# Berinert® (C1 Esterase Inhibitor, Human) (Intravenous)

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## I. Length of Authorization

- Initial*: Prior authorization validity will be provided initially for 12 weeks.
- Renewal*: Prior authorization validity may be renewed every 12 weeks thereafter.

*The cumulative amount of medication(s) the member has on-hand, indicated for the acute treatment of HAE, will be taken into account when authorizing. The authorization will provide a sufficient quantity in order for the member to have a cumulative amount of HAE medication(s) on-hand in order to treat up to 4 acute attacks per 4 weeks for the duration of the authorization.

## II. Dosing Limits

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

- 1000 billable units per 28 days

## III. Initial Approval Criteria [1,23-25]

Prior authorization validity is provided in the following conditions:

**Universal Criteria**

- Must be prescribed by, or in consultation with, a specialist in allergy, immunology, hematology, pulmonology, or medical genetics; **AND**
- Will not be used in combination with another agent indicated for the treatment of acute hereditary angioedema (HAE) attacks; ***NOTE:*** *Requests for combination duplicate therapy may be appropriate in some situations and will be reviewed on a case-by-case basis*; **AND**
- Provider will confirm member has been counseled to avoid possible medication triggers for HAE attacks (e.g. estrogen-containing medications, ACE inhibitors, etc.); **AND**

**Treatment of acute attacks of Hereditary Angioedema (HAE) † ‡ Φ**

- Member has a history of ANY of the following:
  - Recurrent swelling of mucous membranes, skin surface, or subcutaneous tissue

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- Recurrent abdominal attacks with any of the following: severe abdominal pain, nausea, vomiting, or diarrhea
- Episodes of upper airway (e.g., mouth, tongue, larynx, etc.) edema
- Prodromal signs and symptoms (e.g., anxiety, nausea, paresthesia, fatigue, etc.) preceding angioedema attacks
- Positive family history of HAE; **AND**

- Other causes of angioedema have been ruled out (e.g., drug-induced angioedema, mast cell-mediated angioedema, acquired angioedema due to C1-INH deficiency, etc.); **AND**
- Diagnosis of HAE has been confirmed as one of the following:
  - Member has HAE due to C1 inhibitor deficiency (HAE-C1-INH) Type 1 or 2; **AND**
    - Member has a current or historical C1-INH antigenic level below the lower limit of normal as defined by the laboratory performing the test; **OR**
    - Member has a current or historical C1-INH functional level below the lower limit of normal as defined by the laboratory performing the test; **OR**
    - Genetic testing confirms pathogenic mutation(s) in *SERPING1;* **OR**
  - Member has HAE with normal C1INH (HAE-nC1-INH); **AND**
    - Member has a current or historical normal C1-INH level (e.g., C1-INH antigenic level, C1-INH functional level) as defined by the laboratory performing the test; **AND**
      - Member has a confirmed presence of known HAE-causing mutation(s) in the gene(s) for factor XII (*F12),* angiopoietin-1 *(ANGPT1),* plasminogen *(PLG),* kininogen-1 *(KNG1),* myoferlin *(MYOF)*, and/or heparan sulfate-glucosamine 3-O-sulfotransferase 6 (*HS3ST6)*; **OR**
      - Member has recurrent angioedema attacks without hives that are refractory to a 4-week trial of high-dose antihistamines (i.e. second-generation H1 antagonist at four times the standard dose)

† FDA Approved Indication(s); ‡ Compendia Recommended Indication(s); Φ Orphan Drug

## IV. Renewal Criteria [1, 23-25]

Prior authorization validity can be renewed based upon the following criteria:

- Member continues to meet the universal and other indication-specific relevant criteria identified in section III; **AND**
- The member has experienced clinical benefit from therapy (e.g., reduced attack severity, shortened attack duration, etc.); **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: severe hypersensitivity reactions, serious thromboembolic events (arterial or venous), etc.; **AND**
- The cumulative amount of medication(s) the member has on-hand, indicated for the acute treatment of HAE, will be taken into account when authorizing. The authorization will provide a sufficient quantity in order for the member to have a cumulative amount of HAE medication(s) on-hand in order to treat up to 4 acute attacks per 4 weeks for the duration of the authorization.



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## V. Dosage/Administration [1]

| Indication | Dose |
|---|---|
| Treatment of Acute Hereditary Angioedema (HAE) attack | Administer 20 international units (IU) per kg body weight by intravenous injection.<br>***Note:** Members may self-administer Berinert upon recognition of symptoms of an HAE attack after being instructed by their healthcare provider.* |

## VI. Billing Code/Availability Information

HCPCS Code:

- J0597 – Injection, c-1 esterase inhibitor (human), berinert, 10 units; 1 billable unit = 10 units

NDC(s):

- Berinert 500 IU single-dose carton kit (containing a single-dose vial of Berinert and a 10 mL vial of Sterile Water for Injection): 63833-0825-xx
- Berinert 500 IU single-dose vial: 63833-0835-xx

## VII. References


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## Appendix A – Non-Quantitative Treatment Limitations (NQTL) Factor Checklist

Non-quantitative treatment limitations (NQTLs) refer to the methods, guidelines, standards of evidence, or other conditions that can restrict how long or to what extent benefits are provided under a health plan. These may include things like utilization review or prior authorization. The utilization management NQTL applies comparably, and not more stringently, to mental health/substance use disorder (MH/SUD) Medical Benefit Prescription Drugs and medical/surgical (M/S) Medical Benefit Prescription Drugs. The table below lists the factors that were considered in designing and applying prior authorization to this drug/drug group, and a summary of the conclusions that Prime's assessment led to for each.

| Factor | Conclusion |
|---|---|
| Indication | Yes: Consider for PA |
| Safety and efficacy | No: PA not a priority |
| Potential for misuse/abuse | No: PA not a priority |
| Cost of drug | Yes: Consider for PA |

## Appendix 1 – Covered Diagnosis Codes

| ICD-10 | ICD-10 Description |
|---|---|
| D84.1 | Defects in the complement system |

## Appendix 2 – Centers for Medicare and Medicaid Services (CMS)

The preceding information is intended for non-Medicare coverage determinations. Medicare coverage for outpatient (Part B) drugs is outlined in the Medicare Benefit Policy Manual (Pub. 100-2), Chapter 15, §50 Drugs and Biologicals. In addition, National Coverage Determinations (NCDs) and/or Local Coverage Determinations (LCDs) may exist and compliance with these policies is required where applicable. Local Coverage Articles (LCAs) may also exist for claims payment purposes or to clarify benefit eligibility under Part B for drugs which may be self-administered. The following link may be used to search for NCD, LCD, or LCA documents: https://www.cms.gov/medicare-coverage-database/search.aspx. Additional indications, including any preceding information, may be applied at the discretion of the health plan.

Medicare Part B Covered Diagnosis Codes (applicable to existing NCD/LCD/LCA): N/A

| Medicare Part B Administrative Contractor (MAC) Jurisdictions | | |
|---|---|---|
| **Jurisdiction** | **Applicable State/US Territory** | **Contractor** |
| E (1) | CA, HI, NV, AS, GU, CNMI | Noridian Healthcare Solutions, LLC |
| F (2 & 3) | AK, WA, OR, ID, ND, SD, MT, WY, UT, AZ | Noridian Healthcare Solutions, LLC |
| 5 | KS, NE, IA, MO | Wisconsin Physicians Service Insurance Corp (WPS) |
| 6 | MN, WI, IL | Wellpoint Federal |



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| Medicare Part B Administrative Contractor (MAC) Jurisdictions | | |
|---|---|---|
| **Jurisdiction** | **Applicable State/US Territory** | **Contractor** |
| H (4 & 7) | LA, AR, MS, TX, OK, CO, NM | Novitas Solutions, Inc. |
| 8 | MI, IN | Wisconsin Physicians Service Insurance Corp (WPS) |
| N (9) | FL, PR, VI | First Coast Service Options, Inc. |
| J (10) | TN, GA, AL | Palmetto GBA |
| M (11) | NC, SC, WV, VA (excluding below) | Palmetto GBA |
| L (12) | DE, MD, PA, NJ, DC (includes Arlington & Fairfax counties and the city of Alexandria in VA) | Novitas Solutions, Inc. |
| K (13 & 14) | NY, CT, MA, RI, VT, ME, NH | Wellpoint Federal |
| 15 | KY, OH | CGS Administrators, LLC |



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