



### Legal Disclaimers

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Caution should be exercised when interpreting results from separate trials involving separate product candidates. There are differences in the clinical trial design, patient populations, and the product candidates themselves, and the results from the clinical trials of autologous products may have no interpretative value on our existing or future results.

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# Allogene: Singular Focus on Allogeneic Cell Therapy

### SUCCESSFUL TRACK RECORD OF EXECUTION

2018 Allogene Launch

Manufacturing Experience

Clinical Trial Execution

Breaking Down
Allogeneic
Hurdles

Demonstrating Efficacy

Pipeline 2021 Progress













From launch to leadership in less than 3 years

Raised \$1.5B in aggregate capital

Delivered uninterrupted supply for all clinical trials

Newark facility on track to begin production in 2021

Leading the field with 75+ patients treated with AlloCAR T™

2 key Phase 1 datasets presented in 2020 at major medical meetings Controlling Graft vs Host Disease (GvHD) and cell rejection

Creating a window of cell persistence

Complete Responses and Minimal Residual Disease (MRD) responses observed

Proof-of-Concept for proprietary lymphodepletion platform

5 clinical trials planned in 2021, including:

1<sup>st</sup> pivotal trial 1<sup>st</sup> solid tumor trial 1<sup>st</sup> combo trial 1<sup>st</sup> TurboCAR™ trial



# Deep AlloCAR T™ Pipeline Targeting Vast Array of Tumor Types

CATEGORY		PROGRAM	PRE-CLINICAL	PHASE 1	PHASE 2/3 <sup>2</sup>
	CD19	ALLO-501 (NHL) <sup>1</sup>			
	C	ALLO-501A (NHL) <sup>1</sup>			
	⋖	ALLO-715 (MM)			
Hematological Malignancies	BCMA	ALLO-715 + nirogacestat (MM) <sup>3</sup>			
		ALLO-605 (TurboCAR™/MM)			
		ALLO-316 (CD70/AML)			
		ALLO-819 (FLT3/AML)			
		ALLO-316 (CD70/RCC)			
Solid Tumors		DLL3 (SCLC)			
		10 Undisclosed Targets			
Lymphodepletion Agent		ALLO-647 (Anti-CD52 mAb) <sup>4</sup>			

<sup>&</sup>lt;sup>1</sup> Servier holds ex-US commercial rights



<sup>&</sup>lt;sup>2</sup> Phase 3 may not be required if Phase 2 is registrational; Initiation for ALLO-501A Phase 2 trial expected 2H 2021

<sup>&</sup>lt;sup>3</sup> Allogene Sponsored trial in combination with SpringWorks Therapeutics

<sup>&</sup>lt;sup>4</sup> ALLO-647 intended to enable expansion and persistence of allogeneic CAR T product candidates

# Innovating CAR T Therapies to Potentially Expand Access & Reduce Cost



### Cost

- Scalable and efficient manufacturing
- Potential to treat 100+ patients from a single manufacturing run
- Opportunity to reduce ancillary cost of care associated with autologous therapy



### **Innovation**

- Multiplex gene engineering/editing capabilities
- Opportunity for product optimization



### Access

- Potential to treat all eligible patients
- Repeat dosing, if needed
- No need for complex logistics or bridging therapy

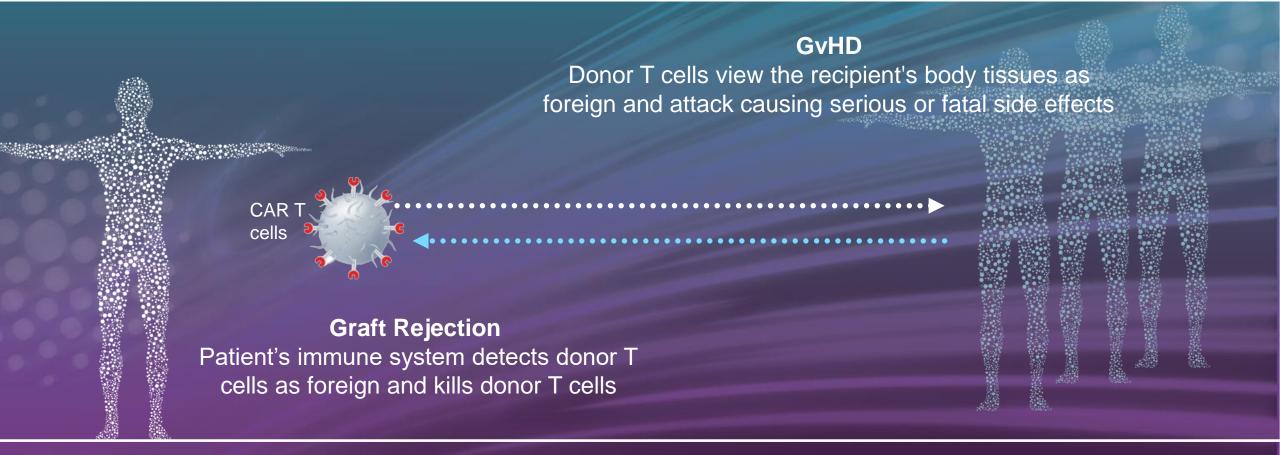


### Speed/Reliability

- "Off the shelf" for on demand treatment
- Less product variability, made from healthy T cells



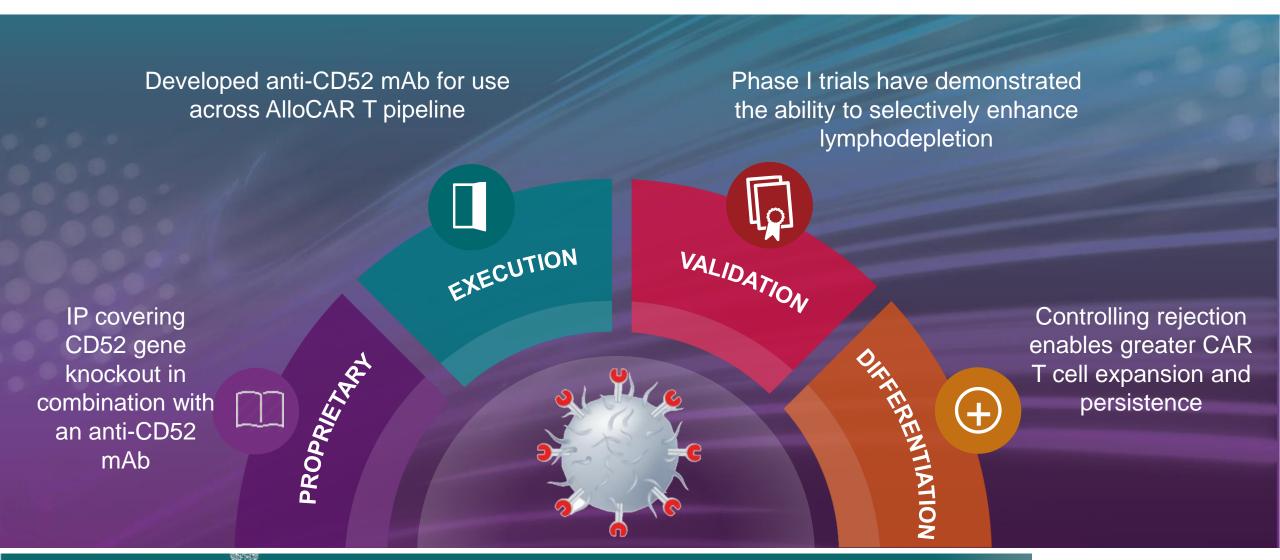
# Defying Immunity: Overcoming GvHD and Graft Rejection



Allogeneic cell therapy engages the fundamental immunological process of Self vs. Non-Self recognition



# ALLO-647 Anchors Novel, Proprietary Approach to Overcoming Rejection





# Industrializing Allogeneic Cell Therapy Production: Strategy





### Industrializing Allogeneic Cell Therapy Production: Infrastructure





### Cell Forge 1 (Newark, CA)

- New state-of-the-art facility
- Designed for clinical and commercial manufacturing, analytical testing and distribution of cell therapies
- Construction complete in 2020, first GMP production 2021

#### **South San Francisco Facilities**

- Manufacturing process and product development
- Analytical methods for process and product understanding and release
- Quality Assurance and Quality Control

#### **External Network**

- Broad CMO and supplier network
- Incorporating external expertise for starting materials, drug substance and drug product manufacturing
- Packaging, labeling, logistics and clinical distribution



# CD19 Program: Advancing ALLO-501A to Potential Pivotal Study Potential to be the First Allogeneic CAR T in Phase 2

ALLO-501 ALPHA Ph1 Trial | Initial Data: ASCO 2020 | Next Update Expected Q2 2021 (ASCO 2021)

### Early efficacy competitive with autologous CAR

T: 83% ORR and 67% CR with higher dose ALLO-647 (N=6)

**Well tolerated**: No GvHD and manageable CRS. Early safety data compared favorably to autologous CAR T

On demand dosing: 5 days from enrollment to treatment vs. 17-54 days for autologous therapies

**Biomarker validation**: Correlations observed with ALLO-647 lymphodepletion, ALLO-501 cell expansion, and tumor response

Further exploration of dosing: Retreatment and consolidated dosing to potentially optimize outcomes

Cell Dose and LD regimen	ALLO-501 <sup>1</sup> ALLO-647 90mg (N=6)	Autologous Ph1 Trials in NHL <sup>2</sup>	Autologous Ph2 Trials in NHL <sup>3</sup>
ORR, n (%)	5 (83%)	64-80%	50-73%
CR, n (%)	4 (67%)	56-60%	32-53%

<sup>&</sup>lt;sup>1</sup> ASCO June 2020; Autologous CAR T naïve patients

<sup>3</sup> Yescarta, Kymriah FDA labeling information and Abramson ASH 2019; Based upon mITT analyses

AE of Interest (≧Gr3)	ALLO-501 Ph1 (N=22)	axi-cel Ph2* (N=101)	tisa-cel Ph2* (N=111)	liso-cel Ph2* (N=269)
Cytokine Release Syndrome	5%	13%	23%	2%
ICANS	-	7///////////	7///////////	
Neurologic Events	///////////////////////////////////////	31%	18%	10%
<b>Graft-versus-Host Disease</b>	-	-	-	-
Infection	9%	23%	25%	12%
Neutropenia	64%	93%	81%	60%
Infusion Reaction	5%**	-	-	-

<sup>\*</sup> Neelapu NEJM 2017, Schuster NEJM 2019, and Abramson ASH 2019

ALLO-501A ALPHA2 Ph1 Trial | Initial Data: Expected Q2 2021 | Anticipate Pivotal Trial Initiation: 2H 2021



<sup>&</sup>lt;sup>2</sup> Kymriah and liso-cel trials include FL and MCL patients; ASH 2015; Schuster, NEJM, 2019; Abramson, ASH 2019

<sup>\*\*</sup> Attributed to ALLO-647

# ALPHA2 Study Design and Endpoints

Phase 1/2, Open-label, Multicenter Dose Escalation and Dose Expansion Study

### **Primary Endpoints**

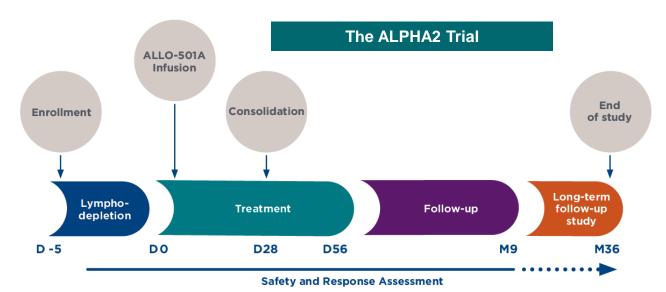
- Safety and dose-limiting toxicity of ALLO-647/Flu/Cy followed by ALLO-501A
- Overall response rate by central imaging review

### **Key Secondary Endpoints**

- Overall response rate by investigator assessment
- ALLO-501A cell kinetics
- ALLO-647 PK

### **Key Eligibility Criteria**

- R/R LBCL
- At least 2 prior lines of therapy, including an anthracycline and anti-CD20 monoclonal antibody
- ECOG 0 or 1
- Prior autologous CAR T allowed if tumor remains CD19+ and patient had a CR ≥ 16 weeks
- Patients with Donor Specific Antibodies are excluded



	DL1 (N=1)	DL2	DL2 Consolidation	DL3
Cell	40 x 10 <sup>6</sup>	120 x 10 <sup>6</sup>	120 x 10 <sup>6</sup> CAR+ T on D0, and D28 for SD or better	360 x 10 <sup>6</sup>
Dose	CAR <sup>+</sup> T cells	CAR+ T cells		CAR <sup>+</sup> T cells

- Lymphodepletion Regimen
  - Fludarabine (Flu), Cyclophosphamide (Cy) and ALLO-647



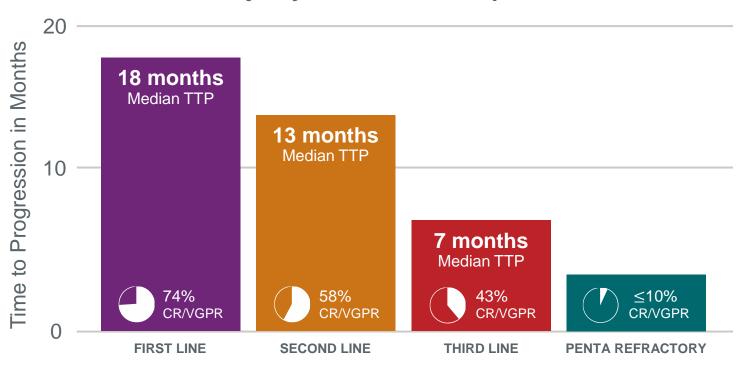
# Why Allogeneic Cell Therapy Matters in Multiple Myeloma



- Multiple myeloma is a progressive disease. Prognosis for patients worsens over time
- Bridging therapy to "control" the disease may increase some cumulative or synergistic toxicities for the patients<sup>2</sup>

Time is of the essence for patients with rapid progression

### Majority of Patients Relapse<sup>1</sup>



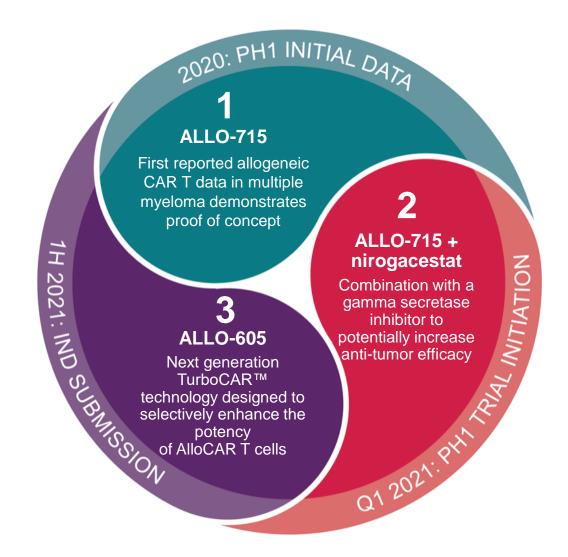
<sup>&</sup>lt;sup>3</sup> Gandhi, et al., Leukemia. 2019 September; 33(9): 2266–2275. doi:10.1038/s41375-019-0435-7; TTP based upon conditional mPFS reported, VGPR based on interpolated values



<sup>&</sup>lt;sup>1</sup>Bird SA, Boyd K. Palliat Care Soc Pract. 2019;13:1-13

<sup>&</sup>lt;sup>2</sup>Zheng Ping-Pin, et al. Drug Discovery Today June 2018; 23:6; 1175-82

# Building an Anti-BCMA AlloCAR T Franchise in Multiple Myeloma





# ALLO-715: First AlloCAR T To Demonstrate Feasibility in Myeloma

ALLO-715 UNIVERSAL Ph1 Trial: *Initial Data Readout: ASH 2020*Next Steps: *Explore further dose escalation and combination with nirogacestat* 

### **Clear benefits associated with an off-the-shelf therapy:**

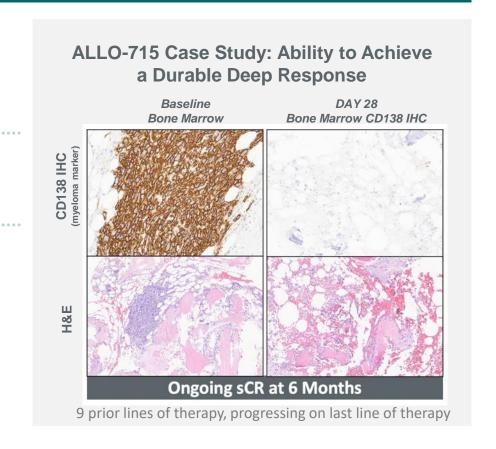
- ~90% of patients treated within 5 days of study enrollment
- Obviates need for bridging therapy prior to dosing

#### Well tolerated across dose levels:

- No GVHD or neurotoxicity (ICANS); manageable grade 1 or 2 CRS
- Infection rate on par with other studies in advanced myeloma

# Dose dependent ALLO-715 activity observed in heavily pretreated, refractory patients

- ALLO-715 cell persistence observed through month 4
- 320M cell dose of ALLO-715 (DL3) with FCA lymphodepletion associated with a 60% Overall Response Rate (ORR)
- 5 of 6 VGPR+ patients assessed for MRD status; all were negative



ASH 2020; image on file



### ALLO-715: Initial Data Creates Pathway for Allogeneic CAR T in MM

### **Initial Safety Compared to BCMA Directed Therapies**

	ALLO-715 Ph1 (N=31) <sup>1</sup>	Ide-CeI 300/450M N=128 <sup>2</sup>	Orva-Cel 300/450/600M N=62 <sup>3</sup>	Cilta-Cel 0.75M/kg N=29 <sup>4</sup>
Cytokine Release Syndrome (CRS)	45%	84%	89%	93%
CRS (Grade ≥3)	0	5.5%	3%	7%
Neurologic Toxicity	0	18%	13%	10%
Neurologic Toxicity (Grade ≥3)*	0	3%	3%	3%
Infection (Grade ≥3)	16%	NR	13%	21%
Neutropenia (Grade ≥3)	52%	89%	90%	100%
Death from AEs	3%	3%	3%	8%

1 ASH 2020; 2 Munshi, ASCO 2020 (Ide-cel); 3 Mailankody, ASCO 2020 (Orva-cel); 4 Madduri, ASH 2020 Abstract

### **Initial Responses Compared to BCMA Directed Therapies**

Cell Dose & LD regimen	ALLO-715 320M & FCA (N=10) <sup>1</sup>	Ide-Cel (BB/BMS) 300/450M N=124 <sup>2</sup>	Orva-Cel (Juno/BMS) 300/450/600M N=62 <sup>3</sup>	Cilta-Cel (JNJ) 0.75M/kg N=97 <sup>4</sup>
ORR, %	60%	73%	92%	95%
VGPR+ Rate, %	40%	53%	68%	88%
MRD- Rate, % (N Evaluated)	100% (4/4)	78% (80/102)	84% (21/25)	94% (49/52)

1 ASH 2020; Responses included 2 subjects with only day 14 assessment and 1 subject who converted from a confirmed PR to VGPR (pending confirmation).; 2 Munshi, ASCO 2020 (Ide-cel); 3 Mailankody, ASCO 2020 (Orva-cel); 4 Madduri, ASH 2020 Abstract



# UNIVERSAL: ALLO-715 + Nirogacestat Cohort

#### **Primary Endpoints**

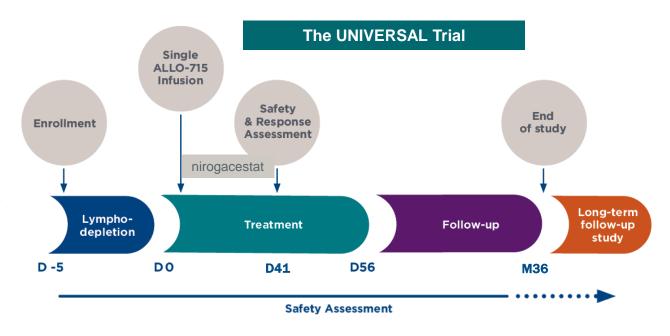
 Safety and tolerability of ALLO-715 in combination with nirogacestat

#### **Secondary Endpoints**

- Anti-tumor activity and cellular kinetics of ALLO-715 in combination with nirogacestat
- ALLO-647 and nirogacestat pharmacokinetics
- Evaluate the expression of BCMA in bone marrow plasma cells with and without nirogacestat

#### **Key Eligibility Criteria**

- Relapsed/Refractory Multiple Myeloma
- ≥ 3 prior therapies including IMiD, PI & anti-CD38
- Refractory to last prior therapy
- ECOG 0 or 1
- No donor-specific antibodies
- No bridging therapy allowed



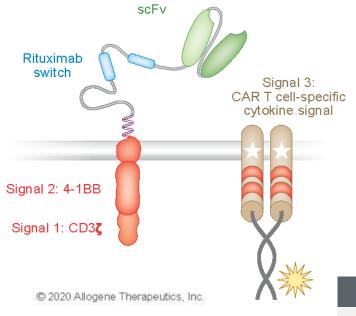
ALLO-715 Dose Escalation: 320 or 480 x 10<sup>6</sup> CAR<sup>+</sup> T cells

IMiD: immunomodulatory imade drug

PI: proteosome inhibitors

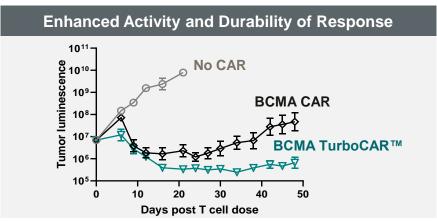


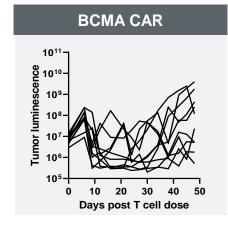
# ALLO-605: First TurboCAR™ Investigational Candidate

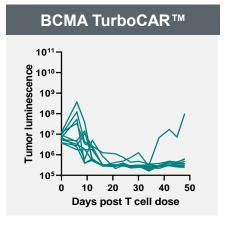


#### ALLO-605 IND Planned in 1H 2021 for the IGNITE Trial

- TurboCAR™ is designed to recapitulate cytokine signaling selectively in CAR T cells
  - Does not stimulate host immune cells which could cause systemic toxicity or reject CAR
  - Delivers survival benefit selectively to CAR T cells
- · Opportunities for development include:
- Improving the efficacy of CAR T cells
- Reducing CAR T cell dose requirement
- Overcoming exhaustion to enable CAR T therapies for solid tumors
- Improved Engraftment and Persistence, and Delayed Exhaustion seen in preclinical studies









# Translating CAR T Success in Hematologic Cancers to Solid Tumors

### **2020 American Cancer Society Statistics**

	Heme Malignancies	Solid Tumors
Incidence	179,000	1,600,000
Deaths	57,000	504,000

### **Worldwide Market for Oncology Drugs in 2018\***

- All drug spend = \$1.2 trillion
- Total cancer drug spend ≈ \$150 billion
- Hematologic cancer drugs ≈ \$31.3 billion

\*IQVIA

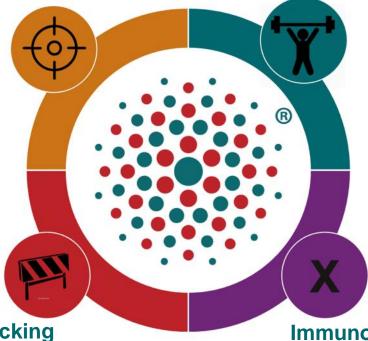
Significant opportunity to expand benefits of CAR T therapy into largest area of unmet need

### **Target Selection/Validation**

- CAR optimization
- Multi-targeting CARs

#### **T Cell Fitness**

- CAR signaling/ TurboCARs™
- Manufacturing improvements



- **Tumor Trafficking**
- Combinations
- CAR T engineering

### Immunosuppressive TME

- Next generation TurboCARs™
- Enhanced/flexible lymphodepletion
- CAR T cell doses, frequencies and administration of cells



# ALLO-316: Investigating an AlloCAR T™ in Renal Cell Carcinoma

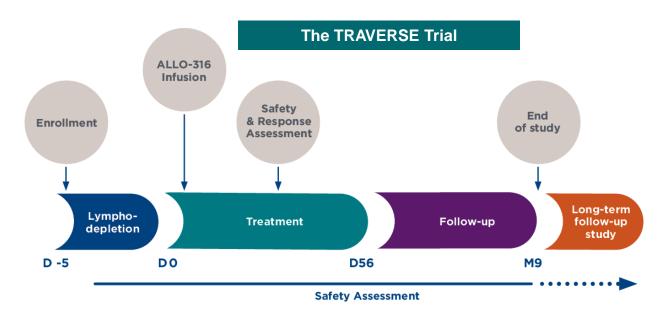
First of Several Solid Tumor Candidates Planned for Clinical Development

# CD70 target selectively expressed in several cancers<sup>1</sup>:

- RCC (80-100% of tumors)
  - High prevalence with limited 'off tumor' expression
- AML (96% of tumors)

#### IND cleared for anti-CD70 candidate ALLO-316:

- ALLO-316 is associated with minimal or no fratricide
- Phase I TRAVERSE trial in RCC to begin in 1H 2021
  - Primary endpoints: Safety and tolerability
  - Secondary endpoints: Anti-tumor efficacy, PK/PD
- Potential second indication in AML targeted for 2021/2022



	DL1	DL2	DL3	DL4
Cell Dose	40 x 10 <sup>6</sup>	120 x 10 <sup>6</sup>	360 x 10 <sup>6</sup>	480 x 10 <sup>6</sup>
	CAR+ T cells	CAR+ T cells	CAR+ T cells	CAR+ T cells



<sup>&</sup>lt;sup>1</sup> Expert Opin Ther Targets. 2008 Mar; 12(3): 341–351. doi: 10.1517/14728222.12.3.341

# Partnerships: Accelerating Development and Positioning for the Future



Global development partner for CD19 with ex-US commercialization rights



Established Allogene Overland
Biopharm joint venture to
develop and commercialize
AlloCAR T™ cell therapies in
greater China





Induced pluripotent stem cells (iPSC)





Preclinical and clinical investigation of AlloCAR T candidates across Allogene's broad portfolio of hematologic and solid tumors



Clinical collaboration to evaluate ALLO-715 in combination with Nirogacestat

**GLOBAL EXPANSION** 

**TECHNOLOGIES** 

RESEARCH



# Allogene Overland Joint Venture: Expanding into Greater China Opportunity to Accelerate Global Development of AlloCAR T Therapies



- Leadership in Allogeneic CAR T
- AlloCAR T assets
  - BCMA
  - CD70
  - FLT3
  - DLL3
- Manufacturing expertise



### **Allogene Overland Biopharm**

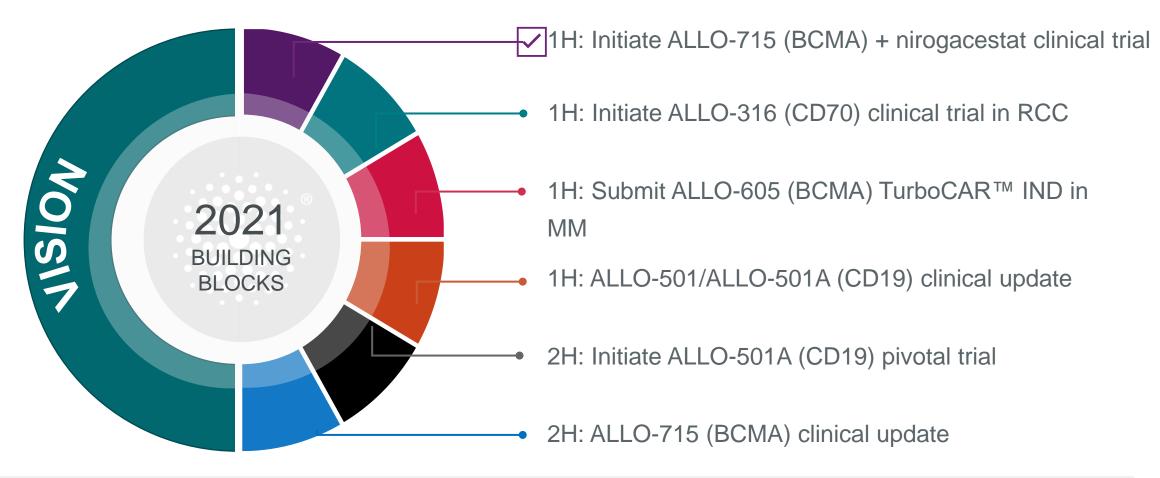
- First-of-its-kind collaboration in allogeneic cell therapy
- Rights to develop AlloCAR T therapies for four targets in Greater China, Taiwan, South Korea and Singapore
- Milestones to Allogene and tiered royalties on net sales



- Backed by Hillhouse Capital
- \$117 million in non-dilutive capital:
  - \$40 million upfront to ALLO
- \$77 million to support joint venture operations



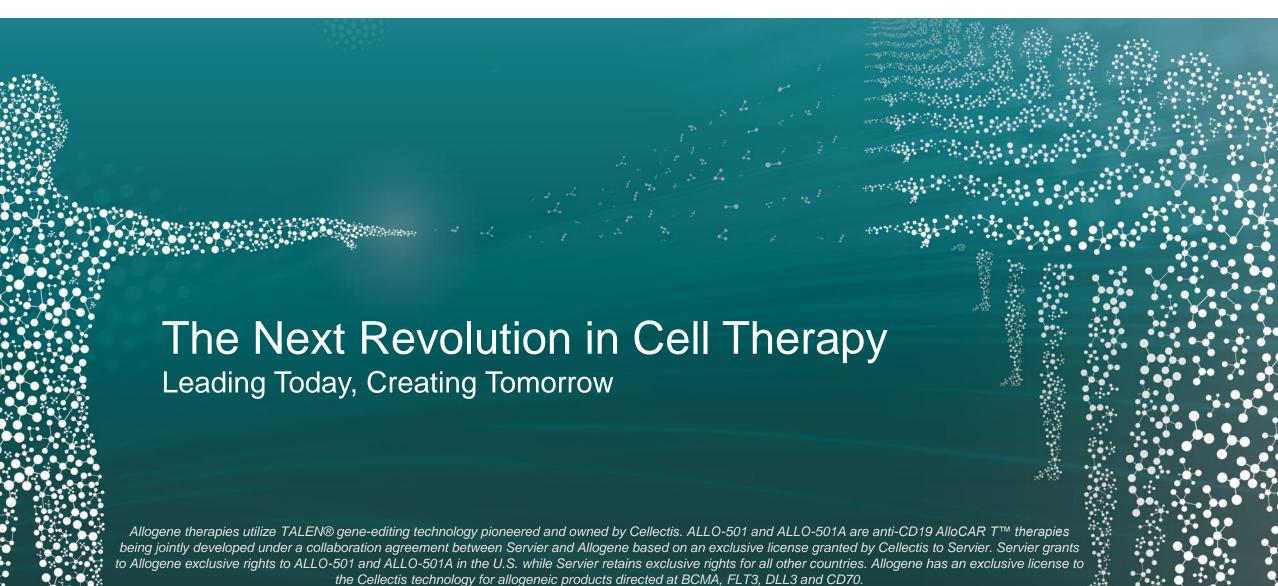
### 2021 Building Blocks to the Allogene Vision



Create and lead the next revolution in cancer treatment by delivering to patients the first AlloCAR  $T^{TM}$  therapies for blood cancers and solid tumors.







# ALLO-501 ALPHA Study (NCT03939026) Design and Endpoints Phase 1, Open-label, Multicenter Dose Escalation Study

### **Primary Endpoints**

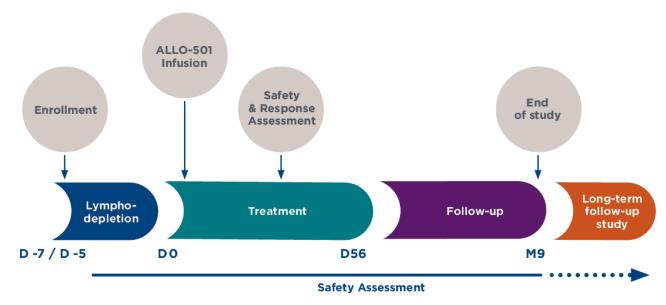
 Safety and dose-limiting toxicity of ALLO-647/Flu/Cy followed by ALLO-501

### **Key Secondary Endpoints**

- Overall response rate
- ALLO-501 cell kinetics
- ALLO-647 PK

### **Key Eligibility Criteria**

- R/R LBCL or FL
- At least 2 prior lines of therapy, including an anti-CD20 monoclonal antibody
- ECOG 0 or 1
- Prior autologous CAR T allowed if tumor remains CD19+
- Patients with Donor Specific Antibodies and rituximab > 15ng/ml were excluded



	DL1	DL2	DL3
Cell Dose	40 x 10 <sup>6</sup>	120 x 10 <sup>6</sup>	360 x 10 <sup>6</sup>
	CAR+ T cells	CAR+ T cells	CAR+ T cells

- Lymphodepletion Regimens
  - LD1: Flu/Cy and ALLO-647 13 mg/d x 3 days
  - LD2/LD3: Flu/Cy and ALLO-647 30 mg/d x 3 days (concomitant/staggered)

Fludarabine (Flu): 30 mg/m2/d x 3 days Cyclophosphamide (Cy): 300 mg/m2/d x 3 days



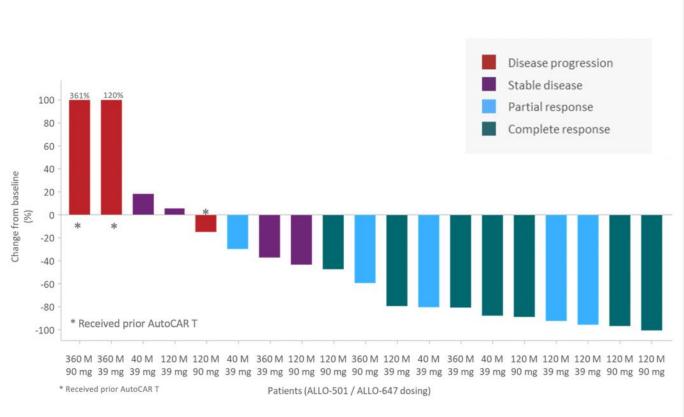
# Phase 1 ALPHA Best Overall Response

Cell Dose	39	mg ALLO-64	7	ALL 39mg	90mg A	LLO-647	All 90mg	All Patients
and LD regimen	40 x 10 <sup>6</sup> CAR <sup>+</sup> cells (N=4)	120 x 10 <sup>6</sup> CAR <sup>+</sup> cells (N=4)	360 x 10 <sup>6</sup> CAR <sup>+</sup> cells (N=3)	ALLO-647 (N = 11)	120 x 10 <sup>6</sup> CAR <sup>+</sup> cells (N=6)	360 x 10 <sup>6</sup> CAR <sup>+</sup> cells (N=2)	All 3011g ALLO-647 (N=8)	(N=19) Rate (95%CI)
ORR, n (%)	3 (75%)	3 (75%)	1 (33%)	7 (64%)	4 (67%)	1 (50%)	5 (63%)	12/19 (63%) (38%, 84%)
CR , n (%)	1 (25%)	1 (25%)	1 (33%)	3 (27%)	4 (67%)	0 (0%)	4 (50%)	7/19 (37%) (16%, 62%)

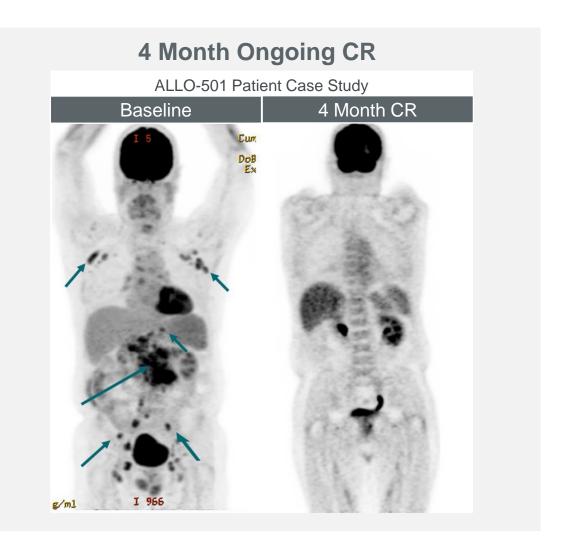
Median follow-up time: 3.8 months (range: 0.7 - 6.1)



### ALLO-501 Demonstrated Meaningful Tumor Reductions

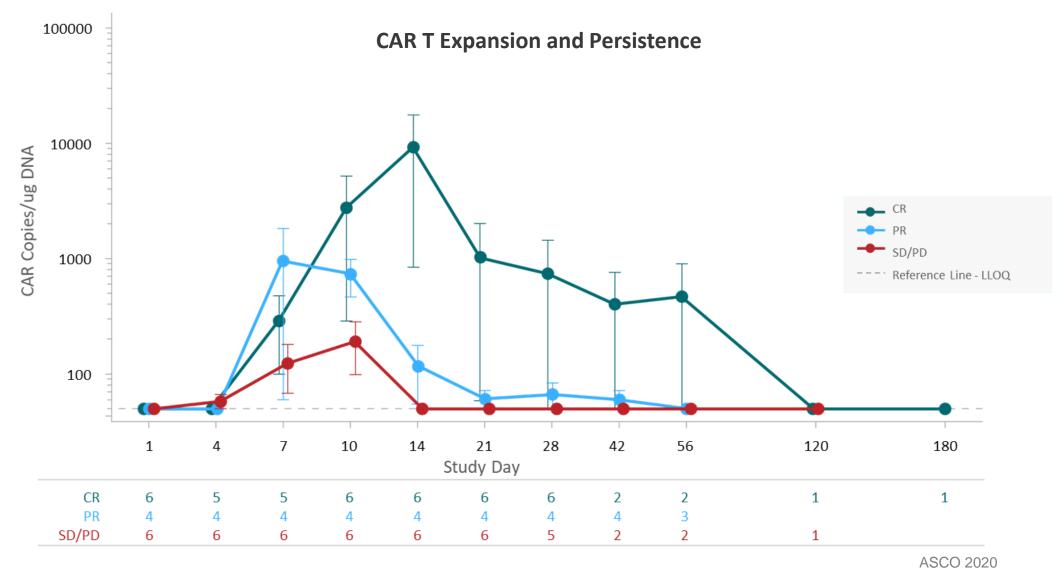








# AlloCAR T Cell Expansion Is Associated with Clinical Response





# ALPHA Trial: De-Risking the CD19 Program

- ☑ Can ALLO-501 be successfully manufactured?
- Can ALLO-501 be safely administered without causing clinically relevant Graft vs. Host Disease?
- Can ALLO-647 be safely administered and allow a sufficient window of lymphodepletion to allow ALLO-501 expansion and persistence?
- ☑ Can ALLO-501 provide complete responses across multiple histologies?

ONGOING Can ALLO-501 provide durable responses?



# Heavily Pretreated Patients with Advanced, Refractory Stage Disease

Median Time from Enrollment to Treatment: **5 Days** 

#### Enrolled (N=35)

4 patients ineligible due to rapidly progressing disease

#### Safety Population (N=31)

5 treated patients yet to reach assessment

#### **Efficacy Population (N=26)**

	Lymp	hodepletion Regi	men		
CAR <sup>+</sup> T Cell Dose	FC	A	CA		
2000	Low Dose ALLO-647	High Dose ALLO-647	Low Dose ALLO-647		
40 x 10 <sup>6</sup> Cells	3	-	-		
160 x 10 <sup>6</sup> Cells	4	-	3		
320 x 10 <sup>6</sup> Cells	6	4	3		
480 x 10 <sup>6</sup> Cells	3	-	-		

Overall median follow-up time = 3.2 Months

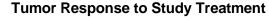
#### All Patients Refractory to Last Line of Therapy

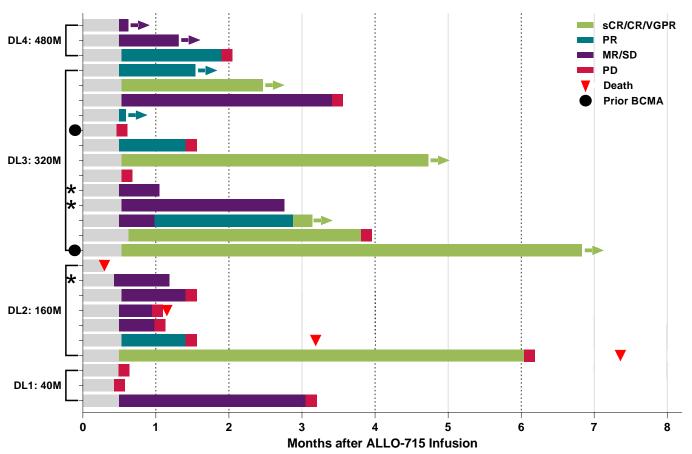
Characteristics		Safety Population (N = 31)
Age, median (range), years		65 (46, 76)
Gender, %	Male	61
	Female	39
ECOG, %	0	48
	1	52
ISS Stage ≥2, %		74
High-risk cytogenetics*, %		48
Extramedullary disease, %		23
High tumor burden <sup>†</sup> , %		39
Time since initial diagnosis, median (range), ye	ears	5.4 (0.9, 20.1)
Number of prior anti-myeloma regimens, media	5 (3 – 11)	
Prior autologous SCT, %	94	
Penta-exposed, %		94





# Objective Responses are Cell Dose-Dependent





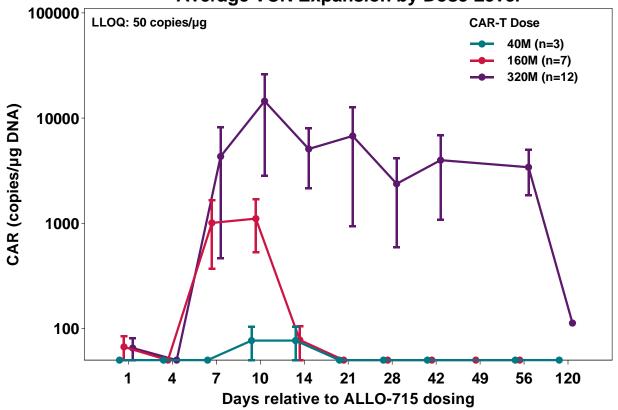
- Median time to response was 16 days
- Increasing response rates as cell dose increases
- 6 out of 9 patients treated with DL3 or DL4 with response remain in response



<sup>\*</sup>Discontinued follow-up on study prior to disease progression.

# AlloCAR T Cell Expansion Increased with ALLO-715 Dose Level



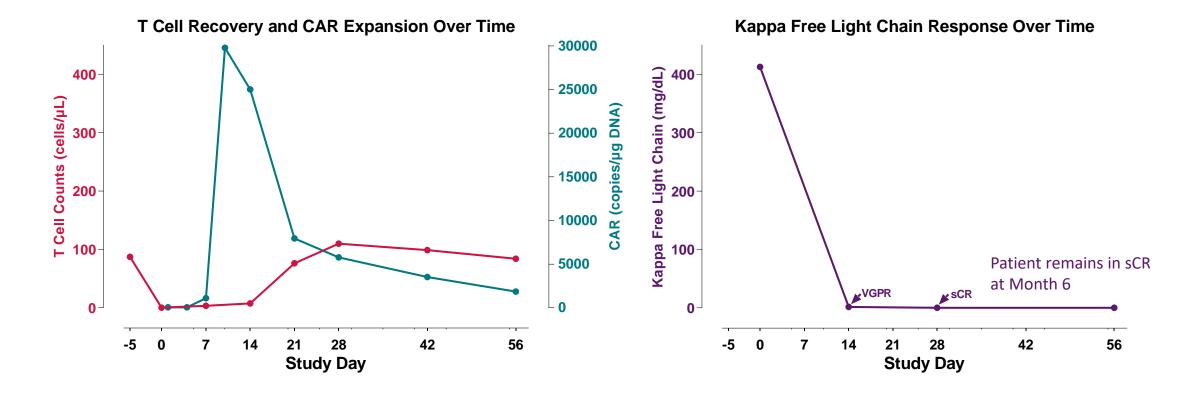


- Cell expansion was observed as early as 7 days
- Improved expansion in patients who received higher cell doses
- Persistence observed out to month 4 in dose level 3
- Patients with CAR T expansion had higher serum levels of IL15 at day 0 and day 14 [data not shown]

As of data cutoff date, limited DL4 vector copy number (VCN) data was available (2 patients with neither patient reaching day 28). Remaining data pending.



### ALLO-715 Case Study: Kinetics of AlloCAR T Cell Persistence, Lymphocyte Count and Response





### UNIVERSAL: De-Risking the Anti-BCMA AlloCAR T Franchise in MM

- ✓ Can ALLO-715 be successfully manufactured?
- Can ALLO-715 be safely administered without causing clinically relevant Graft vs. Host Disease?
- Can ALLO-647 be safely administered and allow a sufficient window of lymphodepletion to allow ALLO-715 expansion and persistence?
- Can an allogeneic CAR T therapy demonstrate a meaningful response in a heavily treated/refractory patient with multiple myeloma?
- ONGOING Can we further enhance the efficacy and durability of allogeneic cell therapy in multiple myeloma?

