

**Developing CD19-Redirected ARTEMIS[®]
T-Cell Therapy for Cancers
and Autoimmune Diseases**

April 10, 2024

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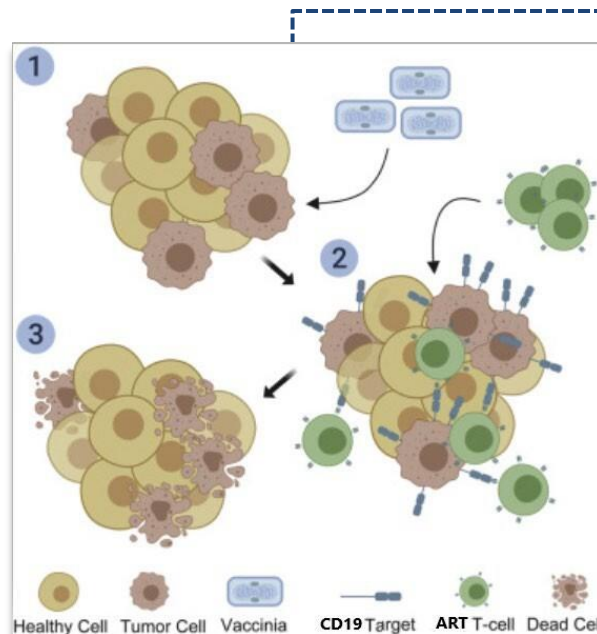
Executive Summary: Background and Opportunity

Recent success treating blood cancers with T-cell immunotherapy

- Juno Therapeutics (Juno) and Kite Pharma (Kite) were leaders in developing T-cell immunotherapy targeting CD19
- FDA approved T-cell immunotherapy targeting CD19 for blood cancer in 2017
- Average price of treatment is around \$400,000 per patient
- Juno and Kite acquired by Bristol-Myers Squibb (Nasdaq: BMY) and Gilead Science (Nasdaq: GILD) for \$9 billion and \$11.9 billion in 2018 and 2017, respectively

Opportunity remains for 2nd generation CD19-targeted T-cell therapies with less toxicity, and autoimmune disease market is wide-open

- Current CD19 T-cell therapy has severe side effects including **Cytokine release syndrome (CRS) and neurotoxicity**
- **Autoimmune disease**, which affects as many as 50 million people in the U.S., have NOT been successfully treated by T-cell immunotherapy



ARTEMIS® vs. CAR-T*

- **Superior efficacy**
- **Enhanced tumor infiltration**
- **Less T-cell exhaustion**
- **Reduced cytokine release syndrome (CRS) and cytokine released toxicities**



- **Estrella's EB103**, which utilizes **ARTEMIS®** T-cell engineering technology, has been validated preclinically and clinically with favorable safety profile and promising efficacy signals.
- The **“Mark and Kill”** approach by combining specially designed oncolytic viruses that label solid tumor cells with CF33-CD19t (“Mark”) and EB103 (“Kill”) is a potential breakthrough for treating solid tumors with T-cell therapy.

* According to head-to-head comparison studies conducted by independent 3rd parties



A clinical-stage biopharmaceutical company developing T-cell therapies with the potential to more effectively treat patients with cancers and autoimmune diseases.

Our Mission

Harness the evolutionary power of the human immune system to transform the lives of patients fighting cancer.



Our Products

- Lead product candidate, EB103, the next-generation **CD19-targeted ARTEMIS® T-cell therapies** with superior efficacy, enhanced tumor infiltration, and less T-cell exhaustion.
- EB104, a **CD19/22 Dual-Targeting ARTEMIS® T-cell therapies** with more efficacy, reducing relapse due to CD19 antigen loss.



Our Partnership

In partnership with Imugene, we combine our **ARTEMIS® technology** with oncolytic virus “CF33-CD19t” to treat solid tumors in a “**Mark and Kill**” strategy.



Estrella's Scientific Advisory Board Includes World-Class Thought Leaders



**Gianpietro Dotti,
MD**



- Research Professor of microbiology and immunology at University of North Carolina
- Director of the Lineberger Comprehensive Cancer Center Immunotherapy Program at University of North Carolina at Chapel Hill

**David Scheinberg,
MD, PhD**



- Physician, scientist, drug developer, entrepreneur, and pioneer of targeted alpha particle therapies
- Memorial Sloan Kettering Cancer Center, Former Chairman of Leukemia Service



Memorial Sloan Kettering
Cancer Center

**Michael Kavanaugh,
MD**



- Associate Clinical Professor of Medicine at University of California, San Francisco
- Former CSO and Head of Research and Non-Clinical Development of CytomX
- Former Senior VP and CSO of Five Prime Therapeutics
- Former VP of Novartis Vaccines & Diagnostics, and ED of Oncology Biologics in Novartis Institutes of Biomedical Research



**Stephan Grupp,
MD, PhD**



- Chief of the Cell Therapy and Transplant Section in the Division of Oncology and Director of Cancer Immunotherapy Program at Children's Hospital of Philadelphia
- Principal Investigator for CD19 CAR-T Kymriah by Novartis



**Randy Schekman,
PhD**



- Cell Biologist at UC Berkeley
- Former editor-in-chief of *The Proceedings of the National Academy of Sciences* and the *Annual Review of Cell and Developmental Biology*
- 2013 Nobel Prize in Physiology or Medicine



Estrella Immunopharma Licensed Patents on CD19-ARTEMIS®



CD19

Issued patents in the US and have 23 applications worldwide



US010301388B2

(12) **United States Patent**
Liu et al. (10) **Patent No.:** US 10,301,388 B2
(45) **Date of Patent:** May 28, 2019

(54) **ANTIBODY AGENTS SPECIFIC FOR HUMAN CD19 AND USES THEREOF** USPC 530/387.1, 387.3; 435/325; 424/93.21
See application file for complete search history.

(71) Applicant: **Eureka Therapeutics, Inc.**, Emeryville, CA (US) (56) **References Cited**

U.S. PATENT DOCUMENTS

(72) Inventors: **Hong Liu**, Emeryville, CA (US);
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(73) Assignee: **Eureka Therapeutics, Inc.**, Emeryville, CA (US)

ARTEMIS®

Four issued patents in the US and 62 applications pending worldwide

(12) **United States Patent**
Lu et al. (10) **Patent No.:** US 10,098,951 B2
(45) **Date of Patent:** Oct. 16, 2018

(54) **ANTIBODY/T-CELL RECEPTOR CHIMERIC CONSTRUCTS AND USES THEREOF** (56) **References Cited**
U.S. PATENT DOCUMENTS

(71) Applicant: **EUREKA THERAPEUTICS, INC.**, Emeryville, CA (US) 3,753,357 A 8/1973 Schwartz
4,199,022 A 4/1980 Senkan et al.

(12) **United States Patent**
Lu et al. (10) **Patent No.:** US 10,464,988 B2
(45) **Date of Patent:** *Nov. 5, 2019

(54) **ANTIBODY/T-CELL RECEPTOR CHIMERIC CONSTRUCTS AND USES THEREOF** C07K 2319/00 (2013.01); C07K 2319/03 (2013.01); C07K 2319/33 (2013.01); C07K 2319/74 (2013.01)

(71) Applicant: **EUREKA THERAPEUTICS, INC.**, Emeryville, CA (US) (58) **Field of Classification Search**
CPC A61K 39/39558; A61K 35/17

(12) **United States Patent**
Liu et al. (10) **Patent No.:** US 10,822,413 B2
(45) **Date of Patent:** Nov. 3, 2020

(54) **CELLS EXPRESSING CHIMERIC ACTIVATING RECEPTORS AND CHIMERIC STIMULATING RECEPTORS AND USES THEREOF** (58) **Field of Classification Search**
None
See application file for complete search history.

(71) Applicant: **EUREKA THERAPEUTICS, INC.**, Emeryville, CA (US) (56) **References Cited**
U.S. PATENT DOCUMENTS

3,753,357 A 8/1973 Schwartz

(12) **United States Patent**
Lu et al. (10) **Patent No.:** US 10,822,389 B2
(45) **Date of Patent:** *Nov. 3, 2020

(54) **ANTIBODY/T-CELL RECEPTOR CHIMERIC CONSTRUCTS AND USES THEREOF** 2317/56 (2013.01); C07K 2317/622 (2013.01); C07K 2317/73 (2013.01); C07K 2319/00 (2013.01); C07K 2319/03 (2013.01); C07K 2319/33 (2013.01); C07K 2319/74 (2013.01)

(71) Applicant: **EUREKA THERAPEUTICS, INC.**, Emeryville, CA (US) (58) **Field of Classification Search**

Estrella Pipeline and Strategy

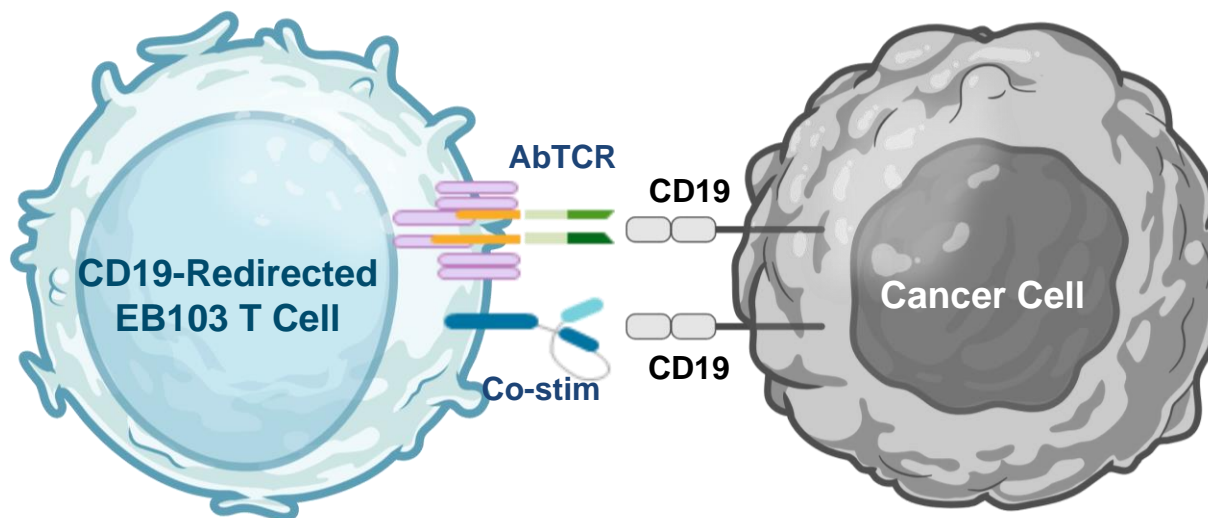


- Our approach is to rapidly advance our lead product candidate **EB103, CD19-Redirected ARTEMIS® T-Cell programs** in relapsed/refractory and high- risk **blood cancers first**.
- We are also developing **EB104, CD19/22 Dual Targeting ARTEMIS® T-Cell Therapy** to treat patients with lower surface CD19 density or a greater prevalence of CD22.
- Meanwhile, in partnership with Imugene we are developing **EB103+ CF33-CD19t** using the **“Mark-and-Kill”** approach to address various types of **solid tumors**.
- For indications beyond cancers, we are working on IND enabling studies of **CD19-Redirected ARTEMIS® T-Cell Programs** for the treatment of autoimmune diseases, starting with systemic lupus erythematosus (SLE).

Indications	Program	Early-Discovery	Late-Discovery	Pre-Clinical	Clinical	Partner	
HEMATOLOGIC MALIGNANCIES							
<i>Diffuse large B cell lymphoma (DLBCL), Follicular Lymphoma (FL), and other types of B-cell Lymphoma</i>	<i>EB103 (CD19)</i>						
<i>Diffuse large B cell lymphoma (DLBCL), Follicular Lymphoma (FL), and other types of B-cell Lymphoma</i>	<i>EB104 (CD19/CD22)</i>						
SOLID TUMORS							
<i>Multiple indications</i>	<i>Combination: EB103 + CF33-CD19t</i>						
AUTOIMMUNE DISEASES							
<i>Systemic Lupus Erythematosus (SLE)</i>	<i>EB201</i>						

A dark blue background featuring a glowing DNA double helix structure on the right side and a network of interconnected nodes and lines on the left side, suggesting a molecular or genetic theme.

Lead Product: EB103 (CD19-Redirected ARTEMIS[®] T-Cell Therapy)



EB103 CD19-Redirected ARTEMIS® T Cells

EB103 Engineered to express ARTEMIS® cell receptors (i.e., the AbTCR and co-stimulatory molecule) on cell surfaces.

Once infused, EB103 T-cells (cell receptors) recognize and bind the CD19-positive cancer cells.

Cell receptors, AbTCR/CD3 complex-mediated signal transduction within the EB103 T-cell is initiated, leading to the activation of the EB103 T-cell.

The second “enhancement” signal is generated when the co-stimulatory molecule expressed on the EB103 T-cells binds to its target, CD19.

EB103 T-cells seek out CD19-positive cancer cells, bind to and destroy them.

Key Unit – AbTCR and Co-stim

The key units of our novel, proprietary CD19-Redirected ARTEMIS® T Cells comprised of an **antibody-T-cell-receptor (AbTCR)** and a **co-stimulatory molecule**:

The Antibody-T-Cell Receptor (AbTCR) serves as the core component featuring:

- A target-binding domain derived from an antibody fragment antigen-binding (Fab) region
- An effector domain derived from portions of a human gamma/delta ($\gamma\delta$) T-cell receptor (TCR)

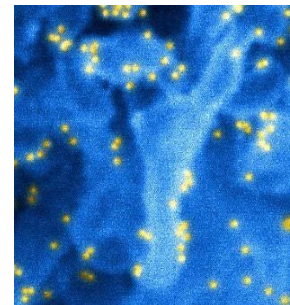
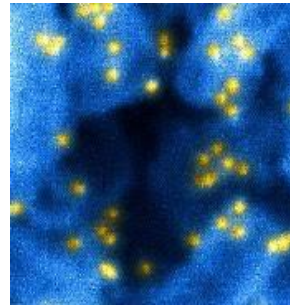
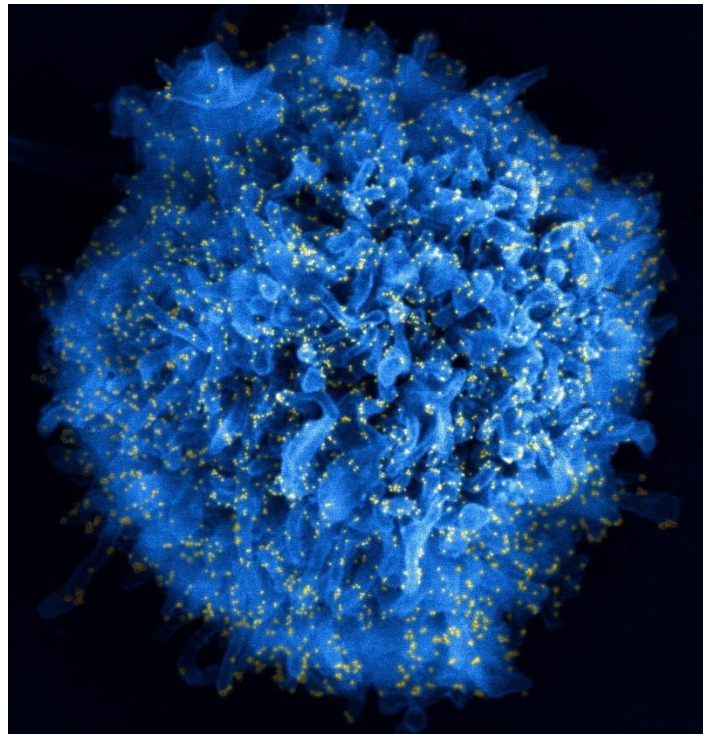
The Co-Stimulatory Molecule is an additional key component featuring:

- A target-binding domain derived from a single-chain variable fragment (scFv)
- A co-stimulatory domain derived from portions of a human co-stimulatory receptor

Both the AbTCR and the co-stimulatory molecule bind to the **CD19 antigen**, a well-validated target commonly overexpressed on blood cancer cells.

ARTEMIS®

Superiority to Conventional CAR-T



ARTEMIS® receptor is primarily localized in microvilli.

*(Collaboration: Alice Liang, Ph.D. Director of Microscopy Laboratory,
NYU Langone Health NYU School of Medicine)*

- ARTEMIS® T-cell therapy is **clinically validated in patients**
- ARTEMIS® vs. CAR-T*
 - ✓ **Superior** efficacy
 - ✓ **Enhanced** tumor infiltration
 - ✓ **Less** T cell exhaustion
 - ✓ **Reduced** Cytokine release syndrome (CRS) and cytokine related toxicities

* According to head-to-head comparison studies conducted by independent 3rd parties

Better Safety and Potent Anti-tumor Efficacy



Stephan A. Grupp, MD, PhD

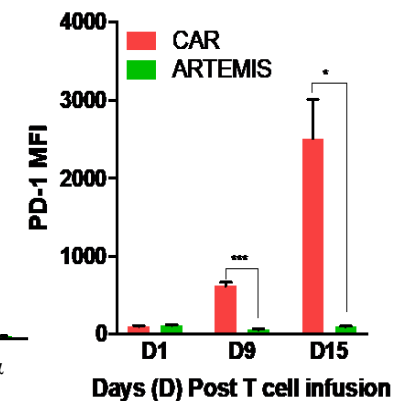
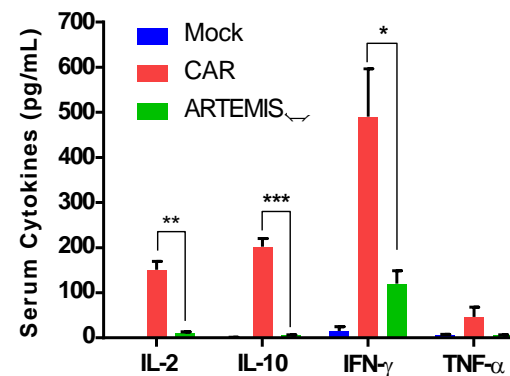
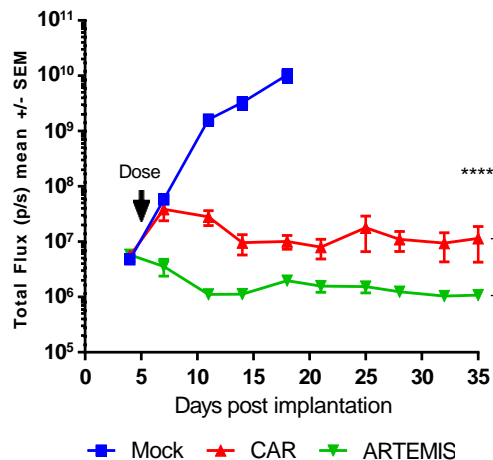
- Delivered CAR T-cell therapy to the first pediatric patient in the world (Emily Whitehead)
- Led the first multicenter global study of Kymriah®, which became the first CAR-T therapy to receive approval from the FDA



- Collaboration research between Eureka Therapeutics, Inc., Estrella's parent company, and Dr. Grupp's team showed CD19 ARTEMIS® T-cell Therapy demonstrated better safety and anti-tumor efficacy.
- The research paper *Xu et al. Cell Discovery (2018) 4:62* published in *Nature* in 2018.

ARTEMIS® vs. CAR-T cells

- Potent anti-tumor activity
- Better safety profile
- Longer durability with less exhausted phenotype



Snapshots of Estrella CD19 Therapy in Lymphoma Patients

- Collaboration research with the First Affiliated Hospital of Xi'an Jiaotong University for exploratory, single-arm, open-label, non-randomized early investigator initiated study (“IIS”).

Abstract # 2870

61th ASH
Annual
Meeting 2019

Summary

- As of Sep 11th, 2019, 9 adult r/r DLBCL patients received autologous ET019003 ARTEMIS™ T cells at the dose of 3x10⁶/kg/infusion. Their clinical responses were followed up for at least 1 month after infusion.
- Repeat dosing of ET019003 T cells is well tolerated.
- Robust expansion of ET019003 T cells in peripheral blood was observed in all subjects. Peak expansion did not correlate with adverse event and response.
- ET019003 T cell therapy demonstrated a favorable safety profile. No more than grade-2 CRS was observed and no patient shows neurotoxicity.
- ET019003 T cells achieved excellent clinical efficacy (ORR: 8/9, best response: CR: 4/9, PR: 4/9).

A novel antibody-TCR (AbTCR) T-cell therapy is safe and effective against CD19-positive relapsed/refractory B-cell lymphoma

Pengcheng He¹ · Haibo Liu¹ · Bryan Zimdahl² · Jie Wang¹ · Minna Luo¹ · Qi Chang² · Fangzhou Tian² · Fan Ni² · Duo Yu² · Huasheng Liu¹ · Limei Chen¹ · Huaiyu Wang¹ · Mei Zhang¹ · Stephan A. Grupp^{3,4} · Cheng Liu²

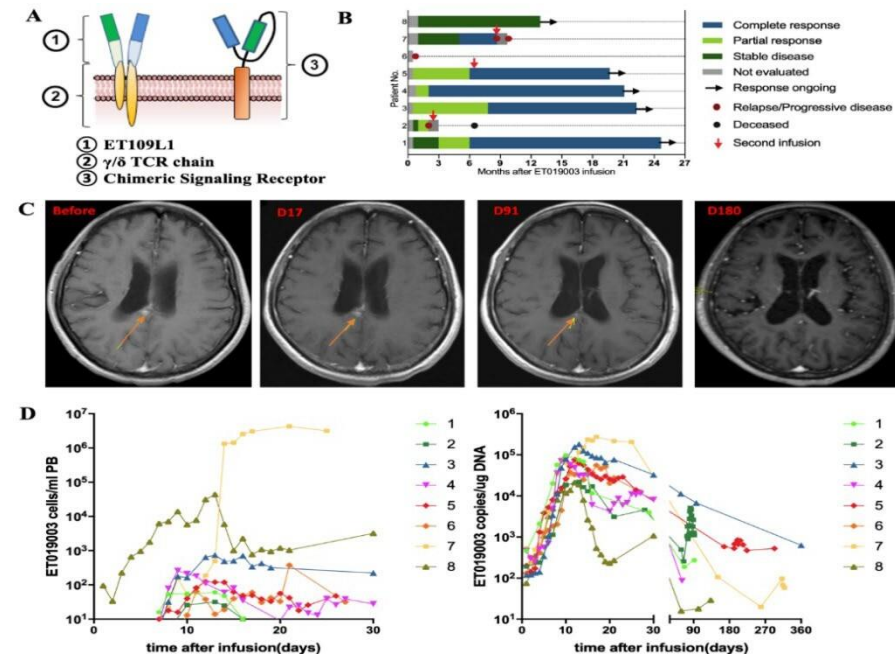
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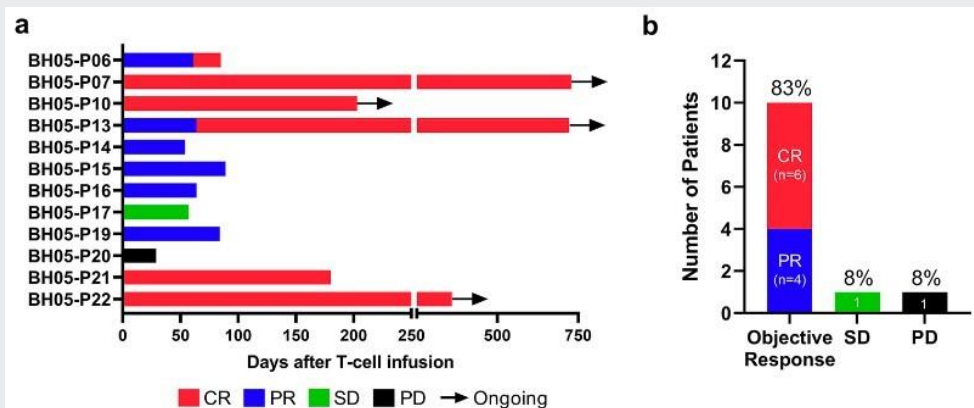
ASH | Annual Meeting & Exposition 2021

826 Novel CD19-Specific γ/δ TCR-T Cells in Relapsed or Refractory Diffuse Large B-Cell Lymphoma

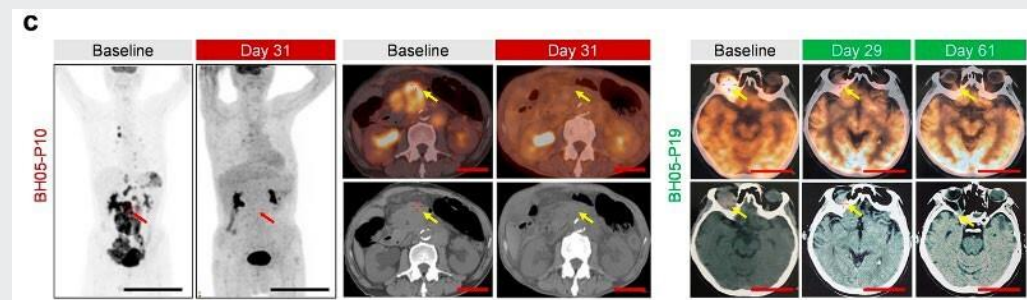
Oral presentation Monday, December 13, 2021: 5:15 PM



- This IIS study ⁽¹⁾ provides data that indicates that EB103 has anti-tumor activity and an attractive safety profile in patients with CD19-positive r/r B-cell lymphoma.

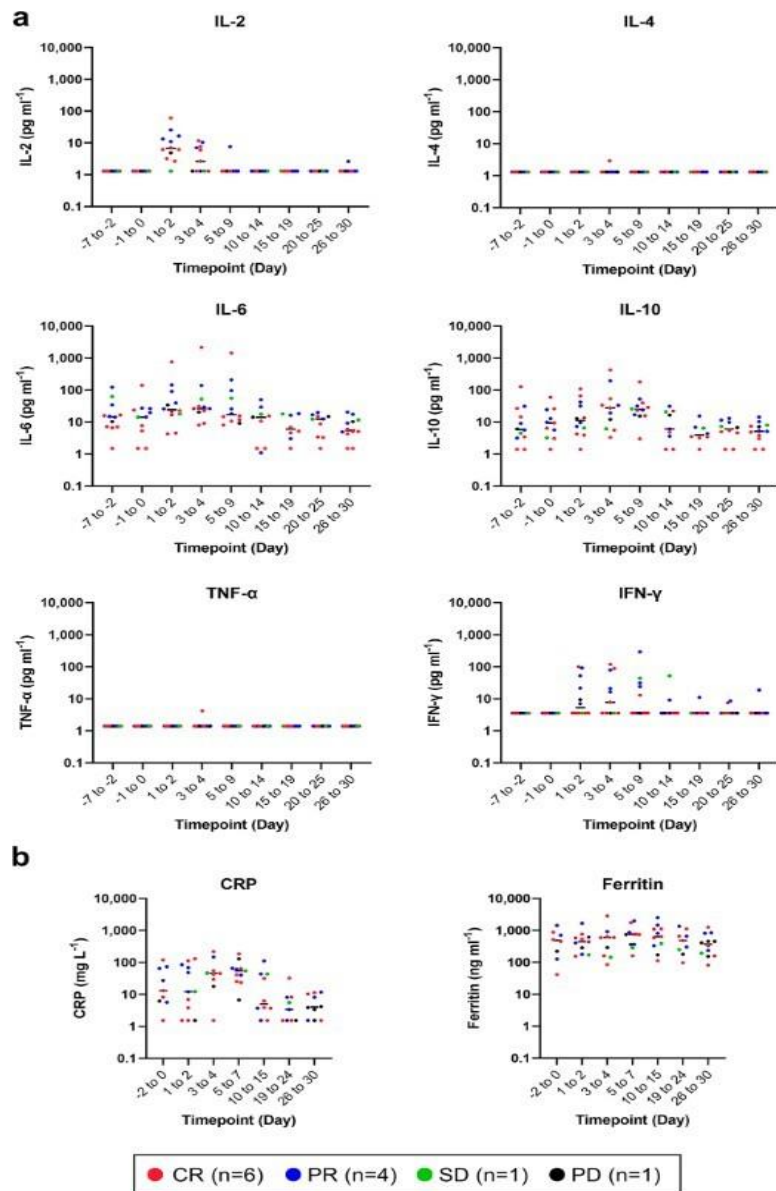


- (a) Treatment response and duration of response after initial infusion of EB103 T-cells. Black arrows indicate ongoing remission and follow-up.
- (b) Best response for the 12 patients. Best response was defined as the best response (i.e., CR > PR > SD > PD) the patient achieved at any time after receiving EB103. CR - complete response, PR - partial response, SD - stable disease, PD - progressive disease.



- (c) Representative radiographic images of two responders (BH05-P10 and BH05-P19) at baseline and the indicated time points after EB103. Red or yellow arrows mark the tumor lesions. Full body images are PET-CT scans. Cross-sectional images are PET scans (top rows) and CT scans (bottom rows). Scale bars: black, 20 cm; red, 6 cm.

(1) Reference: He et al. Journal of Cancer and Clinical Oncology 10 June 2022
A novel antibody-TCR(AbTCR) T-cell therapy is safe and effective against CD19-positive relapsed/refractory B-cell Lymphoma



- **This IIS study ⁽¹⁾ provides data that indicates that EB103 has anti-tumor activity and an attractive safety profile in patients with CD19-positive r/r B-cell lymphoma.**
 - The study enrolled patients from November 2018 to January 2020 ⁽¹⁾ 16 patients were enrolled, and 12 patients were treated.
 - Of the 12 patients treated, six patients (50%) achieved a complete response (“CR”), and four (33%) achieved a partial response (“PR”), with a best objective response rate of 83%.
 - CRs were durable, including two patients with ongoing CRs for 22.7 months and 23.2 months. EB103 was well-tolerated with an attractive safety profile.
 - **No patients experienced severe (grade > 3) CRS, and only one patient experienced ICANS of any grade.** Significant elevations of cytokine levels were not seen, even in patients with marked expansion of EB103 T-cells.



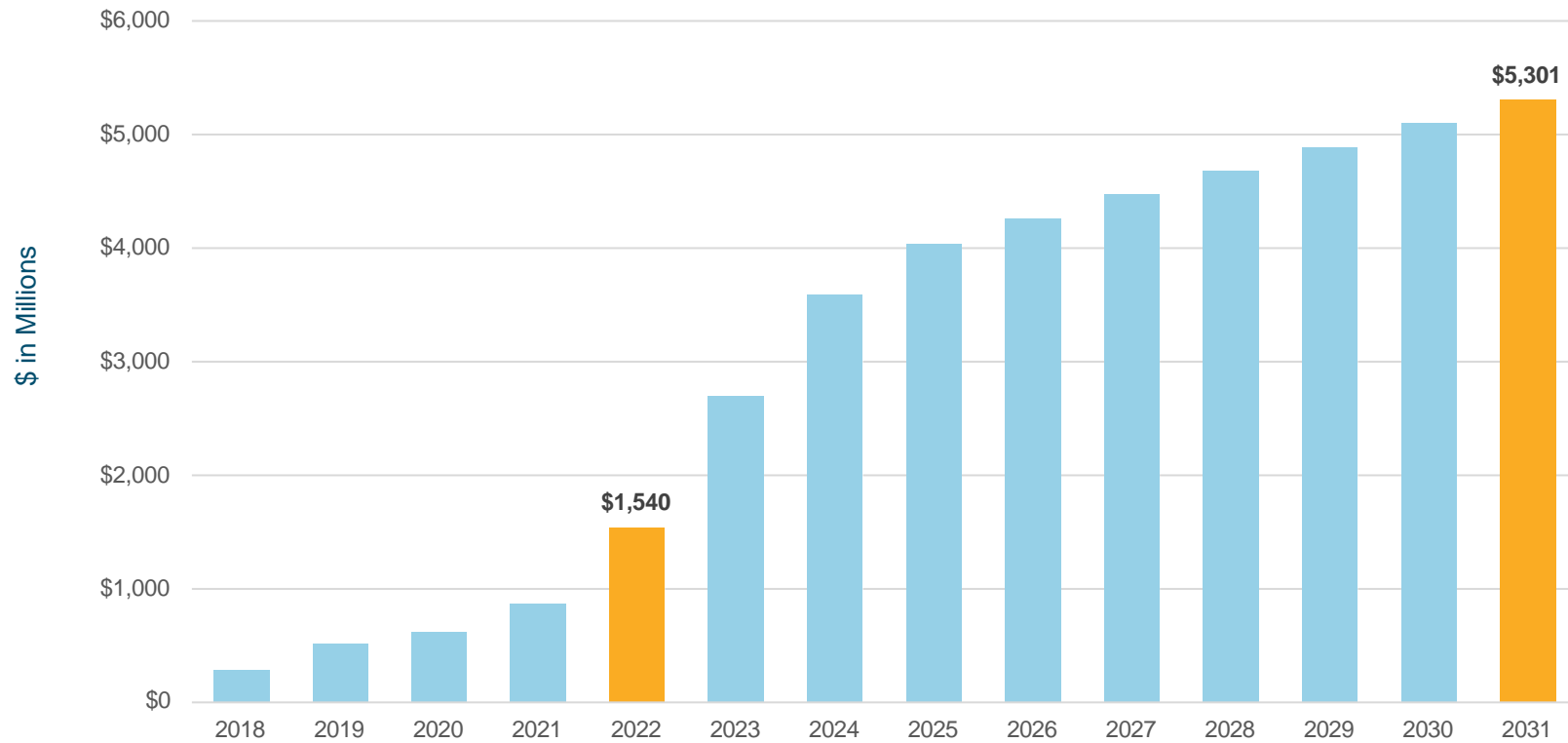
Levels of cytokines and serum inflammatory markers after EB103 T-cell infusion

- (a) Cytokine levels
- (b) Serum c-reactive protein (CRP) and ferritin levels in patients during the first month of EB103. Horizontal lines denote median values. Patients’ best responses are denoted by color of the symbols: CR (red), PR (blue), SD (green), and PD (black). Values less than the limit of detection were recorded as half the lower limit

(1) This IIS was conducted at The First Affiliated Hospital of Xi’an Jiaotong University in China and was registered at as #NCT03642496
 Reference: He et al. Journal of Cancer and Clinical Oncology 10 June 2022
 A novel antibody-TCR(AbTCR) T-cell therapy is safe and effective against CD19-positive relapsed/refractory B-cell Lymphoma

CD-19 Targeted CAR-T Therapies Market

Currently approved CD-19 targeted CAR-T therapies projected sales in lymphoma: **\$5+ billion (2031)**



CD19 CAR-T Product Sales Projection (2018-2031)

CD-19 CAR-T Company Acquisitions

- 1st Generation CD-19 CAR-T Companies Valued at \$9-12B at the Time of Acquisition

FDA approved T-cell therapies are limited to hematologic malignancies



\$11.9 billion
In 2017 by Gilead



\$9 billion in 2018
by Celgene

- Three FDA approved CAR-T therapies historical sales from 2017 to 2022, and projected sales in 2026

Kymriah (\$MM) (Novartis)	2017A	2018A	2019A	2020A	2021A	2022A	2026E
USA	\$6	\$76	\$159	\$205	\$230	\$196	
Worldwide	\$6	\$76	\$278	\$474	\$587	\$536	\$1,090

Yescarta (\$MM) (Gilead/Kite)	2017A	2018A	2019A	2020A	2021A	2022A	2026E
USA	--	\$263	\$373	\$362	\$406	\$1,160	
Worldwide	--	\$264	\$456	\$563	\$695	\$794	\$1,481

Breyanzi (\$MM) (BMS/Juno)	2017A	2018A	2019A	2020A	2021A	2022A	2026E
USA	--	--	--	--	\$84	\$151	
Worldwide	--	--	--	--	\$87	\$182	\$1,515

Treatment	Price (WAC)	Treatment	Price (WAC)
Kymriah (Novartis)	\$373,000-\$475,000	Yescarta (Gilead/Kite)	\$373,000

Treatment	Price (WAC)
Breyanzi (BMS/Juno)	\$410,300

Highlights of EB103 Program

- ARTEMIS® T-cell engineering technology may offer advantages over existing CAR-T technologies, including a potentially improved safety profile and reduced side effects.
- IND clearance in Q1 2023, Phase I trials expected to commence in 1H 2024.

Antibody-based target recognition

- The ability to achieve **high specificity and binding affinity** to intended cancer target when compared to TCRs.

AbTCR includes portions of a human TCR

- The AbTCR associates with the endogenous CD3 complex enabling the AbTCR to use the same activation and regulatory signaling pathways employed by **natural TCRs**.
- This feature may lead to **improved safety** in patients.

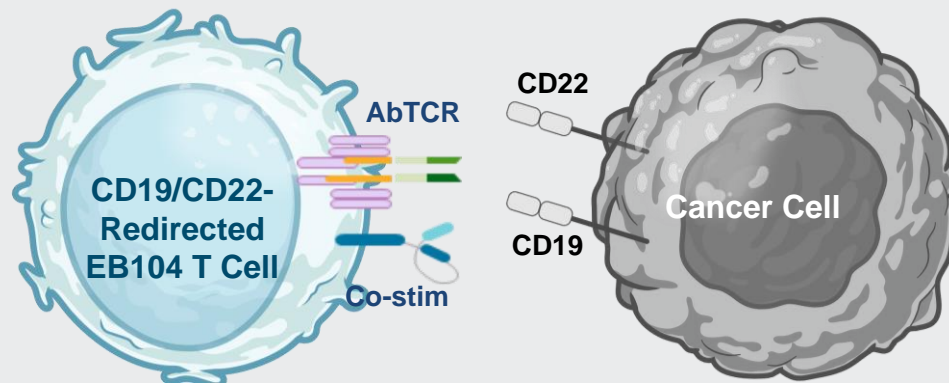
Co-stimulation provided as a separate molecule

- The AbTCR construct does not include an intracellular signaling domain covalently-linked to a co-stimulatory domain, and thus has the potential to eliminate T-cell hyperactivation and consequently, **lower the risk of CRS and ICANS** commonly observed with CAR-T therapy.

A dark blue background with a glowing DNA double helix and a network of interconnected nodes and lines, suggesting a molecular or genetic theme. A light blue horizontal bar is at the top left, and a dark blue horizontal bar is at the bottom right.

EB104 (CD19/22 Dual-Targeting ARTEMIS® T-Cell Therapy)

CD19/CD22-Redirected EB104 T Cell



EB104 Engineered to express ARTEMIS® cell receptors (i.e., the AbTCR and co-stimulatory molecule) on cell surfaces. **AbTCR in EB104** recognize and binds to **both CD19 and CD22 antigen.**

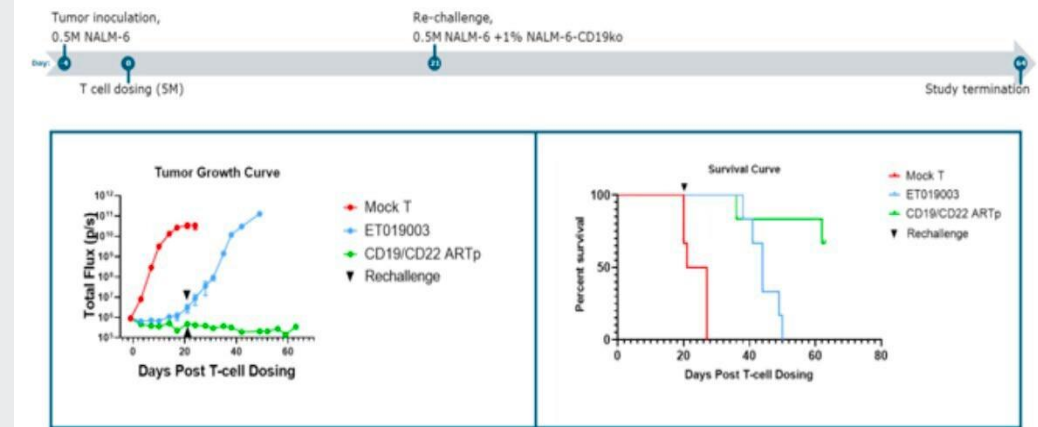
Once infused, EB104 T-cells (cell receptors) recognize and bind the CD19- and CD22-positive cancer cells.

Cell receptors, AbTCR/CD3 complex-mediated signal transduction within the EB104 T-cell is initiated, leading to the activation of the EB104 T-cell.

The second “enhancement” signal is generated when the co-stimulatory molecule expressed on the EB104 T-cells binds to its target, CD19.

EB104 T-cells seek out CD19- and CD22-positive cancer cells, bind to and destroy them.

EB104 Preclinical Data

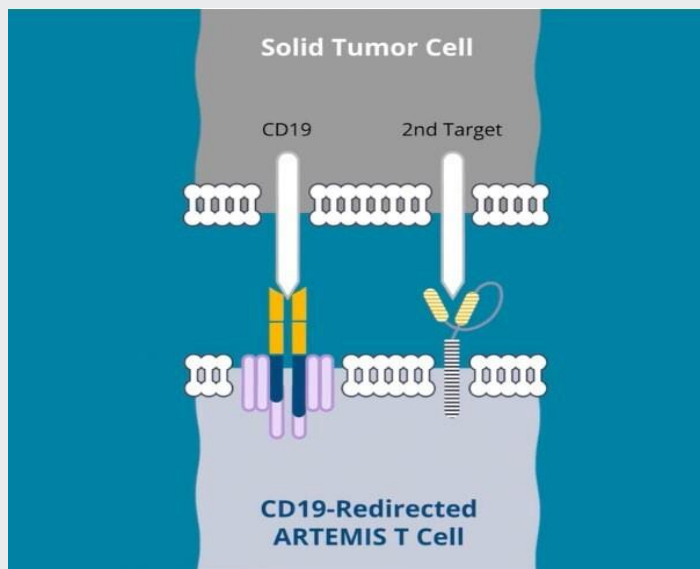


- Results showed that EB104 T-cells have the potential to eradicate Nalm-6 Primary Tumors and Nalm-6-CD19ko re-challenge tumors in the xenograft model, suggesting that EB104 T-cells have the potential to control the growth of tumor cells that do not express CD19.
 - A Nalm-6-CD19ko cell line constructed, with the “knockout” of CD19 gene expression to test the activity of EB104 in mice using NSGTM xenograft models.
 - Inoculated NSGTM mice, that were CD19 positive four days before receiving (i) mock control T-cells, (ii) EB103 T-cells, and (iii) EB104 T-cells. The Primary Nalm-6 Tumors in EB103 and EB104 groups resulted in remission.
 - Then inoculated the NSGTM mice with 1% Nalm-6-CD19ko to mimic diminished CD19 surface expression, creating “re-challenge” tumors.
 - EB103 T-cells not able to control the re-challenge tumors in the EB103 T-cell group, the re-challenge tumors in the EB104 T-cell group resulted in remission.

A dark blue background featuring a glowing DNA double helix structure on the right side and a network of interconnected nodes and lines on the left side, suggesting a molecular or data network.

CF33-CE19t and EB103 (Collaboration with Imugene)

The Mark-and-Kill Approach



To address the lack of solid tumor-specific targets currently available, we use *CF33-CD19t*, an *oncolytic virus*, to force solid tumor cells to express the CD19 protein on the cell surface.

The **CD19-Redirected ARTEMIS® T Cells** can then pursue and kill the **CD19-labeled solid tumors**, offering a potential treatment solution to cancers where there are no inherently abundant solid tumor-specific targets available.

Highlights

- 2nd generation CD19 T-cell Immunotherapy with a “Mark and Kill” strategy
- In a partnership with Imugene, we are investigating the use of EB103 in conjunction with Imugene’s product candidate, CF33-CD19t, to treat **solid tumors** in a “**Mark and Kill**” strategy

Partnership with Imugene (AUX: IMU)

- Founded: 2012
- Headquarter: Melbourne, Australia
- Market Cap: AUD 734 million ⁽¹⁾
- Management:
 - Paul Hopper (Executive Chairman; Serial bioentrepreneur)
 - Leslie Chong (CEO; Former clinical program lead at Genentech)

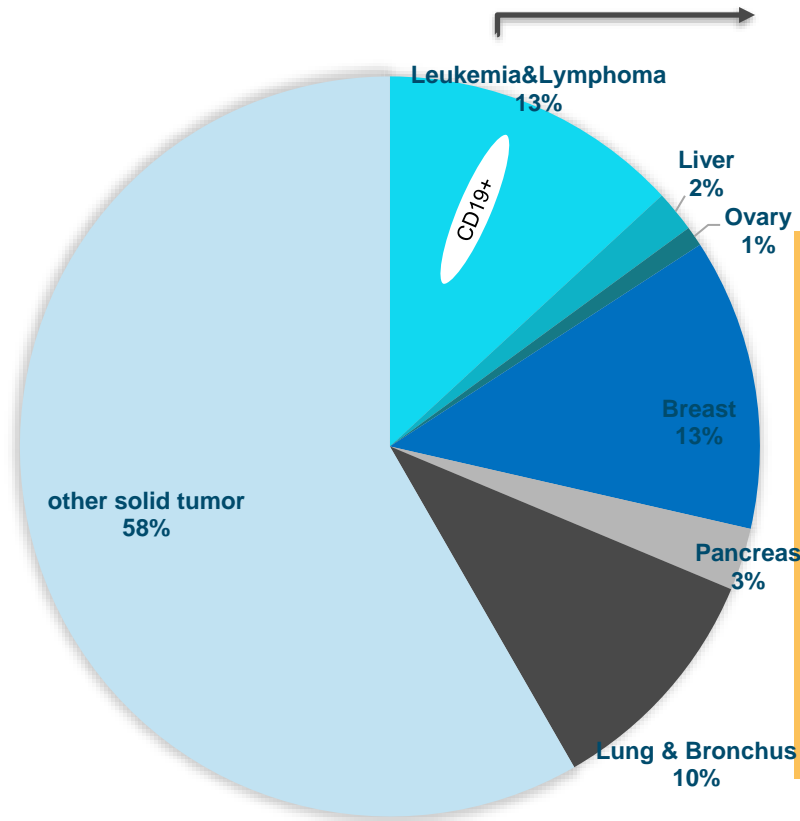
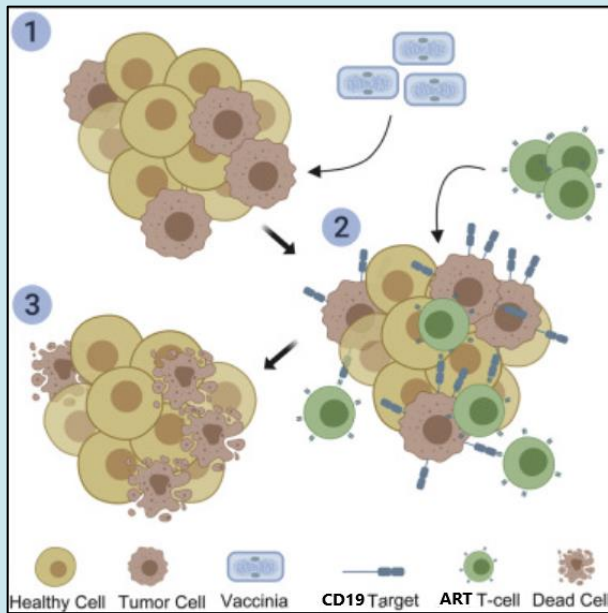
(1) As of April 4th, 2024.

Solid Tumor Market Potentials



Treating Solid Tumor with Estrella's CD-19 T-cell Therapies: "Mark and Kill Strategy"

- CF33 Oncolytic Virus ("Mark" the Tumor-Imugene)
- T-Cell Therapy ("Kill" the Tumor - Estrella)



~ **\$20 billion** combined market capitalization realized by *Juno* and *Kite* with T-cell therapies treating **CD19+** liquid tumors

Solid tumors represent:

- **1.6 million** new cases
- **90%** of total diagnoses in US each year
- **0** FDA approved CAR-T/TCR (T-Cell Receptor) therapies

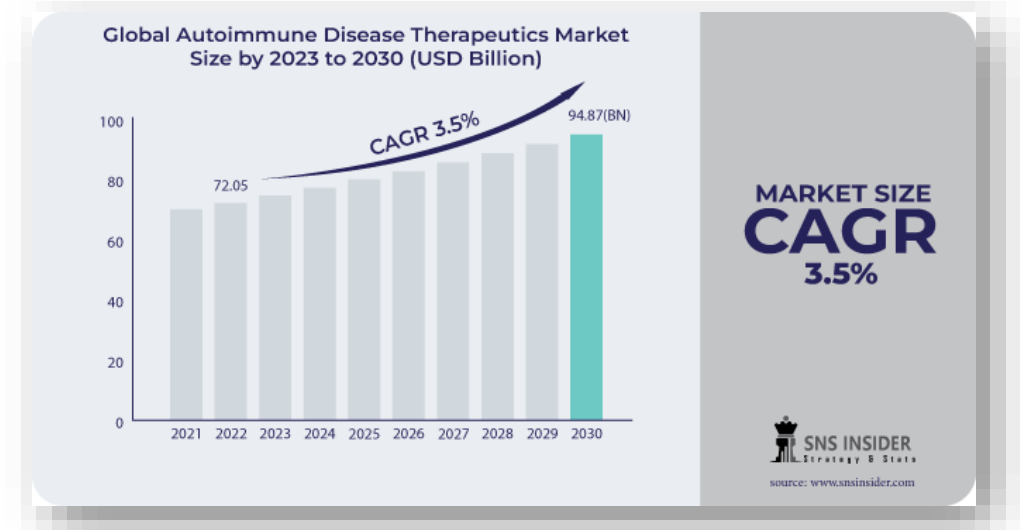
Estimated New Cancer Cases in United States, 2022*

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Going Beyond Cancers: CD19 ARTEMIS® T-Cell Therapy for Autoimmune Diseases

Autoimmune Diseases & SLE Market Overview

- Autoimmune diseases represent a significant health concern globally, characterized by the immune system mistakenly attacking the body's own tissues.
 - Approximately one in ten individuals globally are affected by autoimmune disorders. Over 80 types of autoimmune diseases identified, including systemic lupus erythematosus, rheumatoid arthritis, type 1 diabetes, and multiple sclerosis.
 - In 2022, the global market size for autoimmune disease therapeutics was valued at \$72.05 billion, expected to grow to \$94.87 billion by 2030.
 - North America holds a substantial market share, accounting for over 42.4% of the global revenue in 2021.
- Systemic Lupus Erythematosus (SLE), commonly known as lupus, is a chronic autoimmune disease with symptoms ranging from mild to life-threatening due to systemic inflammation.
 - The prevalence of SLE in the United States is approximately 72.8 per 100,000 person-years.
 - Factors driving this growth include the increasing prevalence of SLE, advancements in medical treatments, and the introduction of new biological therapies.
 - By 2025, the global market size for SLE treatments is expected to reach \$3.8 billion.



Autoimmune Disease Therapeutics Market Size & Share Research [2023-2030], SNS Insider.
Lupus Market Research Report by Type, Treatment, End User, and Region-Forecast till 2032, MRFR, March 2024

CD19 Targeting ARTEMIS® T-Cell Therapy for SLE

Targeted B-cell Depletion

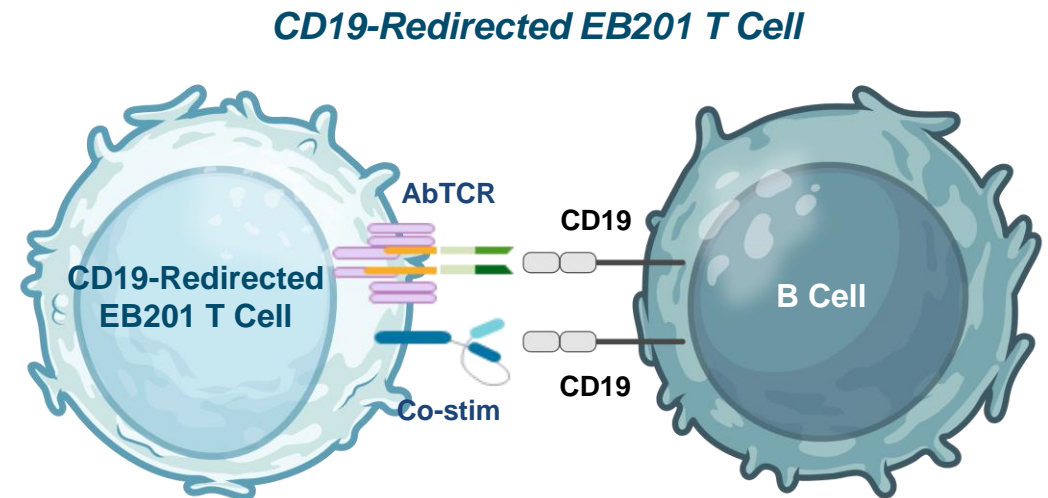
- CD19 CAR-T cells are engineered to recognize and bind to CD19, a protein expressed on the surface of B cells.
- By targeting CD19, ARTEMIS® T cells can specifically eliminate B cells, which are responsible for the production of autoantibodies in autoimmune diseases like SLE.

Autoimmune Regulation

- Reduction in autoantibody-producing B cells leads to a decrease in the systemic autoimmune response.
- The therapy aims to restore immune system balance by removing cells that mistakenly attack the body's own tissues.

Clinical Application

- Initial clinical trials with anti-CD19 CAR-T cells show promise for refractory SLE, where standard treatments are ineffective.
- Anti-CD19 and anti-B cell maturation antigen (BCMA) CAR-T cells have been used in diseases such as SLE and myasthenia gravis with positive outcomes.



Unlike traditional immunosuppressive treatments, EB201 therapy offers the potential for durable remission in certain B-cell mediated autoimmune diseases, such as SLE

Competitive Landscape of CD19 Targeting SLE Programs

EB201 Unique Advantages

- EB201 utilizes an anti-CD19 fully-human binder with the potential of lower immunogenicity. The same binder has been used in ARTEMIS[®] T-cell therapy against r/r B-cell lymphoma IIT in China, with 20 patients' data reported¹.
- Preclinical and early clinical validations suggest CD19-Redirected ARTEMIS[®] T cells outperforms existing FDA-approved CD19 CAR-T therapies in both safety and efficacy, particularly by potentially decreasing risks of CRS and ICANS.
- EB103 program has received IND clearance for blood cancers and is expected to commence Phase I trials in the first half of 2024. EB201 program intends to treat SLE patients with the same ARTEMIS[®] T cell being used in EB103.

CD19-Targeted T-cell Therapy Programs in SLE

Program	Company	Current Phase	Platform
BMS-986353	Bristol Myers Squibb	Phase 1	NEX-T
CABA-201	Cabaletta Bio	IND cleared	CAR-T
KYV-101	Kyverna Therapeutics	Phase 1	CAR-T

¹ Li, C., Zhou, F., Wang, J., Chang, Q., Du, M., Luo, W., Zhang, Y., Xu, J., Tang, L., Jiang, H., Liu, L., Kou, H., Lu, C., Liao, D., Wu, J., Wei, Q., Ke, S., Deng, J., Liu, C., & Mei, H. (2023). Novel CD19-specific γ/δ TCR-T cells in relapsed or refractory diffuse large B-cell lymphoma. *Journal of Hematology & Oncology*

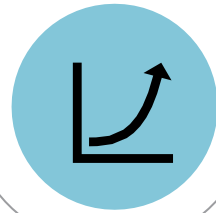
He, P., Liu, H., Zimdahl, B., Wang, J., Luo, M., Chang, Q., Tian, F., Ni, F., Yu, D., Liu, H., Chen, L., Wang, H., Zhang, M., Grupp, S. A., & Liu, C. (2022). A novel antibody-TCR (AbTCR) T-cell therapy is safe and effective against CD19-positive relapsed/refractory B-cell lymphoma. *Journal of Cancer Research and Clinical Oncology*

The potential applicability of EB103 in autoimmune diseases like SLE demonstrates Estrella's innovative approach to leveraging T-cell therapy across a broader spectrum of diseases.

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Summary

Estrella Expects to Achieve High Return Through the Potentially Decreased Risk of its Product Candidates



High Return

- CD19 T cell product average price of \$400K per treatment is expected to be covered by insurance
- Juno and Kite acquisition of \$9 billion and \$11.7 billion respectively, provide the reference for potential company value for blood cancer market
- Potential success to treat autoimmune disease would expand sales and potentially drive Estrella Immunopharma valuation beyond those of Juno and Kite
- Potential for expansion into autoimmune disease market with the same CD19-targeted T-cell therapy



Decreased Risk

- Proven safety of targeting CD19; four CD19 CAR-T therapies have been approved by FDA to date
- Target large blood cancer market: \$2 billion in sales for CD19+ in lymphoma alone as of 2021
- ARTEMIS® technology has undergone independent 3rd party head-to-head comparison studies, suggesting superior safety profile, anti-tumor activity and durability compared to FDA-approved products, Kymriah and Yescarta
- CD19-targeted T-cell therapy has been validated in patients in multiple clinical studies



**Developing CD19-Redirected ARTEMIS®
T-Cell Therapy for Cancers and
Autoimmune Diseases**

