#### **CAR T and Beyond:** Expanding Horizons for Novel Therapies in Hawai'i and Pacific Islands





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#### Introduction



![](_page_1_Picture_2.jpeg)

![](_page_2_Picture_0.jpeg)

#### No relevant financial relationships to disclose

![](_page_2_Picture_2.jpeg)

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![](_page_3_Picture_0.jpeg)

- I. CAR T Therapy Updates
- II. The Future of CAR T Therapy and Beyond
- III. Racial Disparities in Cancer Care
- IV. Cellular Immunotherapy Program in Hawai'i

![](_page_3_Picture_8.jpeg)

![](_page_3_Picture_9.jpeg)

# I: CAR T cell therapy's mechanism of action is unique as it functions as a "living" drug

![](_page_4_Figure_1.jpeg)

![](_page_4_Picture_2.jpeg)

![](_page_4_Picture_3.jpeg)

The-scientist.com

![](_page_4_Picture_5.jpeg)

## I: CAR T cell therapy revolutionized the treatment of certain cancers...

![](_page_5_Picture_1.jpeg)

## I: ... and continues to be an area of active research and innovation

![](_page_6_Figure_1.jpeg)

- 750 active CAR T therapies in development
- In 2022, FDA approved a new CAR T product, and 2 CAR T therapies for use in 2<sup>nd</sup> line

![](_page_6_Picture_8.jpeg)

# I: There are now 6 FDA approved CAR T products available

Product	Year approved	Indications
Kymriah (tisagenlecleucel)	2017	<ul> <li>Children and youthat is refractor</li> <li>Adult patients with the rapy including lymphoma, and</li> </ul>
Yescarta (axicabtagene ciloleucel)	2017	<ul> <li>Adult patients with including DLBCI arising from fol</li> </ul>
Tecartus (Brexucabtagene autoleucel)	2020	<ul> <li>Adult patients v</li> <li>Adults with relationships</li> </ul>
Breyanzi (lisocabtagene maraleucel)	2021	<ul> <li>Adult patients with including DLBCI primary medias</li> </ul>
Abecma (idecabtagene vicleucel)	2021	Adult patients v     least 4 prior line
Carvykti (ciltacabtagene autoleucel)	2022	Adult patients

- oung adults up to age 25 years of age with B-cell precursor acute lymphoblastic leukemia ry or in second or later relapse
- with relapsed or refractory large B-cell lymphoma after two or more lines of systemic ng diffuse large B-cell lymphoma (DLBCL) not otherwise specified, high grade B-cell d DLBCL arising from follicular lymphoma
- with relapsed or refractory large B-cell lymphoma after first-line chemoimmunotherapy, L NOS, primary mediastinal large B-cell lymphoma, high grade B-cell lymphoma, and DLBCL llicular lymphoma
- with relapsed/refractory mantle cell lymphoma apsed/refractory B-cell acute lymphoblastic leukemia
- with relapsed/refractory large B-cell lymphoma after first-line chemoimmunotherapy, IL NOS (including DLBCL arising from indolent lymphoma), high-grade B-cell lymphoma, stinal large B-cell lymphoma, and follicular lymphoma grade 3B
- with multiple myeloma who have not responded to, or whose disease has returned after, at es (different types) of therapy
- with relapsed/refractory multiple myeloma after four or more prior lines of therapy

![](_page_7_Picture_9.jpeg)

### I: But CAR T therapies still present some limitations

![](_page_8_Picture_1.jpeg)

- Use of CAR T therapies is mostly limited to oncology
- Within oncology, CAR T therapies is only effective against certain types of cancers
- All CAR T therapies are autologous

![](_page_8_Picture_6.jpeg)

![](_page_8_Picture_9.jpeg)

# II: Non-oncology CAR T trials account for < 5% of active trials

- Autoimmune disorders
- Allergy & Asthma
- Infectious diseases (i.e. HBV, HCV, HIV, COVID19 etc)
- Cardiac Fibrosis

![](_page_9_Figure_5.jpeg)

Aghajanian et al Nature 2022

![](_page_9_Picture_7.jpeg)

# II: FDA approved CAR T products only targets CD19 & BCMA in hematologic malignancies

![](_page_10_Figure_1.jpeg)

Fredhutch.org

![](_page_10_Picture_3.jpeg)

![](_page_10_Picture_4.jpeg)

### II: Next-generation CAR therapy

![](_page_11_Figure_1.jpeg)

Slide courtesy of Steve Grupp

![](_page_11_Picture_4.jpeg)

# III: Significant disparities exist in access to novel therapies including clinical trials

![](_page_12_Figure_1.jpeg)

Percentage of Representation in Clinical Trials

![](_page_12_Picture_4.jpeg)

![](_page_12_Picture_5.jpeg)

## III: Majority of patients receiving CAR T therapy are non-Hispanic White patients

![](_page_13_Figure_1.jpeg)

Race/ Ethnicity

Ahmed et al ASTCT 2022

![](_page_13_Picture_4.jpeg)

![](_page_13_Picture_5.jpeg)

# III: Lower SES and distance to treating facility are also barriers to receiving CAR T therapy

Time/Income - Time/Income: 120+ minutes : High.. Time/Income - Time/Income: 120+ minutes : Low.. Time/Income - Time/Income: 31-120 min : High... Time/Income - Time/Income: <30 minutes : High.. Time/Income - Time/Income: 31-120 min : Low... Time/Income - Time/Income: <30 minutes : Low..

> Time - Time : 121 minutes+ Time - Time : 31-120 minutes Time - Time : <30 minutes Race - Asian Race - White Hispanic Race - White Race - Non-White Hispanic Race - Other/Unknown Race - Black Payer - Other Payer Payer - Commercial Payer - Medicaid Payer - Uninsured Payer - Medicare Income - Income: >=\$40K Income - Income: <\$40K Gender - Male Gender - Female Age>=65 Age<65

![](_page_14_Figure_3.jpeg)

Ahmed et al ASTCT 2022

![](_page_14_Picture_6.jpeg)

![](_page_14_Picture_7.jpeg)

### III: Hawai'i and the Pacific Islands is the "perfect storm" for propagating health disparities - Distance

![](_page_15_Figure_1.jpeg)

- Geographic isolation poses unique challenges to our medical landscape
- 2,500 miles from nearest city on mainland U.S.
  - Minimum of 5 hour flight time

![](_page_15_Picture_5.jpeg)

![](_page_15_Picture_6.jpeg)

### III: Hawai'i and the Pacific Islands is the "perfect storm" for propagating health disparities – Population/SES

![](_page_16_Figure_1.jpeg)

INCOME

<b>R</b> 79		Japanese	Filipino	Chinese	Total Population	White	Native Hawaiian
e charts are au's 2019	MEDIAN FAMILY INCOME	\$114,825	\$102,324	\$96,993	\$96,462	\$96,345	\$84,699
o combi- e races." ore than viduals can	PER CAPITA	\$40,940	\$27,738	\$33,454	\$36,989	\$39,499	\$25,612

![](_page_16_Picture_6.jpeg)

![](_page_16_Picture_7.jpeg)

![](_page_16_Picture_8.jpeg)

#### III: ...Despite high potential utilization in Hawai'i and the Pacific Islands

Number of new diagnoses across Hawaii Pacific Health medical system

	2022	2021	2020	2019	2018	Average
<b>B-ALL</b>	9	18	11	14	10	12.4
DLBCL	28	26	28	23	18	24.6
Mantle cell lymphoma	2	4	2	3	2	2.6
Multiple Myeloma	23	15	26	25	25	22.8

Internal data

![](_page_17_Picture_4.jpeg)

# IV: Building a cellular immunotherapy program in Hawai'i

![](_page_18_Picture_1.jpeg)

March 26, 2021

![](_page_18_Picture_3.jpeg)

ZUMA-4 – 1<sup>st</sup> CAR T clinical trial Tisagenlecleucel – 1<sup>st</sup> FDA approved CAR T product

May 22, 2023

![](_page_18_Picture_7.jpeg)

![](_page_18_Picture_8.jpeg)

#### IV: Building a cellular immunotherapy program in Hawai'i

![](_page_19_Figure_1.jpeg)

Revised: 9/1/2023

![](_page_19_Picture_4.jpeg)

![](_page_19_Picture_5.jpeg)

# IV: Building a cellular immunotherapy program in Hawai'i

![](_page_20_Picture_1.jpeg)

March 26, 2021

![](_page_20_Picture_3.jpeg)

ZUMA-4 – 1<sup>st</sup> CAR T clinical trial Tisagenlecleucel – 1<sup>st</sup> FDA approved CAR T product

#### Aug 22, 2023

May 22, 2023

![](_page_20_Picture_8.jpeg)

![](_page_20_Picture_9.jpeg)

# IV: A great outcome for our 1<sup>st</sup> CAR T patient in Hawai'i

![](_page_21_Picture_1.jpeg)

![](_page_21_Picture_3.jpeg)

April 2021

Coronal Volume 2/Volume Ex: 13208 93 Se: 14 P: 93.0 DFOV 99.8 cm 5.00 0.00 50 % PET 3,6/

Sept 2023

![](_page_21_Picture_6.jpeg)

![](_page_21_Picture_7.jpeg)

### IV: Thankful for the opportunity to continue our efforts to bring positive impact to our community

![](_page_22_Picture_1.jpeg)

![](_page_22_Picture_2.jpeg)

![](_page_22_Picture_3.jpeg)

#### IV: 5- & 10-year strategic vision/goals

#### Short-term goals:

- Expand clinical trials
- Broaden cellular immunotherapy portfolio
- Enhance community outreach
- Recruitment
- Health disparity assessment

#### Long-term goals:

- Establish biorepository
- Strengthen clinical and translational research capabilities
- National collaboration
- Health equity

![](_page_23_Picture_12.jpeg)

## Thank you!

![](_page_24_Picture_1.jpeg)

![](_page_24_Picture_2.jpeg)