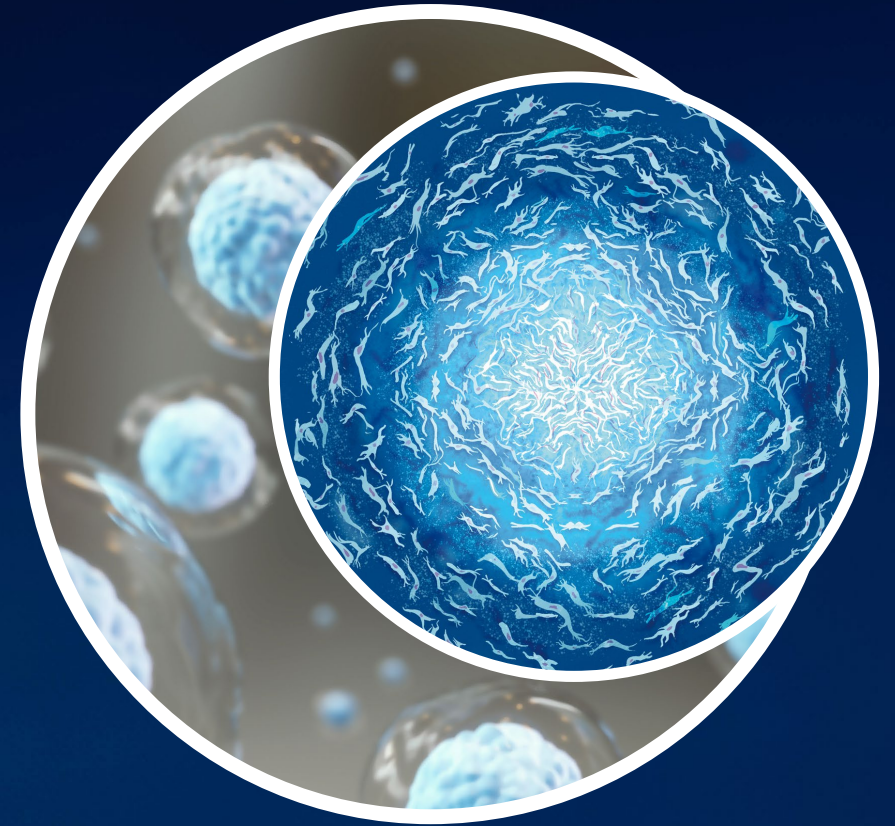




# A Next Generation Stem Cell Therapeutics Company



Investor Presentation  
April 2024

# Important information

## Summary information

This Presentation contains summary information about Cynata Therapeutics Limited and its subsidiaries (CYP) which is current as at 29 April 2024. 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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# Company highlights

## Revolutionary iPSC-based Cymerus™ manufacturing platform

- Effectively **limitless** number of high-quality MSC doses from a **single blood donation**
- Overcomes major obstacle to commercialisation in this highly promising field

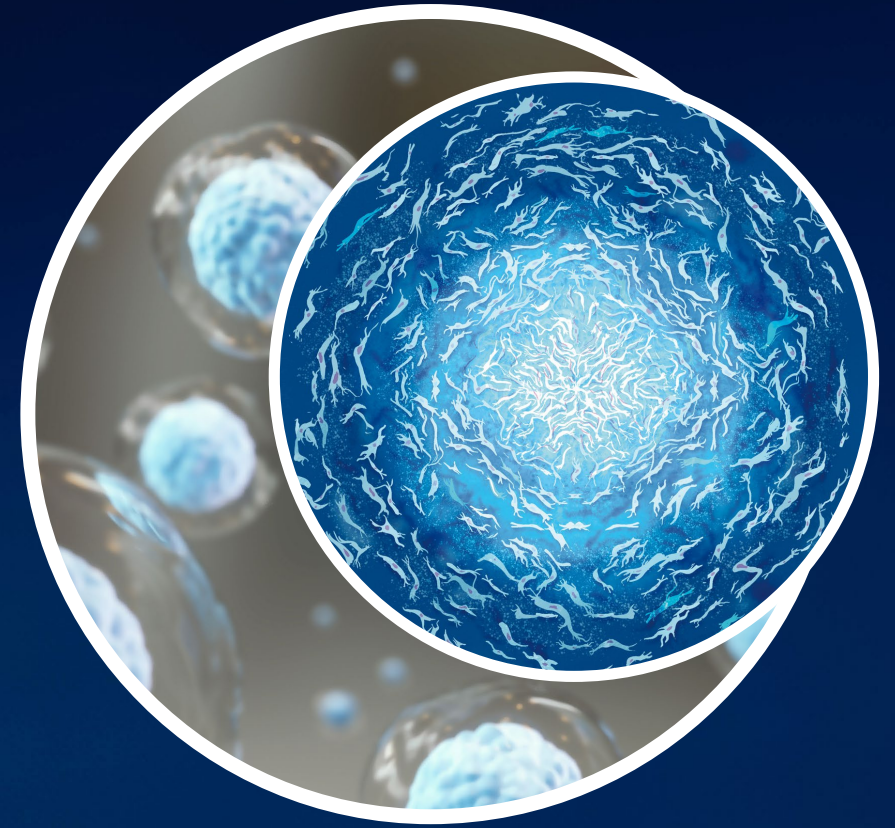
## Compelling clinical data

- **Acute graft versus host disease (aGvHD) Phase 1:** 53% complete response; 87% overall response
- **Diabetic foot ulcer (DFU) Phase 1:** 88% median wound surface area reduction vs 51% in controls<sup>1</sup>

## Rich clinical pipeline

- Three major randomised controlled clinical trial readouts upcoming:  
**DFU** (Phase 1) – early 2025; **aGvHD** (Phase 2) – 2H 2025; and **osteoarthritis** (Phase 3) – early 2026
- New trial in kidney transplantation to commence in Q2 2024

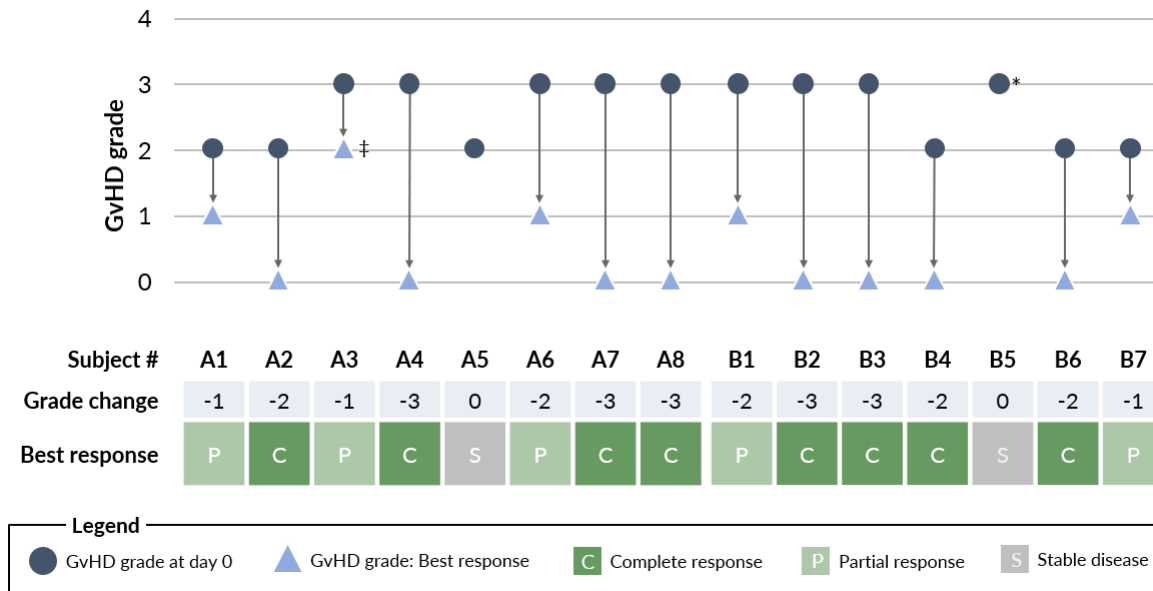
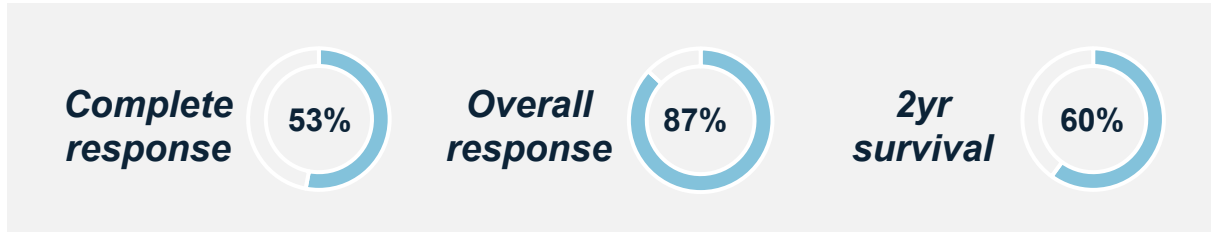
# Compelling Clinical Data



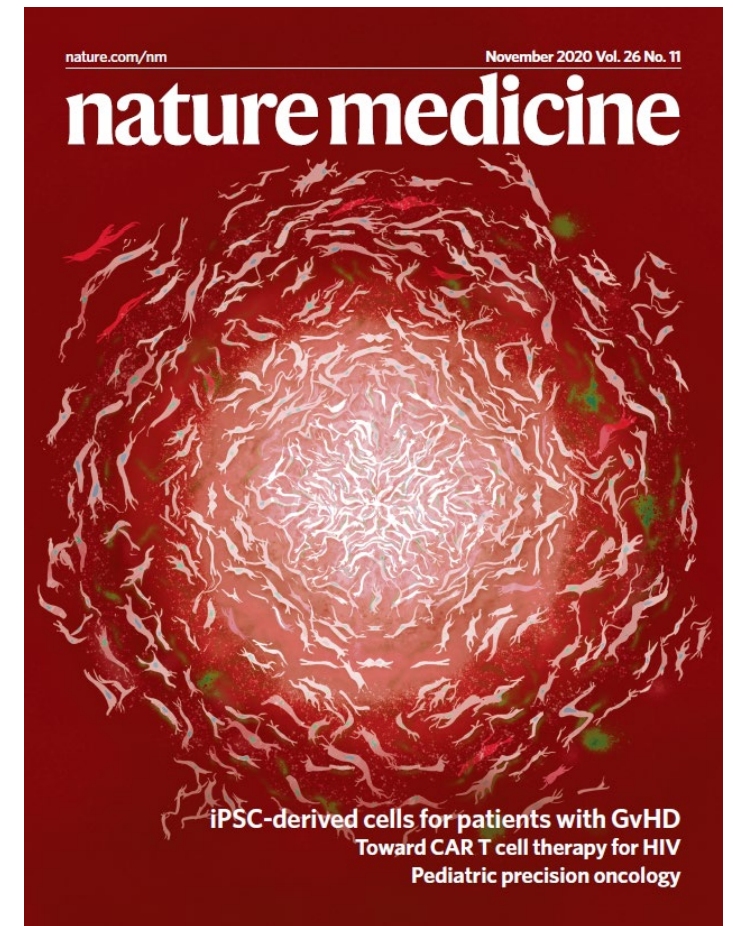
# aGvHD | Phase 1 clinical trial

Product: CYP-001 (Cymerus MSCs for intravenous infusion)

First completed clinical trial worldwide with any iPSC-derived product - published in **Nature Medicine**<sup>1</sup>



No treatment-related serious adverse events or safety concerns identified



- Subjects received  $1 \times 10^6$  cells/kg (max  $1 \times 10^8$  cells) or  $2 \times 10^6$  cells/kg (max  $2 \times 10^8$  cells) by IV infusion on D0 and D7  
 - Eight subjects were enrolled in each cohort, but one subject in Cohort B withdrew prior to infusion of CYP-001  
 ‡ Subject A3 showed a PR at Days 14 and 21 but died due to pneumonia on Day 28; \* Subject B5 withdrew from the trial on Day 22 to commence palliative care  
 1. Bloor et al. Production, safety and efficacy of iPSC-derived mesenchymal stromal cells in acute steroid-resistant graft versus host disease: a phase I, multicenter, open-label, dose-escalation study. Nat Med 2020;26:1720-1725.

# DFU | Phase 1 clinical trial – initial data

Product: CYP-006TK (topical Cymerus MSC wound dressing)

- Ongoing trial in non-healing diabetic foot ulcer (DFU)
- Patients randomised to receive standard of care (SoC) or CYP-006TK for 4 weeks, followed by SoC
- In the first 16 patients enrolled in the trial (8 per group), after 10 weeks' follow-up, the median reduction in wound surface area was:
  - **87.6%** in the active CYP-006TK group
  - compared to **51.1%** in SoC group

Example of ulcer healing in patient treated with CYP-006TK:

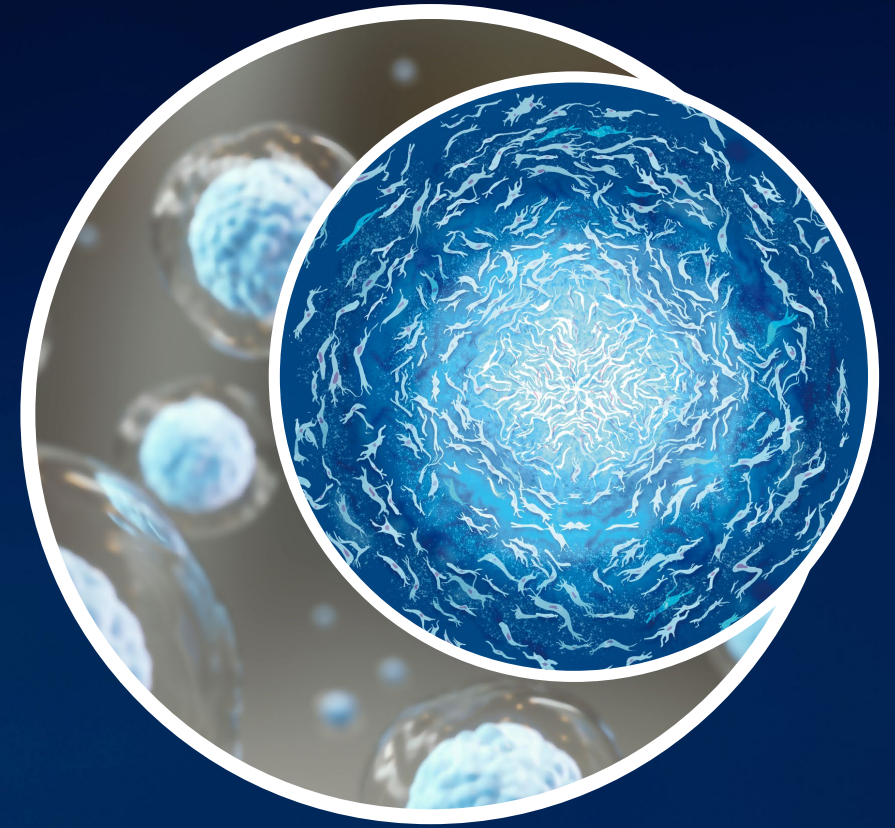
Day 0



Day 28



# Revolutionary iPSC-based Cymerus™ Manufacturing Platform



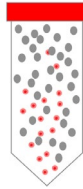
# Conventional MSC process

Ongoing need for new donors



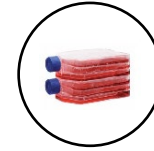
Substantial inter-donor **variability**

MSC isolation



**Small number** of MSCs per donation

Culture expansion



**Extensive** MSC culture expansion required

Major challenges:

- Logistically challenging
- Inter-donor **variability** – **inconsistent** activity in MSCs from different donors
- MSCs undergo **functional changes** during extensive culture expansion

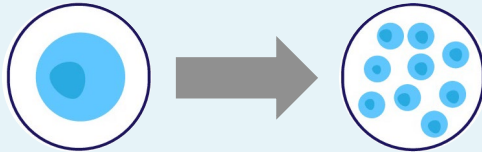
# Cymerus™ iPSC-based process

One donor, one time



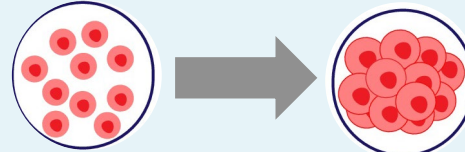
Avoids inter-donor variability

Reprogramming & iPSC expansion



Effectively limitless Expansion potential

Differentiation into MSCs & culture expansion



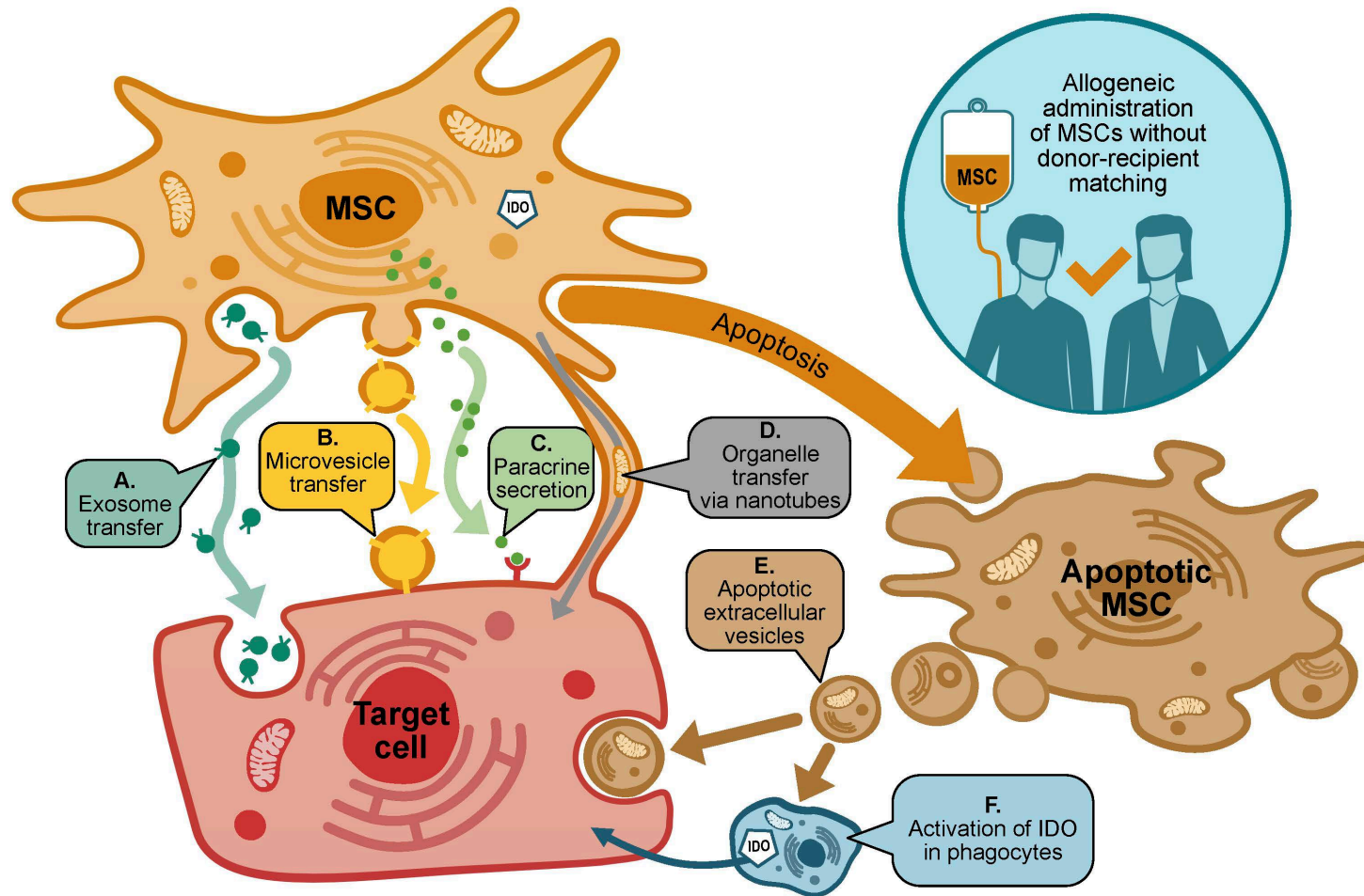
Minimal MSC culture expansion

Advantages of **Cymerus** platform:

- **Effectively limitless** iPSC expansion potential
- **Avoids** need for new donors
- **Avoids** inter-donor variability
- **Avoids** need for extensive MSC expansion
- High level of **consistency**



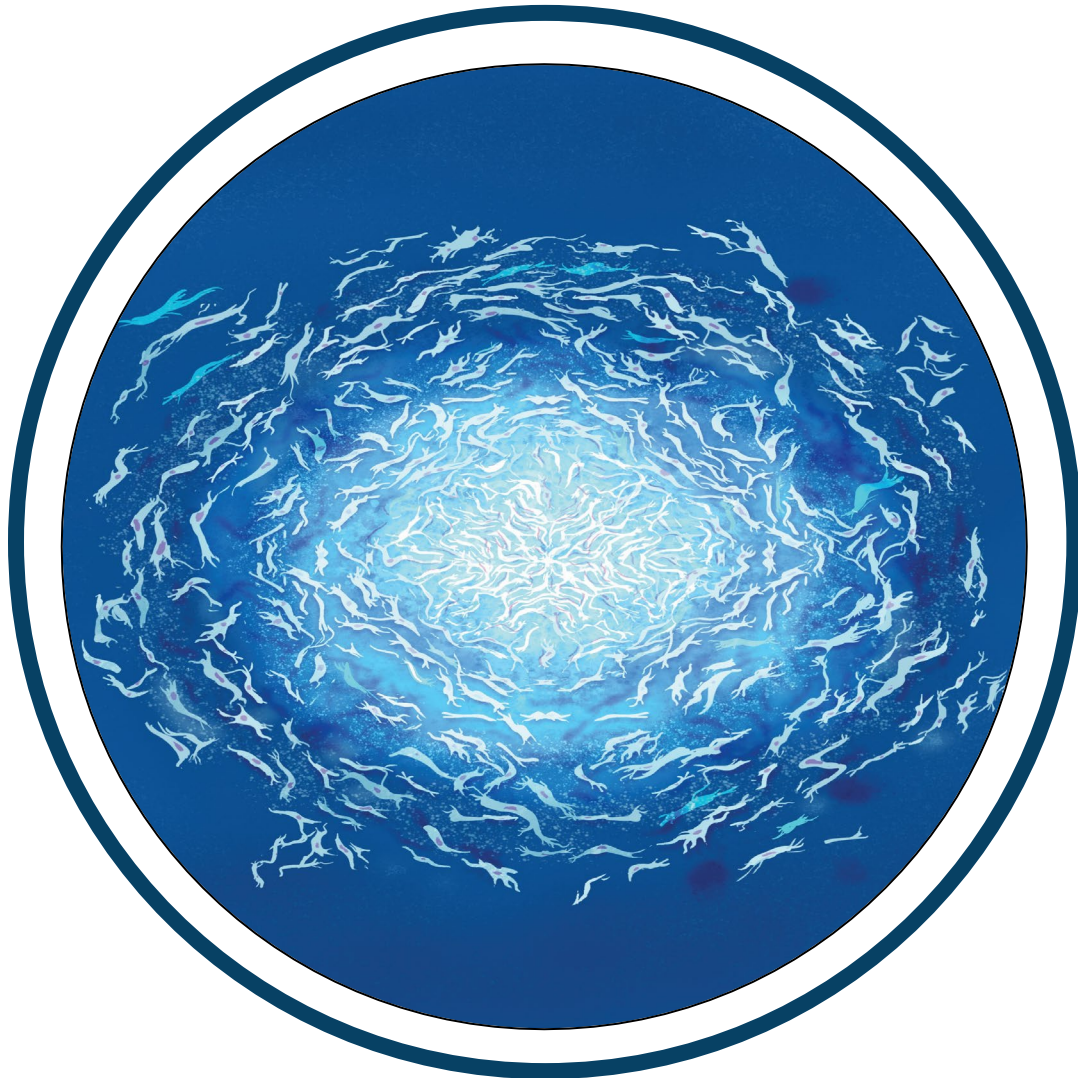
# Why MSCs?



## Mesenchymal stem (or stromal) cells (MSCs):

- promote an **immunomodulatory** environment via multifactorial mechanisms<sup>1</sup>
- the “sensor and switcher of the immune system”<sup>2</sup>
- promote **tissue repair and regeneration**
- can be used **without** donor/recipient matching
- can be **engineered** to express other functional/therapeutic molecules

# Why iPSCs?



## Induced pluripotent stem cells (iPSCs):

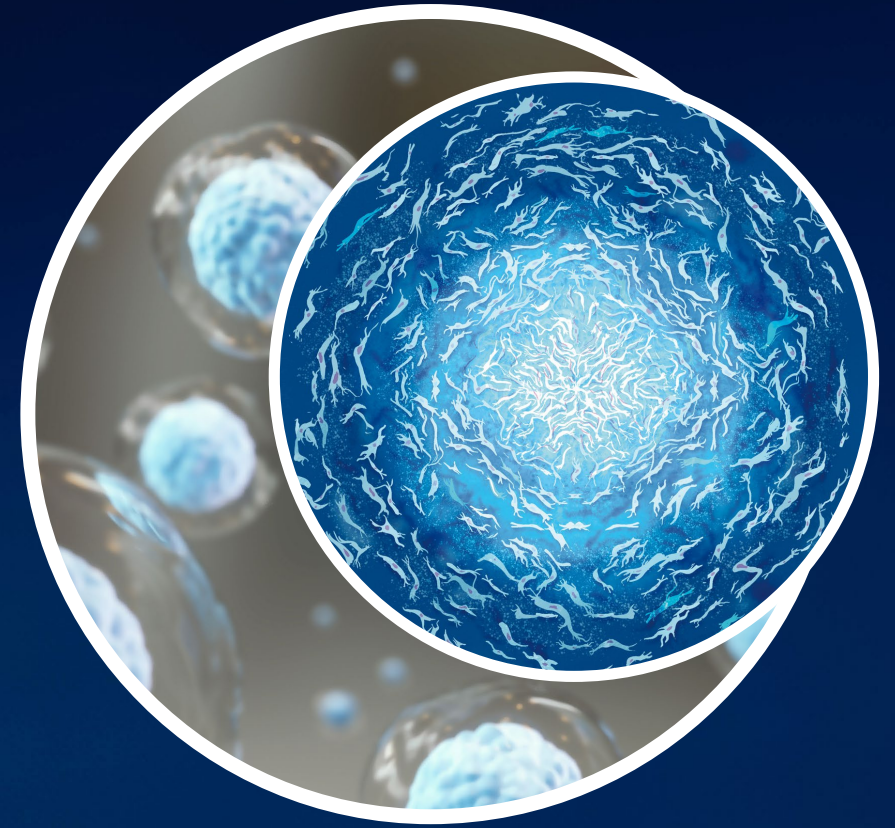
- mature cells from adult donors, reprogrammed to become pluripotent
  - effectively limitless proliferation in cell culture
  - potential to differentiate into any adult cell type (including MSCs)
  - avoids ethical controversy associated with embryonic stem cells
- **ideal** starting material for large scale production of cellular products

# Strategic partnership with Fujifilm





- Fujifilm: one of largest healthcare conglomerates globally, with significant assets in biotechnology sector, bolstered by recent multi-billion dollar investments
- Fujifilm Cellular Dynamics Inc (FCDI: subsidiary of Fujifilm) developed the original iPSC line used in Cynata's Cymerus manufacturing process
- Parties now working towards establishing Cymerus manufacturing process at FCDI with Cynata's progress showcasing Fujifilm's iPSC platform
- Fujifilm holds a 4.5% shareholding in Cynata



Rich Clinical Pipeline  
– Multiple Upcoming Data  
Readouts



# Advanced and diverse clinical pipeline

	Indication	Trial phase	Market opportunity
Cynata Sponsored	 <b>Acute Graft vs Host Disease (aGvHD)</b> CYP-001 <i>(FDA Orphan Designation)</i>	Phase 2 underway	US\$600m <sup>1</sup>
	 <b>Diabetic Foot Ulcers (DFU)</b> CYP-006TK	Phase 1 underway (recruitment complete)	US\$9.6bn <sup>2</sup>
Partnered	 <b>Osteoarthritis (OA)</b> CYP-004 <i>(managed by USYD, funded by NHMRC)</i>	Phase 3 underway (recruitment complete)	US\$11.6bn <sup>3</sup>
	 <b>Renal Transplantation (Renal)</b> CYP-001 <i>(managed and funded by LUMC)</i>	Phase 1 approved	US\$5.9bn <sup>4</sup>

1. Global Graft versus Host Disease Market 2019-2029 (Reflects forecast market in 2026); 2. Zion Market Research, 2019 (represents global treatment market in 2025); 3. Persistence Market Research 2018 research report: "Osteoarthritis Treatment Market: Global Industry Analysis (2012-2016) and Forecast (2017-2025) (Reflect OA market by 2025); 4. Organ Transplant Immunosuppressant Drugs Market in 2026, Grand View Research, Inc., 2019

# aGvHD | Phase 2 clinical trial

## Product

CYP-001 (Cymerus™ iPSC-derived MSCs for intravenous infusion)

## Indication

High risk acute graft versus host disease (aGvHD)<sup>1</sup>

## Study Design

- Randomised controlled trial in ~60 adults (steroids + CYP-001 vs steroids + placebo)
- Primary objective: to assess efficacy of CYP-001 based on Overall Response Rate at Day 28

## Study Conduct

- Clinical sites in USA, Europe and Australia
- Regulatory/ethics approvals secured in Australia, USA, Turkey and EU
- Numerous sites now open for recruitment, with remainder expected to open in Q2 2024
- First patient enrolled – March 2024
- Aiming to complete recruitment by end of calendar year 2024

## Results

Primary evaluation results expected in 2H CY 2025

# DFU | Phase 1 clinical trial

## Product

CYP-006TK (Novel silicone dressing seeded with Cymerus™ iPSC-derived MSCs)

## Indication

Non-healing diabetic foot ulcers (DFU)

## Study Design

- Randomised controlled trial in ~30 adults
- Patients randomised to receive either standard of care or CYP-006TK for 4 weeks, followed by standard of care
- Primary objective is safety; efficacy outcome measures include wound healing, pain & quality of life

## Study Conduct

- Clinical sites in Australia (Adelaide and Perth)
- Recruitment complete (April 2024)
- Last patient visit expected ~September 2024

## Results

- Positive initial results from first 16 patients – median reduction in wound surface area after 10 weeks was **87.6%** in CYP-006TK group compared to **51.1%** in controls (n=8 per group)
- Final results expected Q4 2024 or Q1 2025

# OA | Phase 3 clinical trial<sup>1</sup>

## Product

CYP-004 (Cymerus™ iPSC-derived MSCs for intra-articular injection)

## Indication

Osteoarthritis (OA) of the knee (Kellgren-Lawrence Grade 2-3)

## Study Design

- Randomised, double-blind placebo-controlled trial in ~320 adults
- Each participant receives 3 injections over 12 months; follow-up of 24 months from first dose
- Co-primary endpoints: reduction of knee symptoms and measure of cartilage loss

## Study Conduct

- Trial conducted by University of Sydney, funded by Australian Government NHMRC grant
- Clinical centres in Australia (Sydney and Hobart)
- Recruitment complete (November 2023)
- Last patient last visit expected ~November 2025

## Results

- Results expected in H1 CY 2026



# Renal transplant | Phase 1 clinical trial

## Product

CYP-001 (Cymerus™ iPSC-derived MSCs for intravenous infusion)

## Indication

Prevention of kidney transplant rejection

## Study Design

- ~16 patients to receive CYP-001 after kidney transplantation: cohort 1 (n=3); cohort 2 (n=3); cohort 3 (n=10)
- Trial will evaluate safety (all cohorts) and efficacy of MSCs in facilitating reduction of calcineurin inhibitors (anti-rejection medication; Cohort 3)

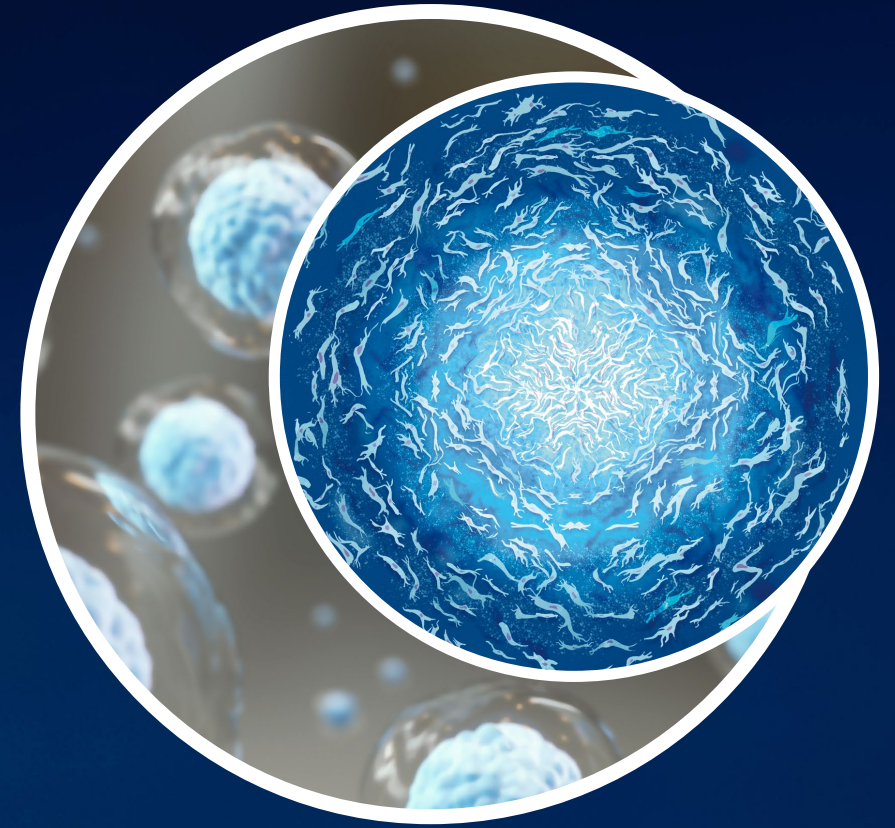
## Study Conduct

- Trial to be conducted and funded by Leiden University Medical Center (LUMC), Netherlands
- Regulatory and ethics approvals in place; final trial start-up activities ongoing
- Aiming to commence recruitment in Q2 2024

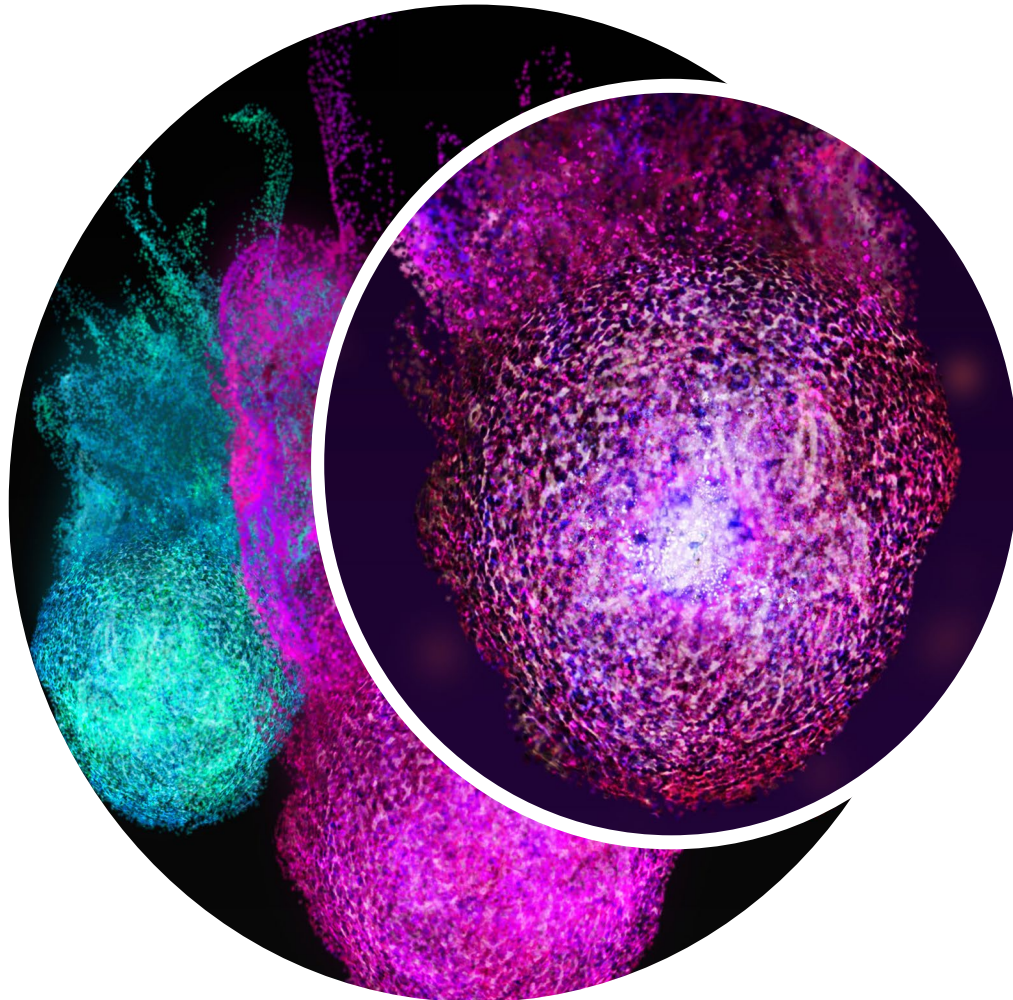
## Results

Results of Cohort 1 anticipated in late 2024

# Strategy, Outlook and Corporate Overview



# Partnering



Partnerships will accelerate our preclinical and clinical development programs



Reinvestment of proceeds to maximise potential of the platform



We will make Cymerus available to other regenerative medicine players – growth of the field in partnership is central to our mission

# Upcoming catalysts\*

Results of three randomised controlled clinical trials expected between early 2025 and early 2026

## 1H 2024

- Renal trial – start of enrolment

## 2H 2024

- Renal trial – results (Cohort A)
- aGvHD trial – completion of enrolment

## 1H 2025

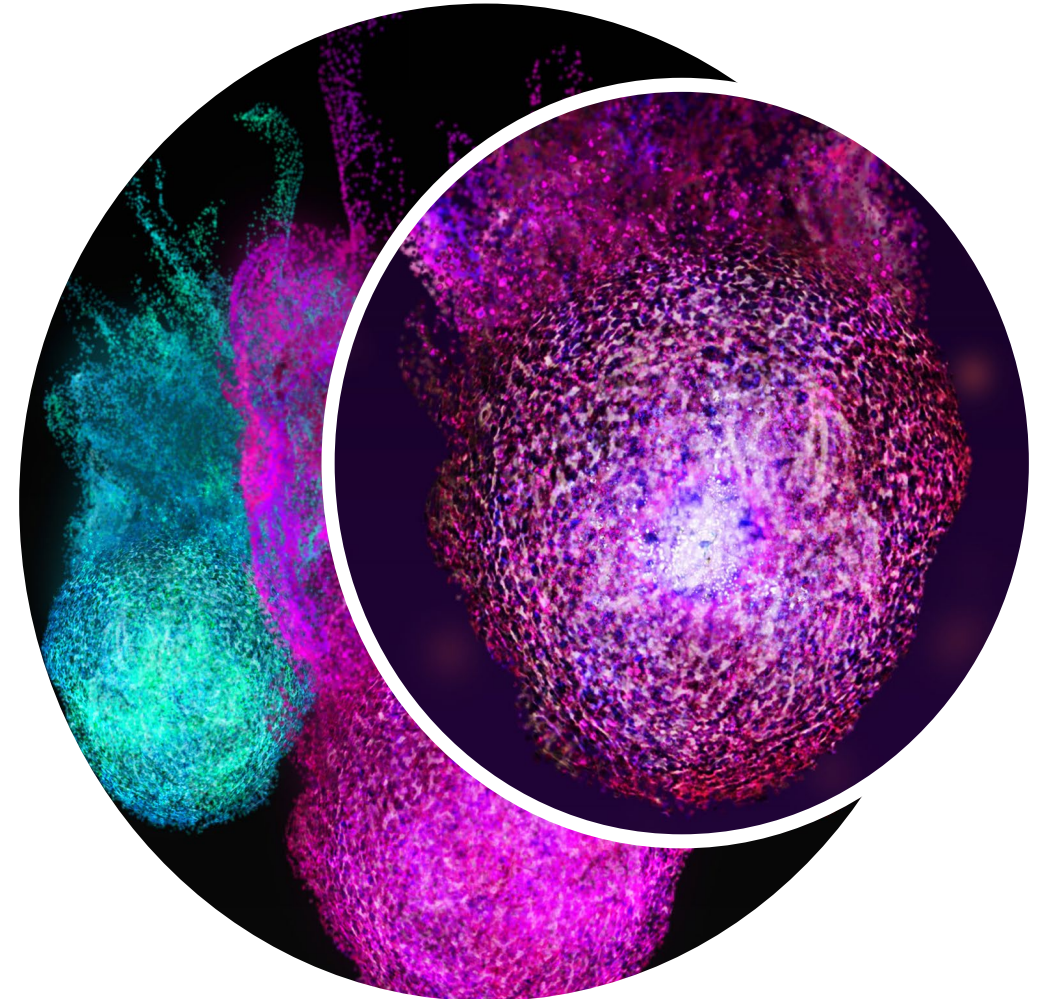
- DFU trial – results (potentially late 2024)

## 2H 2025

- aGvHD trial – results

## 1H 2026

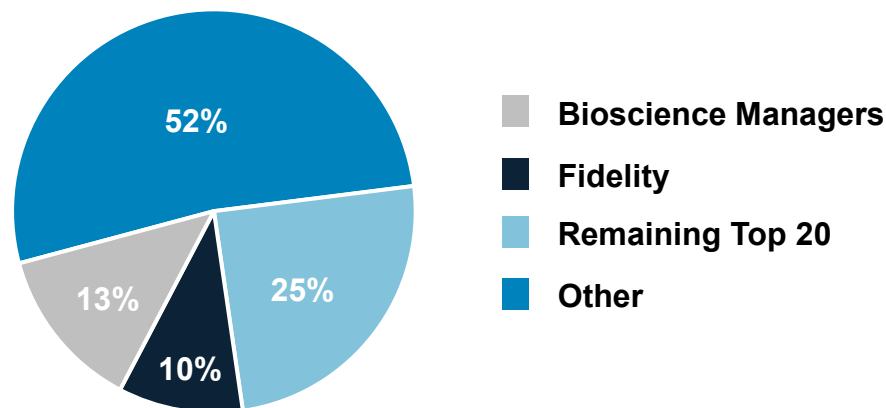
- OA trial - results



# Corporate overview

Cynata has been listed on the Australian Securities Exchange (ASX) since 2013 (Ticker: CYP)

## Shareholder distribution



## Financial information

Share price (29 April 2024) A\$0.21

Shares on issue 179m

**Market capitalisation ~A\$37.6m**

Cash<sup>1</sup> ~A\$9.0m

## Substantial shareholders (>5%)



13.1%

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.



10.0%

Fidelity International is a world leading investment and asset management firm, responsible for total client assets of >US\$750 billion, from clients across Asia Pacific, Europe, the Middle East, South America and Canada.

# Board & Senior Management

Highly skilled and experienced senior leadership team with decades of experience



**Dr Kilian Kelly**  
Chief Executive Officer &  
Managing Director

- 20+ years' experience in biopharma R&D
- Previous roles at Biota Pharmaceuticals, Mesoblast, Amgen & AstraZeneca



**Dr Geoff Brooke**  
Independent Non-Executive Chair

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



**Dr Paul Wotton**  
Independent Non-Executive Director

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



**Ms Janine Rolfe**  
Independent Non-Executive Director

- 20+ years legal, governance and management experience across multiple sectors
- Founder of Company Matters



**Dr Darryl Maher**  
Independent Non-Executive Director

- Former Vice President, R&D and Medical Affairs at CSL Behring
- Former President of Australian Pharmaceutical Physicians Association and Director of Vaccine Solutions



**Mr Peter Webse**  
Company Secretary

- 25+ years company secretarial experience
- Director of Governance Corporate Pty Ltd



**Dr Jolanta Airey**  
Chief Medical Officer

- 25+ years' experience in respiratory, rheumatology, dermatology, biologicals and listed companies
- Previously Director, Translational Development at CSL



**Dr Mathias Kroll**  
Chief Business Officer

- 25+ years' experience in biopharmaceutical industry
- Previously held leadership positions at various institutions, including Bayer, Sanofi-Aventis and GlaxoSmithKline



# Contact Us

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