# US Cost-Effectiveness of Chimeric Antigen Receptor T-Cell Therapy for Patients With Relapsed or Refractory Large B-Cell Lymphoma, Considering Infusion Setting and Payor Claims Data

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# INTRODUCTION

- Chimeric antigen receptor (CAR) T-cell therapies have changed the treatment paradigm for patients with relapsed/refractory (R/R) large B-cell lymphoma (LBCL)
- Three anti-CD19 CAR T-cell therapies are approved by the United States Food and Drug Administration (FDA) for the treatment of adult patients with R/R LBCL after 2 or more lines of systemic therapy:
- Axicabtagene ciloleucel (axi-cel; YESCARTA®)¹,
- Lisocabtagene maraleucel (liso-cel; BREYANZI®)², and
   Tisagenlecleucel (tisa-cel; KYMRIAH®)³

# **OBJECTIVE**

 This payor-perspective model compared lifetime costs and benefits for patients with R/R LBCL treated with CAR T-cell therapy in the United States by therapy (axi-cel versus liso-cel and axi-cel versus tisa-cel) and considered impact of CAR T-cell site of care (inpatient versus outpatient)

#### DATA AND METHODS

#### **Model Structure**

 A decision-tree model compared lifetime direct healthcare costs and benefits between axi-cel and tisa-cel and between axi-cel and liso-cel

#### **Cost Estimation**

- Average per-patient lifetime costs (2020 United States dollars) include
- CAR T-cell-related and other relevant pre-infusion costs (t≤0 days; apheresis, bridging therapy, lymphodepletion chemotherapy, CAR T-cell acquisition/administration)
- Near-term post-infusion costs (0≤t≤90 days; intensive care unit [ICU] and non-ICU inpatient hospitalization, emergency department visits, outpatient/other visits)
- 3. Distant post-infusion costs (t>90 days; pre-/post-progression routine care applied to surviving patients)
- End-of-life costs (t>3 days; palliative/other end-of-life care for newly deceased patients)
- 5. Other post-infusion costs (0<t≤365 days; post-infusion stem cell transplant [SCT] and first-year intravenous immune globulin use)

## Quality-Adjusted Life-Year (QALY) Estimation

- Benefits were determined for each CAR T-cell therapy based on health utilities and partitioned survival models developed using matching-adjusted indirect comparison (MAIC) overall and progression-free survival curves<sup>4,5</sup>
- These matched survival curves were used to model long-term costs and OALYs

# DATA AND METHODS (Continued)

#### Data

#### Table 1. Short-Term Resource Use Model Input Values

	Mean 90-day resource use per patient with				
	CRS	NE	Neither		
CAR T-cell site of care: Inpatient					
ICU, days	2.08	2.32	2.06		
Non-ICU inpatient, days	17.16	16.83	16.97		
ED, visits	0.06	0.12	0.05		
Outpatient/other, visits	5.04	4.70	5.04		
CAR T-cell site of care: Outpatient					
ICU, days	2.35	2.01	1.71		
Non-ICU inpatient, days	7.81	4.76	11.71		
ED, visits	0.01	0.02	0.00		
Outpatient/other, visits	3.38	2.16	6.00		

CAR, chimeric antigen receptor; CRS, cytokine release syndrome; ED, emergency department; ICU, intensive care unit; NE, neurologic event.

Model input values related to short-term post-infusion healthcare resource use (Table 1) and CAR T-cell site of care
were informed by analyses of insurance claims data (Anlitiks All-Payer Claims) for 1,175 adult R/R LBCL patients
with Commercial, Medicaid, Medicare Fee-for-Service, and Medicare Advantage coverage

Table 2. Select Treatment-Related Model Input Values

	Axi-ce	Axi-cel vs		
	Liso-cel	Tisa-cel	Liso-cel	Tisa-cel
CAR T-cell acquisition cost/patient, \$	\$399,000	\$399,000	\$410,300	\$373,000
Receiving bridging therapy, %	0.0%	0.0%	59.1%	91.9%
Receiving post-infusion SCT, %	7.9%	7.9%	7.6%	5.4%
Grade ≥3 AE incidence, %ª				
CRS	9.0%	9.3%	2.6%	17.1%
NE	28.8%	27.1%	14.4%	10.5%
Receiving IVIG, %	30.6%	30.6%	21.0%	30.0%

• Other model input values were based on the ZUMA-1 (axi-cel), TRANSCEND (liso-cel), and JULIET (tisa-cel) clinical

AE, adverse event; axi-cel, axicabtagene ciloleucel; CAR, chimeric antigen receptor; CRS, cytokine release syndrome; IVIG, intravenous immune globulin; liso-cel, lisocabtagene maraleucel; NE, neurologic event;

 Other model input values were based on the ZUMA-1 (axi-cel), TRANSCEND (liso-cel), and JULIET (tisa-cel) clinic studies, and other published literature and publicly available data<sup>6-8</sup> (Table 2)

#### Scenario and Sensitivity Analyses

- Base-case inputs specific to axi-cel were based on ZUMA-1 Cohorts 1 and 2. Scenario analyses using data from ZUMA-1 Cohort 49 and ZUMA-1 Cohort 610 evaluated effects of alternative safety protocols
- In 2 additional scenario analyses, the share of patients who received CAR T-cell therapy in an outpatient site of care varied for all CAR T-cell therapies simultaneously (base: 17%; scenarios: 0% and 34%)
- Deterministic sensitivity analyses were performed to account for uncertainty in model parameters

#### **Results Presentation**

SCT, stem cell transplant; tisa-cel, tisagenlecleucel

 Payor costs and health benefits (both discounted at 3% annually) were used to estimate incremental net monetary benefits (INMBs; QALYs valued at \$150,000) and incremental cost-effectiveness ratios for axi-cel through direct comparisons with liso-cel and tisa-cel owing to use of MAIC-matched input values specific to pair-wise comparisons

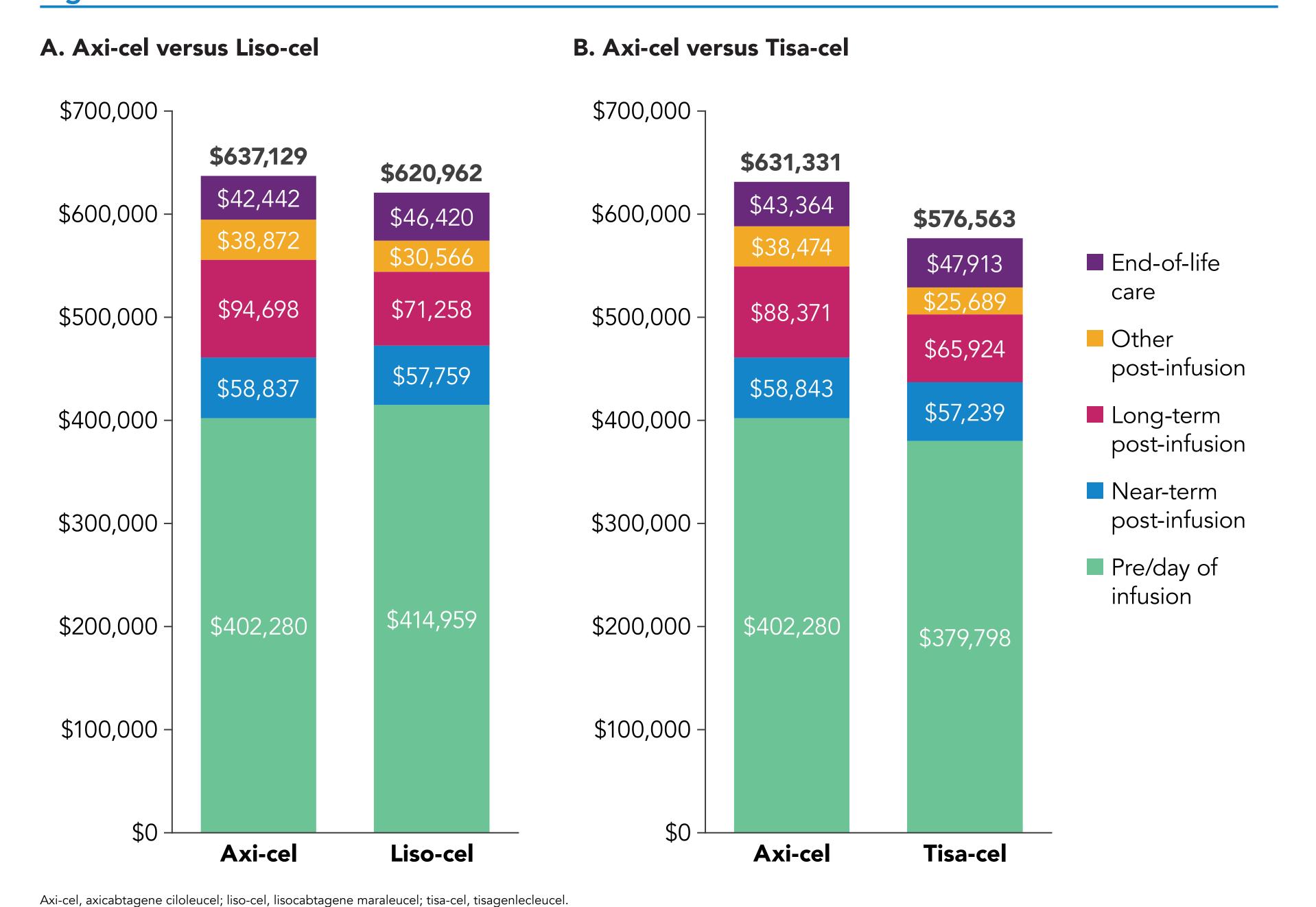
# **RESULTS**

Table 3. Cost-Effectiveness Base Case Results

	Axi-cel versus Liso-cel			Axi-cel versus Tisa-cel		
	Axi-cel [A]	Liso-cel [B]	Δ [A]-[B]	Axi-cel [C]	Tisa-cel [D]	Δ [C]-[D]
Total direct medical costs	\$637,129	\$620,962	\$16,167	\$631,331	\$576,563	\$54,769
Total quality-adjusted life, years	7.705	5.898	1.807	7.240	5.005	2.235
On CAR T-cell therapy, t≤30 days	0.061	0.060	<0.001	0.061	0.060	<0.001
Off therapy, t>30 days	7.644	5.838	1.807	7.180	4.945	2.235
pre-progression	7.452	5.717	1.736	6.954	4.661	2.293
post-progression	0.192	0.121	0.071	0.226	0.284	-0.058
Net monetary benefit	\$518,624	\$263,711	\$254,913	\$454,719	\$174,246	\$280,472
ICER (axi-cel versus comparator)	-	-	\$8,946	-	-	\$24,506

Axi-cel, axicabtagene ciloleucel; CAR, chimeric antigen receptor; ICER, incremental cost-effectiveness ratio; liso-cel, lisocabtagene maraleucel; tisa-cel, tisagenlecleucel.

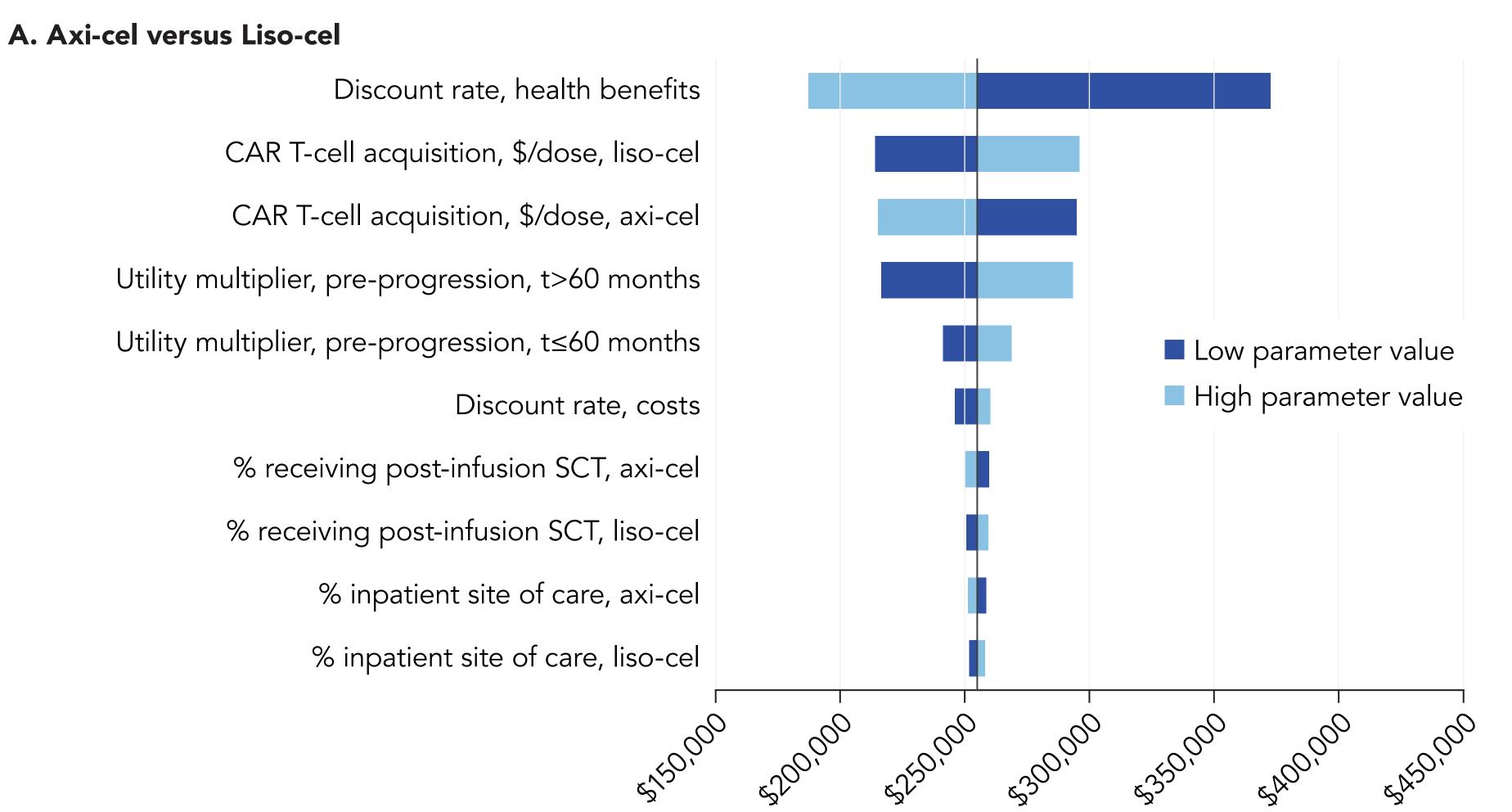
#### Figure 1. Base Case Costs



• Total lifetime direct healthcare costs and QALYs for axi-cel exceeded those for liso-cel and tisa-cel largely due to longer patient lifespans (**Table 3** and **Figure 1**)

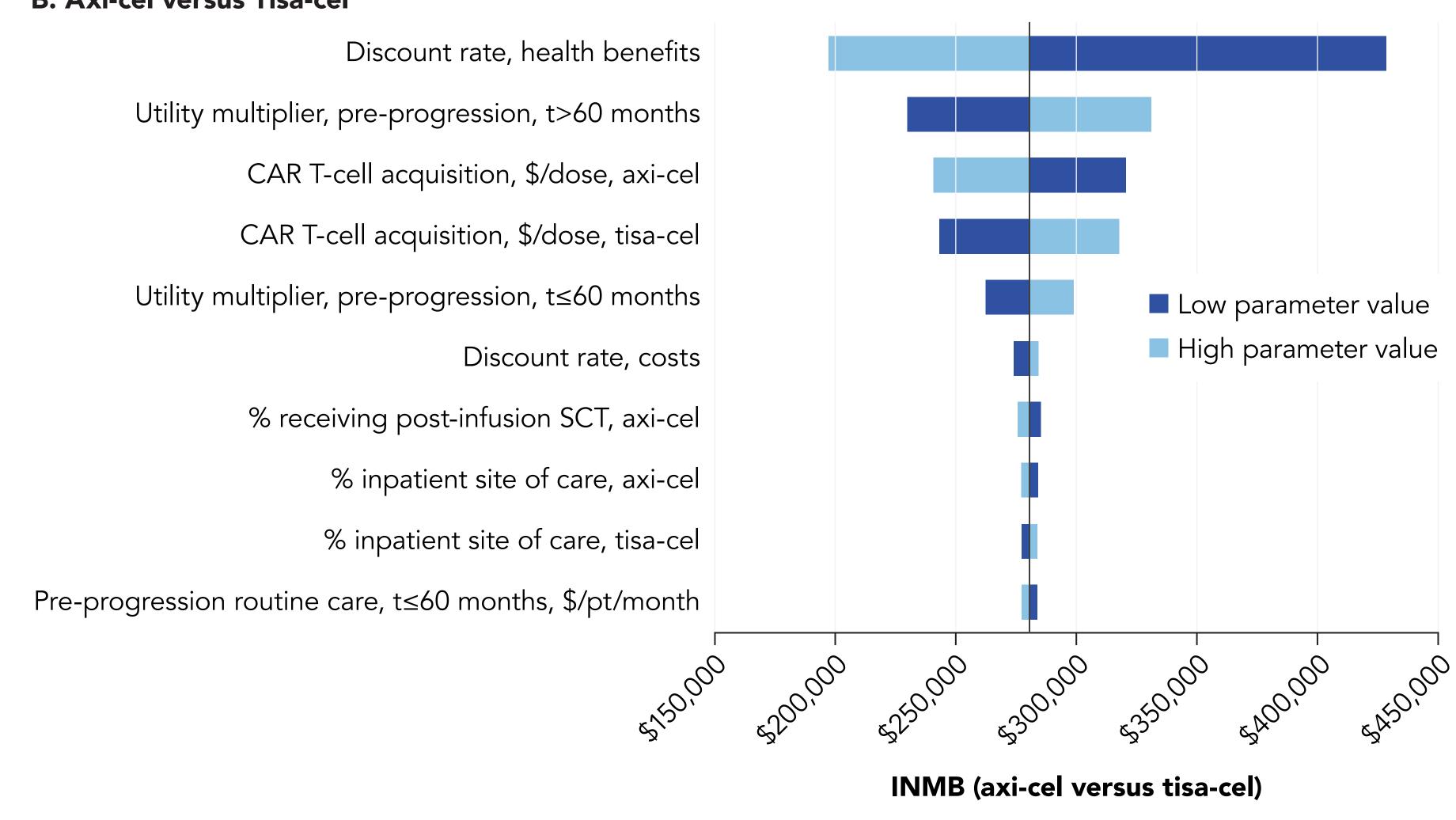
- Base case INMB for axi-cel was \$255,000 compared to liso-cel and \$280,000 compared to tisa-cel
- For axi-cel, incremental costs per QALY gained were \$9,000 versus liso-cel and \$25,000 versus tisa-cel
- Results were not sensitive to alternative outpatient proportions applied to all CAR T-cell therapies or to changes
  in Grade ≥3 cytokine release syndrome/neurologic event rates and other input values for axi-cel consistent with
  different ZUMA-1 cohorts

## Figure 2. Deterministic Sensitivity Analyses



INMB (axi-cel versus liso-cel)

#### B. Axi-cel versus Tisa-cel



 Results were most sensitive to changes in the discount rate applied to health benefits, CAR T-cell acquisition costs, and pre-progression utility multipliers (Figure 2)

Axi-cel, axicabtagene ciloleucel; CAR, chimeric antigen receptor; INMB, incremental net monetary benefits; liso-cel, lisocabtagene maraleucel; pt, patient; SCT, stem cell transplant; tisa-cel, tisagenlecleucel.

 However, in all scenarios and sensitivities, axi-cel was cost-effective versus both comparator therapies at a maximum willingness-to-pay of under \$33,000/QALY

# CONCLUSIONS

- Axi-cel is a cost-effective CAR T-cell option for patients with R/R LBCL compared to tisa-cel and liso-cel
- Treatment site of care does not impact the cost-effectiveness of CAR T-cell treatment

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#### DISCLOSURES

AKCJ: employment with Medicus Economics; consulting or advisory role, through Medicus Economics (a health care consulting firm), for AbbVie, Acadia, Agios, Albireo, Alexion, Apellis, Argenx, Astellas, BeyondSpring, Boehringer-Ingelheim, Biohaven, Celgene, Deciphera, Drexel, F2G, Genentech, Include Health, Kite, Madrigal, Oncopeptides, Penn, Pfizer, PharmaEssentia Rallybio, Reata, Regeneron, Sage, Sanofi, Sarepta, Spark, Sunovion, and Vertex; and research funding from Medicus Economics LLC, which was paid fees by Kite to conduct the research for this analysis. JTS: employment with Kite, and stock or other ownership in Gilead. **SWW:** consulting or advisory role for Kite Pharma, Amgen, Allergan/Abbvie, and Johnson & Johnson. STW: employment with Medicus Economics; consulting or advisory role, through Medicus Economics (a health care consulting firm), for AbbVie, Acadia, Agios, Albireo, Alexion, Apellis Argenx, Astellas, BeyondSpring, Boehringer-Ingelheim, Biohaven, Celgene, Deciphera, Drexel, F2G, Genentech, Include Health, Kite, Madrigal, Oncopeptides, Penn, Pfizer, PharmaEssentia, Rallybio, Reata, Regeneron, Sage, Sanofi, Sarepta, Spark, Sunovion, and Vertex; and research funding from Medicus Economics LLC, which was paid fees by Kite to conduct the research for this analysis. MGB: employment with Medicus Economics; consulting or advisory role, through Medicus Economics (a health care consulting firm), for AbbVie, Acadia, Agios, Albireo Alexion, Apellis, Argenx, Astellas, BeyondSpring, Boehringer-Ingelheim, Biohaven, Celgene, Deciphera, Drexel, F2G, Genentech, Include Health, Kite, Madrigal, Oncopeptides, Penn, Pfizer, PharmaEssentia, Rallybio, Reata, Regeneron, Sage, Sanofi, Sarepta, Spark, Sunovion, and Vertex and research funding from Medicus Economics LLC, which was paid fees by Kite to conduct the research for this analysis. **SJ:** consulting or advisory role, through Medicus Economics (a health care consulting firm), for AbbVie, Acadia, Agios, Albireo, Alexion, Apellis, Argenx, Astellas, Regeneron, Sage, Sanofi, Sarepta, Spark, Sunovion, and Vertex; and research funding from Medicus Economics LLC, which was paid fees by Kite to conduct the research for this analysis **UG:** employment with Thomas Jefferson University; stock or other ownership in Iovance, Pfizer, Moderna, and Gamida; consulting or advisory role for Jazz, Gamida, Mesoblast, and Incyte; and speakers' bureau participation for Jazz, Astellas, Incyte, Kite.

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