

## **Clinical Policy: Dichlorphenamide (Keveyis)**

Reference Number: CP.PMN.261

Effective Date: 03.01.21

Last Review Date: 02.24

Line of Business: Commercial, HIM, Medicaid

[Revision Log](#)

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

### **Description**

Dichlorphenamide (Keveyis<sup>®</sup>) is an oral carbonic anhydrase inhibitor.

### **FDA Approved Indication(s)**

Keveyis is indicated for the treatment of primary hyperkalemic periodic paralysis, primary hypokalemic periodic paralysis, and related variants.

### **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 Keveyis is **medically necessary** when the following criteria are met:

#### **I. Initial Approval Criteria**

##### **A. Hyperkalemic/Hypokalemic Periodic Paralysis and Variants (must meet all):**

1. Diagnosis of primary hyperkalemic or hypokalemic periodic paralysis, or related variants (i.e., Andersen's syndrome, paramyotonia congenita);
2. Age  $\geq$  18 years;
3. Failure of acetazolamide at up to maximally indicated dose, unless contraindicated or clinically significant adverse effects are experienced;
4. If request is for brand Keveyis, member must use generic dichlorphenamide, unless contraindicated or clinically significant adverse effects are experienced;
5. Dose does not exceed 200 mg (4 tablets) per day.

**Approval duration: 2 months**

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

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business:

CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or

2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

## II. Continued Therapy

### A. Hyperkalemic/Hypokalemic Periodic Paralysis and Variants (must meet all):

1. Member meets one of the following (a or b):
  - a. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
  - b. Member is currently receiving medication and is enrolled in a state and product with continuity of care regulations (*refer to state specific addendums for CC.PHARM.03A and CC.PHARM.03B*);
2. Member is responding positively to therapy as evidenced by reduced frequency of paralysis;
3. If request is for brand Keveyis, member must use generic dichlorphenamide, unless contraindicated or clinically significant adverse effects are experienced;
4. If request is for a dose increase, new dose does not exceed 200 mg (4 tablets) per day.

**Approval duration: 12 months**

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

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: 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 policies – CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid, or evidence of coverage documents.

**IV. Appendices/General Information**

*Appendix A: Abbreviation/Acronym Key*

FDA: Food and Drug Administration

*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
acetazolamide (Diamox <sup>®</sup> )	250 to 1,000 mg/day PO in divided doses	1,000 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): hepatic insufficiency, severe pulmonary obstruction, hypersensitivity to dichlorphenamide or other sulfonamides, concomitant use of Keveyis and high dose aspirin
- Boxed warning(s): none reported

*Appendix D: General Information*

- Variants of periodic paralysis include paramyotonia congenita and Andersen syndrome.
- Per the Keveyis prescribing information: primary hyperkalemic periodic paralysis, primary hypokalemic periodic paralysis, and related variants are a heterogeneous group of conditions, for which the response to Keveyis may vary. Therefore, prescribers should evaluate the patient's response to Keveyis after 2 months of treatment to decide whether Keveyis should be continued.

**V. Dosage and Administration**

Indication	Dosing Regimen	Maximum Dose
Primary hyperkalemic periodic paralysis, primary hypokalemic periodic paralysis, and related variants	Initial dose of 50 mg PO QD or BID; titrate based on individual response at weekly intervals up to a maximum recommended daily dose of 200 mg	200 mg/day

**VI. Product Availability**

Tablet: 50 mg

**VII. References**

1. Keveyis Prescribing Information. Hawthorne, NY: Taro Pharmaceuticals U.S.A, Inc.; May 2023. Available at: <https://www.keveyis.com>. Accessed on October 13, 2023.
2. Micromedex<sup>®</sup> Healthcare Series [Internet database]. Greenwood Village, Colo: Thomson Healthcare. Updated periodically. Accessed November 14, 2023.

3. Tawil R, McDermott MP, Brown R, et al. Randomized trials of dichlorphenamide in the periodic paralyses. *Ann Neurol* 2000;47:46-53.
4. Venance SL, Cannon SC, Fialho D, et al. The primary periodic paralyses: diagnosis, pathogenesis and treatment. *Brain* 2006; 129:8.
5. Statland JM, Fontaine B, Hanna MG, et al. Review of the diagnosis and treatment of periodic paralysis. *Muscle Nerve* 2018;54(4):522-530.

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy created; adopted from CP.PCH.04 (policy to be retired); added Medicaid line of business; revised initial approval duration from 3 months to 2 months to align with PI which recommends evaluation of response after 2 months; references to HIM.PHAR.21 revised to HIM.PA.154; no significant changes.	12.15.20	02.21
1Q 2022 annual review: no significant changes; references reviewed and updated.	10.18.21	02.22
Template changes applied to other diagnoses/indications and continued therapy section.	10.07.22	
1Q 2023 annual review: no significant changes; references reviewed and updated.	11.10.22	02.23
Added requirement for use of generic for brand Keveyis requests.	08.22.23	
1Q 2024 annual review: no significant changes; references reviewed and updated.	10.13.23	02.24

**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.

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