# VALUE OF AXICABTAGENE CILOLEUCEL VERSUS CHEMOTHERAPY IN THE LARGE B-CELL LYMPHOMA TREATMENT: HEALTH OUTCOMES BASED ON THE NUMBER OF PATIENTS TREATED IN SPAIN

Córdoba R<sup>1</sup>, López-Corral L<sup>2</sup>, Presa M<sup>3</sup>, Martín-Escudero V<sup>4</sup>, Casado MÁ<sup>3</sup>, Pardo C<sup>4</sup>

¹Hospital Universitario Fundación Jiménez Díaz, Madrid, España; ²Hospital Universitario de Salamanca, España; ³Pharmacoeconomics & Outcomes Research Iberia (PORIB), Madrid, España; ⁴Gilead Sciences, Madrid, España

#### INTRODUCTION

- Diffuse large B-cell lymphoma (DLBCL) is the most common type of malignant lymphoma worldwide<sup>1</sup>. Approximately 30-40% of DLBCL will develop relapsed/refractory (R/R) disease<sup>1,2</sup>, for whom the historical median overall survival (OS) is approximately 6 months<sup>3</sup>.
- Axicabtagene ciloleucel (axi-cel), a chimeric antigen receptor (CAR)-T cell therapy, has demonstrated promising results in terms of survival and response rates<sup>4</sup>, however the actual number of patients treated in Spain differs from epidemiology estimations.

#### **OBJECTIVE**

The study aimed to assess the value of axi-cel versus rituximabbased chemotherapy (CT) in the treatment of DLBCL R/R and primary mediastinal large B-cell lymphoma (PMBCL) after ≥2 lines of systemic therapy, in Spain, based on the number of patients treated.

### **METHODS**

- In order to estimate the life years gained (LYG) and quality-adjusted life years (QA-LYs) for axi-cel and CT over a lifetime horizon, a partitioned survival mixture-cure model (PS-MCM) was used. The model, with monthly duration cycles, comprised the following health states: pre-progression, post-progression and death.
- The efficacy data for axi-cel was obtained from the ZUMA-1 trial after 4-year follow-up<sup>5</sup>. CT OS was derived from the SCHOLAR-1 study<sup>3</sup>, after adjust the baseline characteristics of patients by a propensity score matching. As progression-free survival (PFS) was not collected in SCHOLAR-1, CT PFS was estimated based on PFS/OS axi-cel ratio derived from ZUMA-1.
- Axi-cel OS and PFS curves were extrapolated beyond the follow-up period by a lognormal and Gompertz distributions, respectively. OS curve for CT was adapted to a Gompertz distribution.
- The utility value assigned for pre-progression state (0.72) were obtained from ZUMA-16 and for post-progression state (0.39) from literature<sup>7</sup>. Under a conservative approach, a utility decrement of -0.05 was associated to the axi-cel related toxicities<sup>6</sup>.
- In the base case, the incremental results of axi-cel versus CT were evaluated in a cohort of 183 patients, according to the number of cases treated reported in the "Spanish Plan to Tackle Advanced Therapies" (from May 2020 to June 2021)8.
- In an alternative scenario, the results were analysed in a cohort of 490 patients, according to the number of patients with DLBCL eligible to be treated with axi-cel in Spain<sup>9</sup>.
- Assumptions and parameters were validated by an expert panel in the haemato-oncology field.

#### RESULTS

• In the base case (n=183), over a lifetime horizon, axi-cel versus CT would generate an increase of 162% in LYG (1,313 LYG) and a 165% in QALYs (1,030 QALYs) (Table 1).

Table 1. Base case results (n=183 patients)

| Health Outcomes                 | Axi-cel | Chemotherapy | Incremental |
|---------------------------------|---------|--------------|-------------|
| TOTAL LYG                       | 2,122   | 809          | 1,313       |
| LYG in pre-progression state    | 2,008   | 752          | 1,256       |
| LYG in post-progression state   | 114     | 57           | 56          |
| TOTAL QALYs                     | 1,654   | 624          | 1,030       |
| QALYs in pre-progression state  | 1,619   | 602          | 1,017       |
| QALYs in post-progression state | 44      | 22           | 22          |
| QALY decrements due to AEs      | -9      | 0            | -9          |

AE, adverse event; Axi-cel, axicabtagene ciloleucel; QALY, quality-adjusted life year; LYG, life years gained.

• In the alternative scenario (n=490), the results could increase to 3,515 LYG and 2,759 QALYs with axi-cel versus CT (Table 2).

Table 2. Alternative scenario results (n=490 patients)

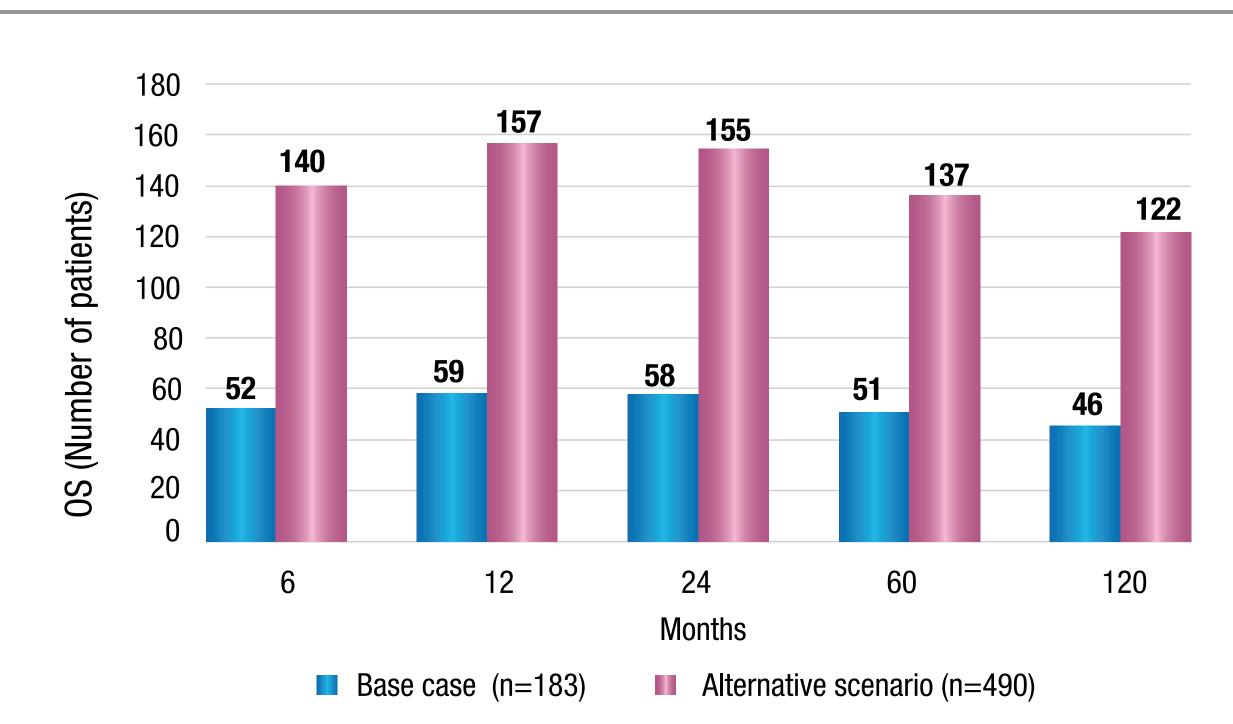
| Health Outcomes                 | Axi-cel | Chemotherapy | Incremental |
|---------------------------------|---------|--------------|-------------|
| TOTAL LYG                       | 5,681   | 2,166        | 3,515       |
| LYG in pre-progression state    | 5,377   | 2,013        | 3,363       |
| LYG in post-progression state   | 304     | 153          | 151         |
| TOTAL QALYs                     | 4,430   | 1,671        | 2,759       |
| QALYs in pre-progression state  | 4,335   | 1,611        | 2,724       |
| QALYs in post-progression state | 119     | 60           | 59          |
| QALY decrements due to AEs      | -25     | 0            | -25         |

AE, adverse event; Axi-cel, axicabtagene ciloleucel; QALY, quality-adjusted life year; LYG, life years gained.

Consequently, treating with axi-cel versus CT the entire population eligible to receive axi-cel (n=490) compared to the currently treated population (n=183), would result in an increase of 2,202 LYG and 1,728 QALYs.

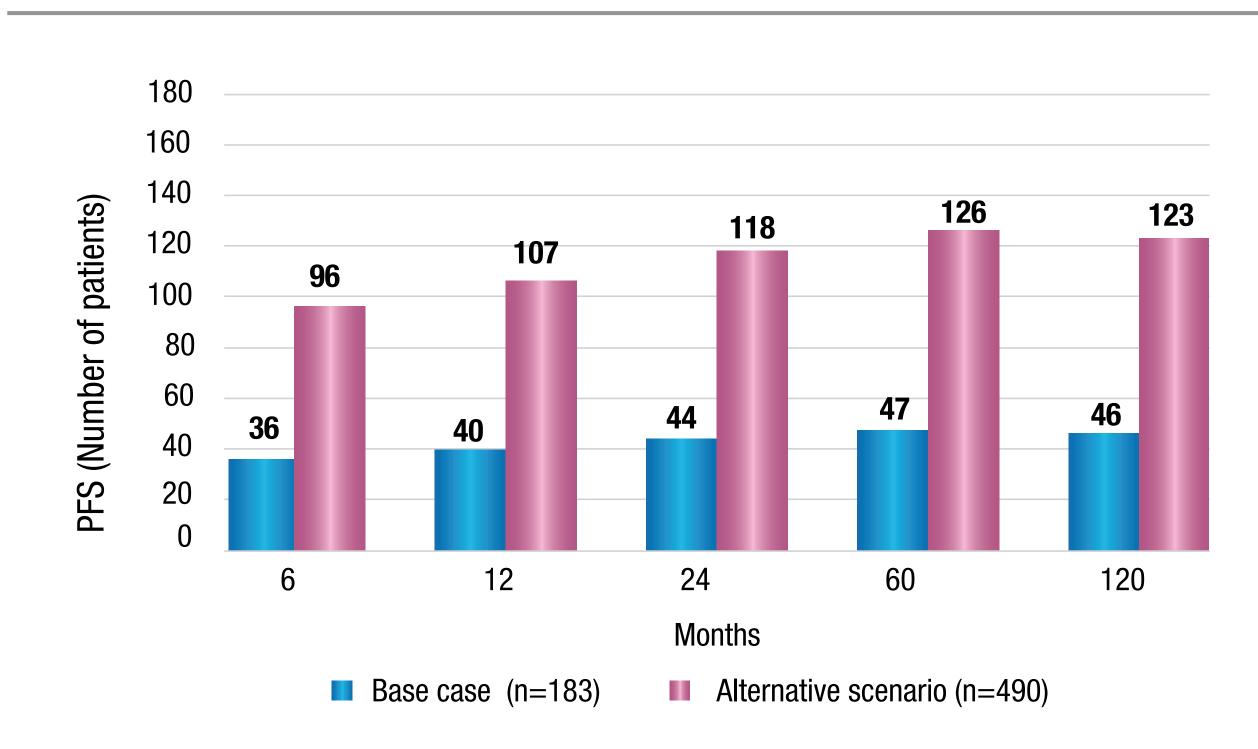
• Considering the modelled efficacy data, axi-cel versus CT would increase, at 5 years, the number of patients in PFS by 195% (n=47 in the base case and n=126 in the alternative scenario) and the number of surviving patients by 194% (n=51 in the base case and n=137 in the alternative scenario) (Figure 1 and Figure 2).

Figure 1. Incremental number of patients in OS\*



\*Incremental number of patients with axi-cel versus CT in OS in the base case considering a cohort of 183 patients, and in the alternative scenario considering 490 patients. OS, overall survival.

Figure 2. Incremental number of patients in PFS\*



\*Incremental number of patients with axi-cel versus CT in PFS in the base case considering a cohort of 183 patients, and in the alternative scenario considering 490 patients. PFS, progression-free survival.

## REFERENCES

1. Tilly H, et al. Ann Oncol. 2015;26:116-25. 2. Friedberg JW, et al. Hematology Am Soc Hematol Educ Program. 2011:2011:498-505

3. Crump M et al. Leukemia & lymphoma. 2018;59(7):1700-9. 4. Locke FL, et al. Lancet Oncol. 2019;20(1):31-42.

6. Lin V, et al. 44th Annual Meeting of the European Society for Blood and

Marrow Transplantation, 2018 7. Chen Q et al. Leukemia & lymphoma. 2018;59(7):1700-9. 8. MoH. https://www.sanidad.gob.es/profesionales/farmacia/Terapias\_Avan-

9. GENESIS-SEFH. https://gruposdetrabajo.sefh.es/genesis/genesis/Enlaces/InformesHosp\_abc.htm?ml=1#A

## DISCLOSURES

5. Jacobson Cet al. Blood. 2021;138:1764

RC has received consulting fees from Abbvie, Astra Zeneca, Beigene, Bristol-Myers Squibb, Genmab, Incyte, Janssen, Kite, Kyowa-Kirin, Lilly, Roche and Takeda; conference fees from Abbvie, Astra Zeneca, Janssen, Kite and Takeda; and research fees from Pfizer. LLC has received consulting and speaking fees from Novartis and Kite. MP and MAC are employees of Pharmacoeconomics & Outcomes Research Iberia (PORIB), a company that received consultancy fees from Gilead for carrying out the project. CP and VME are employees of Gilead Sciences Spain.

## CONCLUSIONS

Since axi-cel would significantly improve health outcomes versus CT, it is crucial to make efforts to increase the number of patients treated with axi-cel in Spain.



