

**Clinical Policy: Alemtuzumab (Lemtrada)**

Reference Number: CP.PHAR.243

Effective Date: 08.01.16

Last Review Date: 05.21

Line of Business: Commercial, HIM, Medicaid

[Coding Implications](#)[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

**Description**

Alemtuzumab (Lemtrada<sup>®</sup>) is a CD52-directed cytolytic monoclonal antibody.

**FDA Approved Indication(s)**

Lemtrada is indicated for the treatment with relapsing forms of multiple sclerosis (MS), to include relapsing-remitting disease and active secondary progressive disease, in adults. Because of its safety profile, the use of Lemtrada should generally be reserved for patients who have had an inadequate response to two or more drugs indicated for the treatment of MS.

Limitation(s) of use: Lemtrada is not recommended for use in patients with clinically isolated syndrome (CIS) because of its safety profile.

**Policy/Criteria**

*Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.*

It is the policy of health plans affiliated with Centene Corporation<sup>®</sup> that Lemtrada is **medically necessary** when the following criteria are met:

**I. Initial Approval Criteria****A. Multiple Sclerosis (must meet all):**

1. Diagnosis of relapsing-remitting or secondary progressive MS;
2. Prescribed by or in consultation with a neurologist;
3. Age  $\geq$  18 years;
4. Failure of all of the following at up to maximally indicated doses, unless clinically significant adverse effects are experienced or all are contraindicated (a, b, c, and d):\*
  - a. Dimethyl fumarate (generic Tecfidera<sup>®</sup>);
  - b. Aubagio<sup>®</sup>;
  - c. Gilenya<sup>®</sup>;
  - d. An interferon-beta agent (Avonex<sup>®</sup>, Betaseron<sup>®</sup>, Rebif<sup>®</sup>, or Plegridy<sup>®</sup>) or glatiramer (Copaxone<sup>®</sup>, Glatopa<sup>®</sup>);

*\*Prior authorization is required for all disease modifying therapies for MS*
5. Lemtrada is not prescribed concurrently with other disease modifying therapies for MS (see Appendix D);
6. Documentation of baseline number of relapses per year and expanded disability status scale (EDSS) score;
7. Dose does not exceed:

- a. First treatment course: 12 mg per day for 5 consecutive days (60 mg total);
- b. Second or subsequent treatment courses: 12 mg per day for 3 consecutive days (36 mg total).

**Approval duration:**

**Medicaid/HIM** – 12 months (*1 treatment course only*)

**Commercial** – 6 months or to the member’s renewal date, whichever is longer (*1 treatment course only*)

**B. Other diagnoses/indications**

1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**II. Continued Therapy**

**A. Multiple Sclerosis** (must meet all):

1. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
2. Member is responding positively to therapy as evidenced by one of the following (a, b, c, or d):
  - a. Member has not had an increase in the number of relapses per year compared to baseline;
  - b. Member has not had  $\geq 2$  new MRI-detected lesions;
  - c. Member has not had an increase in EDSS score from baseline;
  - d. Medical justification supports that member is responding positively to therapy;
3. Lemtrada is not prescribed concurrently with other disease modifying therapies for MS (*see Appendix D*);
4. It has been at least 12 months since completion of the prior treatment course;
5. Dose does not exceed 12 mg per day for 3 consecutive days (36 mg total per treatment course).

**Approval duration:**

**Medicaid/HIM** – 12 months (*1 treatment course only*)

**Commercial** – 6 months or to the member’s renewal date, whichever is longer (*1 treatment course only*)

**B. Other diagnoses/indications** (must meet 1 or 2):

1. Currently receiving medication via Centene benefit and documentation supports positive response to therapy.

**Approval duration: Duration of request or 6 months (whichever is less);** or

2. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**III. Diagnoses/Indications for which coverage is NOT authorized:**

- A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policy – CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid or evidence of coverage documents;
- B. Primary progressive MS.

**IV. Appendices/General Information**

*Appendix A: Abbreviation/Acronym Key*

EDSS: expanded disability status scale

FDA: Food and Drug Administration

MS: multiple sclerosis

*Appendix B: Therapeutic Alternatives*

*This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may not be a formulary agent for all relevant lines of business and may require prior authorization.*

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
Aubagio <sup>®</sup> (teriflunomide)	7 mg or 14 mg PO QD	14 mg/day
Avonex <sup>®</sup> , Rebif <sup>®</sup> (interferon beta-1a)	Avonex: 30 mcg IM Q week Rebif: 22 mcg or 44 mcg SC TIW	Avonex: 30 mcg/week Rebif: 44 mcg TIW
Plegridy <sup>®</sup> (peginterferon beta-1a)	125 mcg SC Q2 weeks	125 mcg/2 weeks
Betaseron <sup>®</sup> (interferon beta-1b)	250 mcg SC QOD	250 mg QOD
glatiramer acetate (Copaxone <sup>®</sup> , Glatopa <sup>®</sup> )	20 mg SC QD or 40 mg SC TIW	20 mg/day or 40 mg TIW
Gilenya <sup>®</sup> (fingolimod)	0.5 mg PO QD	0.5 mg/day
dimethyl fumarate (Tecfidera <sup>®</sup> )	120 mg PO BID for 7 days, followed by 240 mg PO BID	480 mg/day

*Therapeutic alternatives are listed as Brand name<sup>®</sup> (generic) when the drug is available by brand name only and generic (Brand name<sup>®</sup>) when the drug is available by both brand and generic.*

*Appendix C: Contraindications/Boxed Warnings*

- Contraindication(s): hypersensitivity or anaphylactic reactions to alemtuzumab or any of the excipients in Lemtrada, infection with human immunodeficiency virus, active infection
- Boxed warning(s): autoimmunity, infusion reactions, stroke, and malignancies

*Appendix D: General Information*

- Disease-modifying therapies for MS are: glatiramer acetate (Copaxone<sup>®</sup>, Glatopa<sup>®</sup>), interferon beta-1a (Avonex<sup>®</sup>, Rebif<sup>®</sup>), interferon beta-1b (Betaseron<sup>®</sup>, Extavia<sup>®</sup>), peginterferon beta-1a (Plegridy<sup>®</sup>), dimethyl fumarate (Tecfidera<sup>®</sup>), diroximel fumarate (Vumerity<sup>®</sup>), monomethyl fumarate (Bafiertam<sup>™</sup>), fingolimod (Gilenya<sup>®</sup>), teriflunomide (Aubagio<sup>®</sup>), alemtuzumab (Lemtrada<sup>®</sup>), mitoxantrone (Novantrone<sup>®</sup>), natalizumab

(Tysabri<sup>®</sup>), ocrelizumab (Ocrevus<sup>®</sup>), cladribine (Mavenclad<sup>®</sup>), siponimod (Mayzent<sup>®</sup>), ozanimod (Zeposia<sup>®</sup>), and ofatumumab (Kesimpta<sup>®</sup>).

- Lemtrada is available only through a restricted program under a REMS called the Lemtrada REMS Program because of the risks of autoimmunity, infusion reactions, and malignancies.

**V. Dosage and Administration**

Indication	Dosing Regimen	Maximum Dose
Relapsing MS	IV infusion for 2 or more treatment courses: <ul style="list-style-type: none"> <li>• First course: 12 mg/day on 5 consecutive days</li> <li>• Second course: 12 mg/day on 3 consecutive days 12 months after first course</li> <li>• Subsequent courses as needed: 12 mg/day on 3 consecutive days 12 months after any prior course</li> </ul>	See regimen

**VI. Product Availability**

Single-use vial: 12 mg/1.2 mL

**VII. References**

1. Lemtrada Prescribing Information. Cambridge, MA: Genzyme Corporation; September 2020. Available at <http://www.lemtrada.com>. Accessed February 8, 2021.
2. Rae-Grant A, Day GS, Marrie RA, et al. Practice guideline recommendations summary: disease-modifying therapies for adults with multiple sclerosis: report of the Guideline Development, Dissemination, and Implementation Subcommittee of the American Academy of Neurology. *Neurology*. 2018; 90(17): 777-788. Full guideline available at: <https://www.aan.com/Guidelines/home/GetGuidelineContent/904>.

**Coding Implications**

Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

HCPCS Codes	Description
J0202	Injection, alemtuzumab, 1 mg

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Removed MRI requirement. Updated preferencing to require at least one of the highly effective DMTs on formulary (Tecfidera or Gilenya). Removed reasons to discontinue.	07.17	08.17
2Q 2018 annual review: no significant changes; removed HIV contraindication; added HIM; references reviewed and updated.	01.05.18	05.18

Reviews, Revisions, and Approvals	Date	P&T Approval Date
2Q 2019 annual review: for re-auth, removed restriction for a total of 2 treatment courses per updated FDA labeling which allows for 2 or more treatment courses; references reviewed and updated.	02.04.19	05.19
RT4: updated policy with new indications.	11.14.19	
RT4: updated criteria contents with new indication without additional data to consider: secondary progressive MS.	01.06.20	
Updated re-directions per SDC and prior clinical guidance; added COM line of business (CP.CPA.325 retired); revised HIM-Medical Benefit to HIM line of business.	01.21.20	
2Q 2020 annual review: no significant changes; clarified that only 1 treatment course may be approved per authorization; references reviewed and updated.	01.27.20	05.20
Added requirements for documentation of baseline relapses/EDSS and objective measures of positive response upon re-authorization; references reviewed and updated.	05.27.20	08.20
Per November and December SDC and prior clinical guidance, removed redirection to Mayzent; for RRMS modified redirection to require generic dimethyl fumarate, Aubagio, Gilenya, and either an interferon-beta agent or glatiramer.	01.11.21	
2Q 2021 annual review: no significant changes; references to HIM.PHAR.21 revised to HIM.PA.154; updated Appendix C with additional contraindications per revised PI; references reviewed and updated.	02.08.21	05.21

**Important Reminder**

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy,

contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

Providers referred to in this clinical policy are independent contractors who exercise independent judgment and over whom the Health Plan has no control or right of control. Providers are not agents or employees of the Health Plan.

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**Note:**

**For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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