

## Updated Investor Presentation

**Melbourne, Australia; 8 March 2022:** Cynata Therapeutics Limited (ASX: “CYP”, “Cynata”, or the “Company”), a clinical-stage biotechnology company specialising in cell therapeutics, is pleased to release an updated Corporate Investor Presentation which will be used to update existing shareholders, potential investors, and other parties.

The presentation provides an overview of the key competitive advantages of Cynata’s proprietary Cymerus™ technology. Cymerus overcomes manufacturing challenges associated with conventional mesenchymal stem cell (MSC) manufacturing strategies, including inter-donor variability (requiring multiple donors), product inconsistency and quantity limitations.

Cynata has a rich and diverse clinical pipeline with three active clinical trials: a Phase 3 trial in osteoarthritis, the MEND trial in respiratory distress, and a recently commenced trial in diabetic foot ulcers. The Company also recently signed a Strategic Partnership Agreement (SPA) with FUJIFILM, leading on to an agreement with Fujifilm Cellular Dynamics, Inc (a subsidiary of Fujifilm) to produce and supply Cymerus MSCs for clinical and commercial purposes. As part of the SPA, Cynata regained rights to CYP-001 (Cynata’s lead product candidate) in graft-versus-host-disease (GvHD) and planning is now underway for a Phase 2 GvHD clinical trial in the USA.

The presentation is attached to this announcement.

-ENDS-

Authorised for release by Dr Ross Macdonald, Managing Director & CEO

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### About Cynata Therapeutics (ASX: CYP)

Cynata Therapeutics Limited (ASX: CYP) is an Australian clinical-stage stem cell and regenerative medicine company focused on the development of therapies based on Cymerus™, a proprietary therapeutic stem cell platform technology. Cymerus™ overcomes the challenges of other production methods by using induced pluripotent stem cells (iPSCs) and a precursor cell known as mesenchymoangioblast (MCA) to achieve economic manufacture of cell therapy products, including mesenchymal stem cells (MSCs), at commercial scale without the limitation of multiple donors.

Cynata’s lead product candidate CYP-001 met all clinical endpoints and demonstrated positive safety and efficacy data for the treatment of steroid-resistant acute graft-versus-host disease (GvHD) in a Phase 1 trial. Planning for a Phase 2 clinical trial in GvHD is presently underway. Clinical trials of Cymerus products in osteoarthritis (Phase 3), respiratory failure and diabetic foot ulcers (DFU) are currently ongoing. In addition, Cynata has demonstrated utility of its Cymerus technology in preclinical models of numerous diseases, including the clinical targets mentioned above, as well as critical limb ischaemia, idiopathic pulmonary fibrosis, asthma, heart attack, sepsis, acute respiratory distress syndrome (ARDS) and cytokine release syndrome.

**Cynata Therapeutics encourages all current investors to go paperless by registering their details with the designated registry service provider, Automic Group.**



# A Next Generation Stem Cell Therapeutics Company

Investor Presentation (March 2022)

# Investment Highlights

Cynata is a clinical stage biotech developing its proprietary Cymerus™ platform technology for the scalable manufacture of mesenchymal stem cell (MSC) therapeutic products to treat serious disorders



## Unique Manufacturing

**Single donation from a single donor** overcomes suboptimalities in conventional MSC manufacturing



## Strong safety and efficacy

**Positive pre-clinical and clinical data** supporting versatility and efficacy of Cynata's MSCs

**Validation** through corporate partnering



## Multiple clinical trials underway

Rich clinical pipeline:

- **Diabetic Foot Ulcers**
- **Respiratory distress (ARDS)**
- **Osteoarthritis** (Phase 3)

**Phase 2 GvHD** trial to commence in 2022



## Large addressable market

**Combined market opportunity** of clinical trials underway and in planning is **~A\$46bn**



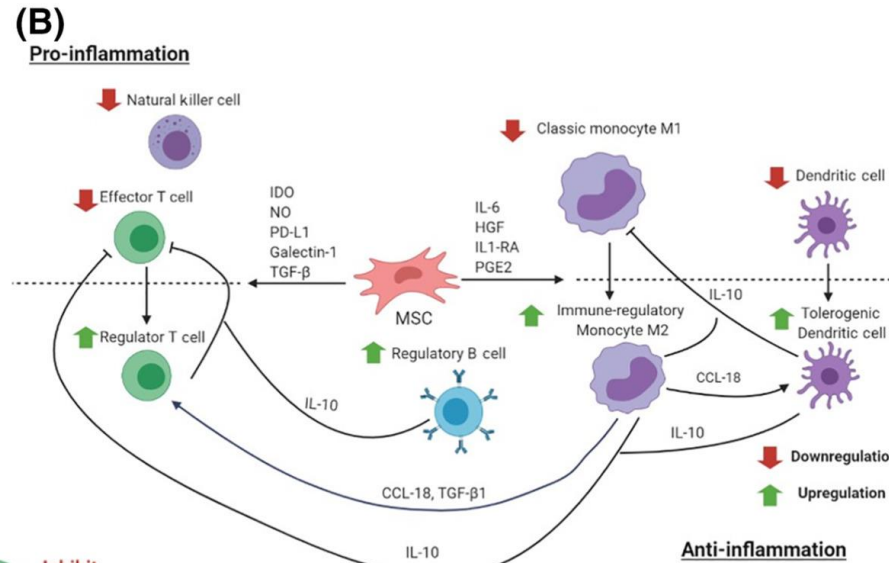
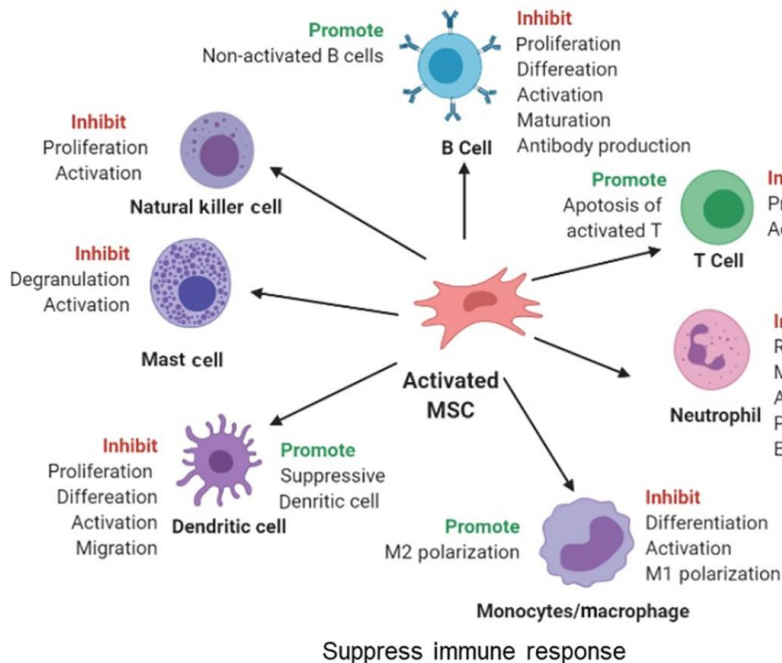
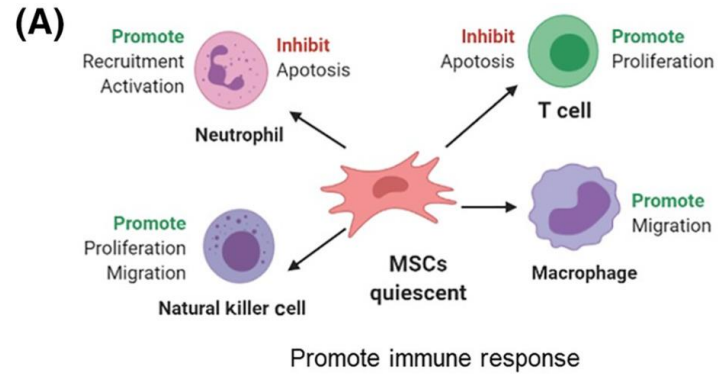
## Significant value upside

**Multiple pathways to commercialisation**, including strategic partnering

Well placed to fund to major catalysts with **~A\$27m<sup>1</sup> in cash**

# Why MSCs?

MSCs play a central co-ordinating role in many of the body's mechanisms of defence, repair and regeneration: the "sensor and switcher of the immune system"<sup>1</sup>



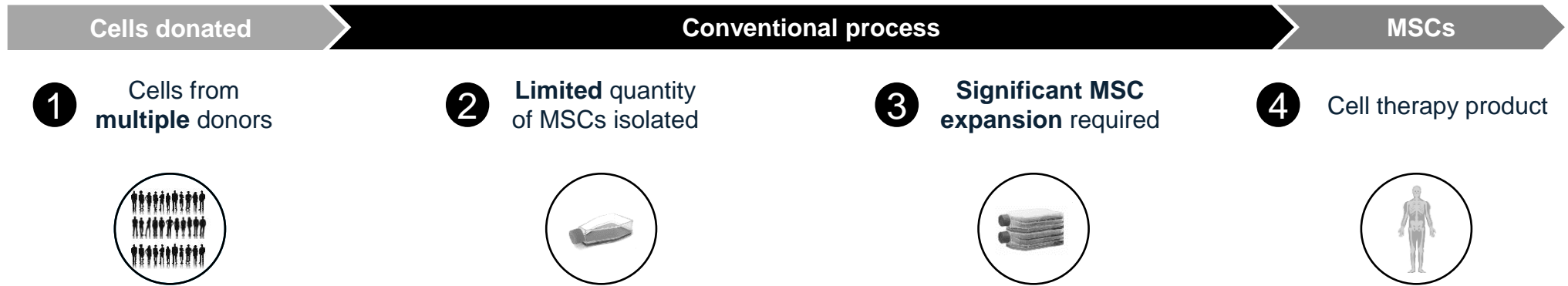
Global interest in MSCs continues to grow

**>1,000<sup>2</sup>**  
clinical trials  
of MSCs have  
been initiated  
in the past  
decade

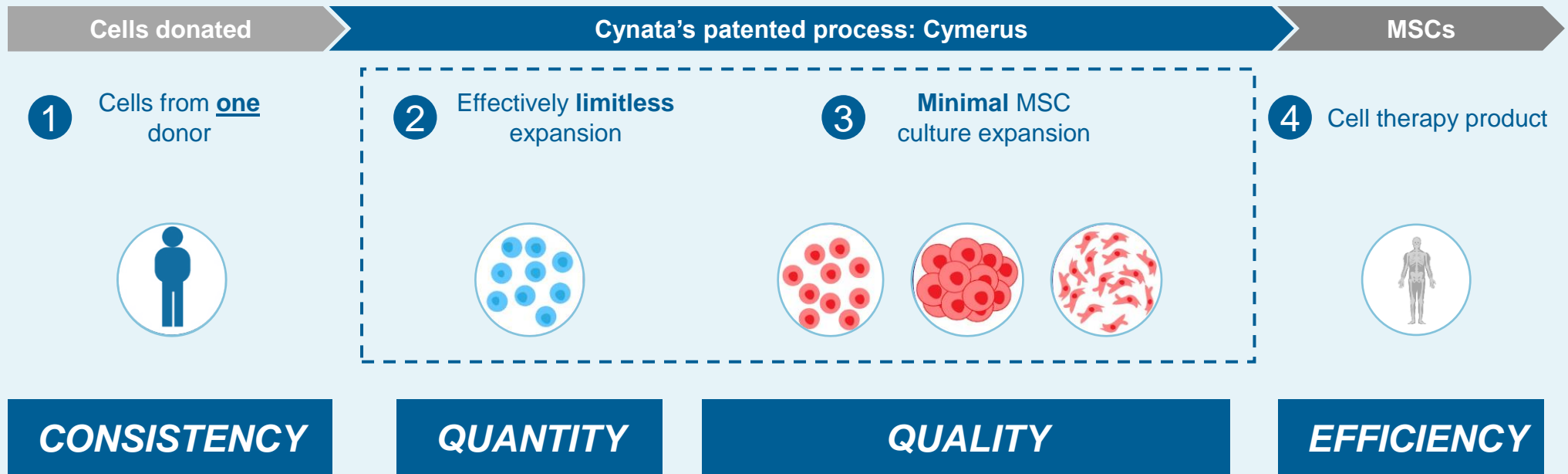
Source: Aldrich E.D. et al.; Stem Cells Transl Med. 2021 Nov;10(11):1500-1515

# Conventional vs. Cynata's Cymerus MSC manufacturing process

*Conventional Processes*



*Cynata's Process*



# Cymerus™ Platform Summary

Cynata is focused on the development of MSC-based therapeutic products using its unique proprietary Cymerus manufacturing technology

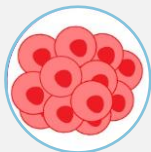
## Cymerus Technology

- A process for generating a range of MSC-based cell therapy products derived from **iPSCs**<sup>1</sup>
- **Patented** under i.p. held by Cynata and by WARF<sup>2</sup>
- MSC products are at the forefront of **next generation cell therapy treatments** for devastating diseases

Patented process: ability to proliferate indefinitely



iPSCs proliferate indefinitely



Create intermediate MCAs



Differentiation into MSCs (end product)

## Key Advantages

*Cynata can manufacture all the MSCs it will ever need from:*



- **Single Blood Donation**
- **Single Donor**

- ✓ Product versatility
- ✓ Manufacturing scalability
- ✓ Cost efficient
- ✓ Product consistency

# Cynata's technology addresses FDA concerns

Cynata's Cymerus process actively addresses current inefficiencies of MSC manufacturing, de-risking clinical development in the US

## Traditional MSC manufacturing is sub-optimal

Other MSC therapy (also targeting GvHD) was not approved by the FDA due to substantial functional variability between lots.

*“Substantial **functional heterogeneity** has been observed between MSC batches **derived from different donors** and **expanded using different tissue culture conditions or duration**, even though all of these batches meet the ISCT criteria for MSCs.”*

- Excerpt from **FDA ODAC Briefing** document for 13 August 2020

## Cynata's technology is optimal



Consistency: No inter-donor variability as only one donor is required (single blood donation)



Scalability: Cynata can produce essentially limitless quantities of MSCs from initial donation



Potency: iPSC-derived manufacturing process does not require excessive culture expansion of MSCs



FDA advisory meeting observations to be leveraged to maximise chance of FDA approval

# Cynata has an advanced and diverse product pipeline

Compelling safety results from Phase 1 GvHD trials<sup>1</sup> and positive preclinical data in each target indication have accelerated the pipeline





# SPA with Fujifilm provides commercial benefits

Cynata executed a Strategic Partnership Agreement with Fujifilm, with Fujifilm remaining very supportive of Cynata and involved in the path to market<sup>1</sup>

## Key SPA terms

- US\$5m fee paid by Fujifilm to Cynata
- Cynata regained all development and commercialisation rights to CYP-001
- First right to provide manufacturing services and product supply
- Fujifilm agreed to further voluntary escrow over their 8.1 million shares in Cynata
- Cynata and Fujifilm Cellular Dynamics, Inc now working towards establishing Cymerus manufacturing process at FCDI

## Strategic benefits for Cynata

- ✓ **Accelerate US development strategy:** With rights to CYP-001 in GvHD regained, Cynata plans on conducting the Phase 2 GvHD trial in the US
- ✓ **Credible company:** Fujifilm is one of the largest conglomerates in the world with a significant network in the biotechnology space
- ✓ **Experienced MSC manufacturer:** Fujifilm Cellular Dynamics Inc (subsidiary of Fujifilm) developed the original iPSC line used in Cynata's Cymerus manufacturing process



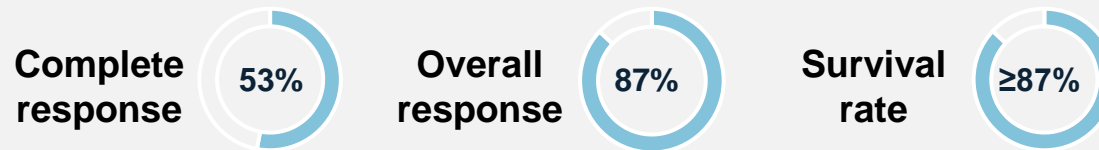
# GvHD | Ground-breaking Phase 1 clinical trial results

Cynata's Phase 1 GvHD trial met all safety and efficacy endpoints and broke ground by being the world's first clinical trial of an allogeneic iPSC-derived product

## Key results<sup>1</sup> demonstrate safety and efficacy of Cymerus MSCs

## Published in prestigious journal<sup>2</sup>

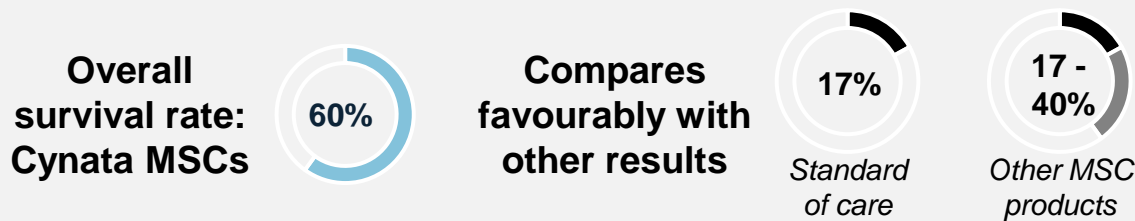
All endpoints achieved  
(Day 100)



Efficacy endpoints were the **same required in a Phase 3 trial**

Response rates were **higher than what we expect would be required in Phase 3** (to support marketing approval)

Outstanding follow-up results  
(Two year)



*Nature medicine is the preeminent peer-reviewed medical journal worldwide*

Current Issue | November 2020



# Accelerate US development strategy for GvHD

Cynata is aiming to progress a Phase 2 GvHD clinical trial in the US, after regaining clinical and commercial rights to CYP-001 in GvHD as part of the SPA with Fujifilm



**Orphan Drug Designation awarded by FDA for CYP-001**



**Phase 1 trial results exhibit strong safety and efficacy**



**SPA with Fujifilm, Cynata regains rights to CYP-001**



**Phase 2 trial design confirmed**



**Engage with the FDA for a Phase 2 trial in the US**








**Commence Phase 2 trial in the US**

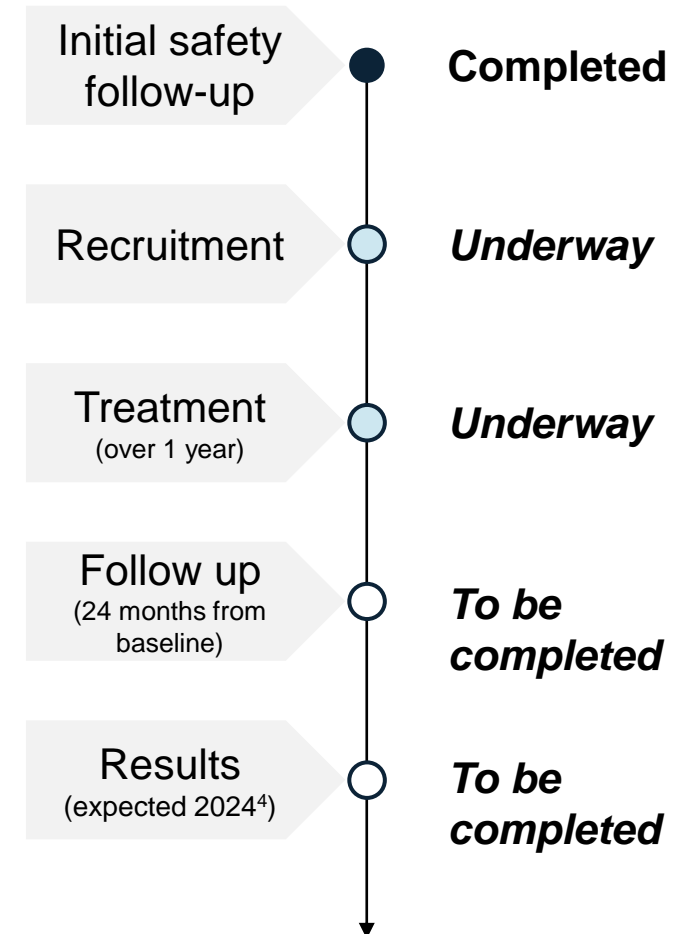


# SCUIpTOR<sup>1</sup> | Osteoarthritis Phase 3 clinical trial

Clinical trial underway, sponsored by the University of Sydney and funded by an NHMRC project grant






 <b>Osteoarthritis</b>	<ul style="list-style-type: none"><li>• <b>Osteoarthritis (OA)</b> occurs when the cartilage in a joint wears away</li><li>• Causes pain, inflammation, swelling and difficulty with movement</li></ul>
 <b>Huge Market Opportunity</b>	<ul style="list-style-type: none"><li>• <b>There is currently no complete cure</b></li><li>• OA estimated to affect &gt;30m Americans</li><li>• Global market of ~<b>US\$11.6bn<sup>2</sup></b></li></ul>
 <b>Strong preclinical data</b>	<ul style="list-style-type: none"><li>• <b>Preclinical research</b> supports efficacy of MSCs</li><li>• Potential to <b>improve the underlying disease</b> as well as alleviating pain</li></ul>
 <b>Substantial external support</b>	<ul style="list-style-type: none"><li>• <b>Funded</b> by the Australian Government NHMRC<sup>3</sup> project grant</li><li>• <b>Led by Professor David Hunter</b>, who is the Florence and Cope Chair of Rheumatology and Professor of Medicine at the University of Sydney</li></ul>
 <b>Trial design</b>	<ul style="list-style-type: none"><li>▪ University of Sydney to <b>enrol 440 patients</b> to participate in the randomised, double-blind placebo-controlled trial</li></ul>

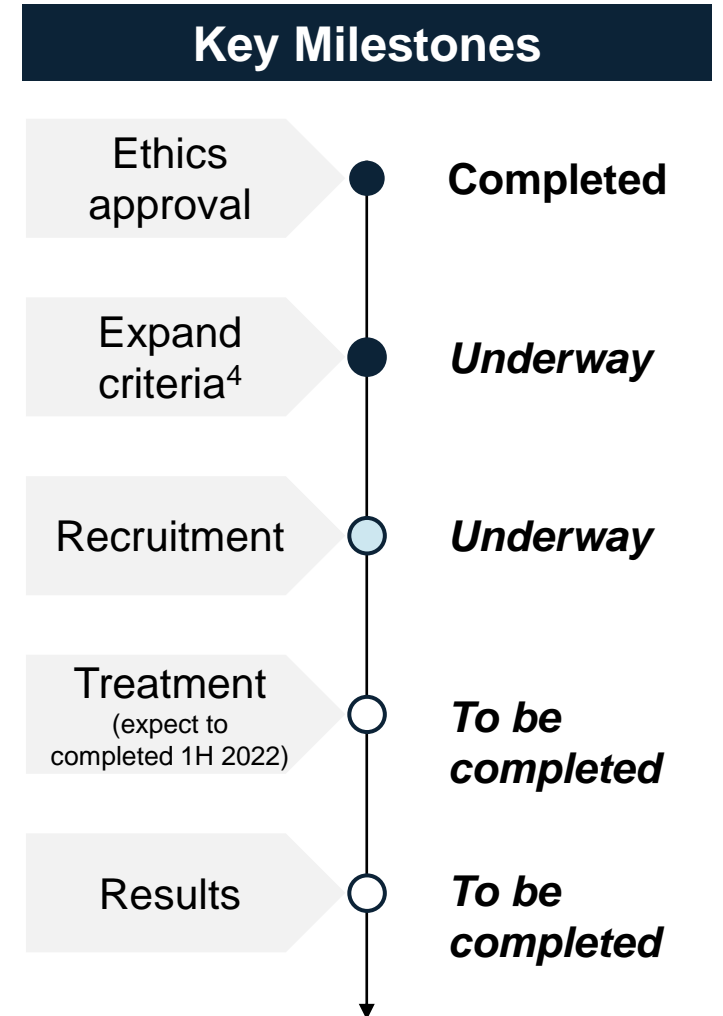
## Key Milestones



# MEND<sup>1</sup> | Clinical trial in respiratory distress

Patient recruitment underway, following expansion of patient population to increase pool of potential subjects

 <b>Respiratory distress</b>	<ul style="list-style-type: none"><li>• <b>Respiratory Failure</b> is a collection of signs and symptoms which collectively can be known as <b>Acute Respiratory Distress Syndrome (ARDS)</b>, a serious complication of e.g. COVID-19</li></ul>
 <b>Huge Market Opportunity</b>	<ul style="list-style-type: none"><li>• <b>Huge potential upside</b>, with combined market greater than <b>US\$8bn<sup>2</sup></b></li></ul>
 <b>Strong preclinical data</b>	<ul style="list-style-type: none"><li>• <b>Preclinical research</b> supports multiple beneficial effects of MSCs, including <b>reducing excessive inflammatory reactions</b></li></ul>
 <b>Substantial external support</b>	<ul style="list-style-type: none"><li>• <b>Collaboration</b> with CPA Research Institute<sup>3</sup> and COVID-19 Stem Cell Treatment Group</li></ul>
 <b>Trial design</b>	<ul style="list-style-type: none"><li>• Seeking to <b>recruit 24 adult patients</b> admitted to intensive care with <b>respiratory failure in Australia</b></li></ul>



1. MEseNchymal covid-19 Trial (MEND)






2. Source: Vasomune Therapeutics company announcement, 2018 (Reflects ARDS global market opportunity of US\$2.5bn); GlobeNewswire, 2020 (Represents Cytokine Release Syndrome (CRS) global market opportunity of US\$0.16m in 2017); GlobalData 2017 (Reflects Sepsis global market opportunity of US\$5.9bn in 2026).

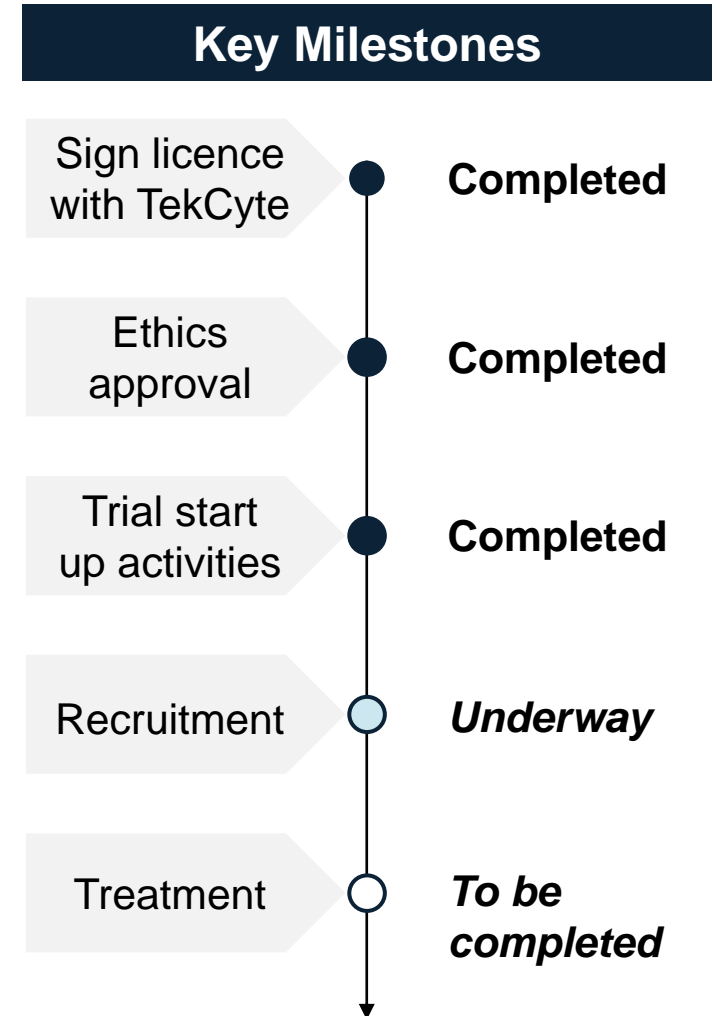
3. CPA = Cerebral Palsy Alliance

4. Ethics committee approval received to expand recruitment criteria (beyond COVID-19)

# DFU | Clinical trial in diabetic foot ulcers

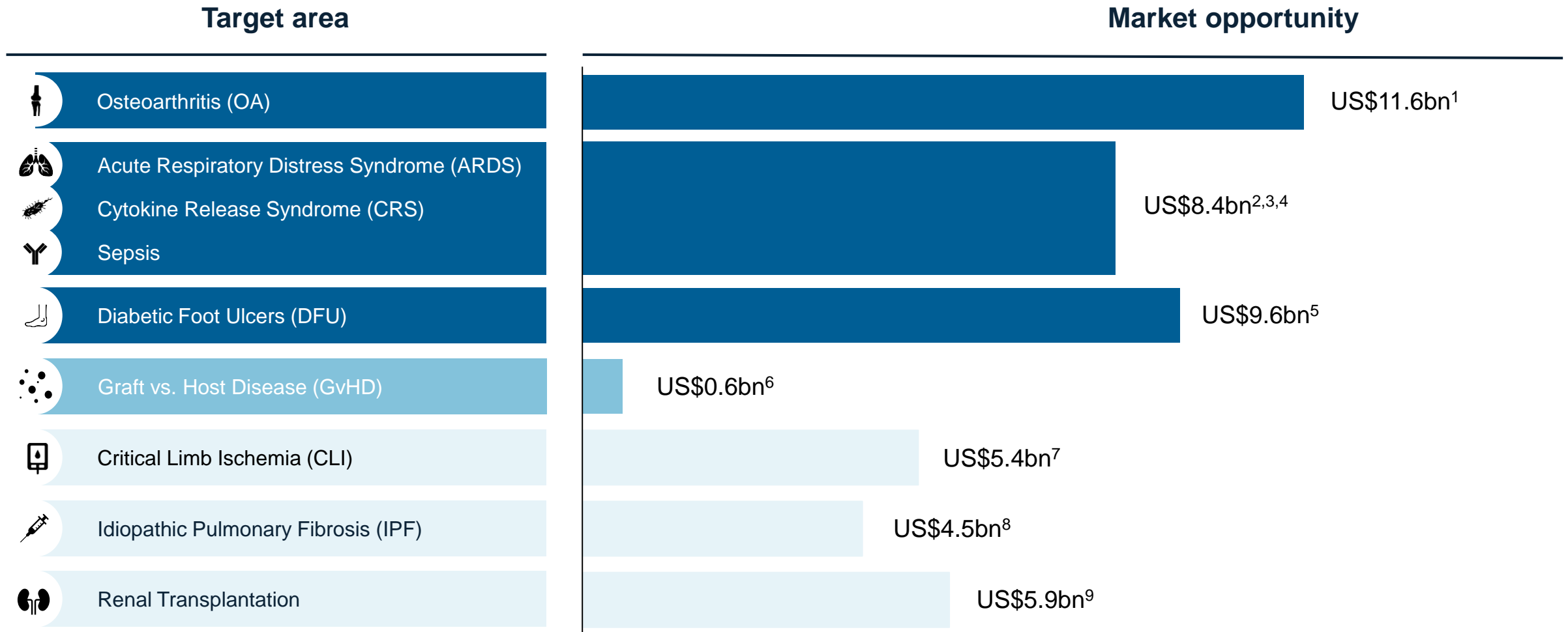
Recruitment has commenced and the trial is open for enrolment

 <b>Diabetic Foot Ulcers (DFU)</b>	<ul style="list-style-type: none"><li>DFU are sores on the feet of patients with diabetes (also known as diabetic wounds)</li></ul>
 <b>Huge Market Opportunity</b>	<ul style="list-style-type: none"><li><b>&gt;400m diabetics globally</b>, with DFU estimated to <b>occur in ~15-25% of patients</b> during their lifetime<sup>1</sup></li><li>Global market is estimated to be <b>~US\$10bn<sup>2</sup></b></li></ul>
 <b>Strong preclinical data</b>	<ul style="list-style-type: none"><li>Positive efficacy data of MSCs in a preclinical model</li><li><b>Cymerus MSCs achieved 86% skin restoration</b> after three days</li></ul>
 <b>Unique competitive positioning</b>	<ul style="list-style-type: none"><li><b>Secured a worldwide exclusive licence agreement with TekCyte</b></li><li>Enables use of polymer-coated dressings that deliver MSCs to DFUs</li></ul>
 <b>Trial design</b>	<ul style="list-style-type: none"><li><b>30 patients</b> with DFU will be randomly assigned to receive CYP-006TK (polymer-coated silicon dressing seeded with Cymerus MSCs) or standard care of treatment, over 4 weeks</li></ul>



# Significant market opportunities

Cynata is targeting attractive market opportunities across a range of target indications



# Near term catalysts

Cynata is in a strong position to advance development of its proprietary Cymerus platform technology

- ❑ **Complete recruitment** of 30 patients in DFU trial
- ❑ **Complete recruitment** of 24 patients in MEND (respiratory failure) trial
- ❑ **Advance US Regulatory strategy**, and...
- ❑ **Commence Phase 2** trial in GvHD
- ❑ **Finalise clinical trial** plans for IPF and renal transplantation
- ❑ **Complete recruitment** of 440 patients in Phase 3 osteoarthritis trial
- ❑ **Progress commercial discussions** and execute further corporate partnership(s)





# Board & management

Highly skilled and experienced senior leadership team with decades of experience in biotechnology



**Dr Geoff Brooke**  
Chairman

- **30+ years experience** in the healthcare investment industry
- Founder and MD of Medvest Inc and GBS Venture Partners



**Dr Ross Macdonald**  
Managing Director / CEO

- **30+ years experience and a track record of success** in pharmaceutical and biotechnology businesses
- Previously CEO of Hatchtech Pty Ltd



**Dr Kilian Kelly**  
Chief Operating Officer

- **15+ years experience** in biopharmaceutical research & development
- Previously Senior Director, Drug Development at Biota Pharmaceuticals, Vice President, Regulatory and Clinical at Mesoblast Limited



**Dr Jolanta Airey**  
Chief Medical Officer

- **25+ years experience** in respiratory, rheumatology, dermatology, biologicals, international markets and listed companies
- Previously Director, Translational Development at CSL Limited and a highly experienced clinician



**Peter Webse**  
Company Secretary

- **23+ years company secretarial experience**
- MD of Platinum Corporate Secretariat Pty Ltd, providing company secretarial and other services



**Dr Paul Wotton**  
Non-Exec Director

- **30+ years experience** in senior positions of life sciences companies
- Previously **President and CEO** of Ocata Therapeutics, Inc and Antares Pharma Inc.



**Dr Stewart Washer**  
Non-Exec Director

- **20+ years of CEO and Board experience** in medical technology, biotech and agri-food companies
- Current Chairman of Orthocell Ltd, and Chairman of Minomic International Ltd



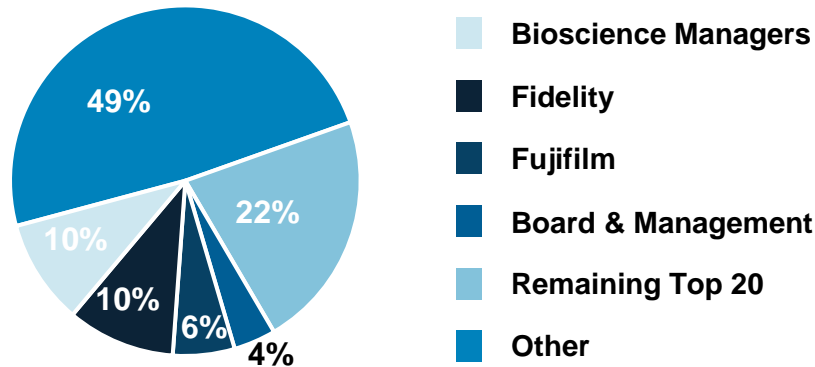
**Dr Darryl Maher**  
Non-Exec Director

- **Vice President** of R&D and Medical Affairs at **CSL Behring Australia**
- He was a **former President** of the Australian Pharmaceutical Physicians Association and a director of Vaccine Solutions

# Corporate overview

Cynata is proud to be supported by major institutional investors and the Company remains well funded and debt free

## Shareholder distribution



## Financial information

Share price (4 March 22)	A\$0.43
Shares on issue	143m
<b>Market capitalisation</b>	<b>~A\$61m</b>
Cash <sup>1</sup>	~A\$27m
<b>Enterprise Value</b>	<b>~A\$34m</b>

## Major institutional shareholders



10.0%

Fidelity International is a world leading investment and asset management firm that invests A\$556.7 billion globally on behalf of clients in Asia-Pacific, UK, Europe, the Middle East and South America.



9.9%

BioScience Managers is an international healthcare investment firm headquarter in Melbourne that finances and enables innovative science and technology with the potential to transform healthcare. They led the December 2020 placement with a \$10m investment into the Company.



5.7%

Fujifilm is a Japanese multinational conglomerate operating in the realms of photography, optics, medical electronics, biotechnology and chemicals. Fujifilm bought into ~8m shares as part of the development and commercialisation partnership agreement with Cynata in January 2017.

# Important information

## Summary information

This Presentation contains summary information about Cynata Therapeutics Limited and its subsidiaries (CYP) which is current at 7 March 2022. This Presentation should be read in conjunction with CYP's other periodic and continuous disclosure information lodged with the Australian Securities Exchange (ASX), which are available at [www.asx.com.au](http://www.asx.com.au).

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