

PHARMACY POLICY STATEMENT

North Carolina Marketplace

DRUG NAME	Breyanzi (lisocabtagene maraleucel)
BENEFIT TYPE	Medical
STATUS	Prior Authorization Required

Breyanzi, approved by the FDA in 2021, is a CD19-directed chimeric antigen receptor (CAR)T-cell therapy for the treatment of relapsed or refractory large B-cell lymphoma. It has accelerated approval status for relapsed or refractory chronic lymphocytic leukemia (CLL) or small lymphocytic leukemia (SLL), follicular lymphoma (FL), and mantle cell lymphoma (MCL). Lymphoma is a cancer of the lymphatic system and white blood cells.

Breyanzi (lisocabtagene maraleucel) will be considered for coverage when the following criteria are met:

Large B-Cell Lymphoma (LBCL)

For **initial** authorization:

1. Member is at least 18 years of age; AND
2. Healthcare facility/provider has enrolled in the Breyanzi REMS; AND
3. Member has a documented diagnosis of large B-cell lymphoma including any of the following:
 - a) Diffuse large B-cell lymphoma (DLBCL) not otherwise specified (including DLBCL arising from indolent lymphoma)
 - b) High grade B-cell lymphoma (HGBCL)
 - c) Primary mediastinal large B-cell lymphoma (PMBCL)
 - d) Follicular lymphoma (FL) grade 3B
 - e) DLBCL arising from follicular lymphoma (transformation FL; TFL)
 - f) Intravascular LBCL
 - g) DLBCL associated with chronic inflammation
 - h) Fibrin-associated DLBCL
 - i) EBV-positive DLBCL, NOS
 - j) T-cell/histiocyte-rich LBCL; AND
4. Member has been treated with first line therapy containing an anthracycline and rituximab (or another CD20-targeted agent); AND
5. Member meets one of the following:
 - a) Relapsed or refractory disease after two or more lines of systemic therapy
 - b) Refractory disease to first-line chemoimmunotherapy (primary refractory) or relapse within 12 months of first-line chemoimmunotherapy
 - c) Refractory disease to first-line chemoimmunotherapy (primary refractory) or relapse after first-line chemoimmunotherapy and ineligible for hematopoietic stem cell transplant (HSCT) due to comorbidities or age; AND
6. Member has an Eastern cooperative oncology group (ECOG) performance status of 0 or 1; AND
7. Member does NOT have any of the following:
 - a) Primary central nervous system (CNS) lymphoma
 - b) Prior CAR T-cell or other genetically-modified T-cell therapy; AND
8. Member has been or will be screened for hepatitis B virus (HBV), hepatitis C virus (HCV), and human immunodeficiency virus (HIV).

9. **Dosage allowed/Quantity limit:**

After 2 or more lines of therapy: A single dose of 50 to 110×10^6 CAR-positive viable T cells

After 1 line of therapy: A single dose of 90 to 110×10^6 CAR-positive viable T cells

If all the above requirements are met, the medication will be approved for 3 months.

For **reauthorization:**

1. Breyanzi will not be reauthorized for continued therapy.

Mantle Cell Lymphoma (MCL)

For **initial** authorization:

1. Member is at least 18 years of age; AND
2. Healthcare facility/provider has enrolled in the Breyanzi REMS; AND
3. Member has a documented diagnosis of relapsed or refractory MCL; AND
4. Member has been treated with 2 or more prior lines of systemic therapy including ALL of the following:
 - a) Alkylating agent
 - b) CD20-targeted drug (e.g., rituximab)
 - c) Covalent Bruton tyrosine kinase inhibitor (BTKi) (i.e., ibrutinib, acalabrutinib, or zanubrutinib); AND
5. Member has an Eastern cooperative oncology group (ECOG) performance status of 0 or 1; AND
6. Member does NOT have any of the following:
 - a) Primary central nervous system (CNS) lymphoma
 - b) Prior CAR T-cell or other genetically-modified T-cell therapy; AND
7. Member has been or will be screened for hepatitis B virus (HBV), hepatitis C virus (HCV), and human immunodeficiency virus (HIV).
8. **Dosage allowed/Quantity limit:** A single dose of 90 to 110×10^6 CAR-positive viable T cells

If all the above requirements are met, the medication will be approved for 3 months.

For **reauthorization:**

1. Breyanzi will not be reauthorized for continued therapy.

Follicular Lymphoma (FL)

For **initial** authorization:

1. Member is at least 18 years of age; AND
2. Healthcare facility/provider has enrolled in the Breyanzi REMS; AND
3. Member has a documented diagnosis of relapsed or refractory FL (grade 1, 2, or 3a [see LBCL section above for grade 3B]); AND
4. Member has been treated with 2 or more prior lines of systemic therapy, including an alkylating agent and CD20-targeted drug (e.g., rituximab); AND
5. Member has an Eastern cooperative oncology group (ECOG) performance status of 0 or 1; AND
6. Member does NOT have any of the following:
 - a) Primary central nervous system (CNS) lymphoma
 - b) Prior CAR T-cell or other genetically-modified T-cell therapy; AND
7. Member has been or will be screened for hepatitis B virus (HBV), hepatitis C virus (HCV), and human immunodeficiency virus (HIV).
8. **Dosage allowed/Quantity limit:** A single dose of 90 to 110×10^6 CAR-positive viable T cells

If all the above requirements are met, the medication will be approved for 3 months.

For **reauthorization**:

1. Breyanzi will not be reauthorized for continued therapy.

Chronic Lymphocytic Leukemia (CLL) or Small Lymphocytic Lymphoma (SLL)

For **initial** authorization:

1. Member is at least 18 years of age; AND
2. Healthcare facility/provider has enrolled in the Breyanzi REMS; AND
3. Member has a documented diagnosis of relapsed or refractory CLL or SLL; AND
4. Member has been treated with 2 or more prior lines of systemic therapy, including a Bruton tyrosine kinase (BTK) inhibitor and a B-cell lymphoma 2 (BCL-2) inhibitor (venetoclax); AND
5. Member has an Eastern cooperative oncology group (ECOG) performance status of 0 or 1; AND
6. Member does NOT have any of the following:
 - a) Primary central nervous system (CNS) lymphoma
 - b) Prior gene therapy; AND
7. Member has been or will be screened for hepatitis B virus (HBV), hepatitis C virus (HCV), and human immunodeficiency virus (HIV).
8. **Dosage allowed/Quantity limit:** A single dose of 90 to 110 × 10⁶ CAR-positive viable T cells

If all the above requirements are met, the medication will be approved for 3 months.

For **reauthorization**:

1. Breyanzi will not be reauthorized for continued therapy.

CareSource considers Breyanzi (lisocabtagene maraleucel) not medically necessary for the treatment of conditions that are not listed in this document. For any other indication, please refer to the Off-Label policy.

DATE	ACTION/DESCRIPTION
05/20/2021	New policy for Breyanzi created.
07/27/2022	Updates to include 2 nd line use in accordance with recent labeling changes and NCCN guidelines. Updated billing code.
12/12/2024	Updated refs. Changed has been screened to has been or will be screened. Added new sections for MCL, FL, and CLL/SLL (label update). LBCL: Added more subtypes that would qualify (per NCCN).

References:

1. Breyanzi (lisocabtagene maraleuce) [package insert]. Bothell, WA; Juno Therapeutics, Inc.; 2024.
2. National Comprehensive Cancer Network. B-Cell Lymphomas (Version 3.2024). https://www.nccn.org/professionals/physician_gls/pdf/b-cell.pdf. Accessed December 12, 2024.
3. National Comprehensive Cancer Network. Chronic Lymphocytic Leukemia/ Small Lymphocytic Lymphoma (Version 1.2025). https://www.nccn.org/professionals/physician_gls/pdf/cll.pdf. Accessed December 13, 2024.
4. Abramson JS, Palomba ML, Gordon LI, et al. Lisocabtagene maraleucel for patients with relapsed or refractory large B-cell lymphomas (TRANSCEND NHL 001): a multicentre seamless design study. *Lancet*. 2020;396(10254):839-852. doi:10.1016/S0140-6736(20)31366-0
5. Abramson JS. Anti-CD19 CAR T-Cell Therapy for B-Cell Non-Hodgkin Lymphoma. *Transfus Med Rev*. 2020;34(1):29-33. doi:10.1016/j.tmr.2019.08.003
6. Kamdar M, Solomon SR, Arnason J, et al. Lisocabtagene maraleucel versus standard of care with salvage chemotherapy followed by autologous stem cell transplantation as second-line treatment in patients with relapsed or refractory large B-cell lymphoma (TRANSFORM): results from an interim analysis of an open-label,

randomised, phase 3 trial [published correction appears in *Lancet*. 2022 Jul 16;400(10347):160]. *Lancet*. 2022;399(10343):2294-2308. doi:10.1016/S0140-6736(22)00662-6

7. Sehgal A, Hoda D, Riedell PA, et al. Lisocabtagene maraleucel as second-line therapy in adults with relapsed or refractory large B-cell lymphoma who were not intended for haematopoietic stem cell transplantation (PILOT): an open-label, phase 2 study [published online ahead of print, 2022 Jul 12]. *Lancet Oncol*. 2022;S1470-2045(22)00339-4. doi:10.1016/S1470-2045(22)00339-4
8. Ernst M, Oeser A, Besiroglu B, et al. Chimeric antigen receptor (CAR) T-cell therapy for people with relapsed or refractory diffuse large B-cell lymphoma. *Cochrane Database Syst Rev*. 2021;9(9):CD013365. Published 2021 Sep 13. doi:10.1002/14651858.CD013365.pub2
9. Morschhauser F, Dahiya S, Palomba ML, et al. Lisocabtagene maraleucel in follicular lymphoma: the phase 2 TRANSCEND FL study [published correction appears in *Nat Med*. 2024 Aug;30(8):2374. doi: 10.1038/s41591-024-03175-4]. *Nat Med*. 2024;30(8):2199-2207. doi:10.1038/s41591-024-02986-9
10. Siddiqi T, Maloney DG, Kenderian SS, et al. Lisocabtagene maraleucel in chronic lymphocytic leukaemia and small lymphocytic lymphoma (TRANSCEND CLL 004): a multicentre, open-label, single-arm, phase 1-2 study. *Lancet*. 2023;402(10402):641-654. doi:10.1016/S0140-6736(23)01052-8

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