", "could", "potential", "possible", "hope" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. The forward-looking statements contained herein represent the judgment of MorphoSys as of the date of this release and involve known and unknown risks and uncertainties, which might cause the actual results, financial condition and liquidity, performance or achievements of MorphoSys, or industry results, to be materially different from any historic or future results, financial conditions and liquidity, performance or achievements expressed or implied by such forward-looking statements. In addition, even if MorphoSys' results, performance, financial condition and liquidity, and the development of the industry in which it operates are consistent with such forward-looking statements, they may not be predictive of results or developments in future periods. Among the factors that may result in differences are MorphoSys' expectations regarding risks and uncertainties related to the impact of the COVID-19 pandemic to MorphoSys' business, operations, strategy, goals and anticipated milestones, including its ongoing and planned research activities, ability to conduct ongoing and planned clinical trials, clinical supply of current or future drug candidates, commercial supply of current or future approved products, and launching, marketing and selling current or future approved products, the global collaboration and license agreement for tafasitamab, the further clinical development of tafasitamab, including ongoing confirmatory trials, and MorphoSys' ability to obtain and maintain requisite regulatory approvals and to enroll patients in its planned clinical trials, additional interactions with regulatory authorities and expectations regarding future regulatory filings and possible additional approvals for tafasitamab as well as the commercial performance of Monjuvi, MorphoSys' reliance on collaborations with third parties, estimating the commercial potential of its development programs and other risks indicated in the risk factors included in MorphoSys' Annual Report on Form 20-F and other filings with the U.S. Securities and Exchange Commission. Given these uncertainties, the reader is advised not to place any undue reliance on such forward-looking statements. These forward-looking statements speak only as of the date of publication of this document. MorphoSys expressly disclaims any obligation to update any such forward-looking statements in this document to reflect any change in its expectations with regard thereto or any change in events, conditions or circumstances on which any such statement is based or that may affect the likelihood that actual results will differ from those set forth in the forward-looking statements, unless specifically required by law or regulation.

The compounds discussed in this slide presentation are investigational products being developed by MorphoSys and its partners and are not currently approved by the U.S. Food and Drug Administration (FDA), European Medicines Agency (EMA) or any other regulatory authority (except for tafasitamab/Monjuvi® and tafasitamab/Minjuvi® in relapsed or refractory DLBCL). The safety and efficacy of these investigational products have not been established and there is no guarantee any investigational product will be approved by regulatory authorities.

Monjuvi® and Minjuvi® are registered trademarks of MorphoSys AG.

# Agenda

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**01**

## **Opening Remarks**

Jean-Paul Kress, M.D., Chief Executive Officer (CEO)

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**02**

## **Phase 3 MANIFEST-2 Study Topline Results**

Tim Demuth, M.D., Ph.D., Chief Research & Development Officer (CR&DO)

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**03**

## **Myelofibrosis in Clinical Practice**

John Mascarenhas, M.D., Professor of Medicine and Director of the Adult Leukemia Program at The Tisch Cancer Institute at Mount Sinai, New York

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**04**

## **Q&A**

Jean-Paul Kress, M.D., Tim Demuth, M.D., Ph.D., Lucinda Crabtree, Ph.D., John Mascarenhas, M.D.

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# MANIFEST-2 Study Results Demonstrate Most Impressive Benefits Seen in Clinical Studies of Patients with Myelofibrosis

## KEY FINDINGS

- Achieved primary endpoint, nearly doubling SVR35
- Significant improvements in key secondary endpoints (absolute change in TSS and TSS50) for intermediate-risk patients
- Strong positive trend in key secondary endpoints for overall population
- Clinically meaningful anemia improvement
- Safety results consistent with prior trials, no new safety signals



## NEXT STEPS

Present detailed findings at 2023 ASH Annual Meeting

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Submit for approval in the U.S. and Europe mid-2024

Pelabresib is an investigational medicine that has not yet been approved by any regulatory authorities.

# Phase 3 MANIFEST-2 Study: One of the Largest Myelofibrosis Studies Ever Conducted

**430** *JAK-inhibitor-naïve myelofibrosis patients randomized, representative of the disease population and aligned with NCCN criteria*

## KEY ENDPOINTS

**PRIMARY:**  
SVR35 at week 24

**SECONDARY:**  
Absolute change in TSS at week 24  
TSS50 at week 24  
(MFSAF v4.0)

## ADDITIONAL ENDPOINTS\*

Duration of  
the splenic  
response

Duration of  
the symptom  
response

Improvement  
in bone  
marrow fibrosis

Hemoglobin  
response

Progression-free  
survival

Overall  
survival

SVR35,  $\geq 35\%$  reduction in spleen volume; TSS, total symptom score; TSS50,  $\geq 50\%$  reduction in total symptom score; MFSAF, Myelofibrosis Symptom Assessment Form

\*Only includes sample of additional endpoints being assessed in Phase 3 MANIFEST-2 study

# Statistically Significant and Clinically Meaningful Improvement in Primary Endpoint: SVR35

POPULATION	SVR35 (Pelabresib + Ruxolitinib)	SVR35 (Ruxolitinib + Placebo)	Difference
<b>All Patients</b> (N = 430)	66%	35%	30.4%* P-value: p<0.001
<b>Intermediate-Risk Patients</b> (N = 400)	66%	36%	29.9%* P-value: p<0.001
<b>High-Risk Patients</b> (N = 30)	64%	25%	39.3% P-value: 0.063

\*Difference calculated using Cochran–Mantel–Haenszel (CMH) common risk difference

# Strong Positive Trend in Key Secondary Endpoints: Absolute Change in TSS and TSS50

POPULATION	Absolute Change in TSS (Pelabresib + Ruxolitinib)	Absolute Change in TSS (Ruxolitinib + Placebo)	Difference	TSS50 (Pelabresib + Ruxolitinib)	TSS50 (Ruxolitinib + Placebo)	Difference
<b>All Patients</b> (N = 430)	-15.99	-14.05	-1.94* P-value: 0.0545	52%	46%	6.0%*** P-value: 0.216
<b>Intermediate Risk Patients</b> (N = 400)	-15.2	-12.7	-2.4* P-value: <0.02	55%	45%	10.05%*** P-value: <0.05
<b>High-Risk Patients</b> (N = 30)	N/A**	N/A**	N/A**	21%	69%	-47.3% P-value: <0.05

\*Least square mean estimate; \*\*No calculation due to missing data rate; \*\*\*Difference calculated using Cochran–Mantel–Haenszel (CMH) common risk difference

# Myelofibrosis in Clinical Practice

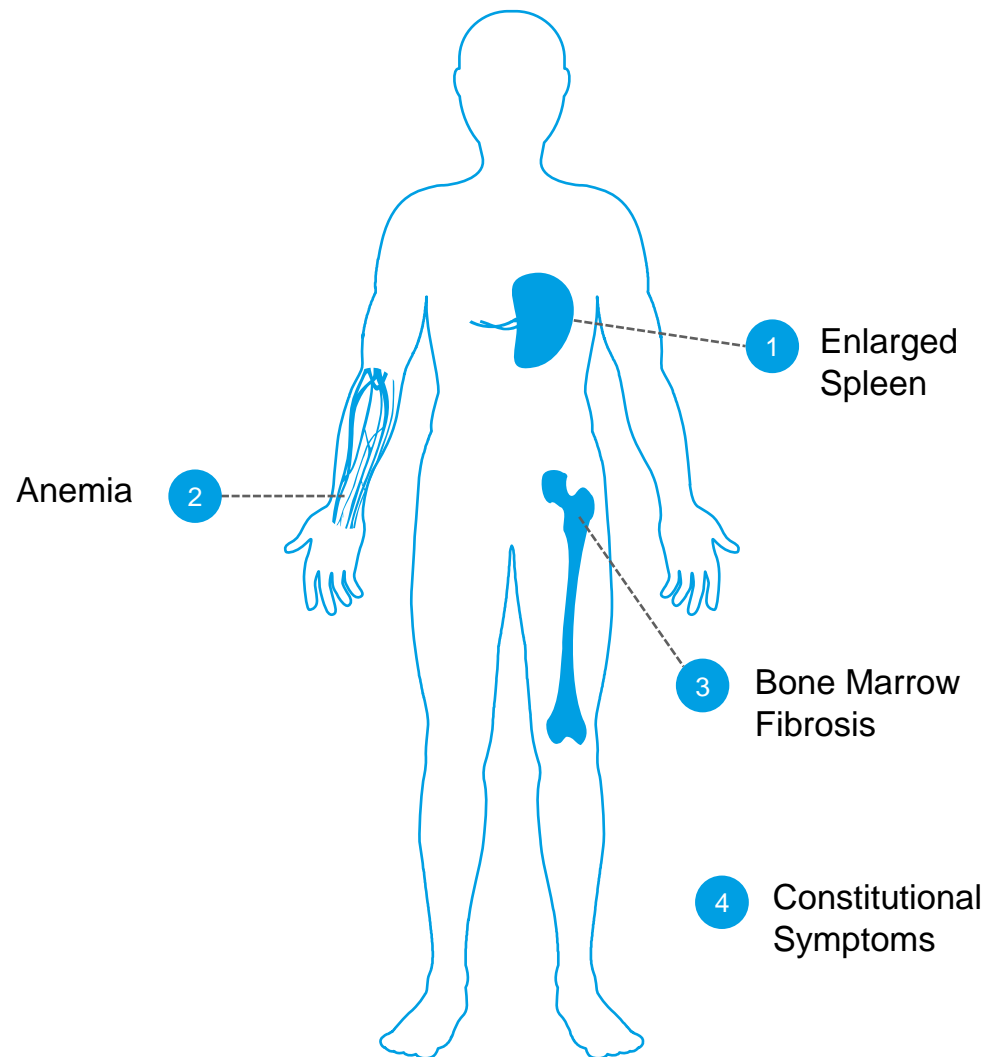


**JOHN MASCARENHAS, M.D.**

Professor of Medicine and Director of the Adult Leukemia Program  
at The Tisch Cancer Institute at Mount Sinai, New York



# Myelofibrosis is a Debilitating, Progressive and Often Deadly Blood Cancer, Characterized by Four Hallmarks



## DIAGNOSIS (DIPSS)

- + Intermediate-risk: **~77% – 86%**
- + High-risk: **~9% – 11%**

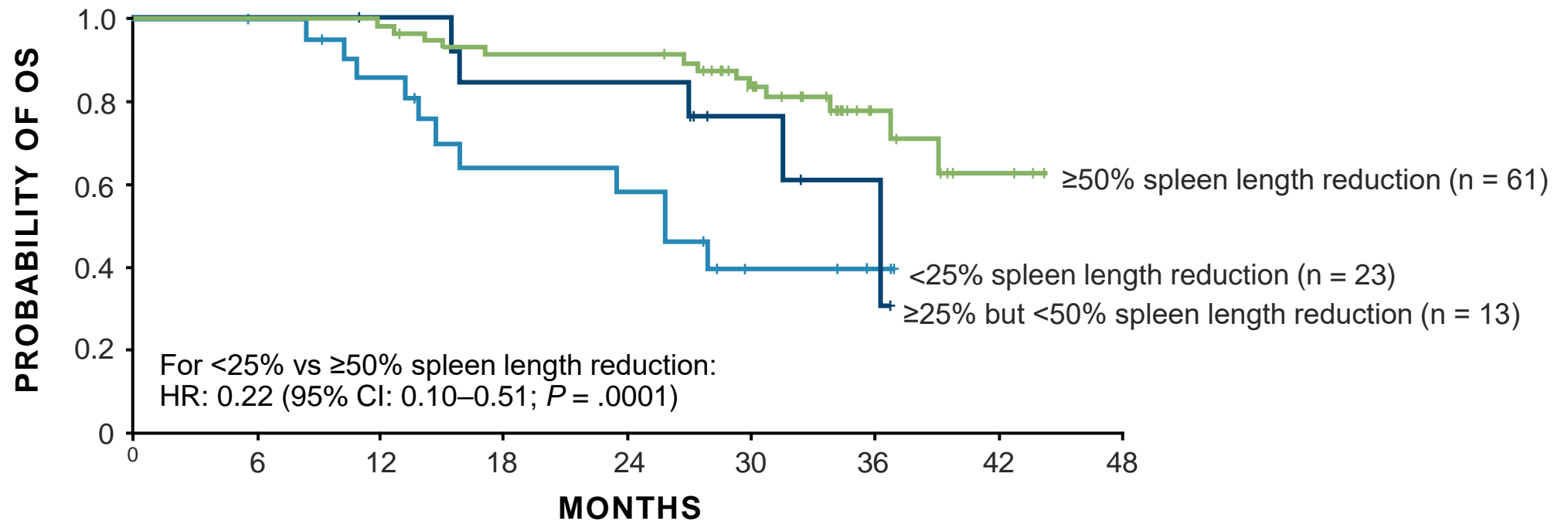
## MEDIAN OVERALL SURVIVAL (DIPSS)

- + Intermediate-risk: **~4 – 14.2 years**
- + High-risk: **~1.5 years**

Passamonti, F et al. Hematological Oncology 2021; Szuber N, et al. Mayo Clinic 2019; Passamonti, F et al. Blood 2010

# Survival Improves With Spleen Length Reduction in Patients Receiving Ruxolitinib

## OPEN-LABEL, SINGLE-ARM PHASE I/II STUDY (N = 107)

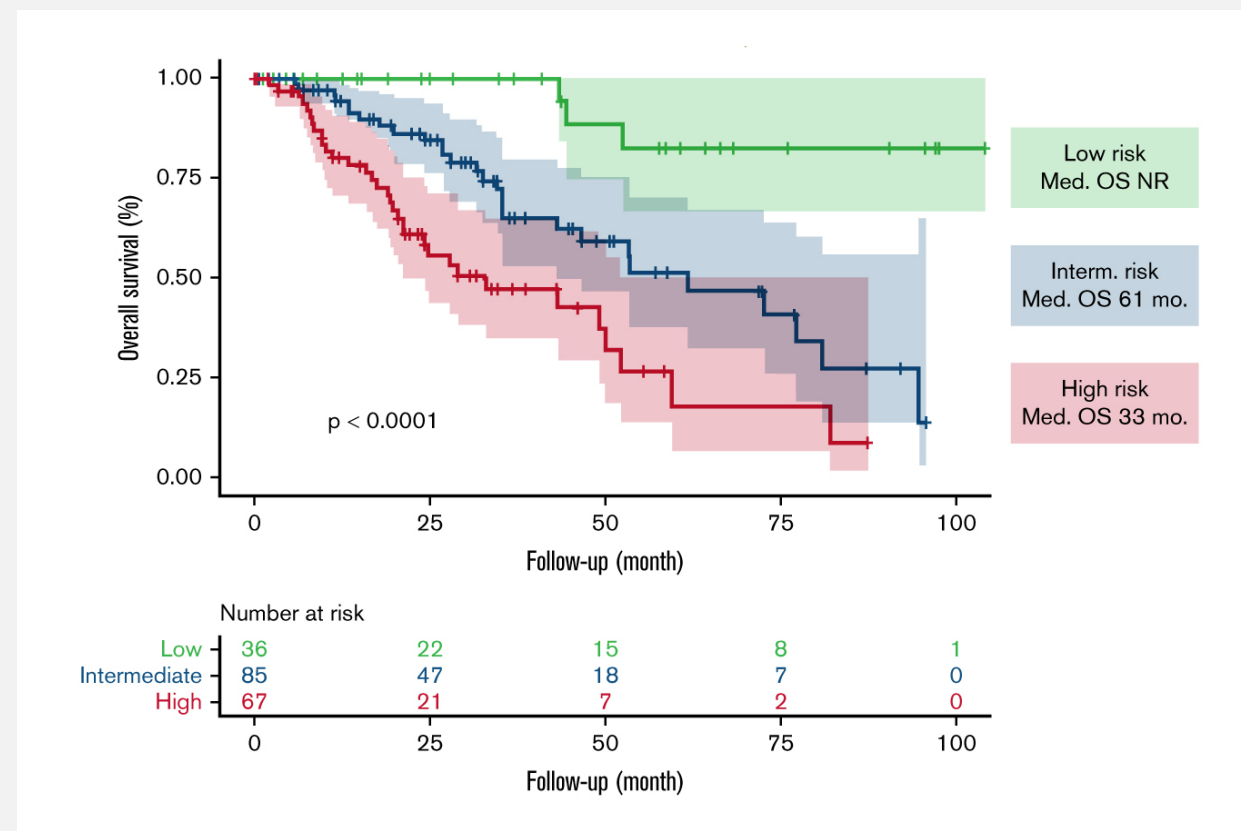


HR, hazard ratio; OS, overall survival. | Verstovsek S, et al. *Blood*. 2012;120:1202-1209.

# RR6: Three Factors Predict Survival Benefit

Risk Factors	Multivariate, HR (95% CI); P-value
RUX dose below 20 mg BID at baseline, 3 months and 6 months	HR = 1.79 (1.07-3.00); P = 0.03
<b>Splenomegaly reduction ≤30% by palpation at 3 months and 6 months</b>	<b>HR = 2.26 (1.40-3.65); P = 0.0009</b>
RBC transfusion need at baseline, 3 months and 6 months	HR = 2.32 (1.19-4.54); P = 0.02

The RR6 model was validated in another cohort of patients (n = 40; P = 0.0276) treated with ruxolitinib at Moffitt Cancer Center



RR6 = response to ruxolitinib after 6 months | Maffioli M. *Blood Adv.* 2022;6 (6):1855-1864.

# MANIFEST-2 Provides Valuable Evidence, Cementing Position of Pelabresib and Ruxolitinib Combination Potential

1. **BET inhibition is rational** and supported by pre-clinical data
2. **Combination of pelabresib and ruxolitinib is clinically active**
  - SVR35 statistically significant and clinically meaningful in overall population
  - Strong numerical improvement in symptom reduction, significant improvement in intermediate-risk patients
3. **Well-tolerated therapy**
  - Safety results consistent with prior trials, no new safety signals
4. **Correlative evidence of biologic disease modification**
  - Hemoglobin level improvement
  - Anemia AE improvement
5. **Paradigm Shift – Combination Therapy:**
  - Support use of combination treatment
  - Start early to prevent patients getting more ill

# Pelabresib and Ruxolitinib Combination Therapy: Potential to Shift Treatment Paradigm in Myelofibrosis



**Most Impressive  
Benefits Seen in  
Myelofibrosis**



**File for  
Approval in the U.S.  
and Ex-U.S.**



**Multi-Billion Dollar  
Market Opportunity**

# 04

## Q&A



**Jean-Paul Kress,  
M.D.**



**Tim Demuth,  
M.D., Ph.D.**



**Lucinda Crabtree,  
Ph.D.**



**John Mascarenhas,  
M.D.**





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**Thank you!**

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