

Real-World Comparative Study of the Effectiveness, Safety, and Health Care Resource Utilization of Lisocabtagene Maraleucel and Bispecific Antibodies in Third-Line or Later Large B-Cell Lymphoma

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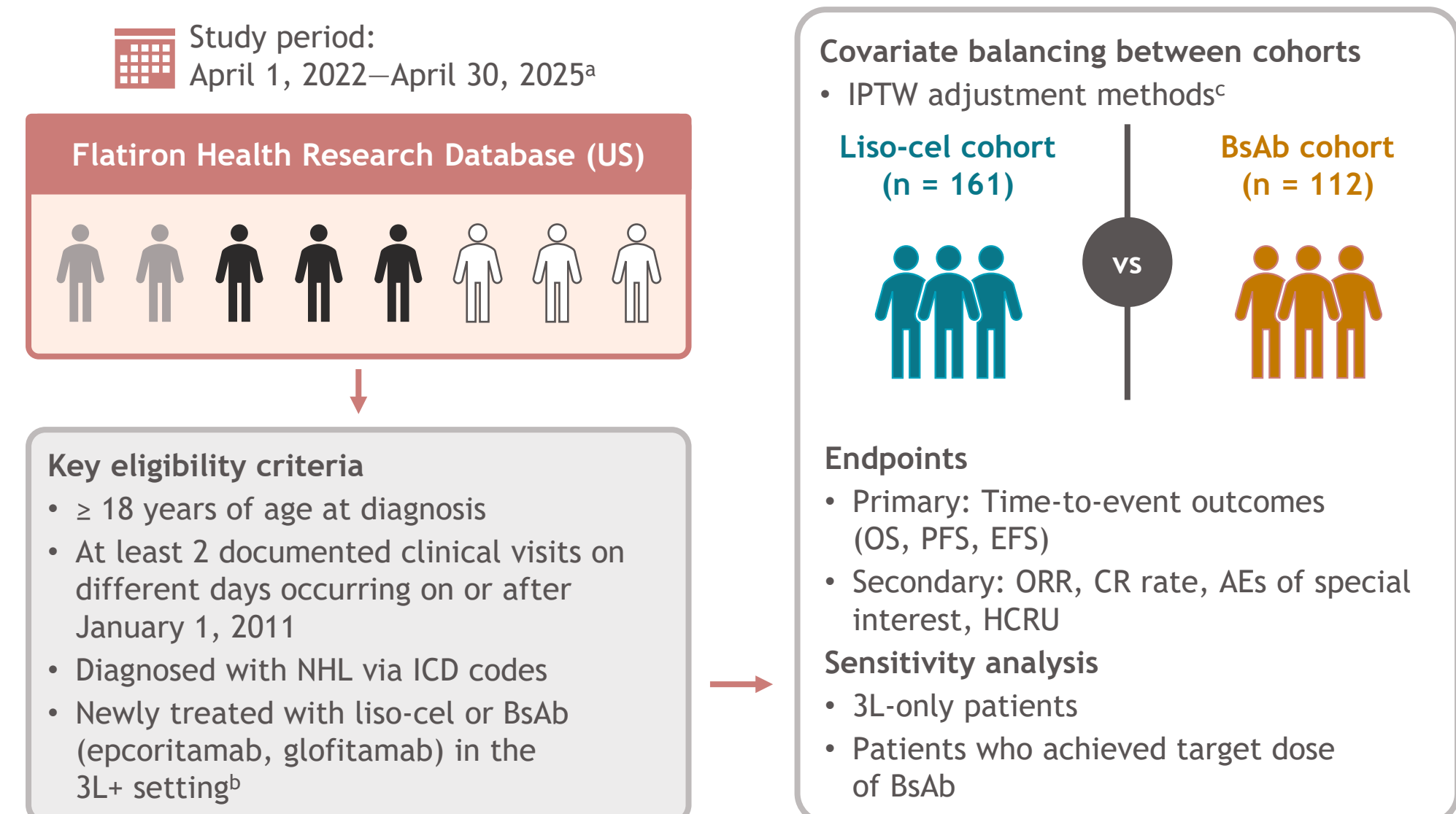
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Background

- There are many therapeutic options available for patients with R/R large B-cell lymphoma (LBCL) in the third-line (3L) or later (3L+) setting, making treatment decisions challenging. Two of the available options include treatment with lisocabtagene maraleucel (liso-cel), an autologous, CD19-directed, 4-1BB CAR T cell product, or with a CD3 × CD20 T-cell–engaging bispecific antibody (BsAb)
- Liso-cel is approved by the FDA and EMA in second-line or later LBCL^{1,2}
- Epcoritamab and glofitamab are BsAbs approved by the FDA and EMA in 3L+ LBCL^{3–6}
- Understanding the comparative effectiveness, safety, and health care resource utilization (HCRU) of liso-cel and BsAb in the real-world clinical setting can inform optimal treatment selection
- Here, we compare the real-world effectiveness, safety, and HCRU among patients with R/R LBCL treated with liso-cel or BsAb in the 3L+ setting

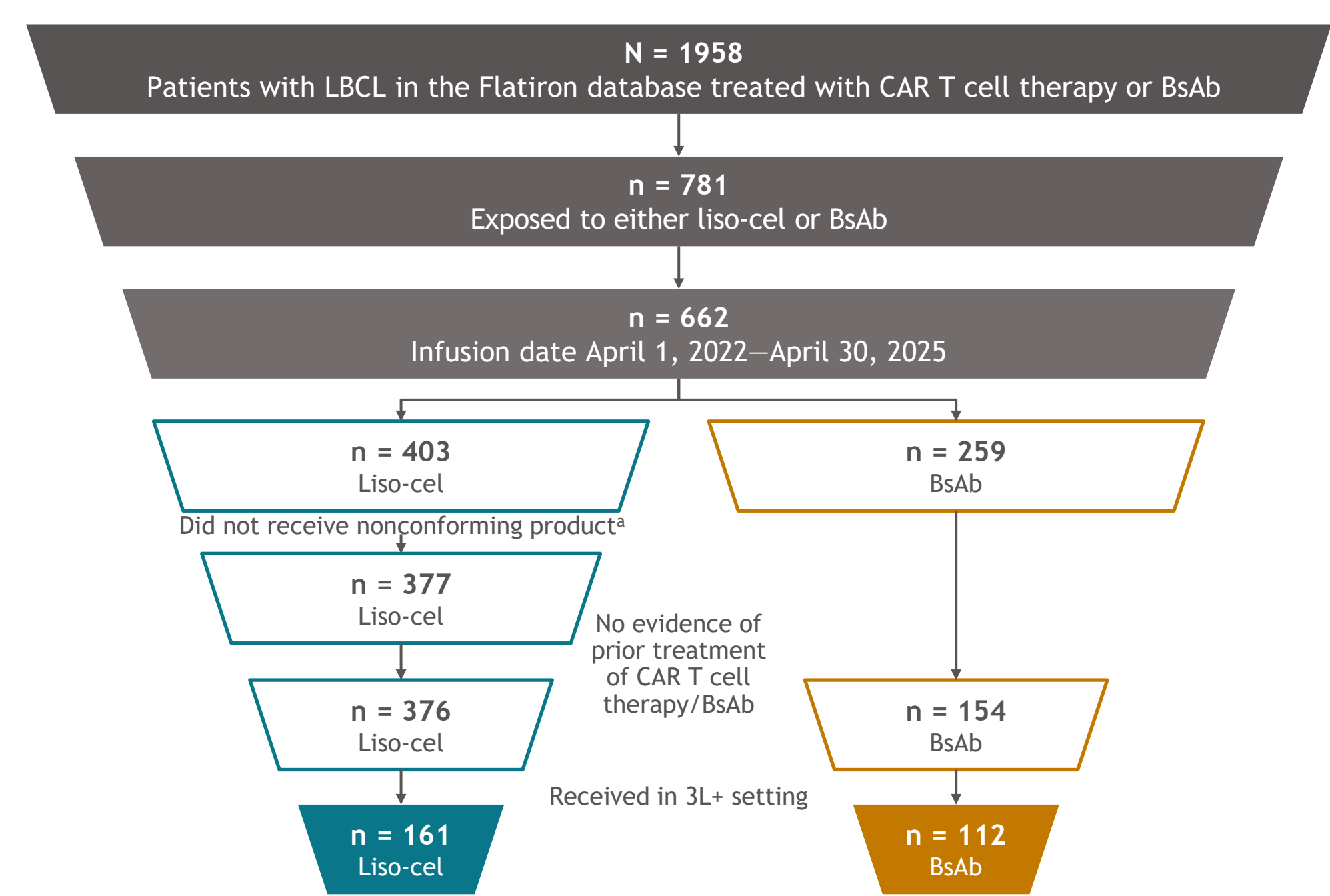
Methods

Figure 1. Study design



Results

Figure 2. Identification of the study cohort



- A total of 273 patients were included, with 161 in the liso-cel cohort and 112 in the BsAb cohort (Figure 2)

Liso-cel was associated with significantly longer OS (unadjusted, $P < 0.001$; adjusted, $P = 0.017$) and PFS (unadjusted, $P < 0.001$; adjusted, $P < 0.001$) than BsAb

Figure 4. Overall (A) and progression-free survival (B)

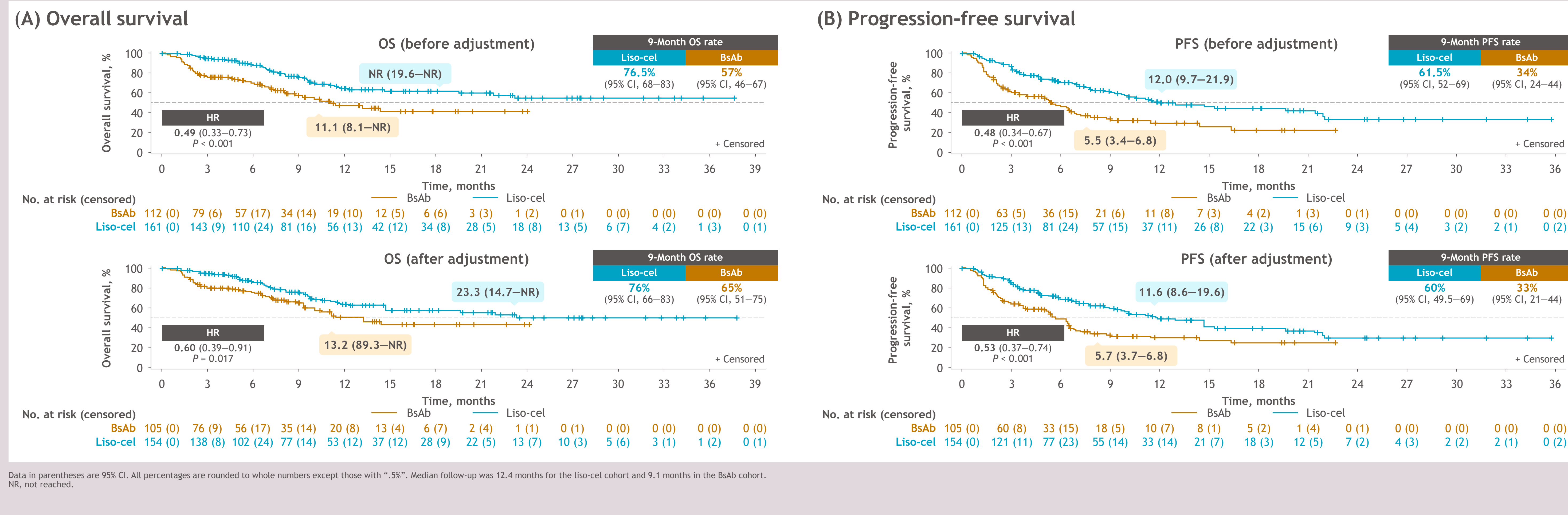
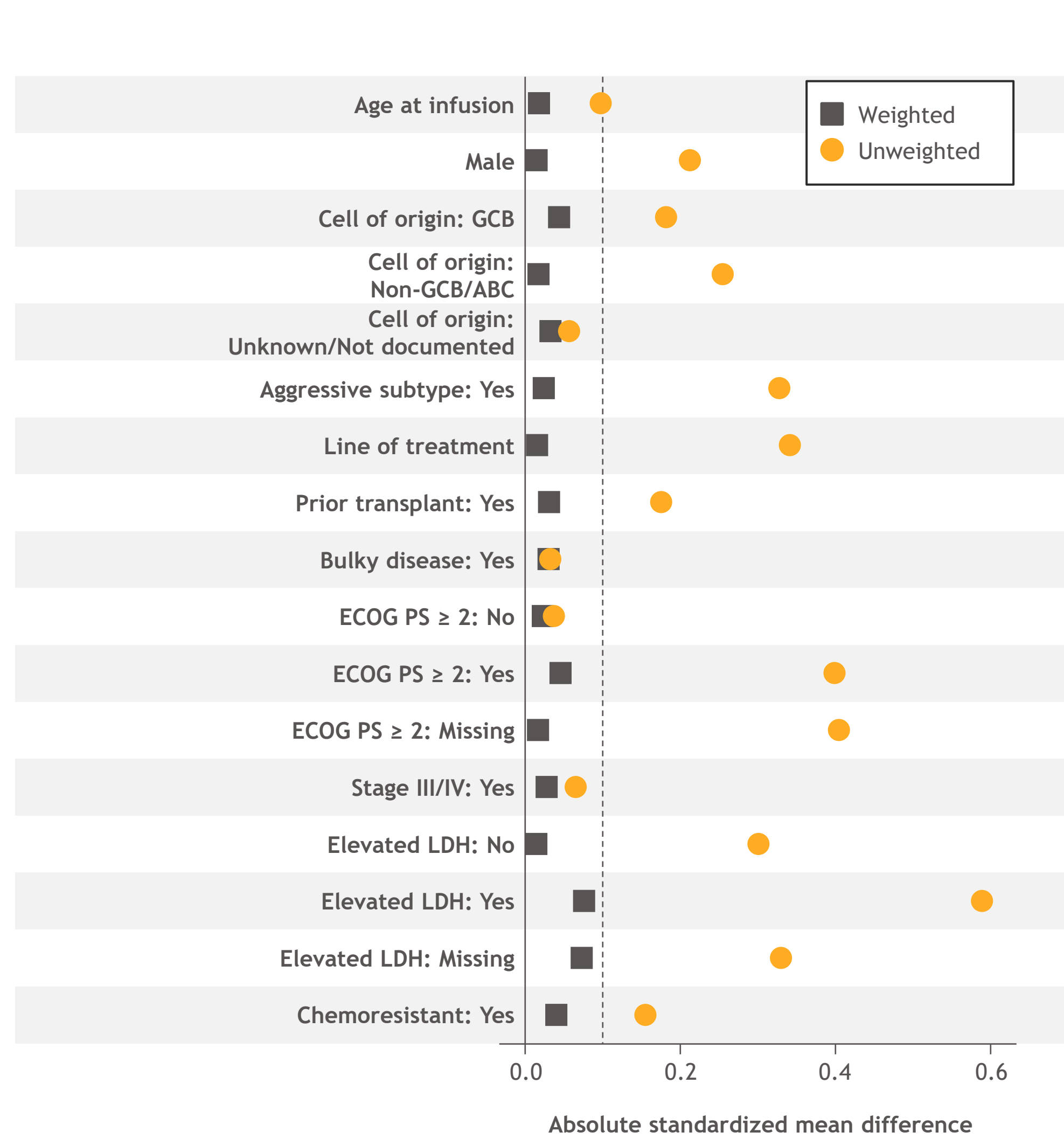


Figure 3. Love diagram for IPTW weighting



ABC, activated B cell; GCB, germinal center B cell.

- After IPTW adjustment, all characteristics were balanced (absolute standardized mean difference < 0.1) (Figure 3)

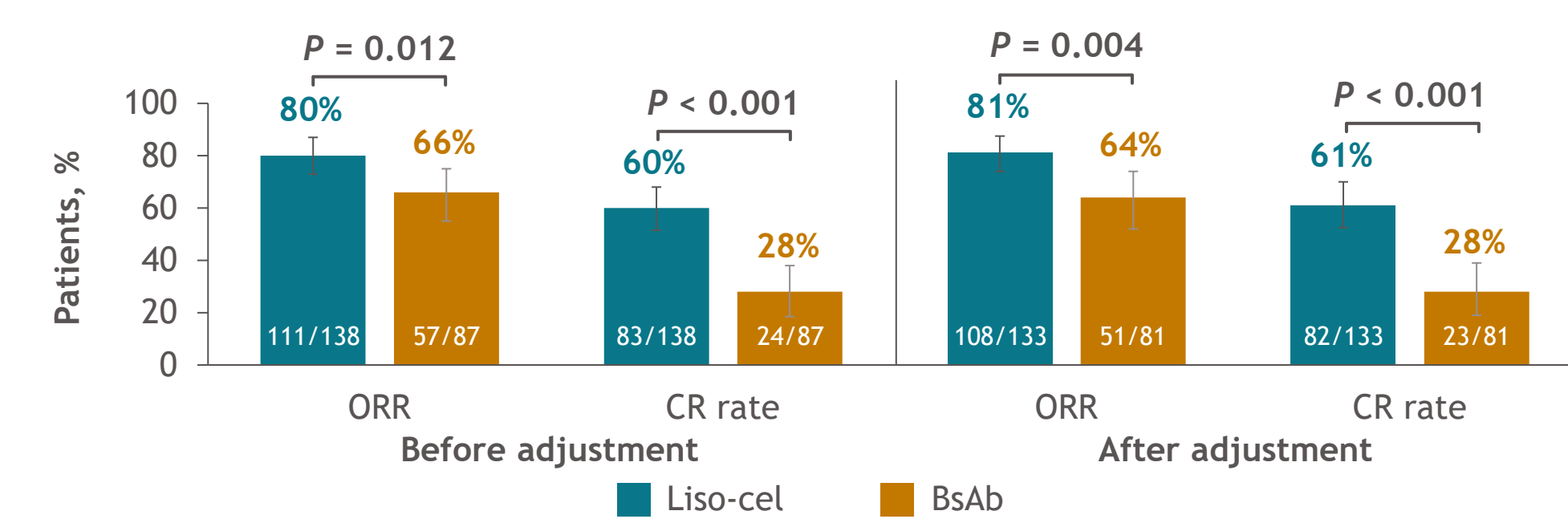
Table 1. Patient demographics and clinical characteristics

	Liso-cel (n = 161)	BsAb (n = 112)	P value
Median (range) age at index, y	72 (62–77)	75 (65–80.5)	0.02
Male, n (%)	95 (59)	76 (68)	0.16
Disease subtype, n (%)			
DLBCL NOS	140 (87)	85 (76)	0.09
Double-hit lymphoma	9 (6)	11 (10)	
High-grade B-cell lymphoma NOS	2 (1)	6 (5)	
Other	10 (6)	10 (9)	
Cell of origin, n (%)			
GCB	61 (38)	52 (46)	0.12
Non-GCB/ABC	56 (35)	26 (23)	
Missing	44 (27)	34 (30)	
ECOG PS, n (%)			0.004
0	28 (17)	23 (21)	
1	60 (37)	42 (37.5)	
2	17 (11)	22 (20)	
3	1 (1)	5 (4)	
4	0	1 (1)	
Missing	55 (34)	19 (17)	
Chemotherapy resistant, n (%)	141 (88)	89 (79)	0.09
CNS involvement, n (%)	4 (2)	5 (4)	0.49
Elevated LDH, n (%)	45 (28)	61 (54)	< 0.0001
Bulky disease, n (%)	106 (66)	74 (66)	1.0
Type of practice, n (%)			< 0.001
Community	25 (16)	39 (35)	
Academic	127 (79)	63 (56)	
Both academic and community	0	10 (9)	
Line of treatment			0.008
3L	113 (70)	60 (54)	
4L+	48 (30)	52 (46)	

All percentages are rounded to whole numbers except those with ".5%". 4L+, fourth line or later; NOS, not otherwise specified.

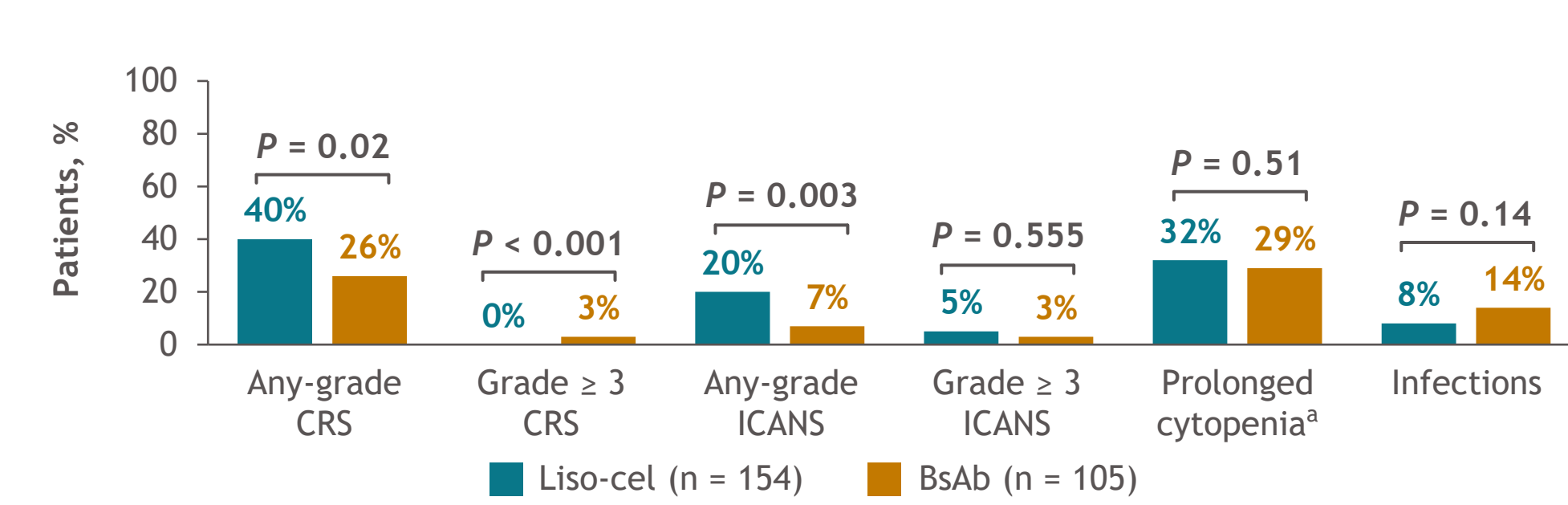
- Most baseline demographics and characteristics were similar between treatment groups (Table 1)
- Statistically significant differences at baseline were observed for age, ECOG PS at infusion, elevated LDH at infusion, practice type, and line of treatment
- In both cohorts, the most common LBCL subtypes were DLBCL NOS (liso-cel, 87%; BsAb, 76%), double-hit lymphoma (liso-cel, 6%; BsAb, 10%), and high-grade B-cell lymphoma NOS (liso-cel, 1%; BsAb, 5%)
- In the liso-cel cohort, 70% received 3L treatment and 30% received 4L+ treatment; in the BsAb cohort, 54% received 3L treatment and 46% received 4L+ treatment
- Median length of follow-up was 12.4 months for the liso-cel cohort and 9.1 months in the BsAb cohort

Figure 5. Response rates^a



- For both unadjusted and adjusted analyses, response rates were significantly higher for liso-cel than BsAb (Figure 5)

Figure 6. Safety (after adjustment)



^aEvidence of grade ≥ 3 thrombocytopenia, neutropenia, and/or anemia during the same line of treatment from 30 days after liso-cel or BsAb infusion to 89 days after liso-cel infusion.

- Incidences of any-grade CRS and ICANS were significantly higher with liso-cel than BsAb (CRS, $P = 0.02$; ICANS, $P = 0.003$), mostly due to grade 1–2 events (Figure 6)
- There was no grade ≥ 3 CRS with liso-cel and low rates of grade ≥ 3 ICANS
- Rates of prolonged cytopenia and infections were similar between cohorts

Table 2. OS and PFS in 3L-only patients and patients who achieved target dose of BsAb (after adjustment)^a

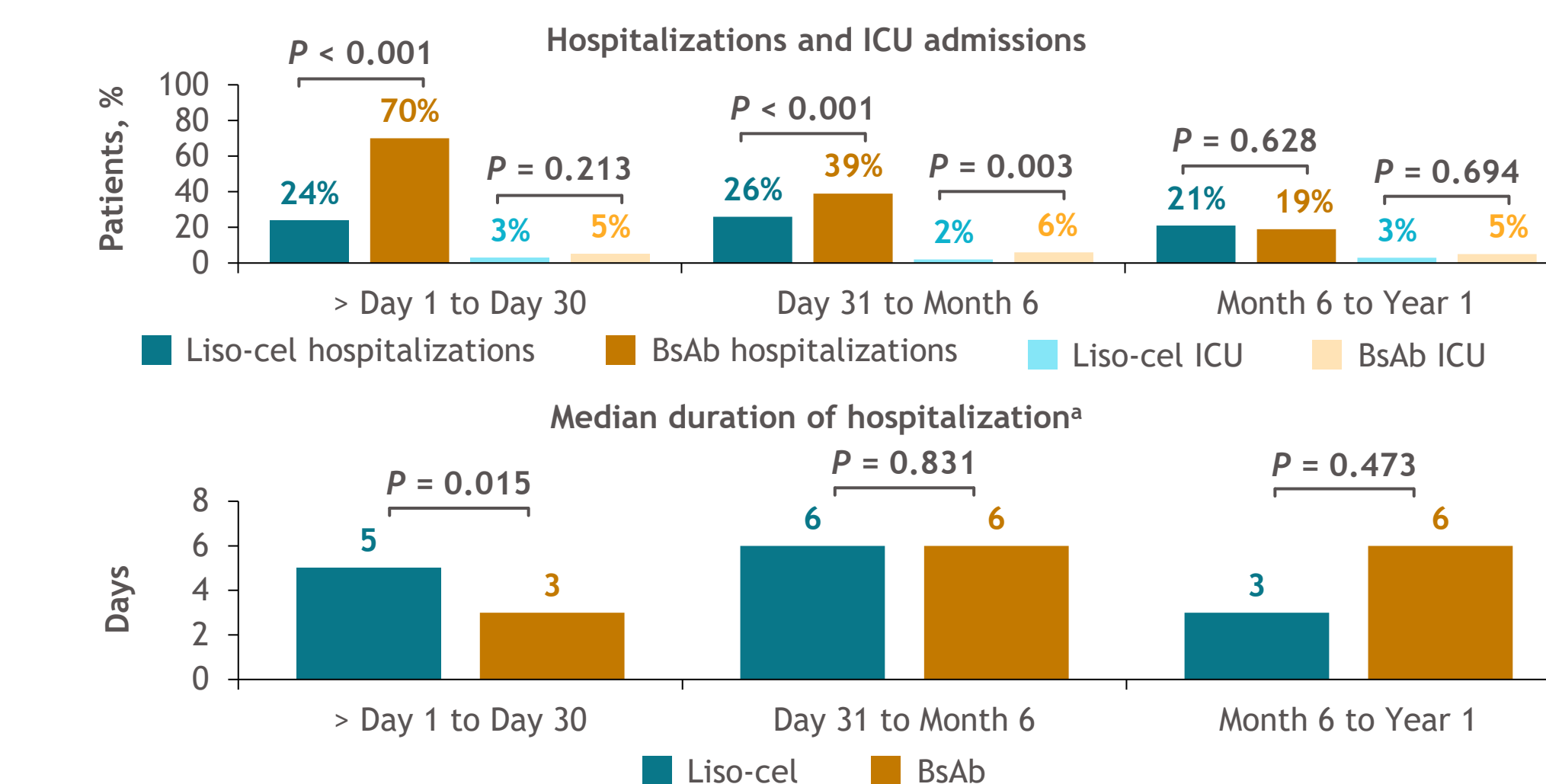
	3L-only patients		HR (95% CI)
	Liso-cel (n = 113)	BsAb (n = 60)	P value
Median OS	NE (19.6–NE)	8.8 (6.6–13.2)	0.33 (0.20–0.56) $P < 0.001$
Median PFS	12.0 (9.3–NE)	5.0 (2.4–7.2)	0.37 (0.24–0.57) $P < 0.001$

	Patients who achieved target dose of BsAb		HR (95% CI)
	Liso-cel (n = 161)	BsAb (n = 89)	P value
Median OS	NE (19.6–NE)	NE (8.3–NE)	0.63 (0.39–1.00) $P = 0.049$
Median PFS	12.0 (9.7–21.9)	6.1 (4.6–13.5)	0.59 (0.40–0.86) $P = 0.006$

^aTarget dose indicates that patients received labeled maintenance after step-up dosing. NE, not evaluable.

- In a sensitivity analysis of 3L-only patients and of those who achieved target dose of BsAb, liso-cel was associated with a significantly longer median OS and PFS (Table 2)
- Adjusted HRs for OS and PFS were approximately similar to or less than those observed in the overall population

Figure 7. Health care resource utilization (after adjustment)



^aCalculated as the duration of hospitalization during a period; if a patient has multiple hospitalizations within the same period, all days are summed during that period.

- There were lower hospitalization and ICU admission rates with liso-cel than BsAb across most time periods (Figure 7)
- The median duration of hospitalization was higher for liso-cel than BsAb over Days 1–30, but similar or possibly lower in the longer term

Conclusions

- In this retrospective, observational, real-world analysis in patients with 3L+ LBCL, liso-cel demonstrated significantly better efficacy than BsAb, with similar HCRU and safety
- Liso-cel was associated with a statistically significant improvement in OS and PFS, and response rates were significantly higher for liso-cel
- The higher rates of CRS and ICANS in patients who received liso-cel were mostly due to grade 1–2 events, suggesting manageable safety; there was no grade ≥ 3 CRS and low incidence of grade ≥ 3 ICANS
- Across most time periods, liso-cel had numerically lower hospitalization and ICU admission rates
- Further data are needed to compare these treatments in earlier lines of therapy

References

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