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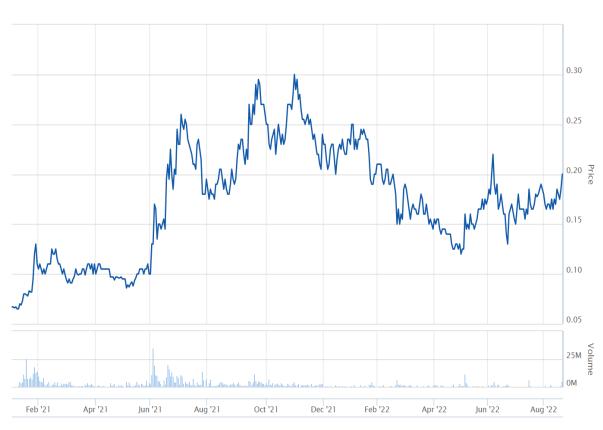
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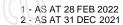


## **Corporate snapshot**



ASX Ticker	PTX
Total Issued Capital	648 M shares
Listed Options	95.4 M
Unlisted Options	12.1 M
Share Price <sup>1</sup>	A\$0.21 (US\$0.15)
Market Capitalisation <sup>1</sup>	A\$141 M (US\$99 M)
	·
Capitalisation <sup>1</sup> Market Cap fully	(US\$99 M) A\$162 M





## **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



2 Targeted Therapies in clinic trials, showing activity



**Upcoming newsflow** from multiple programs



## 4 Innovative Personalised Oncology Assets





# Next-gen CAR-T therapies

### **Targeted therapies**



**OmniCAR** 



#### CD33/CLL-1

Acute Myeloid Leukaemia

#### Her2/EGFRviii

Glioblastoma Multiforme

#### Her2

Breast, Ovarian & Gastric cancers

#### PTX-100

Ras / Rho inhibitor

### PTX-200

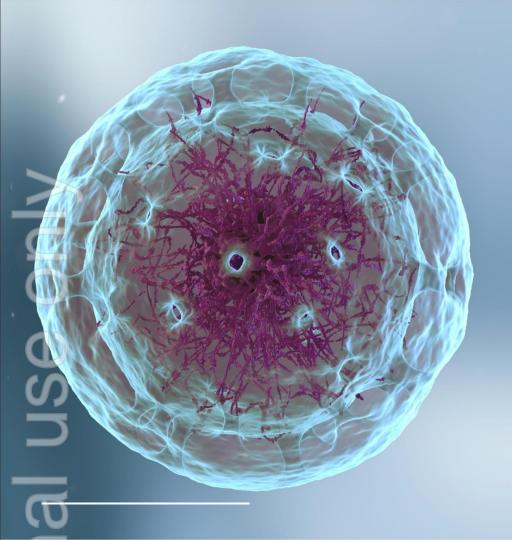
Novel AKT inhibition



# Innovative pipeline in personalised medicine



	Platform	Program	Screening	Preclinical	Phase IB	Phase II
		CD33/CLL-1	AML		•	
	TÂN.	Her2	Breast, Ovarian &	Gastric cancers		
	OmniCAR	Her2/EGFRviii	GBM			
CAR - T		Platform extensions	Various	Thermo Fisher SCIENTIFIC		
	**** O - IID	CellPryme-M	Cell manufacturin	g enhancement		
	: CellPryme	CellPryme-A	Undisclosed			
TARGET	PTX-100		PTCL			
THERAP			AML			



# TARGETED THERAPIES



# PTX-100

FIRST IN CLASS
RAS PATHWAY INHIBITOR

### PTX-100 Phase 1B Summary

- Licensed from



- 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

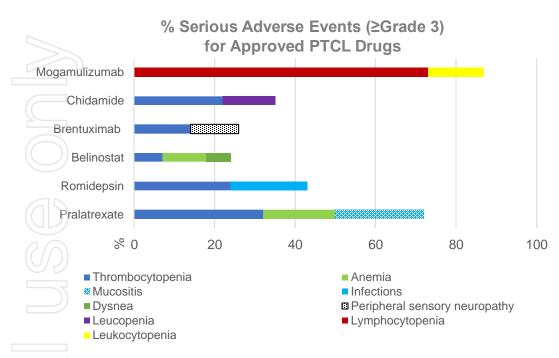




## Favourable safety profile compared to peers



#### **Approved PTCL drugs have troublesome safety profiles**



# PTX-100 HAS AN EXCELLENT SAFETY PROFILE

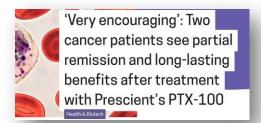
- No serious adverse events related to PTX-100
- Suits fragile patient population
- Good candidate for combination therapy

## **Encouraging activity in TCL**

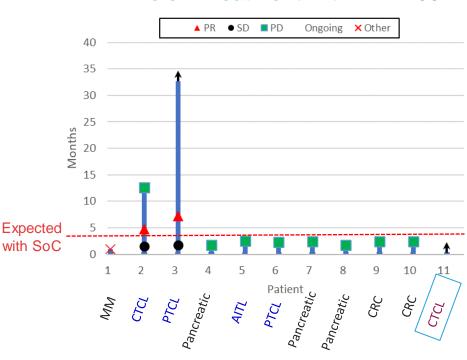


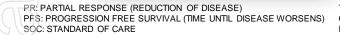
### Early clinical activity

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



#### **Time on Treatment with PTX-100**





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

### **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<sup>®</sup>)
- 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/m<sup>2</sup> as continuous infusion
  - 4 patients with CR/CRi so far
    - 1 patient with PR
- Currently treating expansion cohort at 45 mg/m<sup>2</sup>
  - Granted Orphan Drug Designation by US FDA



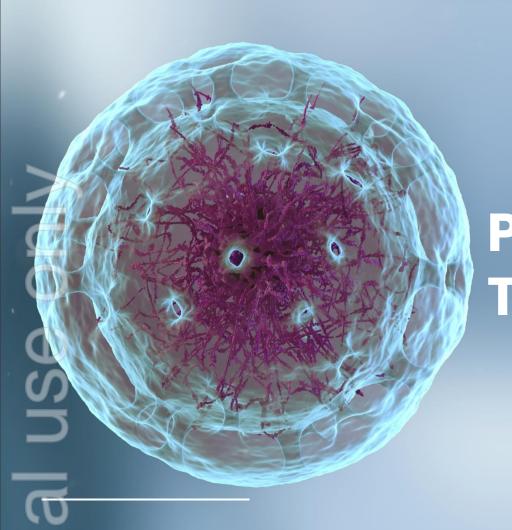


**Principal Investigator** 



Jeffrey E Lancet, M.D.

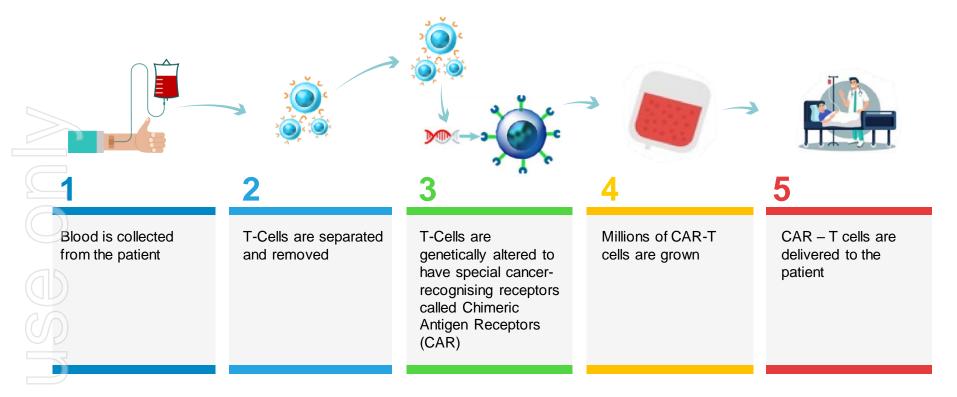




# PLATFORM TECHNOLOGIES

# How does the CAR-T process work?

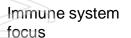




### **Cell Therapy is the future of oncology**









Powerful cancer fighting potential



Highly encouraging data



Long-term remissions



Still a lot to discover

#### **ENDPOINTS** NEWS

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



#### The Guardian

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



98% Overall response rate



#### **CAR-T** is booming GlobalData forecasts 2028 **CAR-T** sales to exceed US\$37bn **TODAY** 2023 GlobalData forecasts Continued positive data flow Kymriah® sales to Some outstanding results exceed US\$1bn 100s of companies targeting different antigens to target 2017 more cancer cells FDA designates CARs a "break-through" therapy First effective Immune cells shown CAR-T cells to protect mice from developed, CD19 2014 cancer, origin of T successfully targeted First CARs cross the cells discovered 2000 regulatory finish line 1960 2012 1990 CAR-T licensed from T cell engineering begins; Penn by Novartis to first-generation CARs develop Kymriah® developed; Antigen-specific T cells used in humans

## Penn is a pioneer and world leader in CAR-T









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

T)	

Safety / Control No control post infusion



Targeting Difficulties with targeting, antigen heterogeneity



**Escape** Difficulties with mutating antigens



Production efficiency Cost prohibitive & slow



**Exhaustion** Cells run out of steam



**Trafficking** Cells cannot find their way



Tumor penetrance Protective layer around tumor



Tumor microenvironment Suppresses immune cells



**Less effective** 

Not sustainable

Too expensive

Don't last



# **CAR-T's key challenges**



		Challenge	OmniCAR	CellPryme-M	
	Safety / Control	No control post infusion	$\checkmark$	-	
	Targeting	Difficulties with targeting, antigen heterogeneity	$\checkmark$	-	Safe
9	Escape	Difficulties with mutating antige	ns 🗸	-	Effective
	Production efficiency	Cost prohibitive & slow	$\checkmark$	-	Sustainable
	Exhaustion	Cells run out of steam	$\checkmark$	$\checkmark$	
Dů!	Trafficking	Cells cannot find their way	$\checkmark$	$\checkmark$	Affordable
**	Tumor penetrance	Protective layer around tumor	$\checkmark$	✓	Enduring
	Tumor microenvironment	Suppresses immune cells	$\checkmark$	$\checkmark$	
					21



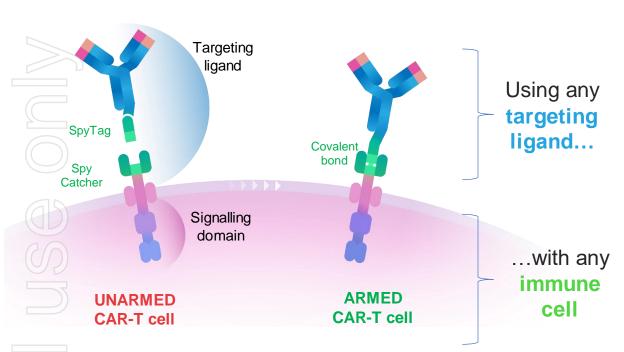


Universal, Next Gen CAR-T Platform

### OmniCAR: flexible, modular CAR platform













Associate Professor Daniel J. Powell, Jr

Professor Andrew Tsourkas





# Complementary platforms to address CAR-T challenges



		Challenge	OmniCAR	CellPryme-M
	Safety / Control	No control post infusion	Tune activity up/down; On/off	-
<b>3</b>	Targeting	Difficulties with targeting, antigen heterogeneity	Target multiple antigens	-
7	Escape	Difficulties with mutating antigens	Sequential targeting	-
	Production efficiency	Cost prohibitive & slow	Far more efficient	✓
	Exhaustion	Cells run out of steam	Longer-lasting cells	✓
III V	Trafficking	Cells cannot find their way	Can direct cells	✓
	Tumor penetrance	Protective layer around tumor	Can overcome	✓
	Tumor microenvironment	Suppresses immune cells	Can overcome	<b>√</b>

### OmniCAR can do what conventional CAR-T cannot \$\frac{1}{2}\$



#### 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





Can be given **any map**; Multiple deployments



Full communication and control at all times, even mid-mission



Send **images** back to base in real time

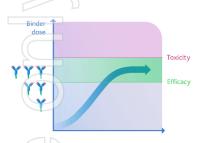


### **OmniCAR: Control Features**



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

### **Dose Titration**



Control activity to safe and efficacious levels

#### On/off switch



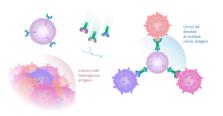
Turn therapy on/off/on without killing or re-administering cells = safety & persistence

### **Target Re-direction**



Re-direct cells from one cancer target to another in vivo

### Multi-Antigen Targeting

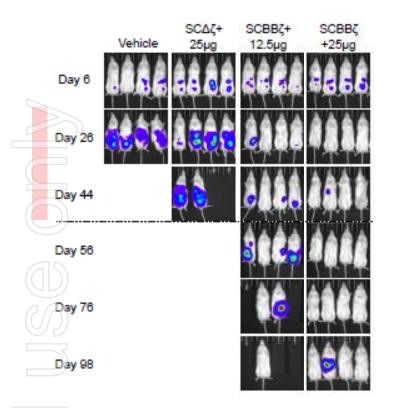


Target multiple cancer antigens simultaneously for thorough cancer killing

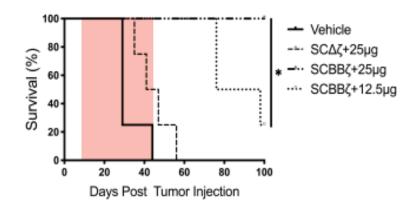


### **Control:** Dose-dependent CAR-T activity



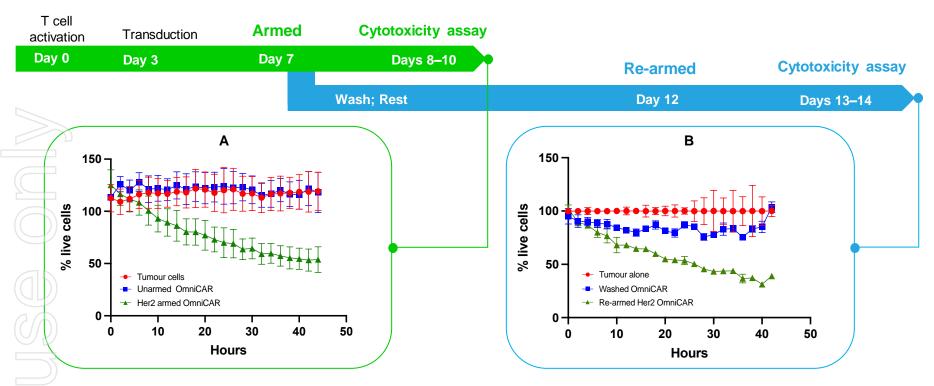


- Ovarian cancer model, using anti-HER2 OmniCAR
- Loading more binder results in proportionate killing of cancer...
- ...and proportionate survival
- Lasting effects even when cease dosing of binder



### OmniCAR cells can be Re-Armed

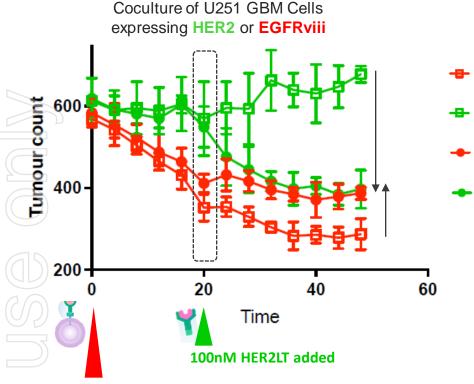




- OmniCAR T cells can be re-armed
- Re-arming results in same levels and kinetics of cytotoxicity as pre-armed
- Another example of **flexible** yet **predictable** activity

### OmniCAR cells can be Redirected





- 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

100nM EGFRviii OmniCAR cells added

## **OmniCAR** manufacturing & COGS advantages



### Conventional CAR-T

6 INDs

6 Manufacturing runs

6 Vectors/Transductions

6 Sets of release testing

No Flexibility

 $= 6 \times (\$\$\$)$ 



1 IND

1 Optimized Manufacture

1 Vector/Transduction

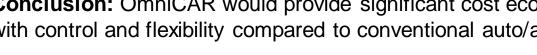
1 Release program

Multiple binders off the shelf

No time delay for subsequent doses

 $= (1 \times $$$) + (6 \times $$)$ 

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

## **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

# 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**



# 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 3<sup>rd</sup> 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



READY FOR CLINICAL TESTING



## **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

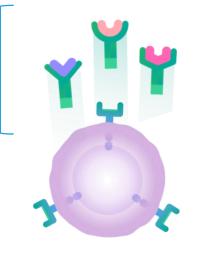


## **CellPryme-M complements OmniCAR**



## OmniCAR

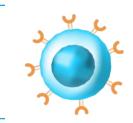
- Multi-targeting
- Redirection
- Control & safety
  - Any target; any cell





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

- Persistence
- Trafficking



Next generation Cell therapies

Current generation cell therapies





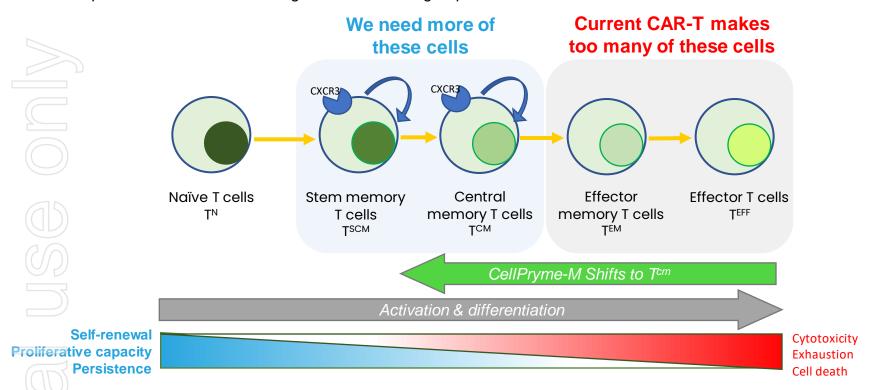
## Complementary platforms to address CAR-T challenges



	Challenge	OmniCAR	CellPryme-M
Safety / Control	No control post infusion	✓	-
Targeting	Difficulties with targeting, antigen heterogeneity	✓	-
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	✓	Cells locate tumors
Tumor penetrance	Protective layer around tumor	✓	Better penetrance
Tumor microenvironment	Suppresses immune cells	$\checkmark$	Less prone to suppression

## More memory cells required for clinical efficacy Prescient

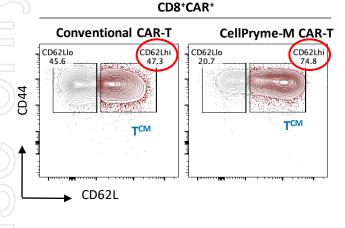
- Clinical efficacy of CAR-T therapy remains dependent on the T cell phenotype
- It is possible to control this during the manufacturing step

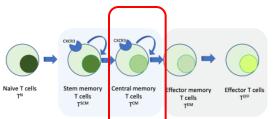


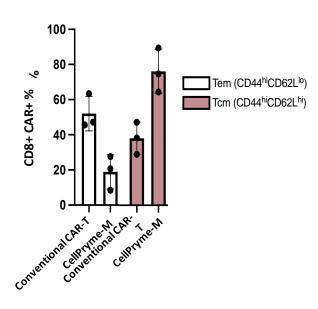
## Greater Persistence: 50% more central memory cells Prescient than conventional CAR-T



## **CellPryme-Mincreases central** memory T cells 1.5-fold within 24hrs



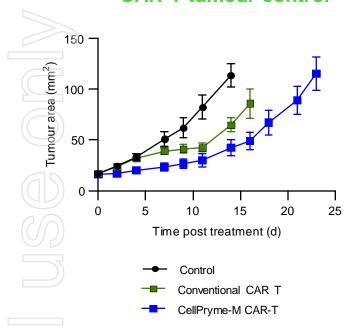




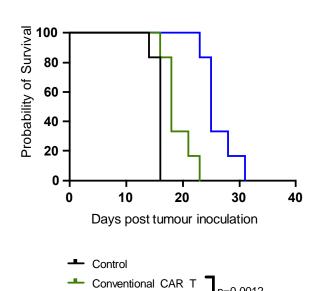
## **CellPryme-M** doubles tumour control and survival



## CellPryme-M nearly doubles CAR-T tumour control



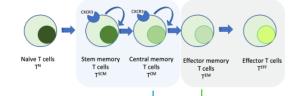
## CellPryme-M doubles survival



CellPryme-M CAR-T

## **Greater Persistence/Less Exhaustion**

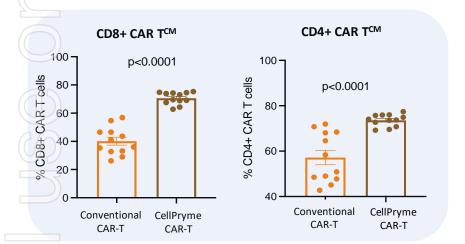




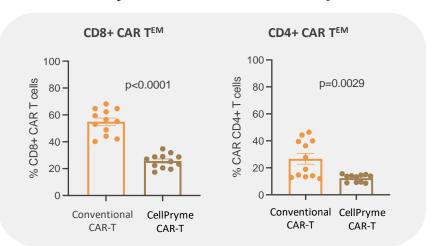
#### **Sustained increase in T<sup>CM</sup>**

#### Sustained decrease in T<sup>EM</sup>

## for both cytotoxic CD8+ and helper CD4+

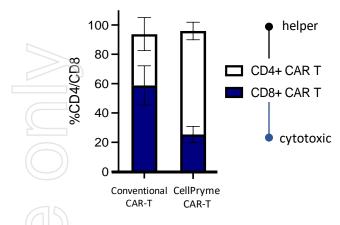


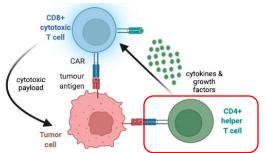
## for both cytotoxic CD8+ and helper CD4+



# 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

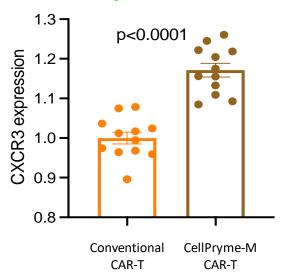
# Trafficking: greater chemokine receptor expression



- 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

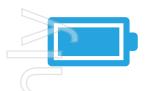
## Chemokine receptor expression on CD8+ cytotoxic CAR-T cells





# 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**

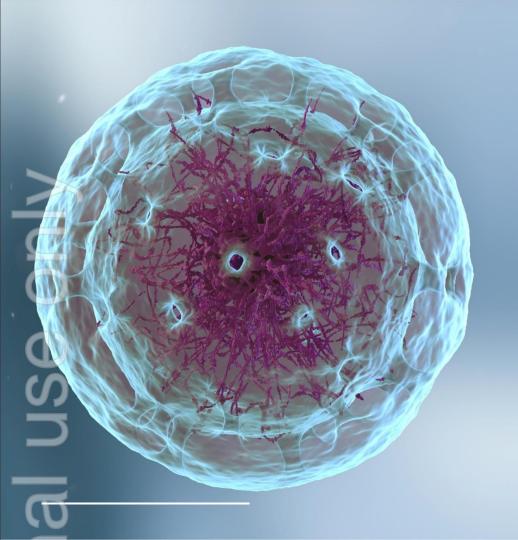
Cells with enhanced self-renewal due to greater genomic stability



#### **Anti-viral**

Cells with potent anti-viral characteristics

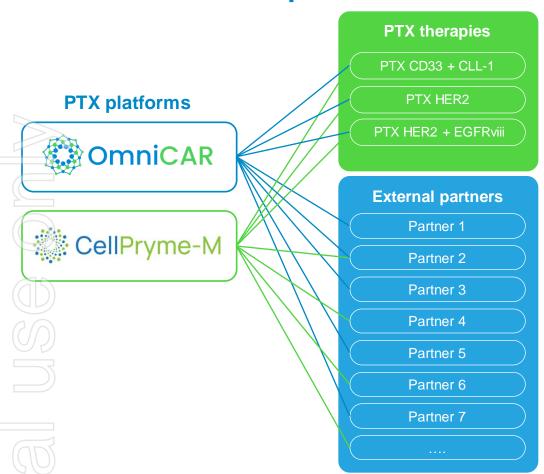




# BUSINESS MODEL & SCOPE

## Prescient's CAR-T platform business model

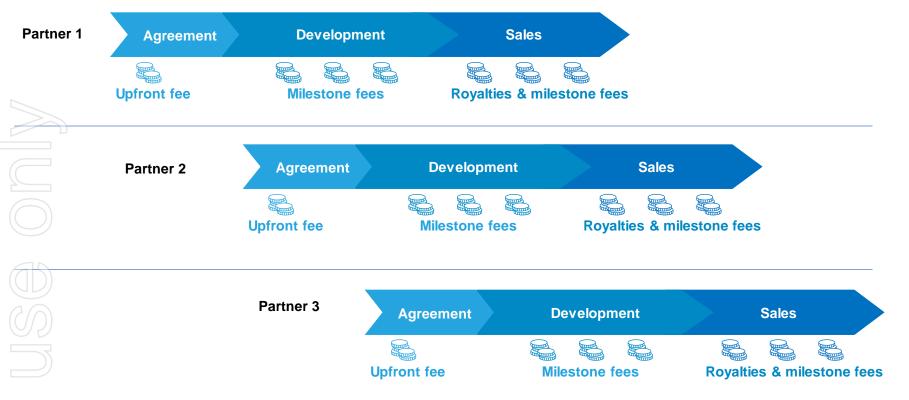




- Huge market
- "Shovels to CAR-T goldrush"
- Diversified risk
- Highly scalable
- Earlier revenue potential

## **Commercial models - Partners**





## **OmniCAR Platform business model**





OmniCAR CD33/CLL1 for **AML** 

OmniCAR Her2
For **Solid Tumours** 

OmniCAR Her2/EGFRviii for **GBM** 





COLLABORATE & LICENSE TO 3RD PARTIES

Partners for TARGETS & BINDERS

Partners for **CELL TYPES** 

Partners for SPECIFIC CONSTRUCTS



## **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



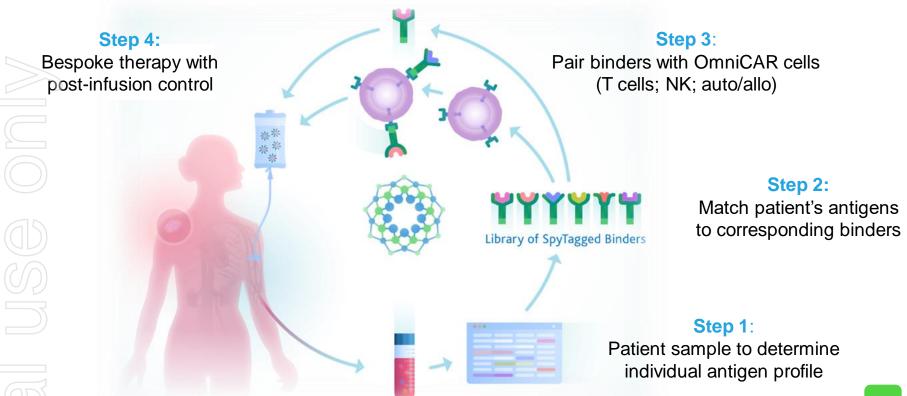
## **EXTERNAL OPPORTUNITIES**

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



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



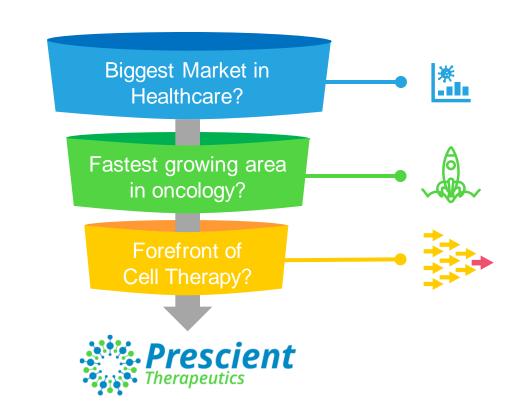




# Summary

## 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



PTX-100 & PTX-200 in clinic

**Top pedigree** 



CellPryme PTX-200

Superior positioning & model



Internal products
+ external partnering



**Shovels to goldrush** 



Highly scalable



Huge & growing market













