Kalbitor® (ecallantide)
(Subcutaneous)

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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 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]:

- 240 billable units per 28 days

III. Initial Approval Criteria [1,22-24]

Prior authorization validity is provided in the following conditions:

- Member is at least 12 years of age; **AND**

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 confirming 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 [1,22-24]

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

- Member must continue 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 serious hypersensitivity reactions, including anaphylaxis, 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

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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.

V. Dosage/Administration [1]

Indication	Dose
Treatment of Acute Hereditary Angioedema (HAE) attack	Administer 30 mg subcutaneously by a healthcare professional in three 10 mg injections. An additional dose of 30 mg may be administered if the attack persists. Not to exceed a total of two 30 mg doses (60 mg) in 24 hours. ***Note:** Kalbitor should ONLY be administered by a healthcare professional.*

VI. Billing Code/Availability Information

HCPCS Code:

- J1290 – Injection, ecallantide, 1 mg; 1 billable unit = 1 mg

NDC:

- Kalbitor 10 mg/mL; carton of 3 single-use vials: 47783-0101-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	Yes: Consider for PA
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

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Medicare Part B Administrative Contractor (MAC) Jurisdictions		
Jurisdiction	**Applicable State/US Territory**	**Contractor**
5	KS, NE, IA, MO	Wisconsin Physicians Service Insurance Corp (WPS)
6	MN, WI, IL	Wellpoint Federal
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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