

# Ruconest® (C1 Esterase Inhibitor [recombinant]) (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 patient 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 patient 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]:**

- 3360 billable units per 28 days

## III. Initial Approval Criteria <sup>1</sup>

Coverage is provided in the following conditions:

- Patient is at least 13 years of age; **AND**

**Universal Criteria** <sup>1,13,19</sup>

- Must be prescribed by, or in consultation with, a specialist in allergy, immunology, hematology, pulmonology, or medical genetics; **AND**
- Patient does not have a history of allergy to rabbits or rabbit-derived products; **AND**
- Will not be used in combination with another agent indicated for the treatment of acute attacks (i.e., Berinert [C1 esterase inhibitor, human], Firazyr [icatibant], Kalbitor [ecallantide]); **NOTE: Requests for combination duplicate therapy may be appropriate in some situations and will be reviewed on a case-by-case basis; AND**
- Confirmation the patient is avoiding the following possible triggers for HAE attacks:
  - Estrogen-containing oral contraceptive agents AND hormone replacement therapy; **AND**
  - Antihypertensive agents containing ACE inhibitors or angiotensin II receptor blockers (ARBs); **AND**
  - Dipeptidyl peptidase IV (DPP-IV) inhibitors (e.g., sitagliptin); **AND**

- Neprilysin inhibitors (e.g., sacubitril); **AND**

### Treatment of acute attacks of Hereditary Angioedema (HAE) † $\Phi$ <sup>1,13,19,20,21</sup>

- Patient has a history of moderate to severe cutaneous attacks (without concomitant hives) OR abdominal attacks OR mild to severe airway swelling attacks of HAE (i.e. debilitating cutaneous/gastrointestinal symptoms OR laryngeal/pharyngeal/tongue swelling); **AND**
- Patient has one of the following clinical presentations consistent with a HAE subtype§, which must be confirmed by repeat blood testing (treatment for acute attack should not be delayed for confirmatory testing):

<b>HAE I (C1-Inhibitor deficiency) § <sup>13,19,20,21</sup></b>
<ul style="list-style-type: none"> <li>• Low C1 inhibitor (C1-INH) antigenic level (C1-INH antigenic level below the lower limit of normal as defined by the laboratory performing the test); <b>AND</b></li> <li>• Low C4 level (C4 below the lower limit of normal as defined by the laboratory performing the test); <b>AND</b></li> <li>• Low C1-INH functional level (C1-INH functional level below the lower limit of normal as defined by the laboratory performing the test); <b>AND</b> <ul style="list-style-type: none"> <li>○ Patient has a family history of HAE; <b>OR</b></li> <li>○ Acquired angioedema has been ruled out (i.e., patient onset of symptoms occurs prior to 30 years of age, normal C1q levels, patient does not have underlying disease such as lymphoma or benign monoclonal gammopathy [MGUS], etc.)</li> </ul> </li> </ul>
<b>HAE II (C1-Inhibitor dysfunction) § <sup>19,21</sup></b>
<ul style="list-style-type: none"> <li>• Normal to elevated C1-INH antigenic level; <b>AND</b></li> <li>• Low C4 level (C4 below the lower limit of normal as defined by the laboratory performing the test); <b>AND</b></li> <li>• Low C1-INH functional level (C1-INH functional level below the lower limit of normal as defined by the laboratory performing the test)</li> </ul>
<b>HAE with normal C1INH (formerly known as HAE III) § <sup>19,20,21</sup></b>
<ul style="list-style-type: none"> <li>• Normal to near normal C1-INH antigenic level; <b>AND</b></li> <li>• Normal to near normal C4 level; <b>AND</b></li> <li>• Normal to near normal C1-INH functional level; <b>AND</b></li> <li>• Repeat blood testing <u>during an attack</u> has confirmed the patient does not have abnormal lab values indicative of HAE I or HAE II; <b>AND</b></li> <li>• Either of the following: <ul style="list-style-type: none"> <li>○ Patient has a known HAE-causing mutation (e.g., mutation of coagulation factor XII gene [F12 mutation], mutation in the angiotensin-converting enzyme 1 gene, mutation in the plasminogen gene, mutation in the kininogen 1 gene, mutation in the myoferlin gene, mutation in the heparan sulfate-glucosamine 3-O-sulfotransferase 6 gene, etc.); <b>OR</b></li> <li>○ Patient has a family history of HAE and documented lack of efficacy of chronic high-dose antihistamine therapy (e.g. <i>cetirizine standard dosing at up to four times daily or an alternative equivalent, given for at least one month or an interval long enough to expect three or more angioedema attacks</i>)</li> </ul> </li> </ul>

† FDA Approved Indication(s); ‡ Compendia Recommended Indication(s);  $\Phi$  Orphan Drug

#### IV. Renewal Criteria <sup>1</sup>

- Patient continues to meet the universal and other indication-specific relevant criteria identified in section III; **AND**
- Significant improvement in severity and duration of attacks has been achieved and sustained; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: severe hypersensitivity reactions (including anaphylaxis), serious thromboembolic events (arterial or venous), etc; **AND**
- The cumulative amount of medication(s) the patient 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 patient 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 <sup>1</sup>

Indication	Dose
Treatment of Acute Hereditary Angioedema (HAE) attack	<p><u>Body weight &lt; 84 kg:</u> Administer 50 U per kg body weight by intravenous injection</p> <p><u>Body weight ≥ 84 kg:</u> Administer 4200 U (2 vials) by intravenous injection</p> <p><i>If the attack symptoms persist, an additional (second) dose can be administered at the recommended dose level. Do not exceed 4200 U per dose. No more than two doses should be administered within a 24-hour period.</i></p> <p><b>**Note:</b> Patients may self-administer Ruconest upon recognition of symptoms of an HAE attack after being instructed by their healthcare provider.</p>

#### VI. Billing Code/Availability Information

HCPCS Code:

- J0596 - Injection, c1 esterase inhibitor (recombinant), ruconest, 10 units; 1 billable unit = 10 units

NDC:

- Ruconest 2100 U single-use 25 mL vial: 71274-0350-xx

#### VII. References

1. Ruconest [package insert]. Warren, NJ; Pharming Healthcare, Inc.; April 2020. Accessed June 2025.
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## 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	National Government Services, Inc. (NGS)
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.

### Medicare Part B Administrative Contractor (MAC) Jurisdictions

Jurisdiction	Applicable State/US Territory	Contractor
K (13 & 14)	NY, CT, MA, RI, VT, ME, NH	National Government Services, Inc. (NGS)
15	KY, OH	CGS Administrators, LLC