IN FRONT OF THE BIGGEST WAVE IN ONCOLOGY

Prescient Therapeutics

August 2022

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Corporate snapshot



ASX Ticker	РТХ	
Total Issued Capital	648 M shares	0.30
Listed Options	95.4 M	
Unlisted Options	12.1 M	
Share Price ¹	A\$0.21 (US\$0.15)	
Market Capitalisation ¹	A\$141 M (US\$99 M)	
Market Cap fully diluted ¹	A\$162 M (US\$114 M)	J W WW 0.05
Cash Position ²	A\$14.7M (US\$11M)	
Top 20 Own	16%	Image: Second

Investment Highlights





World class pedigree.

We license from the best; and work with the best





Many shots on goal for substantial value creation



2 Cell Therapy platforms

Internal & external opportunities





Upcoming newsflow from multiple programs

4 Innovative Personalised Oncology Assets





Innovative pipeline in personalised medicine







TARGETED THERAPIES



PTX-100 FIRST IN CLASS RAS PATHWAY INHIBITOR

PTX-100 Phase 1B Summary

- Phase 1b PK/PD safety study
- Targeting cancers predisposed to Ras & Rho mutations
- Basket trial of:
 - Gastric cancer
 - Pancreatic cancer
 - Colorectal cancer

- Myeloma
- T-cell lymphomas

- Encouraging signal in TCL
- Now expanding the trial in Peripheral T-cell lymphomas (PTCL)
- Granted Orphan Drug Designation by US FDA











Professor H. Miles Prince, AM



Favourable safety profile compared to peers



Approved PTCL drugs have troublesome safety profiles



PTX-100 HAS AN EXCELLENT SAFETY PROFILE

- No serious adverse events related to PTX-100
- Suits fragile patient population
- Good candidate for combination therapy

Encouraging activity in TCL



Early clinical activity

- PRs in 2 patients with aggressive refractory TCL
- Expected PFS of <4 months on SoC
 - r/r CTCL: 12 months (19 cycles)
 - r/r PTCL: >32 months so far (37 cycles, still on therapy)
- Expansion cohort in TCL underway



'Very encouraging': Two cancer patients see partial remission and long-lasting benefits after treatment with Prescient's PTX-100

PR: PARTIAL RESPONSE (REDUCTION OF DISEASE) PFS: PROGRESSION FREE SURVIVAL (TIME UNTIL DISEASE WORSENS) SOC: STANDARD OF CARE

TCL: T CELL LYMPHOMA CTCL: CUTANEOUS T CELL LYMPHOMA PTCL: PERIPHERAL T CELL LYMPHOMA MM: MULTIPLE MYELOMA AITL: ANGIOIMMUNOBLASTIC T CELL LYMPHOMA CRC: COLORECTAL CANCER

Time on Treatment with PTX-100



Now in Expansion Cohort for TCL



- 8 12 patients with r/r T cell lymphoma (esp PTCL)
- Potential bridge to registration study
- Focussing on sweet spot in an area of considerable unmet need
- Shortest path to market

Case Study

- pralatrexate (Folotyn[®])
- Approved for PTCL
 - 5,600 cases/year in US
- US\$450,540 per patient, per year







PTX-200

NOVEL AKT INHIBITION

Phase 1B trial underway: Acute Myeloid Leukemia

- Building upon encouraging Phase 1 results with PTX-200 (monotherapy)
- PI Professor Jeff Lancet at Moffitt, Key Opinion Leader in AML
- 24 patients with cytarabine held constant at 200-400 mg/m² as continuous infusion
 - 4 patients with CR/CRi so far
 - 1 patient with PR
- Currently treating expansion cohort at 45 mg/m²
- Granted Orphan Drug Designation by US FDA





Principal Investigator



Jeffrey E Lancet, M.D.



PLATFORM TECHNOLOGIES

How does the CAR-T process work?





Cell Therapy is the future of oncology





ENDPOINTSNEWS

Carl June: 'We can now conclude that CAR-T cells can actually cure patients'





First patients of pioneering CAR T-cell therapy 'cured of cancer'



Doug Olson still has cancer-killing cells 10 years after infusion. Photograph: AP

thepharmaletter

Janssen gains EC green light for CAR-T therapy Carvykti





Penn is a pioneer and world leader in CAR-T





UNOVARTIS

- Novartis licensed CAR-T technology from Penn in 2012
- Kymriah[®] became the first CAR-T therapy approved by the FDA
- Used for certain blood cancers
- Cost of treatment in excess of \$500,000 per treatment
- GlobalData forecasts Kymriah[®] sales to exceed US\$1 billion in 2023



CAR-T's key challenges



Challenge

	Safety / Control	No control post infusion	
Ø	Targeting	Difficulties with targeting, antigen heterogeneity	Unsafe
100	Escape	Difficulties with mutating antigens	Less effective
	Production efficiency	Cost prohibitive & slow	Not sustainable
	Exhaustion	Cells run out of steam	Too expensive
	Trafficking	Cells cannot find their way	
×	Tumor penetrance	Protective layer around tumor	Don't last
1	Tumor microenvironment	Suppresses immune cells	

CAR-T's key challenges



		Challenge	OmniCAR	CellPryme-M	
	Safety / Control	No control post infusion	\checkmark	-	
Ø	Targeting	Difficulties with targeting, antigen heterogeneity	\checkmark	-	Safe
10	Escape	Difficulties with mutating antiger	ns 🗸	-	Effective
	Production efficiency	Cost prohibitive & slow	\checkmark	-	Sustainable
	Exhaustion	Cells run out of steam	\checkmark	\checkmark	Afferdable
	Trafficking	Cells cannot find their way	\checkmark	\checkmark	Affordable
×	Tumor penetrance	Protective layer around tumor	\checkmark	\checkmark	Enduring
Ĩ	Tumor microenvironment	Suppresses immune cells	\checkmark	\checkmark	





Universal, Next Gen CAR-T Platform

OmniCAR: flexible, modular CAR platform

OmniCAR

Targeting









Associate Professor Daniel J. Powell, Jr

Professor Andrew Tsourkas



ligand Using any targeting SpyTag ligand... Covalent bond Spy Catcher Signalling domain ...with any immune ARMED **UNARMED** cell **CAR-T cell CAR-T cell**





Complementary platforms to address CAR-T challenges



		Challenge	OmniCAR	🔅 CellPryme-M
\bigcirc	Safety / Control	No control post infusion	Tune activity up/down; On/off	-
Ø	Targeting	Difficulties with targeting, antigen heterogeneity	Target multiple antigens	-
	Escape	Difficulties with mutating antigens	Sequential targeting	-
₩,	Production efficiency	Cost prohibitive & slow	Far more efficient	\checkmark
	Exhaustion	Cells run out of steam	Longer-lasting cells	\checkmark
	Trafficking	Cells cannot find their way	Can direct cells	\checkmark
×	Tumor penetrance	Protective layer around tumor	Can overcome	\checkmark
Î	Tumor microenvironment	Suppresses immune cells	Can overcome	\checkmark

OmniCAR can do what conventional CAR-T cannot 🐐



Conventional CAR-T



- Soldier with only one map
- Single weapon
- Only trained to hit one target
- Incapable of redirection
- No communication or control in the field

OmniCAR Can be given **any map**; Armed with any weapon **Multiple deployments** Including several at once Full communication and **control** at all times, even mid-mission Can direct against any target, including simultaneous targets

Send images back to base in real time



OmniCAR: Control Features



Modular and covalent architecture of OmniCAR enables true post-infusion control of CAR functionality



Dose Titration





Target Re-direction



Multi-Antigen Targeting



Control activity to **safe** and efficacious levels Turn therapy on/off/on without killing or re-administering cells = safety & persistence

Re-direct cells from one cancer target to another in vivo Target **multiple cancer antigens simultaneously** for thorough cancer killing

Control: Dose-dependent CAR-T activity





- Ovarian cancer model, using anti-HER2 OmniCAR
- Loading more binder results in proportionate killing of cancer...
- ...and proportionate survival
- Lasting effects even when cease dosing of binder



Powell, DJ et al, JACS; 2020

OmniCAR cells can be Re-Armed





- OmniCAR T cells can be re-armed
- Re-arming results in same levels and kinetics of cytotoxicity as pre-armed
- · Another example of flexible yet predictable activity

OmniCAR cells can be Redirected



Coculture of U251 GBM Cells expressing HER2 or EGFRviii



- U251MG-EGFRviii (no switching)
- U251MG-HER2 (no switching)
- U251MG-EGFRviii (HER2 switching)
- U251MG-HER2 (HER2 switching)

- Rapid cytotoxicity to EGFRviii
- Rapid switching and cytotoxicity against HER2+ tumours upon administration of new binder
- OmniCAR cells can be re-directed to different antigens upon administration of a different SpyTagged binder without new cells

OmniCAR manufacturing & COGS advantages





Conclusion: OmniCAR would provide significant cost economics along with control and flexibility compared to conventional auto/allo CAR-T





Next Gen CAR-T Programs

OmniCAR internal program summary



Targets	Indications	OmniCAR features	Comments
CD33 + CLL-1	Acute Myeloid Leukemia (AML)	 Titration for improved safety Co-arming against CD33 & CLL-1 Sequential targeting 	 Validated targets; expressed on 90%+ of AML blasts & LSCs 1 of 5 programs worldwide; the only next-gen program
HER2	Ovarian; breast & gastric cancers	 Titration for improved safety Persistent binder dosing for improved efficacy TME and checkpoint enhancements 	 Most mature next-gen HER2 CAR-T program Builds on Penn pre-clinical PoC
HER2 + EGFRviii	Glioblastoma multiforme (GBM)	 Titration for improved safety Co-arming against HER2 & EGFRviii Persistent binder dosing for improved efficacy 	 1 of 3 multiple antigen programs in the world Single antigen targeting is inadequate in GBM

OmniCAR progressing towards clinic



- Steady progress across all programs
- OmniCAR AML likely the first program in clinical trials
- Q-Gen Cell Therapeutics appointed as cell manufacturer
 - Clinical grade cells
 - Autologous T cells expressing SpyCatcher
 - Incorporating CellPryme-M for superior phenotype



• Prescient to articulate regulatory path and clinical development details shortly

Thermo Fisher agreement for next version OmniCAR

Thermo Fisher

- Global leader in scientific instrumentation & services
- US\$40 billion revenue
- Expertise in cell & gene technologies and manufacturing

Prescient Therapeutics

Research agreement (MTA) to extend OmniCAR platform:

- Non-viral methods of transduction
 - Greater transduction efficiency
 - Faster
 - Lower COGS
- Automated, closed-end manufacturing
 - Scalable & reproducible
- Gene edits for additional enhancements
- Thermo Fisher carrying entire cost (substantial but undisclosed)

Aims & Outcomes for OmniCAR

Prescient

Further future-proofing OmniCAR platform

- V2 OmniCAR cells that:
 - Can be made in an **automated** process
 - Unmatched reproducibility
 - Faster production time
 - Substantially lower COGS
- Gene-edited OmniCAR cells with functional enhancements
- Seek to incorporate into Prescient's current OmniCAR programs

Positioning OmniCAR for technical & commercial success

- Manufacturing can be easily tech transferred to 3rd parties
- Amenable to decentralised manufacturing
- Ideal for multi-centre treatments:
 - During development
 - Commercial roll-out

Additional potential benefits of the Thermo Fisher agreement



Early access to Thermo Fisher's new, state-of-the-art technologies

• Protocol and process optimization from Thermo Fisher's technical experts

• Regulatory support to enable Prescient's regulatory filings

Ongoing support from Thermo Fisher as OmniCAR programs grow and advance





Cell therapy enhancements

CellPryme: Prescient's newest family member



PROCESS TO ENHANCE CELL THERAPIES

- Current gen and next gen
- Complementary to OmniCAR

2 SYNERGISTIC COMPONENTS

- CellPryme-M
- CellPryme-A (coming soon)

CellPryme-M

Produces superior cells

• Use with any existing CAR-T manufacturing process



CellPryme-M IP FULLY OWNED BY PTX

Developed by PTX in collaboration with Peter Mac

What does CellPryme-M do?



CellPryme-M is a single, rapid manufacturing step that produces a better, more effective cell type:

LONGER LASTING CELLS FOR SUSTAINED TUMOUR KILLING

- 50% more memory T cells
- Doubles helper T cells
- Doubles tumour control

CELLS THAT CAN BETTER LOCATE THE TUMOUR

- Significantly more chemokine receptors for improved trafficking to tumour sites
- Important in solid tumours



CellPryme-M complements OmniCAR





- Multi-targeting
- Redirection
- Control & safety
- Any target; any cell





Process that produces a better <u>cell type</u>

- Persistence
- Trafficking



Next generation Cell therapies Current generation cell therapies

Complementary platforms to address CAR-T challenges



		Challenge	OmniCAR	CellPryme-M
	Safety / Control	No control post infusion	\checkmark	
Ø	Targeting	Difficulties with targeting, antigen heterogeneity	\checkmark	-
	Escape	Difficulties with mutating antigens	\checkmark	
	Production efficiency	Cost prohibitive & slow	\checkmark	Superior cells & yield
	Exhaustion	Cells run out of steam	\checkmark	Longer lasting
	Trafficking	Cells cannot find their way	\checkmark	Cells locate tumors
×	Tumor penetrance	Protective layer around tumor	\checkmark	Better penetrance
Î	Tumor microenvironment	Suppresses immune cells	\checkmark	Less prone to suppression

More memory cells required for clinical efficacy **Prescient**

- Clinical efficacy of CAR-T therapy remains dependent on the T cell phenotype
- It is possible to control this during the manufacturing step



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Greater Persistence: 50% more central memory cells *Prescient* than conventional CAR-T





CellPryme-M doubles tumour control and survival



CellPryme-M nearly doubles CAR-T tumour control



CellPryme-M doubles survival





40

Conventional

CAR-T

CellPryme

CAR-T

0

Conventional

CAR-T

CellPryme

CAR-T

0

Conventional

CAR-T

CellPryme

CAR-T

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Conventional CellPryme

CAR-T

CAR-T

Synergy: CellPryme-M doubles proportion of helper T cells





- Shift towards dominant helper CD4+ CAR T cells
- Helper T cells are known to prevent the exhaustion of cytotoxic CD8+ T cells
 - Some can also have tumour killing ability
- Helper & cytotoxic T cells work in synergy to increase CAR-T persistence





Trafficking: greater chemokine receptor expression

- Effector T cells can downregulate chemokine receptors (CXCR3), limiting the ability of conventional CAR-T cells to locate tumours
- CellPryme-M significantly increases CXCR3 expression on CAR-T cells
- Better trafficking to tumour site
- Better tumour penetrance



Chemokine receptor expression on CD8+ cytotoxic CAR-T cells



CellPryme-M produces CAR-T cell types with ideal characteristics and attributes





Persistence For longevity of effects and continued tumour control



Immune memory

Central memory T cells typically persist 10-20 years and as long as 75 years



Trafficking

CAR-T cells able to find their way to the tumour



Tumour penetrance

Cells that can penetrate solid tumours



Genomic stability

Cells with enhanced self-renewal due to greater genomic stability



Anti-viral Cells with potent anti-viral characteristics



BUSINESS MODEL & SCOPE

Prescient's CAR-T platform business model





- Huge market
- "Shovels to CAR-T goldrush"
- Diversified risk
- Highly scalable
- Earlier revenue potential

Commercial models - Partners





OmniCAR Platform business model





CellPryme-M Next steps and future applications



IN-HOUSE DEVELOPMENT

- PTX will be its own first customer
- Incorporate into internal OmniCAR programs
- Trade secret manufacturing process



EXTERNAL OPPORTUNITIES

- Incorporate into 3rd party programs
- Attractive option for improving existing suboptimal CAR T products
- Haematological malignancies
 → to improve persistence
- Solid tumours
 - \rightarrow to improve trafficking and persistence
- Revenue potential for PTX

The End Game: Personalized "Plug & Play" Cell Therapy Ecosystem







Summary

Top-down analysis is sensible for investors





Oncology*

- 2021: US\$ 280bn
- 2029: US\$ 536bn (8.2% CAGR)

Cell Therapies (CAR-T)

>US\$37bn by 2028^

Prescient Therapeutics

- Next gen platforms
- Scalable
- Controllable
- Any target; any cell
- "Shovels to goldrush" position
- Top pedigree

Investment Thesis Summary

4 blue chip oncology assets

2 next gen platforms



PTX-100 & PTX-200 in clinic



Top pedigree



Superior positioning & model





Shovels to goldrush

Highly scalable









Thank you!

ASX code: PTX

www.ptxtherapeutics.com