

# mesoblast

# Global Leader in Allogeneic Cellular Medicines for Inflammatory Diseases

Presentation at ISCT

Ryoncil® - The First FDA Approved Mesenchymal Stromal Cell Therapy

ASX: MSB; Nasdaq: MESO



#### CAUTIONARY NOTE REGARDING FORWARD-LOOKING STATEMENTS

This presentation includes forward-looking statements that relate to future events or our future financial performance and involve known and unknown risks, uncertainties and other factors that may cause our actual results, levels of activity, performance or achievements to differ materially from any future results, levels of activity, performance or achievements of implied by these forward-looking statements. We make such forward-looking statements pursuant to the safe harbor provisions of the Private Securities Litigation Reform Act of 1995 and other federal securities laws. All statements other than statements of historical facts contained in this presentation are forward-looking statements. Words such as, but not limited to, "believe," "expect," "anticipate," "estimate," "intend," "flangets," "likely," "will," "would," "could," and similar expressions or phrases identify forward-looking statements. We have based these forward-looking statements largely on our current expectations and future events, recent changes in regulatory laws, and financial trends that we believe may affect our financial condition, results of operation, business strategy and financial needs. These statements may relate to, but are not limited to: expectations regarding the safety or efficacy of, or potential applications for, Mesoblast's adult stem cell technologies; expectations regarding the strength of Mesoblast's intellectual property, the timeline for Mesoblast's regulatory approval process, and the scalability and efficiency of manufacturing processes; expectations about Mesoblast's ability to grow its business and statements regarding its relationships; statements oncerning Mesoblast's obsolast's share price or potential market capitalization; and statements concerning Mesoblast's capital requirements and ability to raise future capital, among others. Forward-looking statements should not be read as a guarantee of future performance or results, and actual results may differ from the results anticipated in these forward-looking s

Mesoblast is committed to bringing to market innovative off-the-shelf allogeneic cellular medicines to treat serious and life-threatening inflammatory illnesses

# **Our Mission**

# Global leader in allogeneic cellular medicines for inflammatory diseases

- ✓ World leader in developing allogeneic (off-theshelf) cellular medicines for the treatment of severe and life-threatening inflammatory conditions
- Locations in Australia, the United States and Singapore
- ✓ Listed on the ASX (MSB) and NASDAQ (MESO)
- Developing product candidates for distinct indications based on its remestemcel-L and rexlemestrocel-L stromal cell technology platforms
- ✓ Extensive global intellectual property portfolio with protection extending through to at least 2041 in all major markets
- ✓ FDA-inspected commercial scale manufacturing process and facilities

ONE product FDA approved



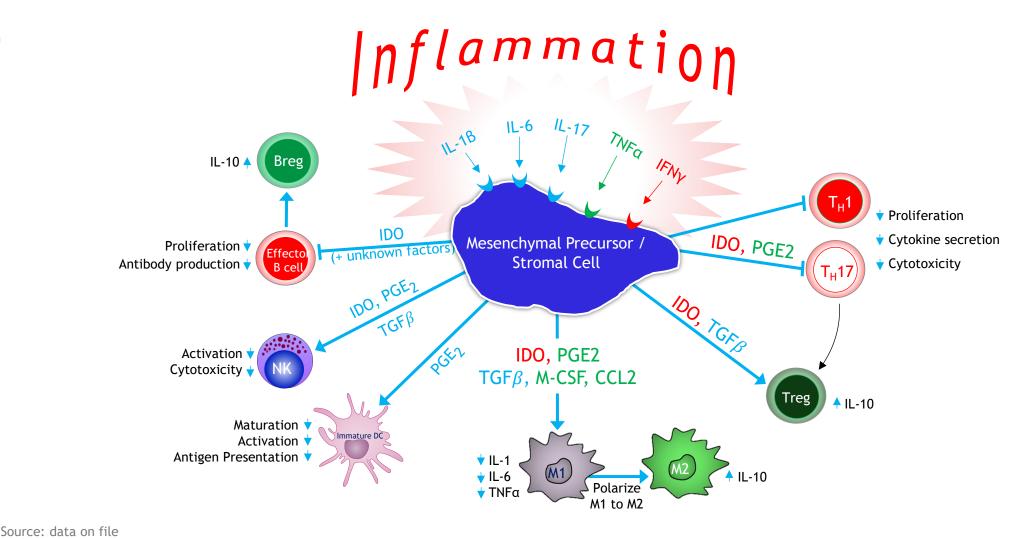
nore than
1,100
patents & applications

Phase 3 trials in **TWO** major indications



## Platform technology: shared mechanism of action across our products

Our mesenchymal precursor/stromal cells respond to and are activated by multiple inflammatory cytokines through surface receptors, resulting in orchestration of an anti-inflammatory cascade





## Mesoblast allogeneic Mesenchymal Precursor / Stromal Cell portfolio

Product	Indication	Phase 2	Phase 3	Regulatory Filing	Approved
RYONCIL® remestemcel-L	Pediatric SR-aGvHD				
	Adult SR-aGvHD		<b>&gt;&gt;&gt;</b>		
RYONCIL® remestemcel-L	IBD / Crohn's		<b>&gt;&gt;</b>		
REVASCOR® rexlemestrocel-L (STRO3+)	Pediatric HLHS		<b>&gt;&gt;&gt;</b>		
	Adult HFrEF End-stage		>:		
	Adult HFrEF Class II/III		<b>&gt;&gt;</b>		
Rexlemestrocel-L (STRO3+)	CLBP		<b>&gt;&gt;&gt;</b>		

SR-aGvHD = Steroid-Refractory Acute Graft versus Host Disease; IBD = Inflammatory Bowel Disease; HLHS = Hypoplastic Left Heart Syndrome HFrEF = Heart Failure with Reduced Ejection Fraction; CLBP = Chronic Low Back Pain;

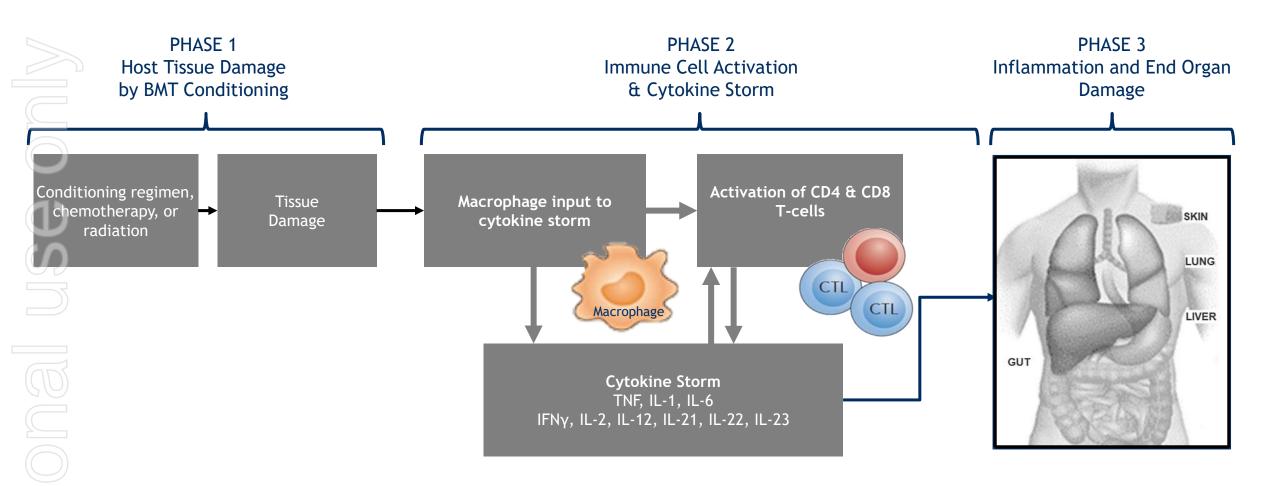
This chart is figurative and does not purport to show individual trial progress within a clinical program Notes:

- JCR Pharmaceuticals Co., Ltd. (JCR), has the right to develop mesenchymal stromal cells (MSCs) in certain fields for the Japanese market, including for the treatment of hematological malignancies, such as Graft vs Host Disease, and for hypoxic ischemic encephalopathy (HIE).
- Grünenthal has an exclusive license to develop and commercialize rexlemestrocel-L for chronic low back pain in Europe and Latin America/Caribbean.
- Tasly Pharmaceuticals has exclusive rights for rexlemestrocel-L for the treatment or prevention of chronic heart failure in China.





# Acute graft versus host disease (aGvHD) is a serious and potentially fatal complication of allogeneic bone marrow transplantation (BMT)





# Opportunity to address critical unmet need in children 2 months and older, including adolescents & teenagers with SR-aGVHD

>30,000 allogeneic BMTs performed globally (>20K US/EU) annually, ~20% pediatric<sup>2,3</sup>

Corticosteroids are first-line therapy for aGvHD

RYONCIL is the only approved therapy for SR-aGvHD in children 2 months and older

~1,500
Children & adolescents
undergo allogeneic BMT
in US annually

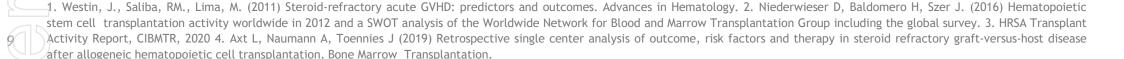
~50%
Incidence of acute
GvHD

~375 pts/year with SR-aGvHD

Approx. 10,000 allogeneic BMTs performed in the US annually

Acute GvHD occurs in ~50% of patients<sup>1</sup>
with approx. half failing to respond to steroids

SR-aGvHD has high mortality<sup>1,4</sup> and significant extended hospital stay costs<sup>2</sup>





### Ryoncil® for treating pediatric patients with SR-aGvHD

The recommended dosage of Ryoncil® for treatment of pediatric SR-aGvHD is 2×106 MSC/kg body weight per intravenous infusion given twice per week for 4 consecutive weeks





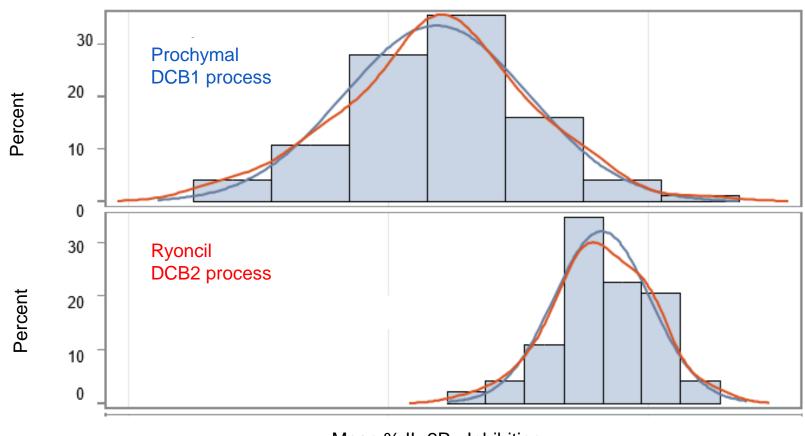






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Potency for Drug Product Lots used in EAP275



Mean % IL-2Rα Inhibition

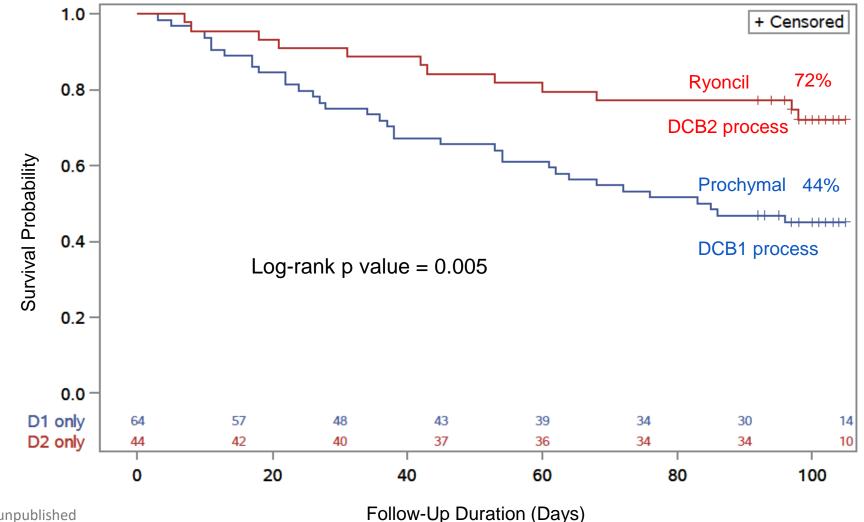
100% of Ryoncil lots released meet FDA approved potency release criteria, compared with less than two thirds of Prochymal lots used in adult GvHD trial 280



# Improvements in manufacturing process give rise to Ryoncil

- a product associated with improved survival in SR-aGvHD





# Ryoncil® delivered high overall response rates at Day 28, a measure that predicts survival in aGvHD¹

### MSB-GVHD001<sup>2,3</sup>

(n=54)
Single-arm, multi-center
Phase 3

Overall Response Rate at Day 28
95% CI 56.4, 82.0

70%

SR-GvHD severity<sup>4</sup> at baseline in GVHD001:

**Grade B: 11%** 

**Grade C: 43%** 

**Grade D: 46%** 

RYONCIL treatment was not discontinued or interrupted in any patient for any laboratory abnormality, and the full course was completed without interruption in more than 85% of patients

Full Prescribing Information at www.ryoncil.com

1. MacMillan ML, Holtan SG, Rashidi A, et al. "Pediatric acute GVHD: clinical phenotype and response to upfront steroids." Bone marrow transplantation vol. 55,1 (2020): 165-171; 2.NCT02336230; 3. Kurtzberg, J. et al. A Phase 3, Single-Arm, Prospective Study of Remestemcel-L, Ex Vivo Culture-Expanded Adult Human Mesenchymal Stromal Cells for the Treatment of Pediatric Patients Who Failed to Respond to Steroid Treatment for Acute Graft-versus-Host Disease. Biol Blood Marrow Transplant 26 (2020) 845-854

m e s o b l a s t

https://doi.org/10.1016/j.bbmt.2020.01.018; 4. International Blood and Marrow Transplantation Registry Severity Index Criteria (IBMTR)

# Ryoncil® long-term survival free from aGvHD

Long-term followup of Ryoncil by the Center for International Blood and Marrow Transplant Research (CIBMTR)

Children from GVHD001

N = 51

88% Grade C/D

Year 2 Survival:

51%

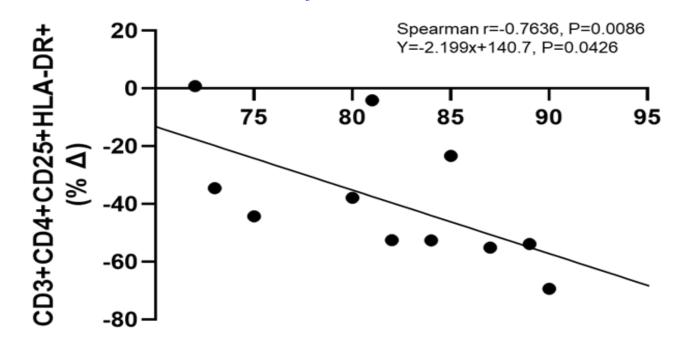
Year 4 Survival:

49%

Only 14% (N=7) died due to aGvHD through 4 years



Ryoncil® lot potency, measured by IL-2Ra Inhibition Assay, correlates with reduction in circulating levels of activated T cells in patients with SR-aGVHD: evidence for immunomodulatory mechanism of action (MOA)



% IL-2Rα Inhibition in vitro

Supporting evidence for Ryoncil immunomodulatory MOA from pharmacodynamic analysis of blood samples in 40 subjects in GVHD001:

- Circulating levels of CD3+CD4+CD25+HLA-DR+ activated T cells were reduced by 64% at Day 180 as compared to baseline.
  - Tumor necrosis factor receptor 1 (TNFR1) levels were reduced by 79% at Day 180 as compared to baseline.
- Suppressor of tumorigenesis 2 (ST2) levels, a biomarker of gut inflammation, were reduced by 75% at Day 180 as compared to baseline.

  mesoblast data unpublished





### **NOW APPROVED**

Mesoblast is excited to announce that an innovative new treatment option is now approved for pediatric patients. Read the press release to learn more about this clinical advancement.

**Visit RYONCIL.com** 

RYONCIL® is the first FDA-approved, offthe-shelf cell therapy for children aged 2 months and older, including adolescents and teenagers, with steroid-refractory acute graft versus host disease (SR-aGvHD), a life-threatening condition with high mortality rates.<sup>1</sup>



Press Release available at www.mesoblast.com

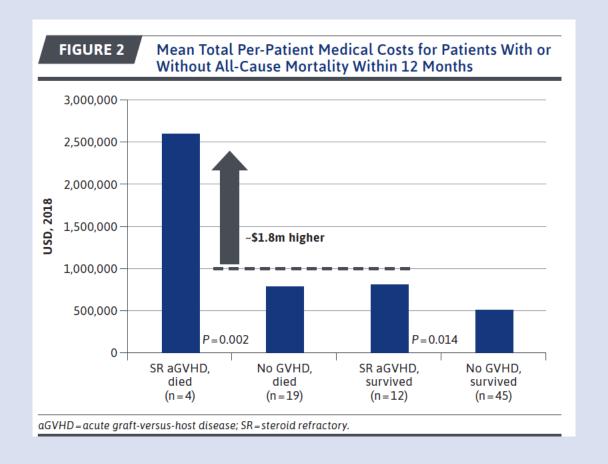
1. Please see the full Prescribing Information at <a href="https://www.ryoncil.com">www.ryoncil.com</a>



### High cost of treating child who dies from SR-aGvHD

The cost of treating a child who dies of SR-aGVHD within 12 months of transplant is:

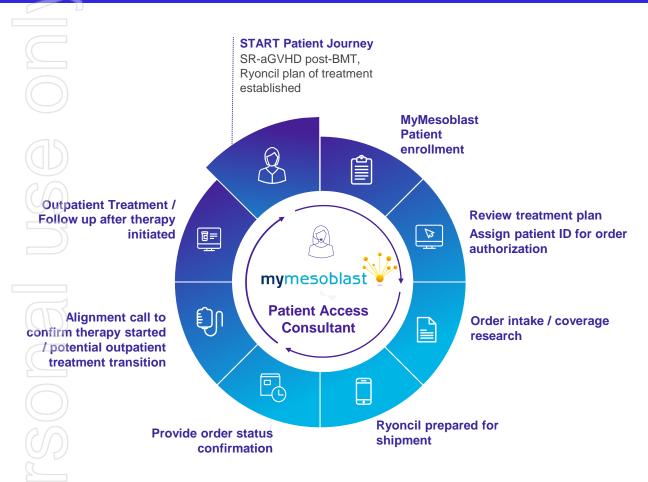
- Approximately \$2.5M
- \$1.8M higher than for those with SR aGvHD who remain alive<sup>1</sup>





### MyMesoblast Mandatory Hub

To assist patients with insurance coverage, financial assistance, and access programs, ensuring that no patient is left behind in receiving this potentially life-saving therapy, a comprehensive patient services hub has been established







### Availability of Ryoncil® for pediatric SR-aGvHD in the U.S. in March 2025

Staged approach based on transplant centers with highest volume and experience with Ryoncil® product

Targeted sales force with experience in bone marrow transplant centers

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15 highest volume centers account for ~50% of patients

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Targeting 45 highest volume centers / 80% of patients





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Thank You

