

Commitment to a Cure

Corporate Presentation
May 2022

NASDAQ: CLLS

**EURONEXT GROWTH: ALCLS.PA** 



# **Forward-Looking Statements**

This presentation contains "forward-looking" statements within the meaning of applicable securities laws, including the Private Securities Litigation Reform Act of 1995. Forward-looking statements may be identified by words such as "at this time," "anticipate," "believe," "expect," "on track," "plan," "scheduled," and "will," or the negative of these and similar expressions.

These forward-looking statements, which are based on our management's current expectations and assumptions and on information currently available to management, including information provided or otherwise publicly reported by our licensed partners, include statements about our research and development projects and priorities, our pre-clinical project development efforts, the timing and progress of clinical trials (including with respect to patient enrollment and follow-up), the timing of our presentation of data, the adequacy of our supply of clinical vials, the timing of completion of construction of our Raleigh, North Carolina manufacturing facility, and operational capabilities at our manufacturing facilities, and the sufficiency of cash to fund operations.

These forward-looking statements are made in light of information currently available to us and are subject to numerous risks and uncertainties, including with respect to the numerous risks associated with biopharmaceutical product candidate development as well as the duration and severity of the COVID-19 pandemic and governmental and regulatory measures implemented in response to the evolving situation.

With respect to our cash runway, our operating plans, including product development plans, may change as a result of various factors, including factors currently unknown to us. Furthermore, many other important factors, including those described in our Annual Report on Form 20-F and the financial report (including the management report) for the year ended December 31, 2021 and subsequent filings Cellectis makes with the Securities Exchange Commission from time to time, as well as other known and unknown risks and uncertainties may adversely affect such forward-looking statements and cause our actual results, performance or achievements to be materially different from those expressed or implied by the forward-looking statements. Except as required by law, we assume no obligation to update these forwardlooking statements publicly, or to update the reasons why actual results could differ materially from those anticipated in the forwardlooking statements, even if new information becomes available in the future.

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## **Mission Statement**

# CELLECTIS IS A CLINICAL-STAGE BIOTECH

Using its Pioneering Gene-editing Platform

TO DEVELOP LIFE-SAVING CELL AND GENE THERAPIES



# **Key Highlights**

ongoing clinical trials



 40+ patients dosed in Cellectis-sponsored trials 150+ patients dosed

 In 5 trials sponsored by Cellectis' licensed partners







\$142M

Cash runway into 2024\*

\*Cash position include cash, cash equivalents and financial assets and restricted cash. Restricted cash was \$5 million as of March 31, 2022, of which \$0,5 million were classified as current financials assets



GMP facilities operational since mid-2021

 End-to-end manufacturing autonomy (buffers, DNA, mRNA, vectors to the final UCART product candidate) Meaningful Milestones
Expected Over the Next 12
Months

- Clinical data updates
- New IND with innovative therapy



# Allogeneic CAR T-Cell Pipeline





<sup>1</sup> ALLO-501 and ALLO-501A are exclusively licensed to Servier and under a joint clinical development program between Servier and Allogene. The ALPHA and ALPHA2 studies targets Diffuse Large B-Cell Lymphoma (DLBCL) and Follicular Lymphoma (FL) indications, which are subtypes of NHL 2 Phase 3 may not be required if Phase 2 is registrational.

<sup>3</sup> ALLO-715 and ALLO-605 target BCMA which is a licensed target from Cellectis. ALLO-715 and ALLO-605 utilize TALEN® gene-editing technology pioneered and owned by Cellectis. Allogene has an exclusive license to the Cellectis technology for allogeneic products directed at the BCMA target. Allogene holds global development and commercial rights for this investigational candidate.

<sup>4</sup> Allogene sponsored trial in combination with SpringWorks Therapeutics.

<sup>5</sup> ALLO-316 targets CD70 which is a licensed target from Cellectis. ALLO-316 utilize TALEN® gene-editing technology pioneered and owned by Cellectis. Allogene has an exclusive license to the Cellectis technology for allogeneic products directed at the CD70 target. Allogene holds global development and commercial rights for this investigational candidate.

# **Diversified Partnerships with Industry Leaders**



Research collaboration and exclusive 2020 worldwide license agreement to develop TILS Undisclosed Financials gene-edited TILs

Worldwide research collaboration and license agreement to develop gene-edited iPSC-derived NK and CAR-NK cells

allogeneic CAR T-cell targets

**iPSC-derived** NK

**BCMA** 

**CD70** 

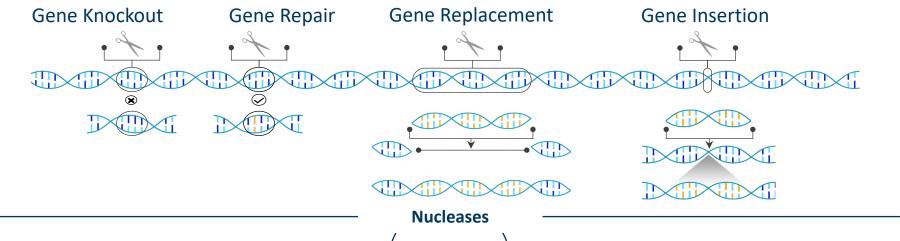
**Milestones** 

\$20M Upfront Convertible Note Up to \$805M in Development & Sales Milestones + Single-Digit Royalties on Sales

+ High Single-Digit Royalties on Sales



# **Powerful and Comprehensive Gene Editing Platform**

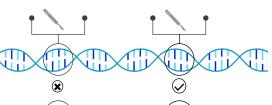


# **Experts in Gene Editing:**

- √ 30+ years for Cellectis' founders
- ✓ 22 years doing gene editing
- √ 10 years developing TALEN®



**Base editors** 



Gene Knockout



Gene Repair

# **Proprietary electroporation technology**



# Why TALEN®?

# **SAFE**

Avoids unintended genetic modifications

# **PRECISE**

Targets editing at the desired site (up to the base pair)



# **EFFICIENT**

High rates of gene editing for Knock Outs and Knock Ins



# Why TALEN®?

## **TALEN® CRISPR** In clinic since 2016 >> In clinic since 2018 170+ patients dosed **Every 7 base pairs Every 64 base pairs Euchromatin, heterochromatin Euchromatin only** >> 32 base pairs >> ~20 base pairs No chromotrypsis reported **Chromotrypsis reported** >> mRNA only **RNP**

**Scattered** 

**Strong for CLLS** 

**MATURITY** 

**PRECISION** 

**SAFETY** 

**GENOME OUTREACH** 

**RECOGNITION SITE** 

**VECTORIZATION** 

### Our Focus in 2022

# Generate clinical data from our 3 ongoing trials

To support determination of recommended Phase 2 dose (RP2D) and lymphodepletion for:

- UCART22 in r/r B-ALL patients
- UCART123 in r/r AML patients
- UCARTCS1 in r/r MM patients

# File IND & Initiate Phase 1 for

UCART20x22 in r/r NHL patients

# Manufacture (in-house) and release clinical batches of

- UCART22
- UCART20x22



# Cellectis Allogeneic CAR T-Cell Programs UCART Platform



# Allogeneic CAR T-cell Therapies are the Future

# **Scalable manufacturing**



Reduced cost:

1 batch = 100s doses

# "Off-the-shelf" availability



**Treatment** 

Decision

Allogenei UCART

Therapy

Immediately available to patients

# **Efficiency & robustness**



Potency and consistency may improve with healthy donor T-cells

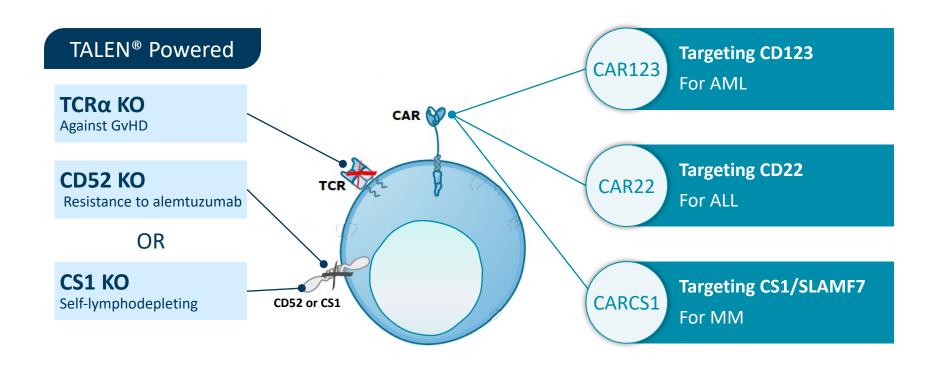
# **Market access**



Available to all patients irrespective of condition



# **Cellectis' Clinical-Stage Candidate UCART Products**





# **UCART22 – BALLI-01 Trial Design**

Open Label Dose-escalation and Dose-expansion Study to Evaluate the Safety, Expansion, Persistence and Clinical Activity of UCART22 in Patients with Relapsed or Refractory CD22+ B-cell Acute Lymphoblastic Leukemia

#### **Dose Escalation**

Determine MTD and/or RP2D

**Dose Expansion** 

LD regimen: FC or FCA

Up to 30 pts; mTPI design; 2-4 pts/cohort

Up to 53 pts; binomial exact study design; LD regimen: FC or FCA

#### **OBJECTIVES**

#### PRIMARY/SECONDARY:

- Safety & tolerability
- MTD/RP2D
- Response (NCCN criteria; investigator assessed)

#### **EXPLORATORY**

- UCART22 expansion and persistence, VCN and chimerism in WB and BM
- Immune reconstitution

# KEY ELIGIBILITY CRITERIA

- Patients aged 15 years to 70 years
- Adequate organ function
- ECOG PS ≤1
- B-ALL blast CD22 expression ≥70%
- Received ≥1 standard chemotherapy regimen and ≥1 salvage regimen

DOSE LEVELS		
DL-1	1 ×10⁴ cells/kg	
DL1	1 ×10 <sup>5</sup> cells/kg	
DL2	1 ×10 <sup>6</sup> cells/kg	
DL3	5 ×10 <sup>6</sup> cells/kg	
F: 30 mg/m2/d x4d; C: 1 g/m2/d x3 d; F: 30 mg/m2/d x3 d; C: 500 mg/m2/d x3 d A: 20 mg x3d		



# **UCART22 Administration Shows Encouraging Tolerable Safety Profile**

#### PATIENT CHARACTERISTICS (N=12)

Median age: 30 (20-61)

WHO classification:

B-ALL with recurrent genetic abnormalities:
 7 (58%);

CRFL2 rearrangement: 4 (33%)

Median prior lines of therapy: 3 (2-6)

Prior blinatumomab: 8 (73%)

Prior inotuzumab: 5 (45%)

Prior CD19 CART: 3 (27%)

#### **SAFETY: FCA Cohorts (N=6)**

- 0 dose limiting toxicity
- 0 ICANS (immune effector cell associated neurotoxicity)
- 0 severe UCART22-related TEAEs (treatment emergent adverse events)
- 3 patients with mild to moderate CRS (cytokine release syndrome)
- 1 patient with GII GvHD; skin only\*

\*not confirmed by biopsy; in context of re-activation of prior allogeneic bone marrow donor

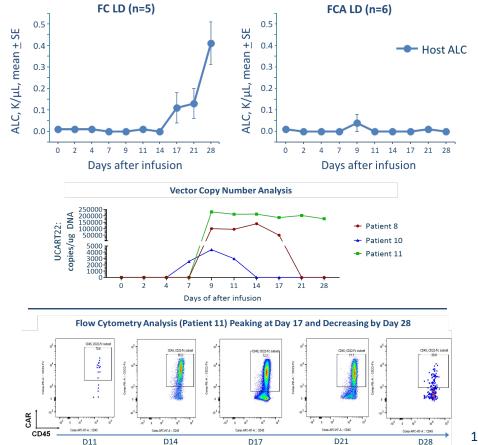


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# **UCART22** Expansion Associated with Encouraging Anti-leukemic Activity

#### **EFFICACY: FCA Cohorts (N=6)**

- Host lymphocytes on average remained suppressed
- UCART22 expansion was observed and was associated with anti-leukemic activity
- 2/6 patients achieved blast reductions to < 5% by day 28
  - 1 pt in DL2: 0.4% BM blast
  - 1 pt in DL2i: 0% BM blast



# UCARTCS1 - MELANI-01 Study Schema

Phase I, Open Label Dose-Escalation Study to Evaluate the Safety, Expansion, Persistence and Clinical Activity of UCARTCS1, Administered in Patients with Relapsed or Refractory Multiple Myeloma



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#### **Primary and Secondary**

- Safety and tolerability
- MTD and Efficacy

#### **Exploratory Objectives**

- CS1 expression on MM cells
- UCARTCS1 expansion and persistence
- Changes in serum biomarkers; immune cell reconstitution

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- Patients with confirmed MM (IMWG criteria) relapsed after prior MM therapy
- ECOG PS <2</li>
- No prior investigational drug or CAR therapy targeting CS1
- Adequate organ function

DOSE LEVELS		
DL-1	3 ×10⁵ cells/kg	
DL1	1×10 <sup>6</sup> cells/kg	
DL2	3 ×10 <sup>6</sup> cells/kg	
DL3	9 ×10 <sup>6</sup> cells/kg	
F: 30 mg/m2/d x 4d; C: 1 g/m2/d x3 d		

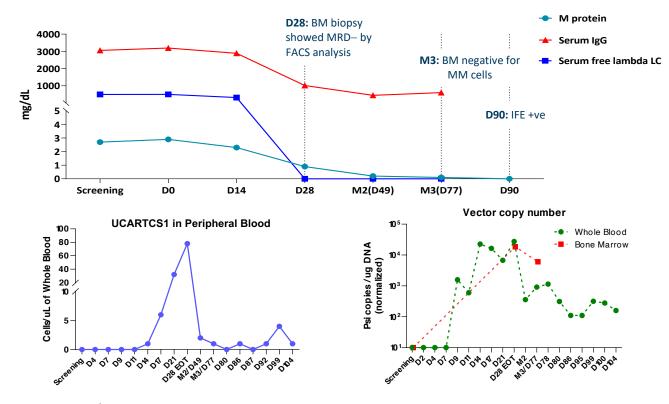


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\*Lymphodepletion: Fludarabine 30mg/m2/day, Day -5 to -2; Cyclophosphamide  $1g/m^2/day$ , Day -4 to -2.

# Preliminary Data Validate CS1 as a Target for CAR T in Multiple Myeloma

DL1 patient MRD-Neg VGPR





# UCART123 – AMELI-01 Trial Design

Phase I, Open Label Dose-escalation and Dose-expansion Study to Evaluate the Safety, Expansion, Persistence and Clinical Activity of UCART123 in Patients with Relapsed or Refractory Acute Myeloid Leukemia

#### **Dose Escalation**

**Determine MTD** and/or RP2D

## **Dose Expansion**

Up to 28 pts; mTPI design; 2-4 pts/cohort

18-37 pts; Simon's two-stage design

#### **OBJECTIVES**

#### **Primary and Secondary**

- Safety & tolerability
- Establish MTD and identify RP2D
- Efficacy

#### **Exploratory Objectives**

- UCART123 expansion, trafficking, and persistence
- Profile cytokine, chemokine, growth factor, and C-reactive protein levels postinfusion

#### **KEY ELIGIBILITY CRITERIA**

- Patients with relapsed or primary refractory AML (>5% bone marrow blasts)
- Patients with CD123+ blast cells
- PS of ≤1 and adequate organ function
- Identified donor and transplant strategy prior to LD (dose-escalation)

DOSE LEVELS		
DL-1	1.25×10 <sup>5</sup> cells/kg	
DL1	2.5×10 <sup>5</sup> cells/kg	
DL2	6.25×10 <sup>5</sup> cells/kg	
DL3	3.30×10 <sup>6</sup> cells/kg	
DL4	5.05×10 <sup>6</sup> cells/kg	
F: 30 mg/m2/d x 4d; C: 750 g/m2/d x 3d;		

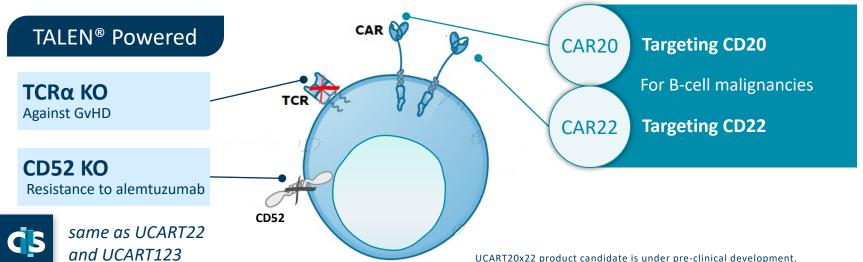
F: 30 mg/m2/d x 4d; C: 750 g/m2/d x3 d; A: 12 mg/d x4d



# UCART20x22 - A Dual Allogeneic CAR T-cells for B-cell Malignancies

# Why UCART20x22?

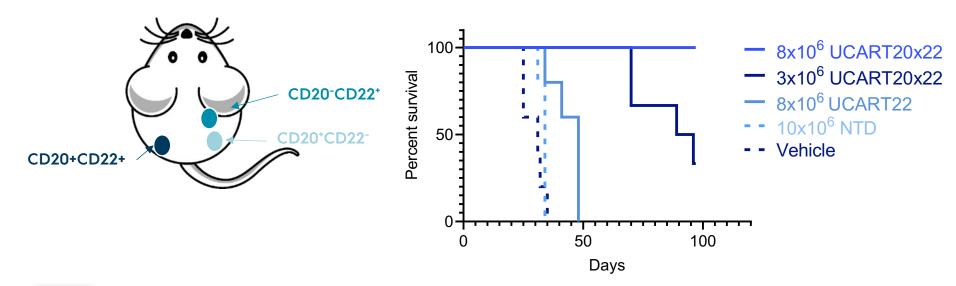
- Strong alternative to CD19 (highly competitive/crowded)
- CD22 and CD20 are validated targets in B-cell malignancies
- Dual targeting designed for better killing & to prevent target escape
- Strong in vitro and in vivo preclinical results & fast to develop



# UCART20x22 - Efficient Activity in Vivo Against Multiple Target Combinations

Single mouse carrying subcutaneous lymphoma tumors expressing different antigen combinations

Efficient *in vivo* clearance of tumors expressing one or two antigens (CD20 and/or CD22) in a dose dependent manner, starting at low dose





# **Cellectis' UCART Platform Takeaways**

# 170+ patients administered UCART derived from Cellectis' technology

- **1. GVHD:** TCR $\alpha$  KO results in safe, non-alloreactive UCART
- 2. Engraftment: CD52 KO + alemtuzumab provides a safe, effective & controllable therapeutic window
- 3. Persistence: Redosing feasible; encouraging results in enhanced activity in both NHL and ALL
- **4. Safety:** Profile on par with approved autologous CAR T therapies
- **5. Efficacy:** Anti-tumor activity consistent with autologous products



# CELLECTIS ORGANIZATION



# From UCART Discovery to Patients' Bedside





Paris, France

HQ, PD/AD, Starting Materials

# 55,000 sq ft. facility

- ✓ Process & analytical development
- Raw materials manufacturing
- ✓ QC labs
- Warehouse
- ✓ Cryogenic Storage rooms



# Raleigh, North Carolina

UCART – Clinical & potential for Commercial

# 82,000 sq ft. facility

- ✓ Cell therapy GMP manufacturing
- QC labs
- Warehouse
- ✓ Cryogenic Storage rooms



# New York, New York

Innovation, Clinical Development

# 25,000 sq ft. facility

- ✓ Gene Editing platform TALEN®
- √ I/O discovery platform
- Gene therapy discovery platform
- Clinical development



# **Cellectis 2022 Expected Milestones**

## UCART22

Enroll DL3 with FCA preconditioning and start dosing with products manufactured in-house

# **UCART123**

Enroll DL2 and DL2i with FCA preconditioning

# **UCARTCS1**

Enroll DL1 with FC preconditioning

## UCART20x22

File IND and initiate Phase 1 trial in patients with R/R NHL

# **GMP FACILITIES**

Release batches of UCART22 and UCART20x22

## **PARTNERS**

Updates from licensed partners Servier, Allogene, lovance and Cytovia



# **Key Takeaways – Why Cellectis?**



# **Innovative Allogeneic CAR-T**

Breaking Paradigms with Life-Saving Therapies



# **End-to-end In-house Manufacturing**

Owning Manufacturing is Owning the Product



**Best-In-Class Gene Editing Platform** 

Safe, Powerful & Comprehensive



# **Strong Partnerships**

Diversified Indications Leading to Financial Upsides



# **THANK YOU**

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