ASX ANNOUNCEMENT **8 October 2024**

Cynata to Present at Alliance for Regenerative Medicine Meeting

Melbourne, Australia; 8 October 2024: Cynata Therapeutics Limited (ASX: **"CYP", "Cynata",** or the **"Company"**), a clinical-stage biotechnology company specialising in cell therapeutics, is participating in the *Cell and Gene Meeting on the Mesa* in Phoenix, Arizona.

This annual conference is run by the Alliance for Regenerative Medicine (ARM), which represents more than 400 members across 25 countries, including emerging and established biotechnology companies, academic and medical research institutions, and patient organisations.

The conference program features expert-led panels, extensive partnering capabilities, exclusive networking opportunities, and 100+ dedicated presentations by the leading publicly traded and privately held companies in the space. The conference attracts more than 2,000 attendees from over 150 companies.

At 10:45am local time today (8 October 2024), Dr Kilian Kelly, Cynata's Chief Executive Officer and Managing Director, will present on the Company's Cymerus™ iPSC[1]-derived MSC[2] technology and clinical development programs. Cynata will also be participating in partnering meetings throughout the conference.

A copy of the presentation is attached.

-ENDS-

Authorised for release by Dr Kilian Kelly, CEO & Managing Director

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Lauren Nowak, Media Contact, +61 (0)400 434 299, lauren@littlebigdeal.au

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. A Phase 2 clinical trial in GvHD under a cleared US FDA IND, as well as trials of Cymerus products in osteoarthritis (Phase 3 – patient enrolment completed) and diabetic foot ulcers (DFU – patient enrolment completed) are currently ongoing, while a trial in renal transplant is expected to commence in the near future. In addition, Cynata has also demonstrated utility of its Cymerus technology in preclinical models of numerous diseases, including 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.

[1] iPSC = induced pluripotent stem cell
[2] MSC = mesenchymal stem (or stromal) cell

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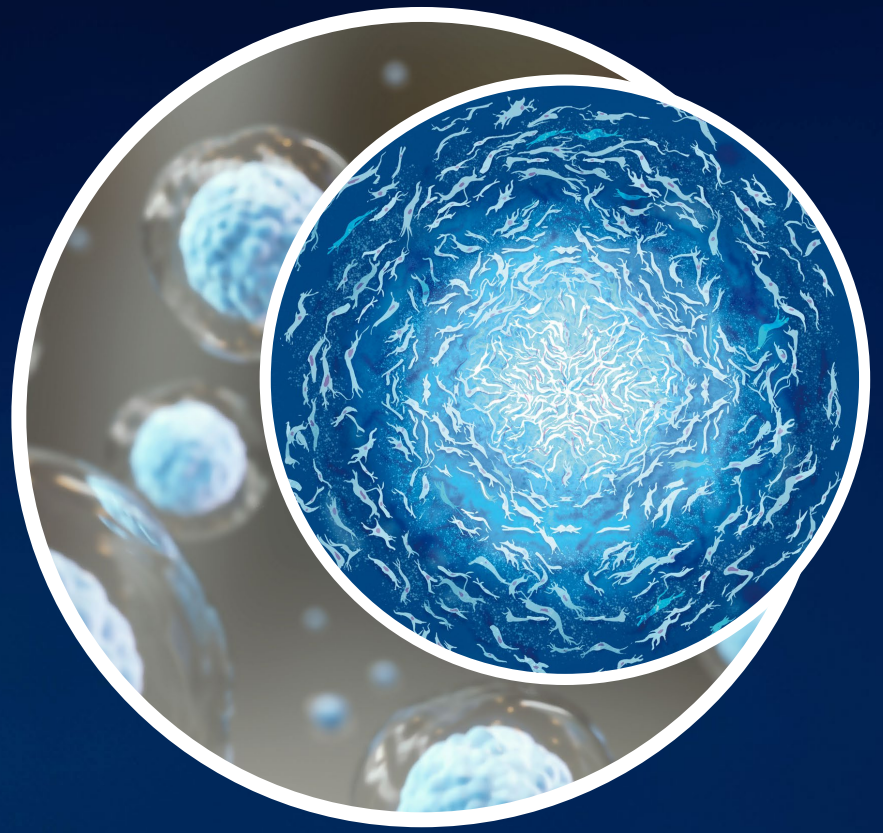
A Clinical Stage Next Generation Stem Cell Therapeutics Company



CELL GENE MEETING ON THE MESA

Phoenix, Arizona, 8 October 2024

Important information

Summary information

This Presentation contains summary information about Cynata Therapeutics Limited and its subsidiaries (**CYP**, or **Cynata**) which is current as at 4 October 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.

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Company highlights

Revolutionary Cymerus™ manufacturing platform

- Mesenchymal stem cells (**MSCs**)[1] have shown potential to treat a wide range of illnesses[2]
- However, standard manufacture requires ongoing supply of donors and extensive MSC culture expansion → challenges with consistency, potency and scale
- The induced pluripotent stem cell (**iPSC**)-based Cymerus™ platform overcomes these challenges by enabling production of an **effectively limitless** number of **consistent** MSC doses **from a single blood donation**

Cynata leads the burgeoning iPSC-derived therapy field

- **First completed iPSC clinical trial** worldwide
- US FDA **Orphan Drug Designation**[3] and cleared **IND**[4]
- Compelling clinical data in **acute graft versus host disease (aGvHD)**[5] and **diabetic foot ulcer (DFU)**[6]
- **Four active clinical programs** (including ongoing **Phase 2** and **Phase 3** trials)
- **Three randomised controlled clinical trial readouts** upcoming between **late 2024** and **early 2026**

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1. Also known as mesenchymal stromal cells
2. Zhou J, Shi Y. Cell Mol Immunol 20, 555–557 (2023)
3. CYP-001 granted Orphan Drug Designation for treatment of aGvHD – qualifies Cynata for incentives including extended marketing exclusivity, tax credits and fee waivers
4. IND = Investigational New Drug application – the clearance required from FDA to conduct clinical trials
5. Completed Phase 1 clinical trial in steroid-resistant aGvHD;
6. Initial data in first 16 patients (n=8 per group) after 10 weeks; final results in all 30 patients expected in Q4 2024/Q1 2025

Advanced and diverse clinical pipeline

Indication	Trial phase	Upcoming catalysts*	Market opportunity
Cynata Sponsored			
Acute Graft vs Host Disease (aGvHD) CYP-001 **FDA Orphan Designation**	**Phase 2 ongoing**	**Enrolment completion – Q4 2024** **Results – 2H 2025**	US$600m[1]
Diabetic Foot Ulcers (DFU) CYP-006TK	**Phase 1 ongoing** **(enrolment complete)**	**Results – Q4 2024/Q1 2025**	US$9.6bn[2]
Partnered			
Osteoarthritis (OA) CYP-004 *(managed by USYD, funded by NHMRC)*	**Phase 3 ongoing** **(enrolment complete)**	**Results – 1H 2026**	US$11.6bn[3]
Kidney Transplantation CYP-001 *(managed and funded by LUMC)*	**Phase 1/2 approved**	**Enrolment start – Q4 2024** **Cohort 1 results – Q1 2025**	US$5.9bn[4]

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

USYD = University of Sydney; NHMRC = National Health and Medical Research Council; LUMC = Leiden University Medical Center
* Timing of events is approximate, based on the Company's information as at the date of this presentation, and subject to change

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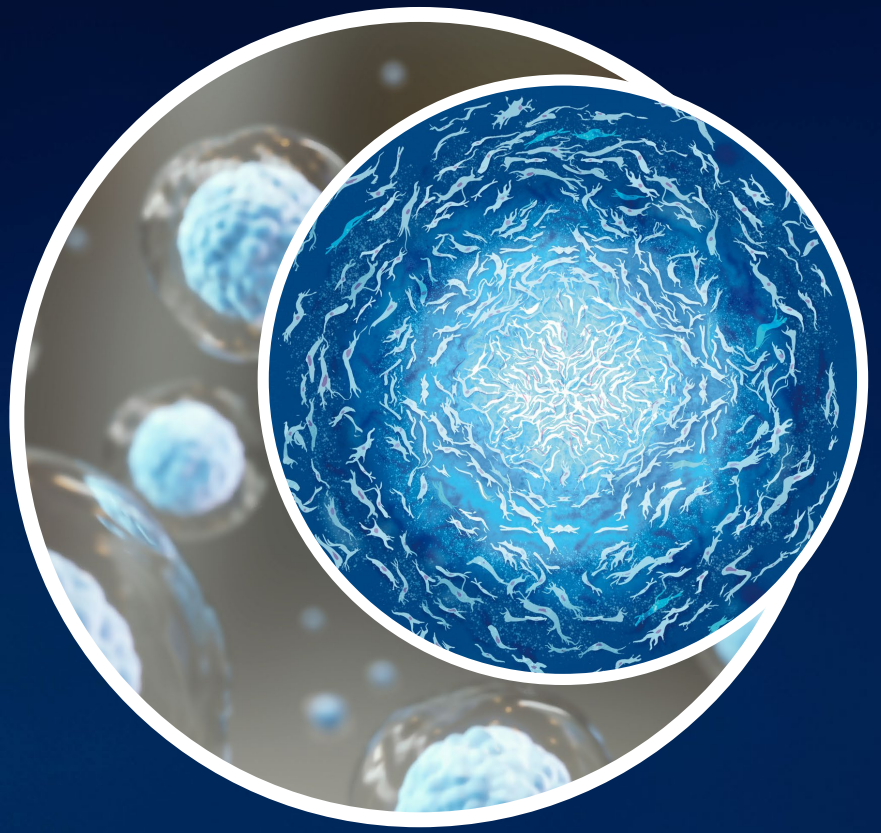
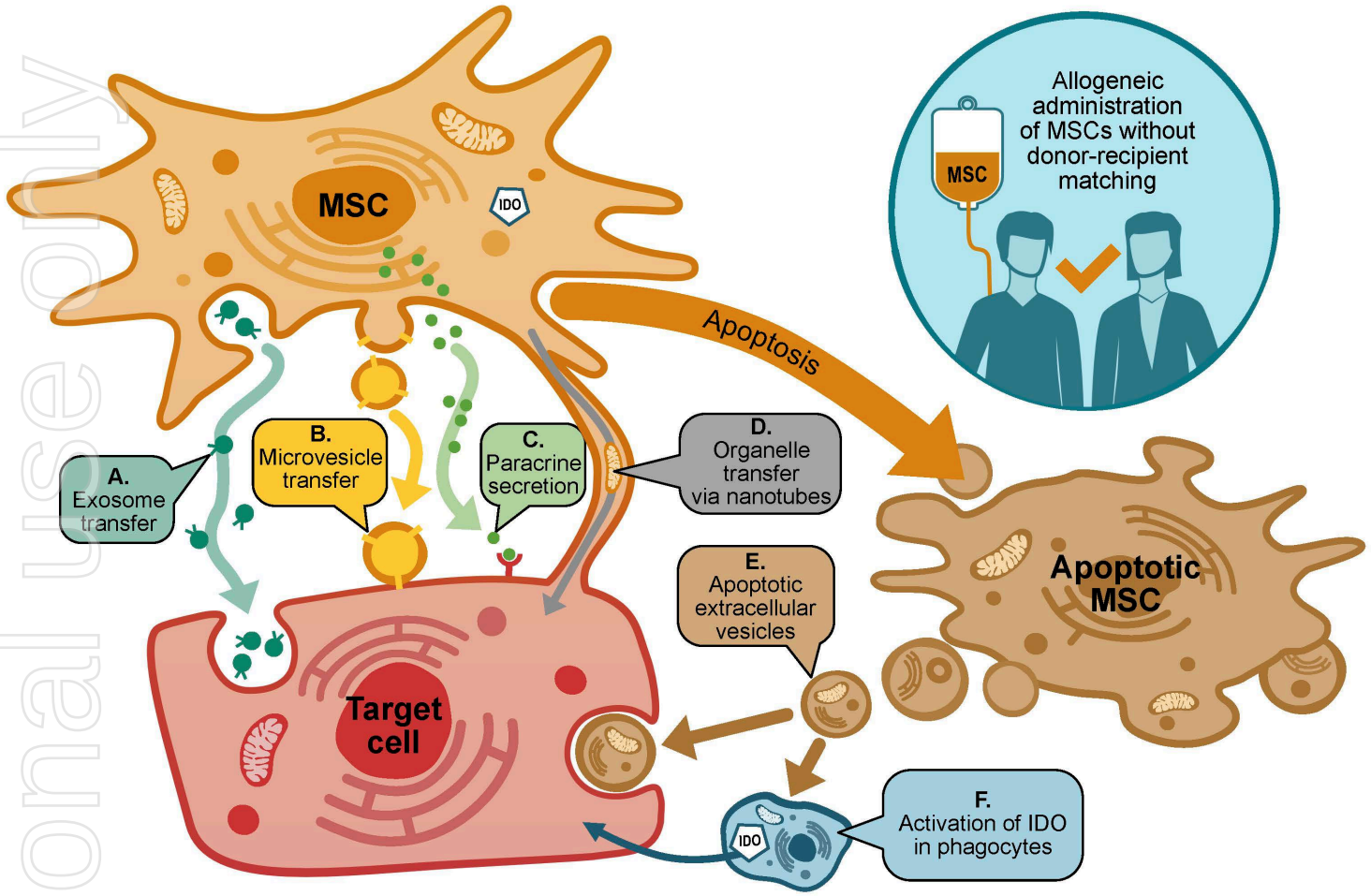
Revolutionary iPSC-based Cymerus™ Manufacturing Platform



Therapeutic potential of MSCs

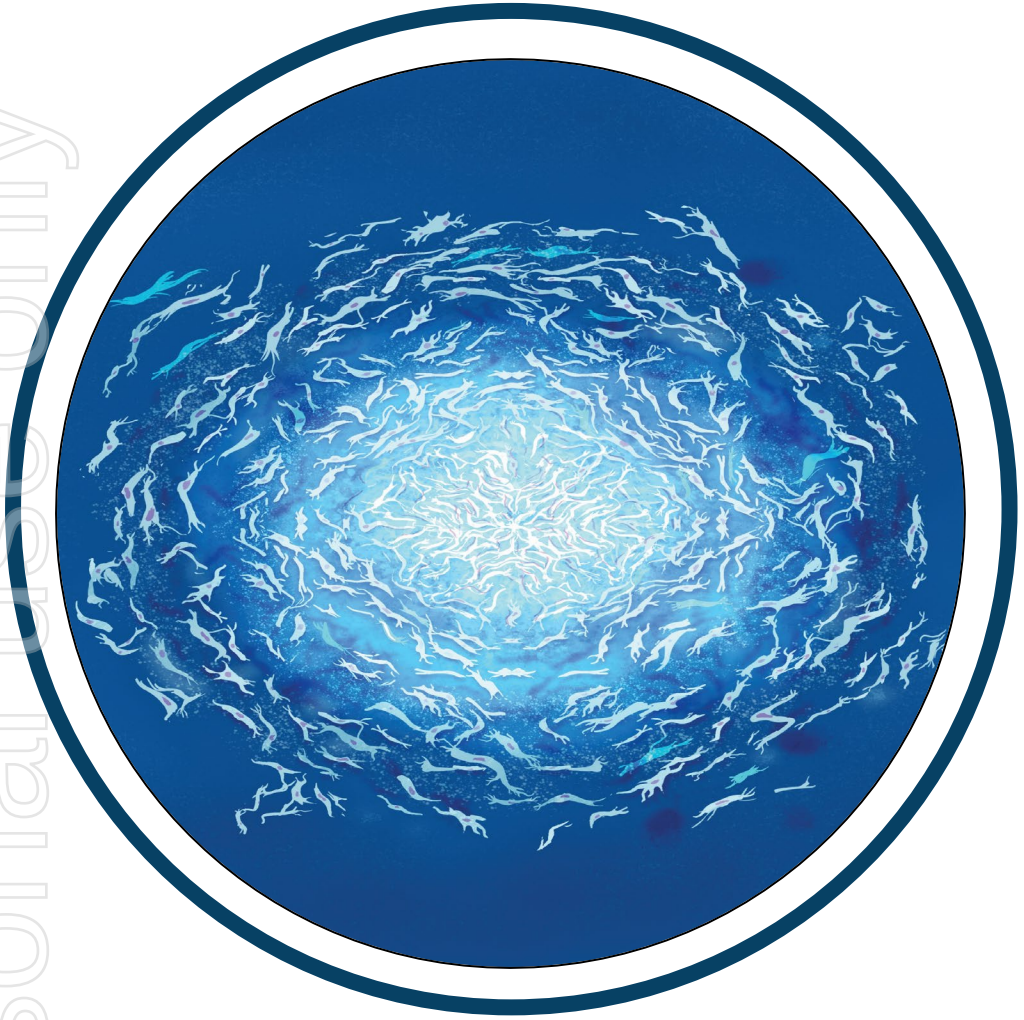


Mesenchymal stem cells[1] (MSCs):

- Promote an **immunomodulatory** environment[2]
- The "sensor and switcher of the immune system"[3]
- Promote **tissue repair** and **regeneration**
- Can be used **without** matching donors to recipients
- Can be **engineered** to express other functional/therapeutic molecules
- However, with conventional manufacturing methods, there are consistency, potency and scalability challenges

1. Also known as mesenchymal stromal cells
2. Kelly and Rasko, Front. Immunol. 12:761616 (2021)
3. Sarsenova et al, Front. Immunol.13:1010399 (2022)

Advantages of iPSC-based platform



Induced pluripotent stem cells (iPSCs):

- Mature **adult** cells **reprogrammed** to become **pluripotent**, which means:
  - Effectively **limitless** proliferation capacity
  - Potential to differentiate into any adult cell type (including MSCs)
- Similar properties to embryonic stem cells … but iPSCs are derived from **adult donors**, so they **avoid** ethical controversy associated with embryonic stem cells

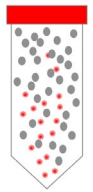
→ iPSCs are **ideal** starting material for commercial production of cellular products

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:

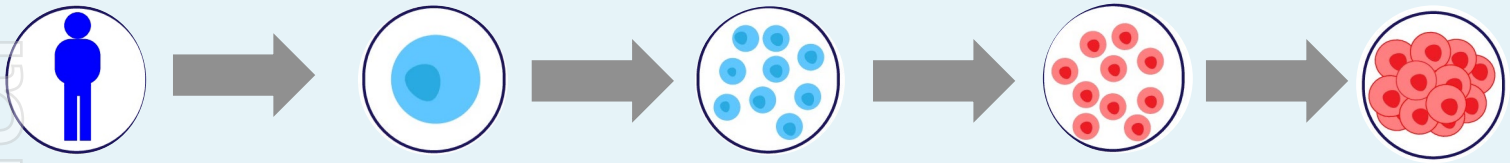
- MSCs undergo **functional changes** and **loss of potency** during extensive culture expansion
- Continuously finding and testing new donors is **logistically challenging**
- Inter-donor **variability** – **inconsistent** activity in MSCs from different donors

Cymerus™ iPSC-based process

One donor, one time

Reprogramming & iPSC expansion

Differentiation into MSCs & culture expansion



Avoids inter-donor variability

Effectively **limitless** expansion potential

Minimal MSC culture expansion

Robust patent protection

Advantages of **Cymerus™** platform:

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

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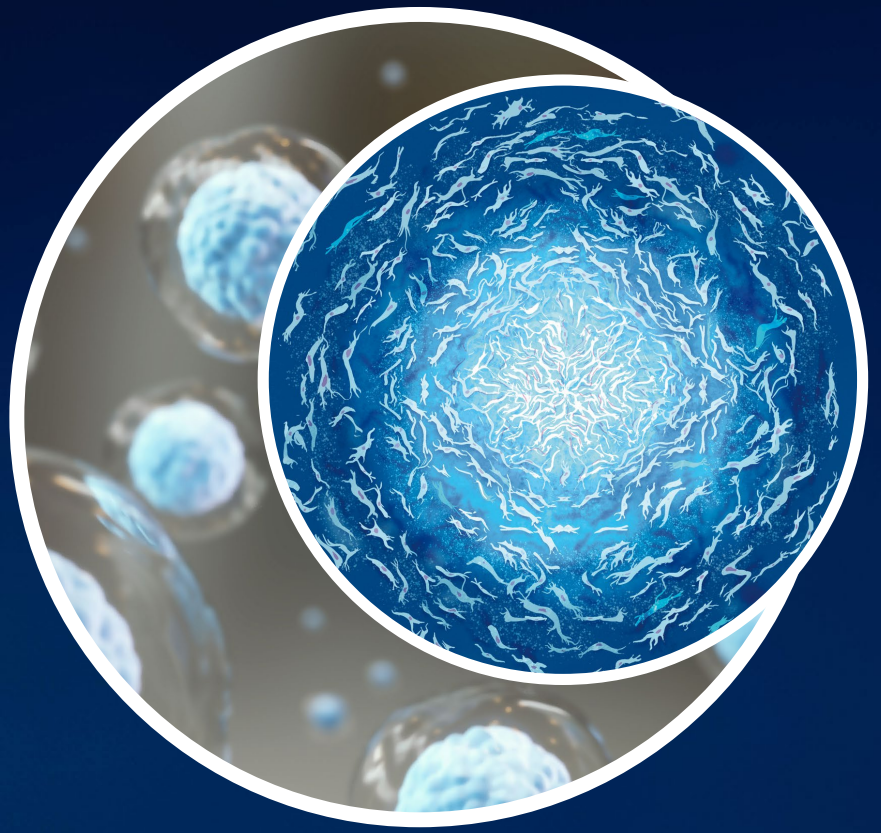
Compelling Clinical Data



CYP-001: Two *Nature Medicine* publications

- CYP-001 has been granted **Orphan Drug Designation** by the US FDA for the treatment of GvHD
- Phase 1 trial of CYP-001 was the first completed clinical trial worldwide with **any iPSC-derived product**

nature medicine
LETTERS
https://doi.org/10.1038/s41591-020-1050-x
Nature Medicine 26, 1720–1725 (2020)

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

Adrian J. C. Bloor[1,2], Amit Patel[1], James E. Griffin[3], Maria H. Gilleece[4], Rohini Radia[5], David T. Yeung[6,7], Diana Drier[8], Laurie S. Larson[8], Gene I. Uenishi[9], Derek Hei[10], Kilian Kelly[11], Igor Slukvin[9] and John E. J. Rasko[12,13,14]

nature medicine
Nature Medicine 30, 1556–1558 (2024)
https://doi.org/10.1038/s41591-024-02990-z

Two-year safety outcomes of iPS cell-derived mesenchymal stromal cells in acute steroid-resistant graft-versus-host disease

Kilian Kelly[1], Adrian J. C. Bloor[2], James E. Griffin[3], Rohini Radia[4], David T. Yeung[5,6] & John E. J. Rasko[7,8,9]

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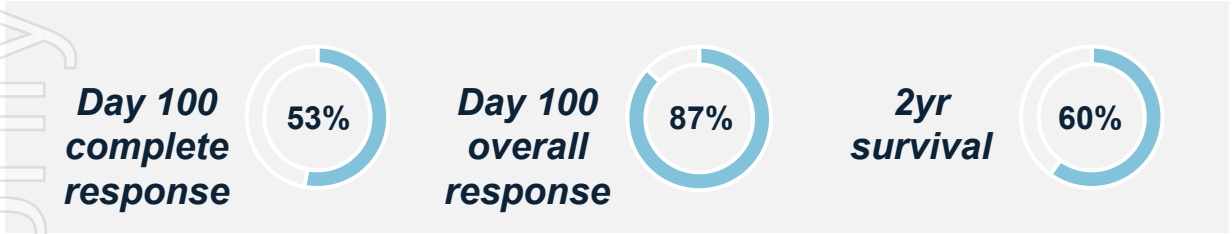
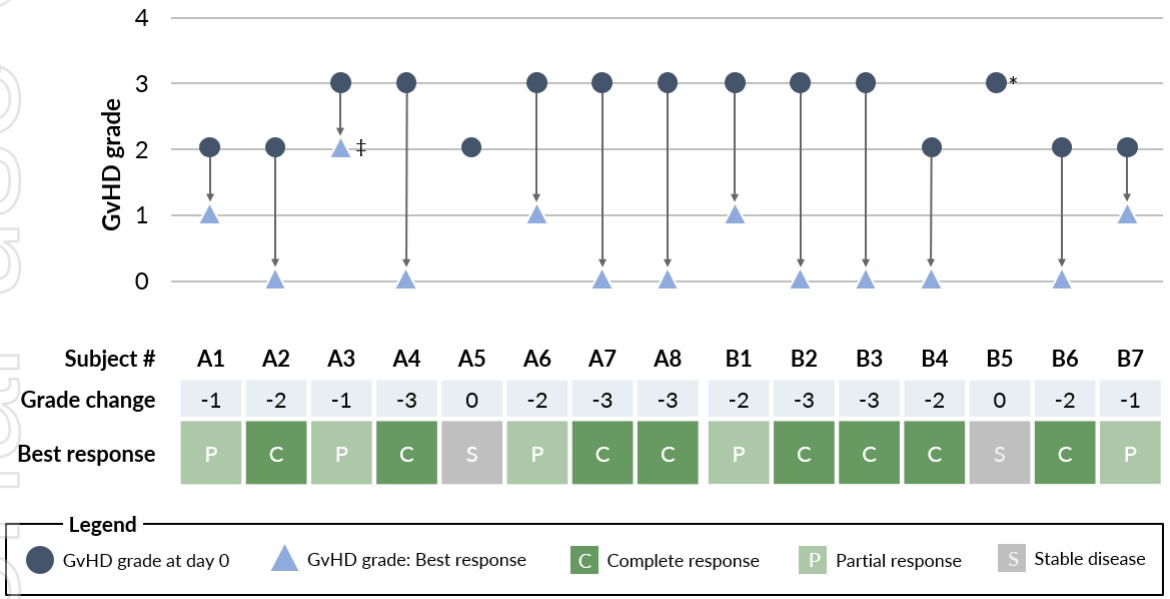
1. CYP-001 is Cynata's Cymerus iPSC-derived MSC product candidate for intravenous infusion

aGvHD | Phase 1 clinical trial - results

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

Trial conducted in 15 patients with **steroid-resistant aGvHD (SR-aGvHD)**





- CYP-001 was shown to be **safe and well tolerated**, with **sustained outcomes up to 2 years** after the first infusion
- **No serious adverse events or other safety concerns related to CYP-001**
- Very encouraging **response rates** and **overall survival**

- Subjects received $1x10^6$ cells/kg (max $1x10^8$ cells) or $2x10^6$ cells/kg (max $2x10^8$ 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

For further information: https://clinicaltrials.gov/study/NCT02923375

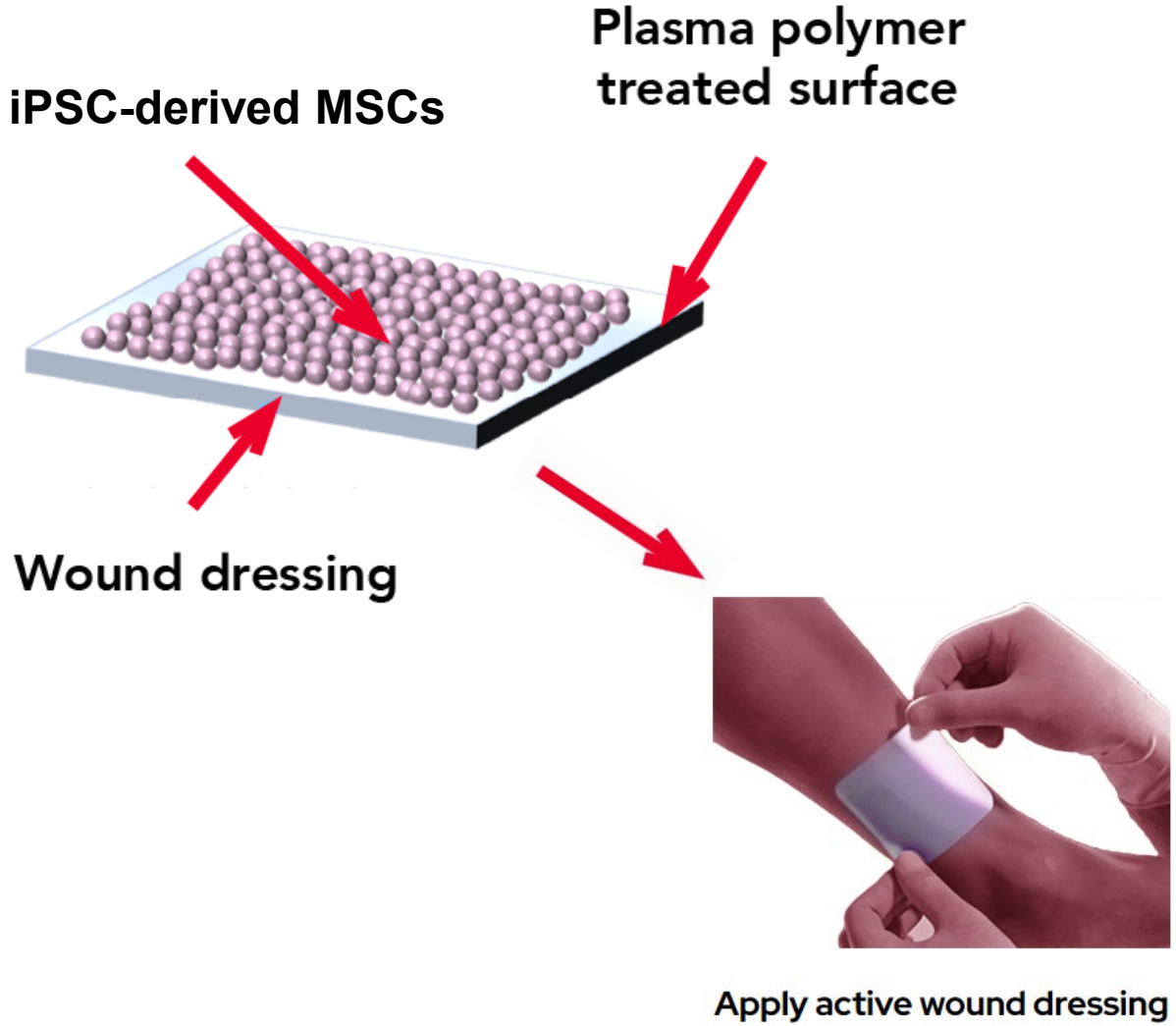
Ph1 SR-aGvHD results compared to other therapies

	CYP-001 (Ph 1)	Ruxolitinib (Ph 3)	"Best available therapy" controls (Ph 3)
Day 28 Overall Response	67%	62%	39%
Day 56-60* Overall Response	73%	40%	22%
Overall Survival	60% after 2 years	38% after 18 months	36% after 18 months
Safety	No safety concerns related to CYP-001 identified	Serious adverse reactions to ruxolitinib are common	Several other agents investigated for GvHD have poor safety profiles

Notes:
- Ruxolitinib is approved for treatment of SR-aGvHD in most jurisdictions
- Comparisons are for illustrative purposes only; data taken from different clinical trials with different sample sizes (BAT: n=155; Rux: n=154; CYP-001: n=15)
- D28/D56-60 time points used for response rate comparison as D28/D56 were the only response rate time points reported in the ruxolitinib/best available therapy clinical trial (NCT02913261); Overall Response at Day 56-60 refers to Day 56 response for ruxolitinib and best available therapy, and Day 60 response for CYP-001.

CYP-006TK – a novel topical MSC product

- CYP-006TK utilises a proprietary surface-coating, optimised for the delivery of MSCs directly to the wound bed
- Technology exclusively licenced to Cynata by Tekcyte Limited (agreement for Cynata to acquire this IP outright announced 1 July 2024)



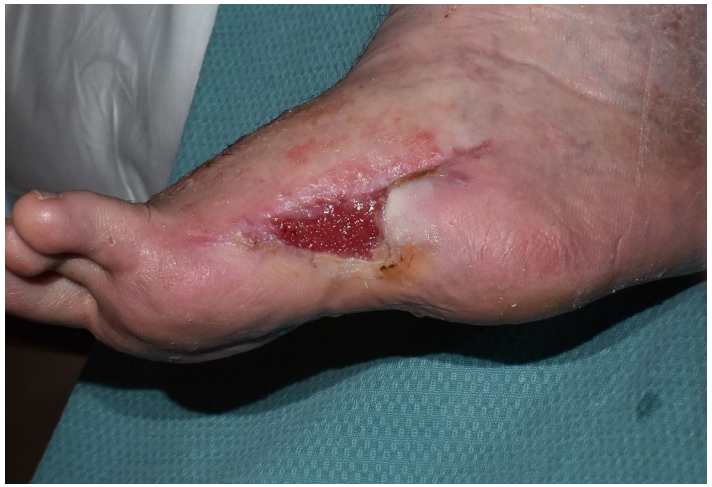
DFU | Phase 1 clinical trial – initial data

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

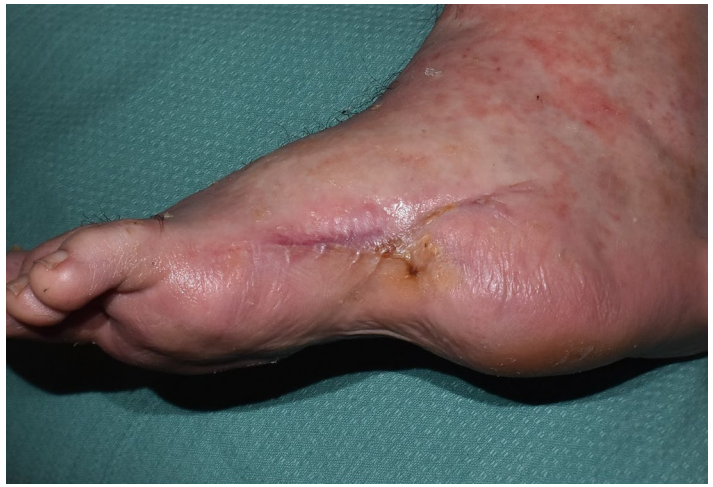
- Ongoing trial in non-healing diabetic foot ulcer (DFU)
- Patients randomised to receive standard of care (SoC) or CYP-006TK for 4 weeks, followed by SoC
- In the first 16 patients enrolled in the trial (8 per group), after 10 weeks' follow-up, the median reduction in wound surface area was:
  - **87.6%**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

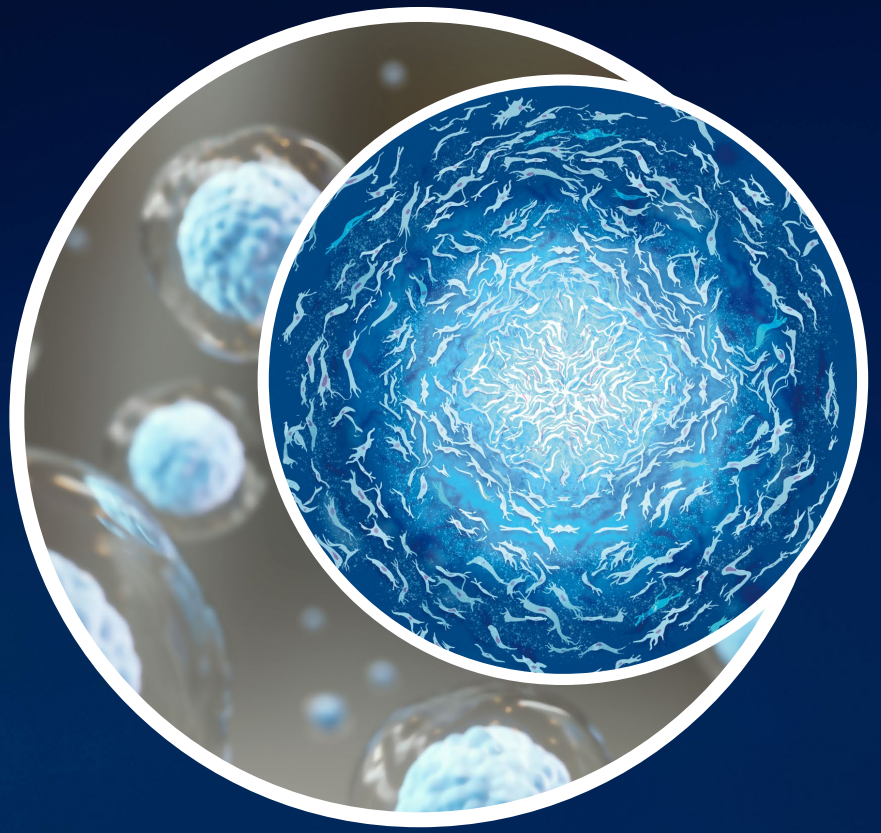


Day 28



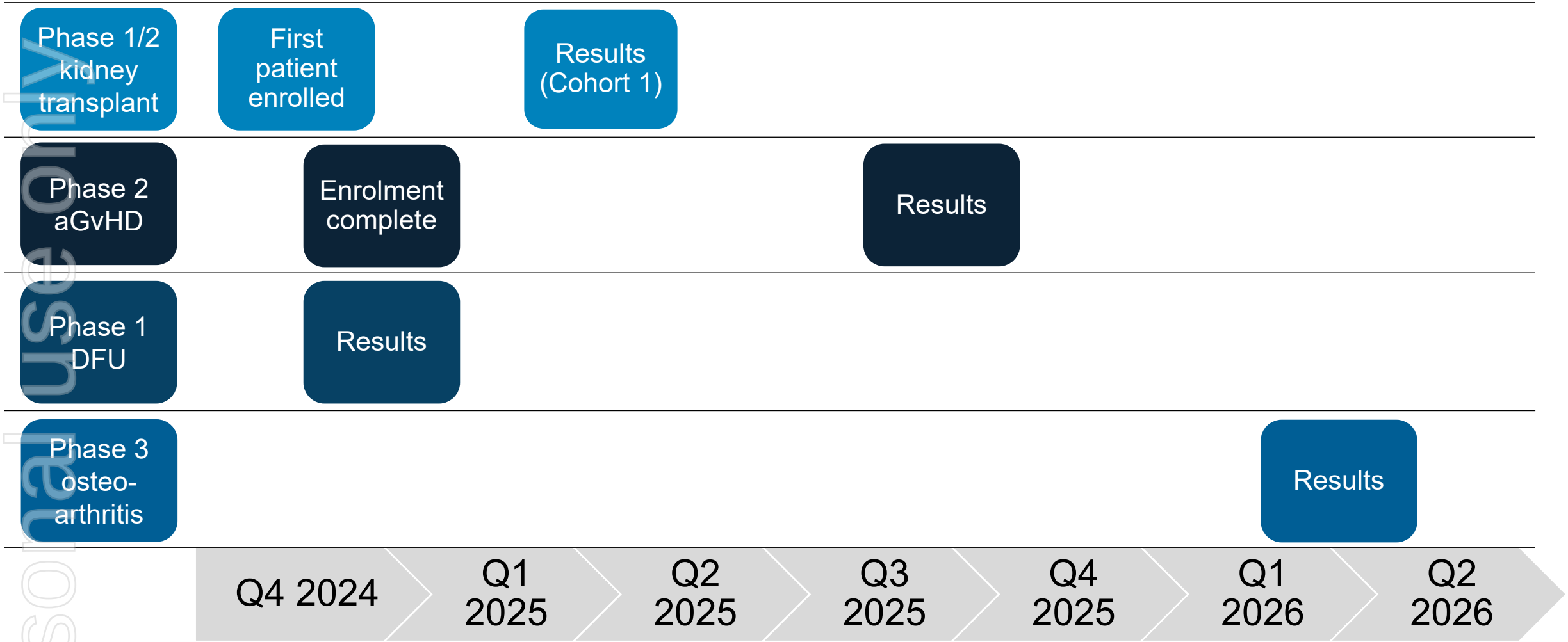
For further information: https://clinicaltrials.gov/study/NCT05165628

Outlook



Upcoming catalysts*

Results of three randomised controlled clinical trials expected between late 2024 and early 2026



* Timing of events is approximate, based on the Company's information as at the date of this presentation, and subject to change

Summary

Next generation stem cell company	• Leading platform technology in burgeoning stem cell sector • Diverse and highly credentialed leadership team with proven experience
Scalable manufacturing	• Cymerus™ manufacturing technology protected by robust patent portfolio • Enables scalable production of consistent MSCs from a single donation from a single donor, overcoming major challenges with conventional approaches
Compelling clinical data	• Very encouraging safety and efficacy results from aGvHD clinical trial (CYP-001) • Promising initial data from ongoing DFU clinical trial (CYP-006TK)
Rich clinical pipeline	• Broad pipeline with four active clinical programs • FDA orphan drug designation & cleared IND for ongoing Phase 2 aGvHD clinical trial • Patient enrolment complete in DFU & OA clinical trials • Commencement of kidney transplantation clinical trial imminent
Significant growth potential	• Global estimated market opportunity across targeted indications of ~US$28bn[1] • Focus on indications with significant unmet need • Proactive B-2-B outreach to drive partnering strategy

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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

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