# BRINGING THE PROMISE OF CELL THERAPY TO LIFE

Corporate Update January, 2024



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# "We can now conclude that CAR-T cells can CURE

patients with leukemia"

Dr Carl June, MD
Richard W. Vague Professor in Immunotherapy
Director of the Center for Cellular Immunotherapies at the
Perelman School of Medicine



### CHIMERIC THERAPEUTICS (ASX: CHM) CORPORATE PROFILE

#### Market Information

Share Price (AUD) (January 17, 2024)	\$0.029
 52 Week Range (AUD) 	\$0.025- \$0.086
Shares on Issue	727 M
Market Cap (AUD) (January 17, 2024)	~\$21M

Current Approximate Cash Position (AUD)

~\$15M

(Including R&D rebate, Lind Placement and Entitlement Offer Receipts as announced in January 2024)



Bringing
the Promise
of Cell
Therapy to
Life

# 2 CLINICAL TRIALS WITH POSITIVE PHASE 1 RESULTS

- Positive CHM 1101 Phase 1A data in Recurrent Brain Cancer
- Positive CHM 0201 Phase 1A data in Acute Myeloid Leukemia and Advanced Colorectal Cancer
- Areas of high unmet medical need

### ADVANCING CLINICAL DEVELOPMENT

- 4 clinical trials in solid tumours and blood cancers in 2024
- Multiple clinical catalysts in the next 12-18 months

### EXPERIENCE AND EXPERTISE

- Industry leading team of experienced cell therapy experts
- Proven development and commercialization expertise

### **KEY CLINICAL CATALYSTS**

### **ACROSS THE 3 PLATFORM TECHNOLOGIES**

#### 2023 Clinical Achievements

#### **2024 Clinical Milestones**

CHM 1101 (CLTX CAR T)

- Ph. 1A Dose Escalation Complete in GBM
- Ph. 1A Positive Preliminary Data in GBM
- Ph. 1B 1st Patient Treated in GBM.

- Ph. 1B Dose Expansion Initiation in GBM
- Ph. 1B Preliminary Dose Expansion Data

CHM 2101 (CDH17 CAR T) FDA IND Clearance for Ph. 1A Basket
 Trial in Colorectal Cancer, Gastric
 Cancer and Neuroendocrine Tumors

- Ph. 1A Basket Trial Initiation
- Ph. 1A Preliminary Dose Escalation Data

CHM 0201 (CORE NK)

- Ph. 1B ADVENT AML Trial Initiation
- Ph. 1B CHM 0201 + Vactosertib
   1st Patient Treated

- Ph. 1B ADVENT AML 1st Patient Treated
- Ph. 1B ADVENT AML Dose Confirmation Cohort Completed
- Phase 1B ADVENT AML Preliminary Data



### CHIMERIC: 3 NOVEL PLATFORM TECHNOLOGIES

### **CHM 1101** (CLTX CAR T)

2020

First in class CLTX CAR T for brain cancer and other solid tumours

Positive Phase 1A Clinical Trial in Recurrent Glioblastoma

Ongoing Phase 1B Clinical Trial in Recurrent Glioblastoma



### **CHM 2101** (CDH17 CAR T)

2021

First in class CDH17 CAR T for gastrointestinal cancers

FDA IND clearance Nov 23

Phase 1A Clinical Trial in Colorectal Cancer, Gastric Cancer and Neuroendocrine Tumours



### **CHM 0201** (CORE NK)

2022

Potentially best in class NK cell platform for blood cancers and solid tumours

Positive Phase 1A Clinical Trial in Colorectal Cancer and AML

Ongoing Phase 1B Clinical Trial at MDACC in AML

Phase 1B Clinical Trial at CWRU in AML/ Colorectal Cancer

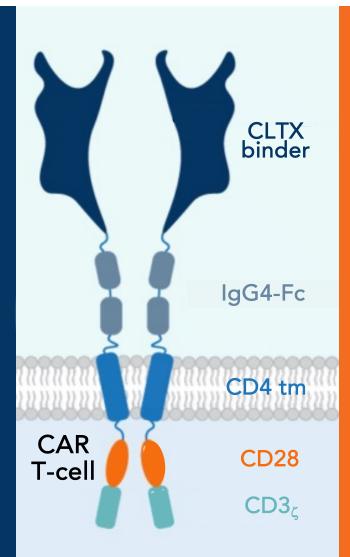




# CHM 1101 (CLTX CAR T) FIRST IN CLASS CAR T OPTIMIZED FOR SOLID TUMORS

### NOVEL TARGET

- First in class Chlorotoxin (CLTX) binder targets MMP2
- MMP2 is a biologically relevant target, contributing to metastasis and tumor cell survival
- High MMP2 target frequency (70%+) across a broad range of solid tumors
- Selective tumor binding and established safety from diagnostic utilization



### OPTIMIZED CONSTRUCT

- ✓ Novel, natural toxin Chlorotoxin (CLTX) binder with a compact, stable, peptide structure
- ✓ Optimized hinge and costimulatory domains selected to maximize potency and minimize exhaustion
- ✓ Construct tailored to promote immune synapse formation
- Designed to overcome solid tumor challenges with a manufacturing process that enriches for Tn/ mem cells

# POSITIVE PHASE 1A CLINICAL DATA IN RECURRENT, LATE-STAGE BRAIN CANCER

### DISEASE CONTROL RATE

55%

Disease Control Rate (DCR) in heavily pretreated patients

Exceeding historical disease control rates of 20-37%<sup>1</sup>

### SURVIVAL

### ~10 months

Median survival in patients that achieved disease control

14+ months

Survival in two patients that achieved disease control

~7 month survival expectation after first recurrence<sup>2</sup>

#### **SAFETY**

#### Generally, well tolerated

- No Dose Limiting Toxicities
- No Cytokine Release Syndrome
- No Tumour Lysis Syndrome

### HOW IS GLIOBLASTOMA (GBM) TREATED TODAY? TREATMENT BECOMES MORE CHALLENGING AS THE DISEASE PROGRESSES

**PROGRESSION** 

**OR DISEASE** 

**RECURRENCE** 

Treatment in front line is the standard of care combination of surgery, radiation and temozolomide.

Patients are generally expected to progress or recur after 7-8 months<sup>1</sup>

Treatment in subsequent lines of therapy becomes progressively more difficult with reduced survival expectations.

On initiation of 2<sup>nd</sup> line therapy, expected survival is approximately 7 months<sup>2</sup>

1st LINE THERAPY

STANDARD OF CARE

**Surgery + Radiation + Temozolomide** 

2<sup>nd</sup> Line **Therapy** 

3rd Line **Therapy** 

4th Line Therapy

5th Line Therapy

15

**NO STANDARD** OF CARE

1. Jeffree R (2020) Australian Journal of General Practice. Volume 49, Issue 4, April 2020. 2. Gallego O.

DIAGNOSIS

Nonsurgical treatment of recurrent glioblastoma. Curr Oncol. 2015 Aug;22(4):e273-81

CHIMERICTHERAPEUTICS

Glioblastoma Survival Expectation = ~ 15 Months

### CHM 1101 PHASE 1A PATIENT DEMOGRAPHICS TREATING 4<sup>TH</sup> LINE PATIENTS

46%

of patients received CHM 1101 as 4<sup>th</sup> line therapy

#### **CHM 1101 Patients**

2 <sup>nd</sup> Line Therapy			5 <sup>th</sup> Line Therapy	
0%	36%	46%	18%	

Approved therapies in recurrent Glioblastoma treated 2<sup>nd</sup> line patients

Age	Median Range	57 years 37-73 years
Sex	Male Female	55% 45%
Race	Caucasian Black Asian Hispanic	73% 9% 9% 9%
# of Prior Surgeries	0 1 2 3	18% 36% 36% 9%
Pathology	Grade 4 Glioblastoma Grade 4 Astrocytoma	91% 9%
IDH	Wild type Mutant	91% 9%



### **CHM 1101 RESULTS IN CONTEXT**

### 55% DISEASE CONTROL IN LATE LINE PATIENTS





\*Temodal\* temozolomide

**2**<sup>nd</sup> **Line** Treatment<sup>1</sup>

**20%**Disease Control<sup>1</sup>

4<sup>th</sup> Line

**CHM 1101** 

**Treatment** 

55%
Disease Control

**2nd Line** Treatment<sup>2</sup>

37%
Disease Control<sup>2</sup>

CHM 1101 was studied in median 4<sup>th</sup> line patients and demonstrated a Disease Control Rate (DCR) higher than NCCN approved and recommended therapies studied in 2<sup>nd</sup> line



- 1. The Lancet Oncology: Volume 20, Issue 1, 1-164, e1-e65
- 2. Journal of Clinical Oncology 28, no. 12 (April 20, 2010) 2051-2057

### SURVIVAL EXPECTATIONS FOR PATIENTS WITH RECURRENT GBM

		Therapy	Line of Therapy	Overall Survival
	APPROVED THERAPIES	SURGERY	2 <sup>nd</sup> Line <sup>1</sup>	5.75 months <sup>1</sup>
		TUMOUR TREATING FIELDS	3 <sup>rd</sup> Line <sup>2</sup>	6.6 months <sup>2</sup>
		BEVACIZUMAB	2 <sup>nd</sup> Line <sup>3</sup>	7.75 months <sup>3</sup>
		TEMOZOLOMIDE	2 <sup>nd</sup> Line <sup>4</sup>	3.7 months <sup>4</sup>
J.	INVESTIGATIONA L THERAPIES	REGORAFENIB	2 <sup>nd</sup> Line <sup>5</sup>	6.5 months <sup>5</sup>
		GALUNISERTIB + LOMUSTINE	2 <sup>nd</sup> Line <sup>6</sup>	6.7 months <sup>6</sup>

~7 months
survival
demonstrated
from first
recurrence with
approved and
investigational
therapies

<sup>6.</sup> Neuro-Oncology, Volume 18, Issue 8, August 2016, Pages 1146–1156



<sup>1.</sup> Curr Oncol. 2015 Aug;22(4):e273-81

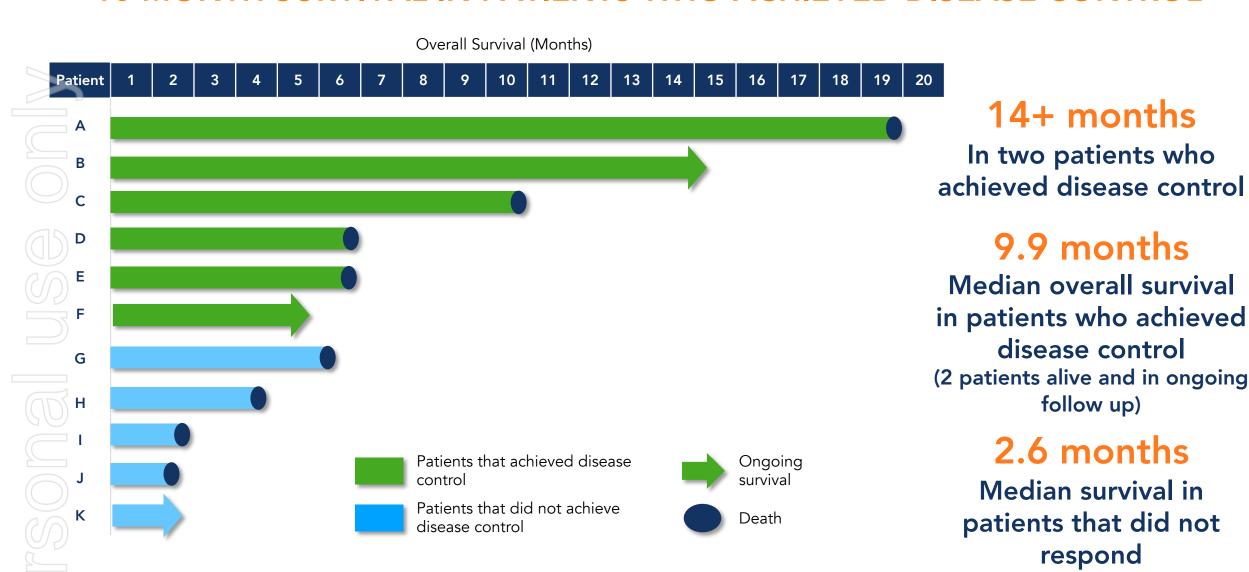
Eur J Cancer. 2012 Sep;48(14):2192-202.

Journal of Clinical Oncology, 2009. 27(5): p. 740-5 Curr Oncol. 2015 Aug;22(4):e273-81

<sup>5.</sup> Lancet Oncol. 2019, 20, 110-119

### CHM 1101 PHASE 1A SURVIVAL

### ~10 MONTH SURVIVAL IN PATIENTS WHO ACHIEVED DISEASE CONTROL



CHIMERICTHERAPEUTICS

## CHM 1101 PHASE 1A PATIENTS WITH LONG TERM (14+ MONTHS) SURVIVAL





### CHM 1101 PHASE 1A SAFETY A MANAGEABLE SAFETY PROFILE AT ALL DOSE LEVELS

No Dose Limiting Toxicities (DLT's)

No Cytokine Release Syndrome (CRS) No Tumour Lysis Syndrome (TLS) Grade 3 (serious)
non-hematological
events were not
considered treatment
related:

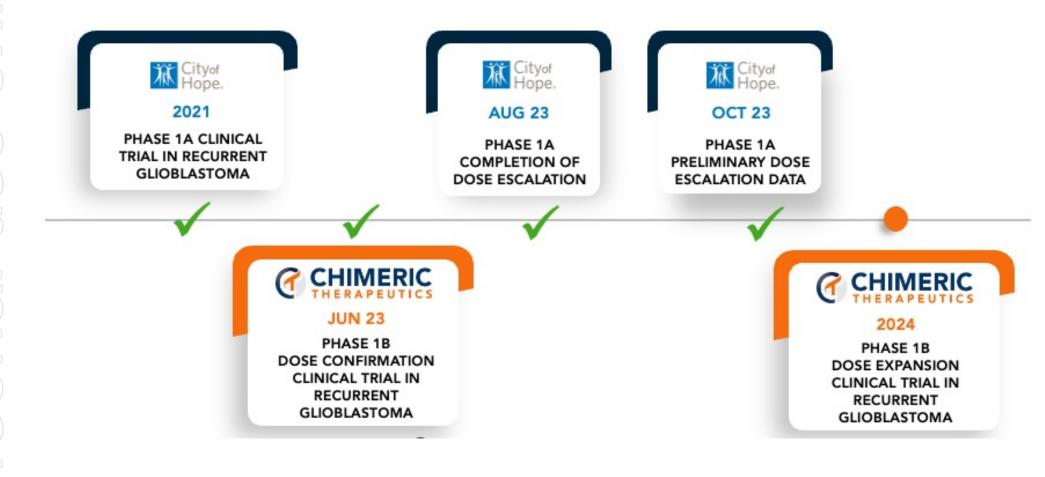
- Cerebral Edema X3\*
- Adrenal Insufficiency\*
- Headache\*
- Confusion\*
- Syncope
- Fatigue
- Ataxia

\*associated with GBM disease progression



### CHM 1101 NEXT STEPS ADVANCING IN PHASE 1B CLINICAL DEVELOPMENT

Based upon the safety and efficacy signal demonstrated in the Phase 1A City of Hope clinical trial, CHM 1101 will advance to Dose Expansion in the Chimeric Phase 1B Clinical Trial in 2024





### CHM 1101 SUMMARY

### POSITIVE PHASE 1A DATA IN LATE STAGE GLIOBLASTOMA

- ✓ 4<sup>th</sup> line, difficult to treat patients
- √ 55% Disease Control Rate
- √ ~10 months survival for patients that achieved disease control
- ✓ Two patients with survival beyond 14 months
- Manageable safety profile at all dose levels

#### **ADVANCING TO PHASE 1B DOSE EXPANSION**

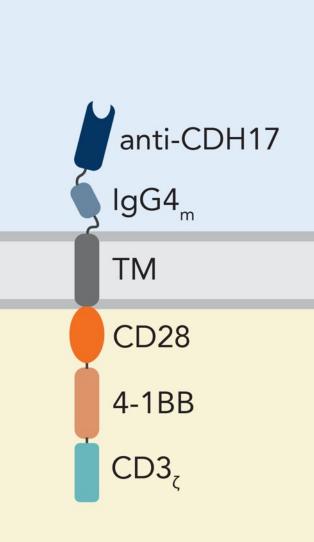
✓ Phase 1B recruiting at Sarah Cannon Research Institute



# CHM 2101 (CLTX CAR T) FIRST IN CLASS CAR T DESIGNED FOR GI CANCERS

### NOVEL TARGET

- First in class anti- Cadherin-17 (CDH17) binder
- CDH17 is a biologically relevant target, contributing to metastasis and tumor cell survival
- ✓ High CDH17 expression rate in GI cancers; ≥ 95% in lead indications, ≥ 50-65% in other tumors
  - ✓ Masked from CAR T recognition on normal cells, CDH17 expression pattern allows selective targeting of tumors by CHM 2101



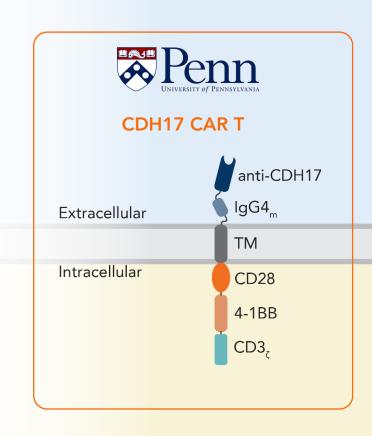
### OPTIMIZED CONSTRUCT

- ✓ Single domain VHH antibody binder with high affinity for human CDH17
- ✓ Highly compact, stable VHH antibodies less prone to aggregation than scFvs enabling complete tumor eradication while sparing normal tissues
- ✓ Superior 3<sup>rd</sup> Generation CAR designed for solid tumors with dual CD28 and 4-1BB costimulatory domains
- ✓ Hinge domain empirically selected for maximum potency and length adjusted to tighten immune synapse for target engagement

### CHM 2101: FDA IND CLEARANCE

### A FIRST-IN-CLASS 3RD GENERATION CAR T FOR GASTROINTESTINAL TUMOURS

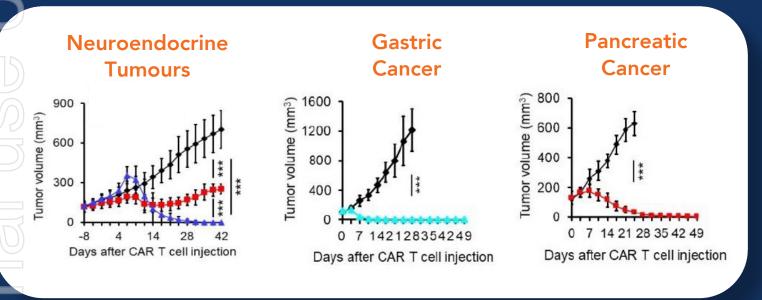
- FDA clearance for Chimeric IND for a Phase 1/2 clinical program in Colorectal Cancer, Gastric Cancer and Neuroendocrine Tumours
- First-in-class CAR T cell therapy licenses from world renowned cell therapy center, the University of Pennsylvania after a decade of optimization
- Broad applicability of the CDH17 CAR T to address unmet medical needs in gastrointestinal (GI) cancers
- Clinical trial to be initiated in 2024 at leading US institutions

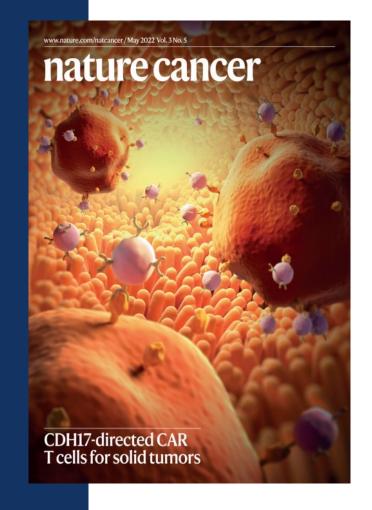




### CHM 2101 (CDH17 CAR T) POTENT IN VIVO EFFICACY

CHM 1101 induced complete eradication of tumours with no relapse in seven mouse xenograft tumour models





Source: Feng et al., Nature Cancer, 2022





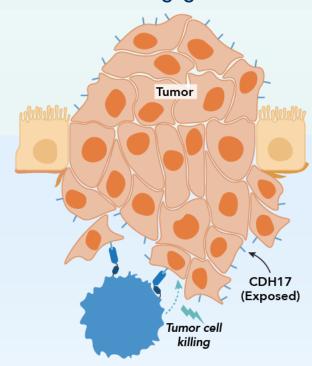
### CHM 2101 (CDH17 CAR T)

### CAR T CELLS UNABLE TO ACCESS NORMAL CELLS DUE TO TIGHT JUNCTIONS

#### **Normal Cells** In normal cells, CDH17 is The tight junctions between epithelial cells are thought to hidden beneath tight junctions prevent CAR T cells from accessing that reinforce the barriers of CDH17 on normal intestinal normal cells. epithelial cells.. CDH17 (Hidden) Intestinal epithelial cells Basement membrane Tight junction **CAR T** CAR Basal side of intestine

#### **Cancer Cells**

In cancer, loss of cell polarity causes CDH17 to be exposed on the tumor cell surface, enabling CAR T cells to engage and kill tumor cells.

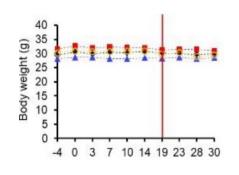


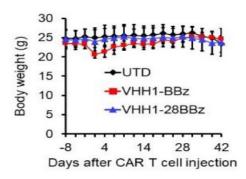


### **CHM 2101: SAFETY IN PRECLINICAL MODELS**

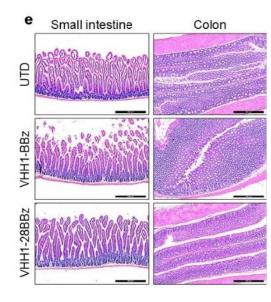
### DESPITE BROAD EXPRESSION OF CDH17 ON NORMAL INTESTINE, MICE TREATED WITH CHM 2101 DEMONSTARTED NO TREATMENT RELATED TOXICITY

#### No body weight loss



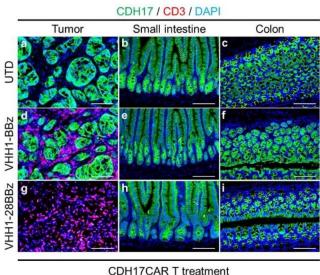


#### No damage to normal tissue



No tissue damage was also observed for stomach, pancreas, heart, liver, and kidney.

### No infiltration of normal intestine by CHM 2101 cells







### CHM 2101 (CDH17 CAR T)

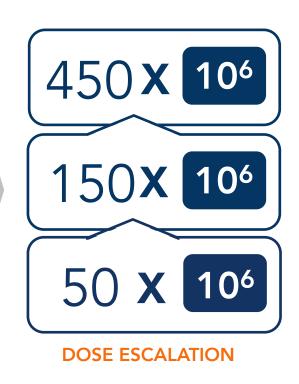
### PHASE 1A CLINICAL TRIAL IN RELAPSE / REFRACTORY GI CANCERS

Accelerated development through a Phase 1A basket trial design with patient eligibility in 3 tumour types

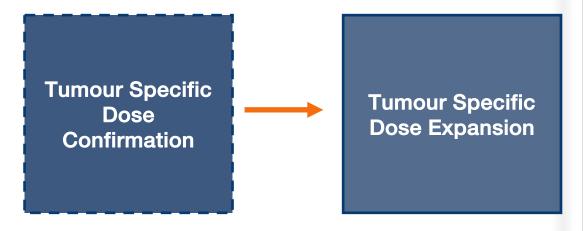
Colorectal Cancer

Neuroendocrine Tumours

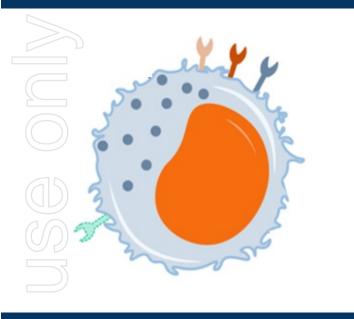
Gastric
Cancer



Upon signal confirmation, dose confirmation and expansion in tumour specific cohorts



### CHM 0201



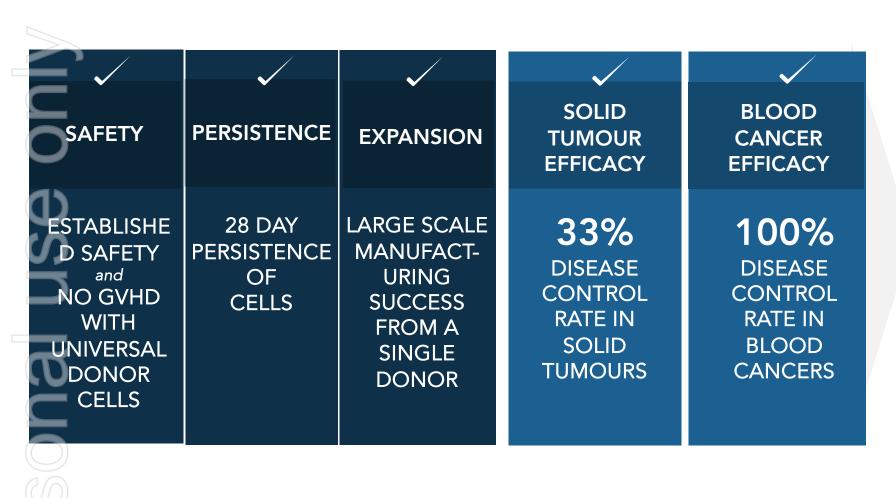
BEST-IN-CLASS NK CELL FOUNDATION A platform technology that can be leveraged to develop multiple new therapies

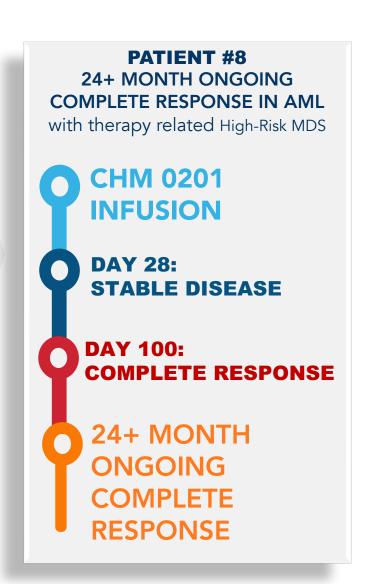
Broad applicability across 10+ disease areas including blood cancers and solid tumours

**Encouraging** Phase 1A **Clinical Trial Results** across multiple key endpoints in Acute Myeloid Leukemia and Colorectal Cancer

### CHM 0201: COMPLETED PHASE 1A

### ENCOURAGING CLINICAL OUTCOMES WITH A 24+ MONTH ONGOING COMPLETE RESPONSE





### CHM 0201 for BLOOD CANCERS AND SOLID TUMOURS ADVANCING NOVEL PHASE 1B CLINICAL COMBINATIONS



### CHM 0201 + VACTO Phase 1B

### PAUSED DUE TO STAFF RESOURCING

Acute Myeloid Leukemia and Colorectal Cancer

First Study to Investigate NK

Cells + Vactosertib





#### **ONGOING ENROLLMENT**

FDA IND Clearance

Acute Myeloid Leukemia

First NK Combination Study in Front Line AML

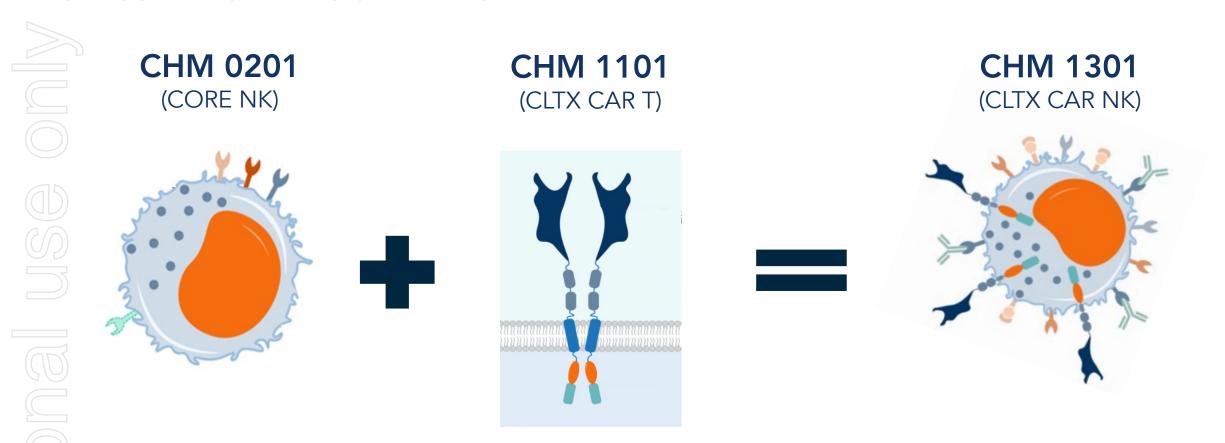


Making Cancer History®



### **CHM 1301: CLTX CAR NK DEVELOPMENT**

LEVERAGING PORTFOLIO SYNERGIES TO BUILD UPON A A BEST-IN-CLASS NK CELL FOUNDATION

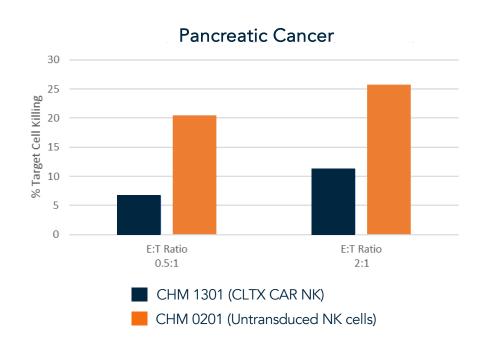


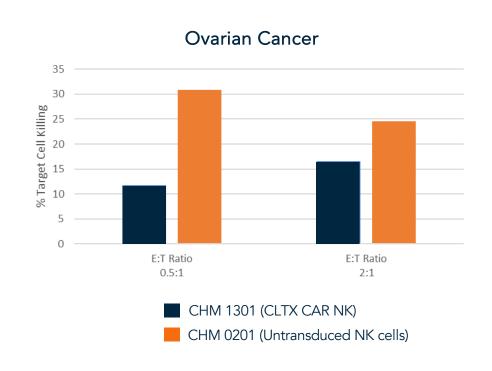


### CHM 1301: CLTX CAR NK

#### UP TO 300% ENHANCED CELL KILLING IN TWO NEW TUMOUR TYPES

CHM 1301 demonstrated ~2-3x increased potency against Ovarian and Pancreatic cancer cells when compared to CHM 0201







# CHIMERICTHERAPEUTICS VALUE PROPOSITION FOR INVESTORS

1.

### ADVANCED, UNDERVALUED PORTFOLIO

Chimeric's pipeline and portfolio are highly undervalued in relation to other cell therapy companies

2.

# ADVANCING DEVELOPMENT TO VALUE REALIZATION

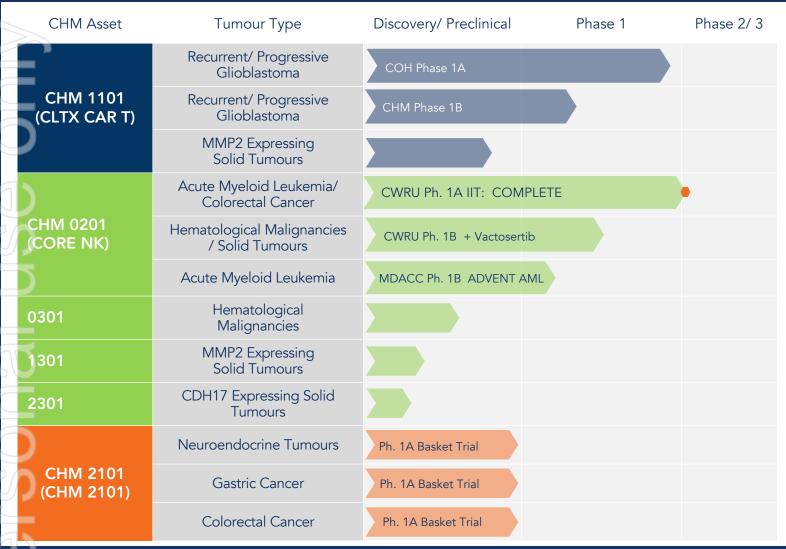
Chimeric's assets are entering Phase 1B, where big pharma M&A is now focused 3.

### NAVIGATING THE CHALLENGES

Chimeric is responding to todays biotech environment with program prioritization, cash preservation and business development

### 1. ADVANCED, UNDERVALUED PORTFOLIO

Chimeric has an industry leading cell therapy portfolio with novel technology platforms in clinical stage development





Novel technology platforms



Assets with positive clinical data



Ongoing clinical trials

MDACC: MD Anderson Cancer Centre/ COH: City of Hope / CWRU: Case Western Reserve University

### **CHIMERIC: INNOVATION UNDERVALUED IN TODAYS MARKET**

	CHIMERIC	THERAPEUTICS	Prescient Therapeutics Cell Therapy Pipeline Only	#carina biotech
Assets with Positive Clinical Data	2	No Clinical Data	No Clinical Data	No Clinical Data
Enrolling Clinical Trials	4	No Clinical Trials	No Clinical Trials	1
Patients Treated	25+	No Patients Treated	No Patients Treated	1
FDA IND's	6	No FDA IND's	No FDA IND's	1
Share Price/ Market Cap.	0.033 / \$24M As of Jan. 8/ 2024	<b>0.125 / \$114M</b> As of Jan. 8/ 2024	0.061/ \$49M As of Jan. 8/ 2024	Private / \$41M Valuation



### VALUE REALIZATION OPPORTUNITY FOCUSES ON PHASE 1B/ 2 CHIMERIC IS ADVANCING DEVELOPMENT TO BIG PHARMA'S FOCUS

In today's market, big pharma M&A has shifted its focus from preclinical assets to assets that are clinically de-risked with Phase1A/B data

As development advances to enhance M&A opportunity, funding requirements become more significant.

**PHASE** 1A/B **CLINICAL TRIALS** 

BIG **PHARMA** PHASE OF INTEREST

3-4 years

Establishing safety, efficacy and the recommended dose \$\$\$

**PHASE** 

2/3

**REGISTRATION** 

**TRIAL** 

Registration trial to support commercial approval

**REGULATORY APPROVAL** 

1 year

Approval by global regulatory authorities to commercialize

**ASSET** DISCOVERY

3-5 years

Discovery and lead optimisation of assets

1-2 years

PRE

**CLINICAL** 

In vitro and in vivo experiments with lead asset

3-5 years

\$\$\$\$\$

### 2. VALUE REALIZATION OPPORTUNITY FOCUSES ON PHASE 1B/ 2

#### RECENT M&A HIGHLIGHTS BIG PHARMA'S FOCUS ON CLINICALLY DE-RISKED ASSETS

Date	Acquirer / Investor	Licensee	Phase of Development	Total Deal Value
August 2023	astellas	POSEIDA THERAPEUTICS	1B	\$50M USD (equity investment)
July 2023	Pfizer	CARIBOU BIOSCIENCES®	1B	\$25M (equity investment)
May 2023	Janssen PHARMACEUTICAL COMPANIES OF Goldmon Goldmon	CBMG Cellular Biomedicine Group	1B / 2	\$245M USD+
October 2022	GILEAD	ARCELLX	2	\$225 USD



### 3. NAVIGATING THE CHALLENGES

To enable clinical advancement, Chimeric has responded to the challenges of todays market with program prioritization and cash preservation while creating value through business development





### CHIMERIC LEADERSHIP TEAM

### **GLOBAL EXPERTS IN CELL THERAPY DEVELOPMENT & COMMERCIALIZATION**

#### **EXPERIENCE**

Years of Cell Therapy Experience

#### **EXPERTISE**

Development Programs

### **PROVEN**

75 Of the FDA Approved CAR T Cell Therapies















### CELLULAR IMMUNOTHERAPY SCIENTIFIC ADVISORY BOARD

### WORLD RENOWNED SCIENTISTS AND CLINICIANS



Chair of the Cellular Therapeutics Cross Disciplinary Group, Mayo Clinic

Dr Lin is a pioneer in cellular immunotherapy having participated in many of the first in human CAR T cell therapy trials and multiple Ph. 2 cellular immunotherapy clinical trials.



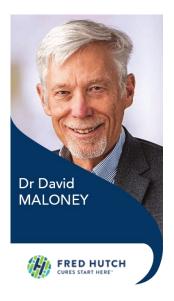
Director, the David and Etta Jones Center for Cellular Therapy, University of Chicago

Dr Bishop is a widely recognized as an expert in hematopoietic stem cell transplant and cellular therapy research and patient care, with a focus on leukemias and lymphomas.



Director of Translational Research, Immune Effector Cell Therapies Dana-Farber Cancer Institute

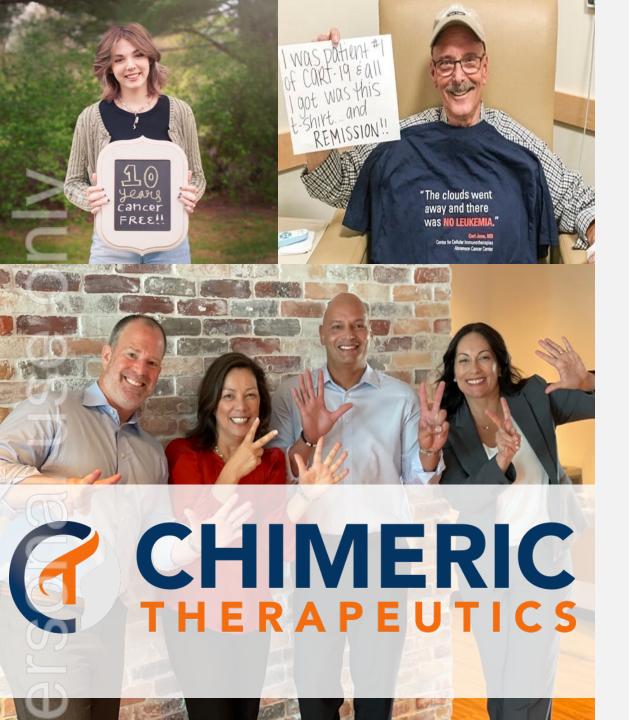
Dr Smith specializes in Cellular immunotherapy and hematological malignancies with a focus in multiple myeloma.



Medical Director, Cellular Immunotherapy and the Bezos Family Immunotherapy Clinic Fred Hutchinson Cancer Center

Dr Maloney has been a clinical investigator in over 15 cellular therapy clinical trials ranging from phase 1, first in human trials to commercially approved CAR T cell therapies.





- POSITIVE CLINICAL DATA in GBM and AML
  Two assets with positive Phase 1A clinical data
- ADVANCING CLINICAL DEVELOPMENT
  Four clinical trials in solid tumours and blood cancers in 2024
  - NEAR TERM MILESTONES
- Multiple clinical catalysts in next 12-18 months
- IMPACTFUL BUSINESS DEVELOPMENT
  Collaborative clinical trials with limited CHM
  funding with success in securing non-dilutive
  funding
- Experienced team with significant cell therapy development and commercialization expertise

#### **CONTACT INFORMATION**

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# SAPPENDIX.

### GLIOBLASTOMA (GBM)

#### MOST COMMON AND DEADLY OF ALL PRIMARY BRAIN TUMOURS

# WHAT IS GLIOBLASTOMA (GBM)?

Glioblastomas (also called GBM) are cancerous grade 4 brain tumours

Glioblastomas are fast growing and the most common malignant primary brain tumour

### WHO GETS GBM?

~1000 new cases of GBM are diagnosed in AUS each year<sup>1.</sup>

Average age at diagnosis is 62 years

Diagnosis is more common in males than females

# WHAT ARE THE RISK FACTORS FOR GBM?

The exact cause of GBM is unknown

The majority of patients have no family history or identifiable risk factors

# WHAT IS THE PROGNOSIS FOR GBM?

Without Treatment: ~3 months survival<sup>2</sup>.

With Treatment:
Less than 15 months<sup>3.</sup>

5-year Survival:
4.6% in Australia<sup>4.</sup>

Grochans, S.; Cybulska, A.M.; Simi 'nska, D.; Korbecki, J.; Kojder, K.; Chlubek, D.; Baranowska-Bosiacka, I. Epidemiology of Glioblastoma Multiforme–Literature Review. Cancers 2022, 14, 2412. https://doi.org/10.3390/cancers14102412



Jeffree R (2020) Australian Journal of General Practice. Volume 49, Issue 4, April 2020 "Current Management of Cerebral Gliomas"

<sup>2.</sup> Schapira AH (2007). Neurology and clinical neuroscience. Philadelphia: Mosby Elsevier. p. 1336. ISBN 978-0323070539.

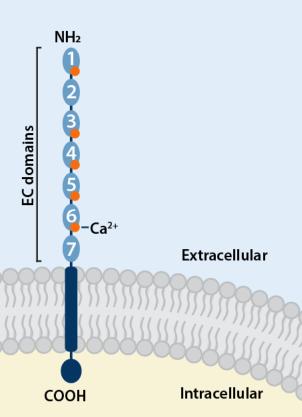
### CDH17: AN IDEAL TUMOR TARGETING ANTIGEN KEY TO CANCER PROLIFERATION

CDH17 (Cadherin-17) is an **oncogenic driver** of tumor formation and cancer metastasis, most specifically in gastrointestinal tumors.

Overexpression of CDH17 has been shown to be correlated with poor prognosis and the promotion of metastasis.

Inhibition of CDH17 has resulted in reduced proliferation and increased apoptosis of cancer cells

CDH17 is a member of the cadherin superfamily. Cadherins are calcium-dependent cell-cell adhesion molecules which play important roles in organ development, the maintenance of tissue integrity and cancer development.



Structural Features of Cadherin-17 (CDH17)

**CHIMERICTHERAPEUTICS**