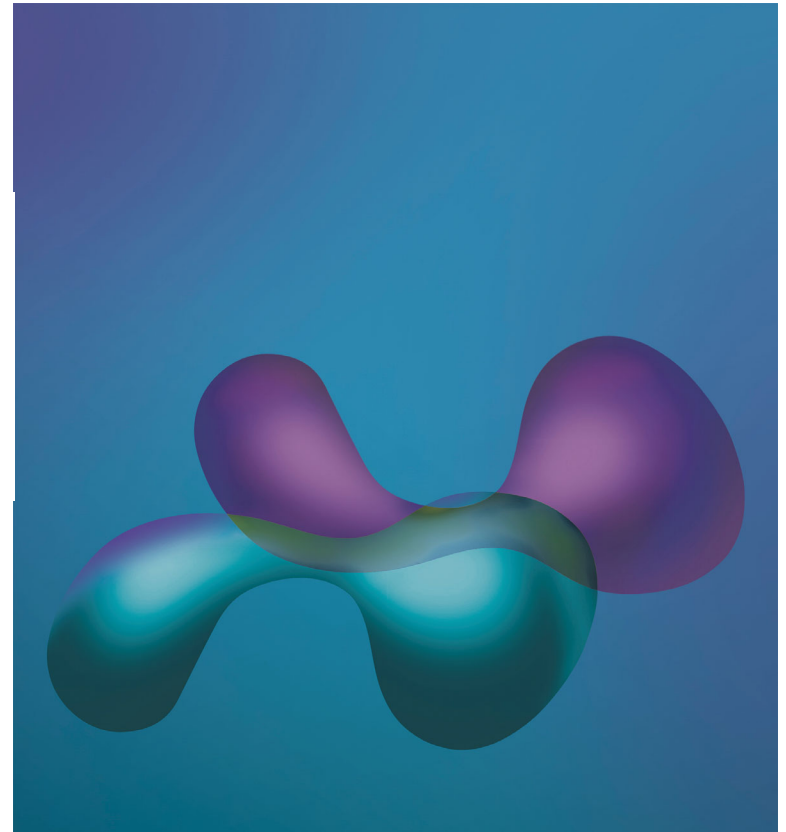




# EFFICACY AND SAFETY OF TABELECLEUCEL IN PATIENTS WITH EPSTEIN-BARR VIRUS-ASSOCIATED LEIOMYOSARCOMAS (EBV<sup>+</sup> LMS)

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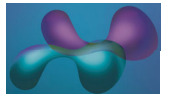


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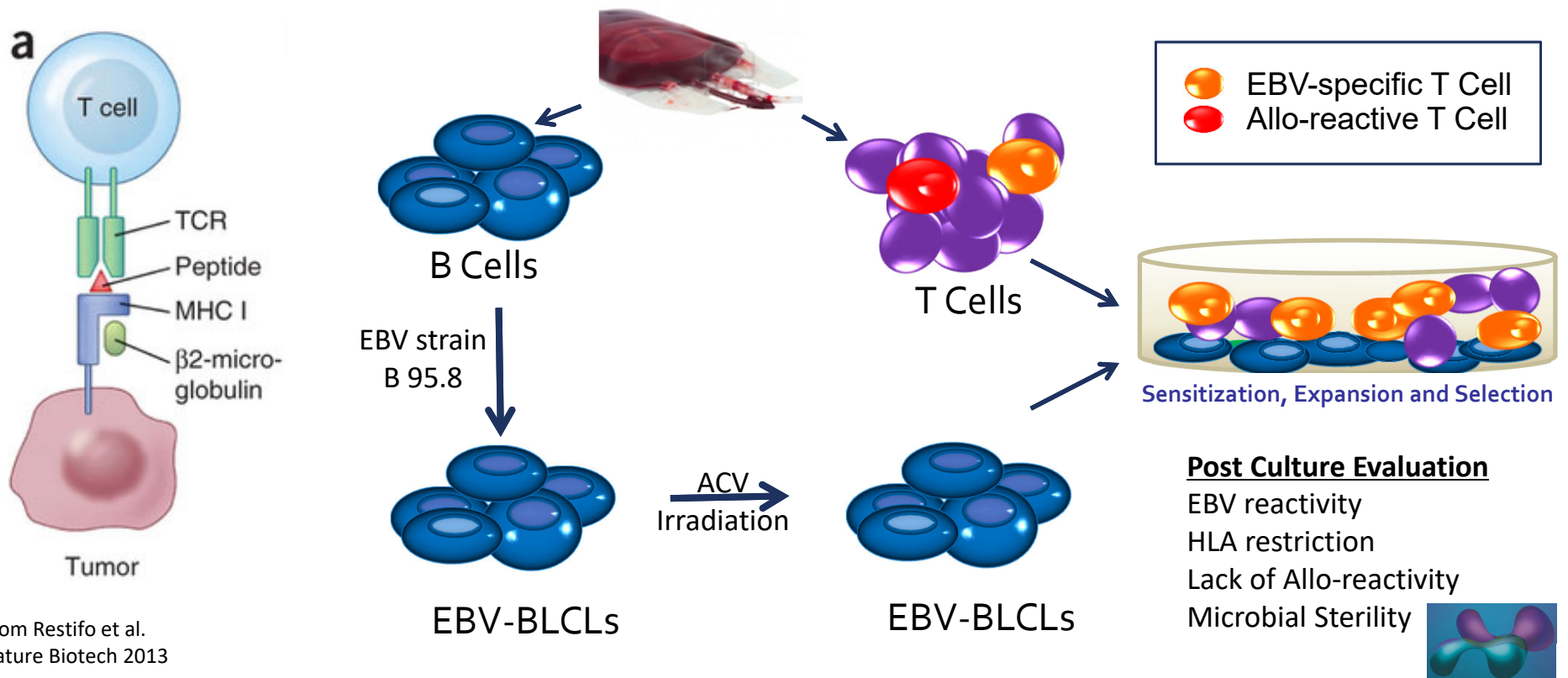
## Disclosures

- I have no personal disclosures.
- Atara Biotherapeutics holds the license and IND to develop and commercialize the third party EBV cell therapy program (Tabelecleucel) I will be discussing today.
- MSKCC and several investigators at MSK hold an interest in Atara Biotherapeutics.
- Akshay Sudhindra, Wei Ye, Minoti Hiremath, and Willis Navarro are employees and stockholders of Atara Biotherapeutics.



# Tabelecleucel Manufacturing

Investigational, off-the-shelf, genetically unmodified, allogeneic T-cell immunotherapy that targets EBV antigens



# EBV<sup>+</sup> Leiomyosarcoma Subset

## Disease Background

|                              |  |
|------------------------------|--|
| <b>Incidence</b>             | <ul style="list-style-type: none"> <li>Very Rare EBV<sup>+</sup> Disease</li> <li>0.7/1000 pt-yrs in SOT recipients<sup>1</sup></li> </ul>                           |
| <b>Tumor Character</b>       | <ul style="list-style-type: none"> <li>Stromal cell soft tissue sarcoma</li> <li>Typically aggressive</li> <li>Exhibit EBV latency II pattern</li> </ul>             |
| <b>Established Therapies</b> | <ul style="list-style-type: none"> <li>Responds poorly to radiation and chemotherapy resulting in limited treatment options and poor outcomes<sup>2</sup></li> </ul> |
| <b>Survival</b>              | <ul style="list-style-type: none"> <li>Median OS 19.9, 7.6, and 0.9 months for patients with 0, 1 or 2, and ≥3 risk factors<sup>2*</sup></li> </ul>                  |

<sup>1</sup>Stubbins RJ et al. Transpl Infect Dis. 2018:e13010.

<sup>2</sup>Wang Z et al. Cancer Med. 2016;5(12):3437-3444.

\*Hemoglobin <10 g/dL, BMI<30 kg/m<sup>2</sup>, albumin <3.5 g/dL, and neutrophilia.

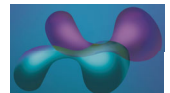


## Patient Demographics

|                           |   |
|---------------------------|---|
| <b>Number of Patients</b> | 12 <sup>a,b</sup>   |
| 95-024                    | 5   |
| 11-130                    | 3   |
| 201                       | 4   |
| <b>Age Group</b>          |   |
| <16 / ≥ 16 years          | 8 (66.7%) / 4 (33.3%)   |
| <b>Group</b>              |   |
| SOT                       | Heart (3), kidney (2), small bowel (1 <sup>b</sup> ), multi-visceral (1 <sup>a</sup> ), HCT followed by lung and kidney (1) |
| Other ID                  | 3   |

<sup>a</sup>1 patient treated on both 95-024 and 11-130.

<sup>b</sup>1 patient received autologous EBV-CTLs and tabellecleucel.



# Tabelecleucel Treatment and Safety Results - EBV<sup>+</sup> LMS Patients

## Tabelecleucel Treatment

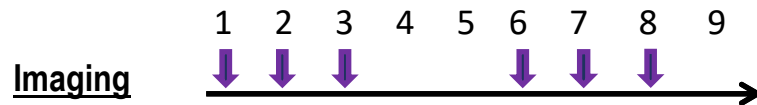
## Safety Data

### HLA Matching

- Tabelecleucel was matched for at least 2/8 HLA alleles at high resolution and restricted by an HLA allele shared by the patient

### Tabelecleucel Administration

- 1 dose per week x 3 of  $1-2 \times 10^6$  cells/kg/dose every 4-6-week cycle



- Performed before the 1<sup>st</sup> dose of each cycle

### Key Assessments

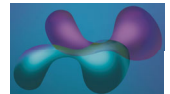
- Efficacy: Overall Response Rate as determined by RECIST 1.1  
Overall Survival
- Safety: Studies are ongoing, so safety assessments reported herein are based on SAEs collected by the data snapshot of Nov 16<sup>th</sup>, 2018

### Tabelecleucel Administration

Median number of cycles of tabelecleucel: 6 (range: 1 - 15) months

Median treatment duration: 7.5 (range: 1.4 – 52.5) months

- Treatment-emergent SAEs (TESAEs)
  - 6 of 12 patients (50%) with TESAEs
- Events occurred once, except pyrexia (n = 2)
- The events observed in this subset of patients are reflective of a population of ill patients
- No TESAE was assessed as related to treatment
- No TESAEs led to treatment discontinuation
- No specific safety trends were observed

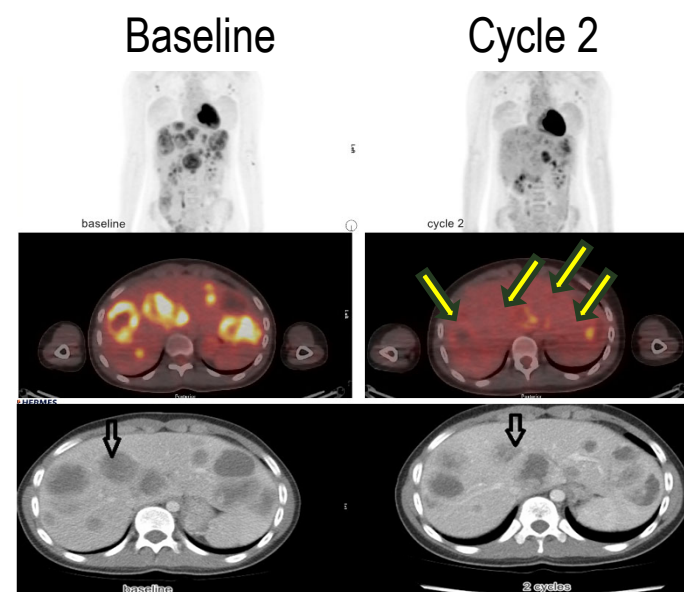


# Response to Tabelecleucel in Patients Treated for EBV<sup>+</sup> LMS (RECIST 1.1)

| N                           | 12                       |
|-----------------------------|--------------------------|
| ORR (95% CI)                | 16.7% (2.1%,48.4%)       |
| CR                          | 0                        |
| PR                          | 2                        |
| SD                          | 8                        |
| PD                          | 0                        |
| NE*                         | 2                        |
| Median OS (months)          | 77.4 (95% CI: 18, NE)    |
| Median Follow-Up (months)   | 22.5 (Range: 2.8, 109.3) |
| Longest Time to PD (months) | 15.5 (2.8 – 65.4)        |

- Immune response criteria were not used.
- Metabolic responses were observed (by PET) in 3 of 4 (75%) patients on study 201.
- \*Not Evaluable – 1 patient received tabelecleucel and autologous product.  
1 patient was too early to evaluate.

## 15 yo female w/ LMS post- heart transplant



Patient switched tabelecleucel product due to limited availability and eventually progressed at Cycle 3.



## Conclusions

1. This analysis represents one of the larger prospective studies of patients with EBV<sup>+</sup> LMS.
2. Tabelecleucel available as an off-the-shelf cellular therapy to treat patients with EBV<sup>+</sup> LMS, a disease for which there are no established treatment options.
3. The combination of CT-based and metabolic responses to tabelecleucel, in the context of prolonged survival, demonstrate that tabelecleucel may provide clinical benefit in this typically radiation- and chemotherapy-resistant disease.
4. The safety profile of tabelecleucel appears consistent with a favorable risk profile and our previous observations.

