

## Medical Policy:

### Breyanzi® (lisocabtagene maraleucel)

POLICY NUMBER	LAST REVIEW	ORIGIN DATE
MG.MM.PH.324	March 28, 2024	April 5, 2021

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The treating physician or primary care provider must submit to EmblemHealth, or ConnectiCare, as applicable (hereinafter jointly referred to as “EmblemHealth”), the clinical evidence that the member meets the criteria for the treatment or surgical procedure. Without this documentation and information, EmblemHealth will not be able to properly review the request preauthorization or post-payment review. The clinical review criteria expressed below reflects how EmblemHealth determines whether certain services or supplies are medically necessary. This clinical policy is not intended to pre-empt the judgment of the reviewing medical director or dictate to health care providers how to practice medicine. Health care providers are expected to exercise their medical judgment in rendering appropriate care. Health care providers are expected to exercise their medical judgment in rendering appropriate care.

EmblemHealth established the clinical review criteria based upon a review of currently available clinical information (including clinical outcome studies in the peer reviewed published medical literature, regulatory status of the technology, evidence-based guidelines of public health and health research agencies, evidence-based guidelines and positions of leading national health professional organizations, views of physicians practicing in relevant clinical areas, and other relevant factors). EmblemHealth expressly reserves the right to revise these conclusions as clinical information changes and welcomes further relevant information. Each benefit program defines which services are covered. The conclusion that a particular service or supply is medically necessary does not constitute a representation or warranty that this service or supply is covered and/or paid for by EmblemHealth, as some programs exclude coverage for services or supplies that EmblemHealth considers medically necessary.

If there is a discrepancy between this guideline and a member's benefits program, the benefits program will govern. Identification of selected brand names of devices, tests and procedures in a medical coverage policy is for reference only and is not an endorsement of any one device, test or procedure over another. In addition, coverage may be mandated by applicable legal requirements of a state, the Federal Government or the Centers for Medicare & Medicaid Services (CMS) for Medicare and Medicaid members. All coding and web site links are accurate at time of publication.

EmblemHealth may also use tools developed by third parties, such as the MCG™ Care Guidelines, to assist us in administering health benefits. The MCG™ Care Guidelines are intended to be used in connection with the independent professional medical judgment of a qualified health care provider and do not constitute the practice of medicine or medical advice. EmblemHealth Services Company, LLC, has adopted this policy in providing management, administrative and other services to EmblemHealth Plan, Inc., EmblemHealth Insurance Company, EmblemHealth Services Company, LLC, and Health Insurance Plan of Greater New York (HIP) related to health benefit plans offered by these entities. ConnectiCare, an EmblemHealth company, has also adopted this policy. All of the aforementioned entities are affiliated companies under common control of EmblemHealth Inc.

## Definitions

Breyanzi, a CD19-directed genetically modified autologous T-cell immunotherapy, is indicated for the treatment of adults with relapsed or refractory large B-cell lymphoma after two or more lines of systemic therapy, including diffuse large B-cell lymphoma (DLBCL) not otherwise specified (including DLBCL arising from indolent lymphoma), high-grade B-cell lymphoma, primary mediastinal large B-cell lymphoma, and follicular lymphoma grade 3B.

## Length of Authorization

Coverage is available for one Breyanzi treatment course (single-dose) per lifetime.

## Dosing Limits [Medical Benefit]

### Quantity Limitations:

Sufficient quantity will be provided for one-time treatment:

1. Dosing is based on the number of chimeric antigen receptor (CAR)-positive viable T cells.

2. A single dose of BREYANZI contains 50 to 110 × 10<sup>6</sup> CAR-positive viable T cells (consisting of 1:1 CAR-positive viable T cells of the CD8 and CD4 components), with each component supplied separately in one to four single-dose 5 mL vials .
3. Each mL contains 1.5 × 10<sup>6</sup> to 70 × 10<sup>6</sup> CAR-positive viable T cells.
4. The infusion volume is calculated based on the concentration of cryopreserved drug product CAR-positive viable T cells concentration. The volume may differ for each component infused.

**Max Units (per dose and over time) [HCPCS Unit]:**

1 billable unit (1 infusion of up to 110 million autologous anti-cd19 CAR-positive viable T-cells)

## Guideline

### I. INITIAL APPROVAL CRITERIA

#### 1. B-Cell Lymphoma

- A. Patient has one of the following diagnoses (i, ii, iii, iv, v, vi, vii, viii, or ix):
  - i. Large B-cell lymphoma; **OR**
  - ii. Diffuse large B-cell lymphoma, not otherwise specified
    - a) May include DLBCL arising from indolent lymphoma; **OR**
  - iii. High-grade B-cell lymphoma; **OR**
  - iv. Primary mediastinal large B-cell lymphoma; **OR**
  - v. Follicular lymphoma grade 3B; **OR**
  - vi. Acquired immunodeficiency syndrome (AIDS)-related diffuse large B-cell lymphoma; **OR**
  - vii. Human herpesvirus-8 (HHV8)-positive diffuse large B-cell lymphoma; **OR**
  - viii. Primary effusion lymphoma; **OR**
  - ix. Post-transplant lymphoproliferative disorders; **AND**
- B. Patient is ≥ 18 years of age; **AND**
- C. The medication is prescribed by or in consultation with an oncologist; **AND**
- D. The administering facility has been certified to administer Breyanzi and is enrolled in Breyanzi REMS program; **AND**
- E. Patient meets **ONE** of the following:
  - i. refractory disease to first-line chemoimmunotherapy or relapse within 12 months of first-line chemoimmunotherapy; **OR**
  - ii. refractory disease to first-line chemoimmunotherapy or relapse after first-line chemoimmunotherapy and are not eligible for hematopoietic stem cell transplantation (HSCT) due to comorbidities or age; **OR**
  - iii. relapsed or refractory disease after two or more lines of systemic therapy; **AND**
- F. Patient has received lymphodepleting chemotherapy prior to infusion of Breyanzi; **AND**
- G. Patient has not been previously treated with CAR-T therapy;
 

*Note; Current FDA-approved CAR-T therapies include Breyanzi, Kymriah® (tisagenlecleucel suspension for intravenous infusion), Tecartus™ (brexucabtagene suspension for intravenous infusion), and Yescarta® (axicabtagene suspension for intravenous infusion); **AND***
- H. The requested use is supported by FDA-approved prescribing information OR the National Comprehensive Cancer Network (NCCN) Clinical Practice Guidelines (NCCN Guidelines®) and/or NCCN Drugs & Biologics Compendium (NCCN Compendium®) with a recommendation of category level 1 or 2A.

#### Limitations/Exclusions:

1. Breyanzi is not indicated for the treatment of patients with primary central nervous system lymphoma.

## Applicable Procedure Codes

Code	Description
Q2054	Lisocabtagene maraleucel, up to 110 million autologous anti-cd19 car-positive viable t cells, including leukapheresis and dose preparation procedures, per therapeutic dose

## Applicable NDCs

Code	Description
73153-0900-01	Breyanzi (lisocabtagene maraleucel) suspension for intravenous infusion

## ICD-10 Diagnoses

Code	Description
C82.40	Follicular lymphoma grade IIIb, unspecified site
C82.41	Follicular lymphoma grade IIIb, lymph nodes of head, face, and neck
C82.42	Follicular lymphoma grade IIIb, intrathoracic lymph nodes
C82.43	Follicular lymphoma grade IIIb, intra-abdominal lymph nodes
C82.44	Follicular lymphoma grade IIIb, lymph nodes of axilla and upper limb
C82.45	Follicular lymphoma grade IIIb, lymph nodes of inguinal region and lower limb
C82.46	Follicular lymphoma grade IIIb, intrapelvic lymph nodes
C82.47	Follicular lymphoma grade IIIb, spleen
C82.48	Follicular lymphoma grade IIIb, lymph nodes of multiple sites
C82.49	Follicular lymphoma grade IIIb, extranodal and solid organ sites
C83.00	Small cell B-cell lymphoma, unspecified site
C83.01	Small cell B-cell lymphoma, lymph nodes of head, face and neck
C83.02	Small cell B-cell lymphoma, intrathoracic lymph nodes
C83.03	Small cell B-cell lymphoma, intra-abdominal lymph nodes
C83.04	Small cell B-cell lymphoma, lymph nodes of axilla and upper limb
C83.05	Small cell B-cell lymphoma, lymph nodes of inguinal region and lower limb
C83.06	Small cell B-cell lymphoma, intrapelvic lymph nodes
C83.07	Small cell B-cell lymphoma, spleen
C83.08	Small cell B-cell lymphoma, lymph nodes of multiple sites
C83.09	Small cell B-cell lymphoma, extranodal and solid organ sites
C83.30	Diffuse large B-cell lymphoma, unspecified site
C83.31	Diffuse large B-cell lymphoma, lymph nodes of head, face, and neck
C83.32	Diffuse large B-cell lymphoma, intrathoracic lymph nodes
C83.33	Diffuse large B-cell lymphoma, intra-abdominal lymph nodes
C83.34	Diffuse large B-cell lymphoma, lymph nodes of axilla and upper limb
C83.35	Diffuse large B-cell lymphoma, lymph nodes of inguinal region and lower limb
C83.36	Diffuse large B-cell lymphoma, intrapelvic lymph nodes
C83.37	Diffuse large B-cell lymphoma, spleen
C83.38	Diffuse large B-cell lymphoma, lymph nodes of multiple sites
C83.39	Diffuse large B-cell lymphoma, extranodal and solid organ sites
C83.90	Non-follicular (diffuse) lymphoma, unspecified, unspecified site
C83.91	Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of head, face, and neck

C83.92	Non-follicular (diffuse) lymphoma, unspecified, intrathoracic lymph nodes
C83.93	Non-follicular (diffuse) lymphoma, unspecified, intra-abdominal lymph nodes
C83.94	Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of axilla and upper limb
C83.95	Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of inguinal region and lower limb
C83.96	Non-follicular (diffuse) lymphoma, unspecified, intrapelvic lymph nodes
C83.97	Non-follicular (diffuse) lymphoma, unspecified, spleen
C83.98	Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of multiple sites
C83.99	Non-follicular (diffuse) lymphoma, unspecified, extranodal and solid organ sites
C82.40	Follicular lymphoma grade IIIb, unspecified site
C82.41	Follicular lymphoma grade IIIb, lymph nodes of head, face, and neck
C82.42	Follicular lymphoma grade IIIb, intrathoracic lymph nodes
C82.43	Follicular lymphoma grade IIIb, intra-abdominal lymph nodes
C82.44	Follicular lymphoma grade IIIb, lymph nodes of axilla and upper limb
C82.45	Follicular lymphoma grade IIIb, lymph nodes of inguinal region and lower limb
C82.46	Follicular lymphoma grade IIIb, intrapelvic lymph nodes
C82.47	Follicular lymphoma grade IIIb, spleen
C82.48	Follicular lymphoma grade IIIb, lymph nodes of multiple sites
C82.49	Follicular lymphoma grade IIIb, extranodal and solid organ sites
C82.50	Diffuse follicle center lymphoma, unspecified site
C82.51	Diffuse follicle center lymphoma, lymph nodes of head, face, and neck
C82.52	Diffuse follicle center lymphoma, intrathoracic lymph nodes
C82.53	Diffuse follicle center lymphoma, intra-abdominal lymph nodes
C82.54	Diffuse follicle center lymphoma, lymph nodes of axilla and upper limb
C82.55	Diffuse follicle center lymphoma, lymph nodes of inguinal region and lower limb
C82.56	Diffuse follicle center lymphoma, intrapelvic lymph nodes
C82.57	Diffuse follicle center lymphoma, spleen
C82.58	Diffuse follicle center lymphoma, lymph nodes of multiple sites
C82.59	Diffuse follicle center lymphoma, extranodal and solid organ sites
C85.20	Mediastinal (thymic) large B-cell lymphoma, unspecified site
C85.21	Mediastinal (thymic) large B-cell lymphoma, lymph nodes of head, face, and neck
C85.22	Mediastinal (thymic) large B-cell lymphoma, intrathoracic lymph nodes
C85.23	Mediastinal (thymic) large B-cell lymphoma, intra-abdominal lymph nodes
C85.24	Mediastinal (thymic) large B-cell lymphoma, lymph nodes of axilla and upper limb
C85.25	Mediastinal (thymic) large B-cell lymphoma, lymph nodes of inguinal region and lower limb
C85.26	Mediastinal (thymic) large B-cell lymphoma, intrapelvic lymph nodes
C85.27	Mediastinal (thymic) large B-cell lymphoma, spleen
C85.28	Mediastinal (thymic) large B-cell lymphoma, lymph nodes of multiple sites
C85.29	Mediastinal (thymic) large B-cell lymphoma, extranodal and solid organ sites
C85.80	Other specified types of non-Hodgkin lymphoma, unspecified site
C85.81	Other specified types of non-Hodgkin lymphoma, lymph nodes of head, face and neck
C85.82	Other specified types of non-Hodgkin lymphoma, intrathoracic lymph nodes
C85.83	Other specified types of non-Hodgkin lymphoma, intra-abdominal lymph nodes
C85.84	Other specified types of non-Hodgkin lymphoma, lymph nodes of axilla and upper limb
C85.85	Other specified types of non-Hodgkin lymphoma, lymph nodes of inguinal region of lower limb
C85.86 O	Other specified types of non-Hodgkin lymphoma, intrapelvic lymph nodes
C85.87	Other specified types of non-Hodgkin lymphoma, spleen

C85.88	Other specified types of non-Hodgkin lymphoma, lymph nodes of multiple sites
C85.89	Other specified types of non-Hodgkin lymphoma, extranodal and solid organ sites
C91.10	Chronic lymphocytic leukemia of B-cell type not having achieved remission
C91.12	Chronic lymphocytic leukemia of B-cell type in relapse
D47.Z1	Post-transplant lymphoproliferative disorder (PTLD)
C85.80	Other specified types of non-Hodgkin lymphoma, unspecified site

## Revision History

Company(ies)	DATE	REVISION																								
EmblemHealth & ConnectiCare	3/28/2024	Annual Review: Updated dosing limits																								
EmblemHealth & ConnectiCare	7/24/2023	<p>Annual Review:</p> <p><b>B-Cell Lymphoma:</b> Initial Criteria: Added: "Acquired immunodeficiency syndrome (AIDS)-related diffuse large B-cell lymphoma; OR Human herpesvirus-8 (HHV8)-positive diffuse large B-cell lymphoma; <b>OR</b> Primary effusion lymphoma; <b>OR</b> Post-transplant lymphoproliferative disorders; <b>AND</b>"</p> <p>Removed:</p> <p>"Patient has a relapsed or refractory disease, defined as progression after 2 or more lines of systemic therapy; <b>AND</b>"</p> <p>Added "Patient meets <b>ONE</b> of the following:</p> <ul style="list-style-type: none"> <li>i. refractory disease to first-line chemoimmunotherapy or relapse within 12 months of first-line chemoimmunotherapy; <b>OR</b></li> <li>ii. refractory disease to first-line chemoimmunotherapy or relapse after first-line chemoimmunotherapy and are not eligible for hematopoietic stem cell transplantation (HSCT) due to comorbidities or age; <b>OR</b></li> <li>iii. relapsed or refractory disease after two or more lines of systemic therapy; <b>AND</b>"</li> </ul> <p>Removed ICD-10 Z51.12; Added ICD-10 codes:</p> <table border="1"> <tr> <td>C82.40</td> <td>Follicular lymphoma grade IIIb, unspecified site</td> </tr> <tr> <td>C82.41</td> <td>Follicular lymphoma grade IIIb, lymph nodes of head, face, and neck</td> </tr> <tr> <td>C82.42</td> <td>Follicular lymphoma grade IIIb, intrathoracic lymph nodes</td> </tr> <tr> <td>C82.43</td> <td>Follicular lymphoma grade IIIb, intra-abdominal lymph nodes</td> </tr> <tr> <td>C82.44</td> <td>Follicular lymphoma grade IIIb, lymph nodes of axilla and upper limb</td> </tr> <tr> <td>C82.45</td> <td>Follicular lymphoma grade IIIb, lymph nodes of inguinal region and lower limb</td> </tr> <tr> <td>C82.46</td> <td>Follicular lymphoma grade IIIb, intrapelvic lymph nodes</td> </tr> <tr> <td>C82.47</td> <td>Follicular lymphoma grade IIIb, spleen</td> </tr> <tr> <td>C82.48</td> <td>Follicular lymphoma grade IIIb, lymph nodes of multiple sites</td> </tr> <tr> <td>C82.49</td> <td>Follicular lymphoma grade IIIb, extranodal and solid organ sites</td> </tr> <tr> <td>C83.00</td> <td>Small cell B-cell lymphoma, unspecified site</td> </tr> <tr> <td>C83.01</td> <td>Small cell B-cell lymphoma, lymph nodes of head, face and neck</td> </tr> </table>	C82.40	Follicular lymphoma grade IIIb, unspecified site	C82.41	Follicular lymphoma grade IIIb, lymph nodes of head, face, and neck	C82.42	Follicular lymphoma grade IIIb, intrathoracic lymph nodes	C82.43	Follicular lymphoma grade IIIb, intra-abdominal lymph nodes	C82.44	Follicular lymphoma grade IIIb, lymph nodes of axilla and upper limb	C82.45	Follicular lymphoma grade IIIb, lymph nodes of inguinal region and lower limb	C82.46	Follicular lymphoma grade IIIb, intrapelvic lymph nodes	C82.47	Follicular lymphoma grade IIIb, spleen	C82.48	Follicular lymphoma grade IIIb, lymph nodes of multiple sites	C82.49	Follicular lymphoma grade IIIb, extranodal and solid organ sites	C83.00	Small cell B-cell lymphoma, unspecified site	C83.01	Small cell B-cell lymphoma, lymph nodes of head, face and neck
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EmblemHealth & ConnectiCare	4/6/2022	Transferred policy to new template, updated procedure code.	
EmblemHealth & ConnectiCare	4/5/2021	New Policy	

## References

1. Breyanzi<sup>®</sup> intravenous suspension [package insert]. Bothell, WA. Juno Therapeutics, Inc. Updated February 9, 2021. Available at: <https://dailymed.nlm.nih.gov/dailymed/drugInfo.cfm?setid=594bb413-af3b-4b97-afb3-bfe2b174f2ed>. Accessed February 26, 2021.
2. Breyanzi<sup>®</sup> intravenous suspension. IBM Micromedex<sup>®</sup> [database online]. Greenwood Village, CO. Truven Health Analytics. Available at: <https://www.micromedexsolutions.com>. Updated February 12, 2021. Accessed February 26, 2021.