

**Orladeyo (berotralstat)**  
**Effective 01/01/2023**

|                              |  |  |  |
|------------------------------|--|--|--|
| <b>Plan</b>                  | <input type="checkbox"/> MassHealth UPPL<br><input checked="" type="checkbox"/> Commercial/Exchange  | <b>Program Type</b>                        | <input checked="" type="checkbox"/> Prior Authorization<br><input checked="" type="checkbox"/> Quantity Limit<br><input type="checkbox"/> Step Therapy |
| <b>Benefit</b>               | <input checked="" type="checkbox"/> Pharmacy Benefit<br><input type="checkbox"/> Medical Benefit     |  |  |
| <b>Specialty Limitations</b> | This medication has been designated specialty and must be filled at a contracted specialty pharmacy. |  |  |
| <b>Contact Information</b>   | <b>Medical Benefit</b><br><b>Pharmacy Benefit</b>  | Phone: 833-895-2611<br>Phone: 800-711-4555 | Fax: 888-656-6671<br>Fax: 844-403-1029   |
| <b>Exceptions</b>            | N/A  |  |  |

### Overview

Orladeyo (berotralstat) is a plasma kallikrein inhibitor that inhibits plasma kallikrein proteolytic activity, controlling excess bradykinin generation to prevent attacks of hereditary angioedema (HAE) in adults and pediatric patients 12 years and older.

### Coverage Guidelines

Authorization may be granted for members new to the plan who are currently receiving treatment with Orladeyo, excluding when the product is obtained as samples or via manufacturer's patient assistance programs.

#### OR

Authorization may be granted for members when ALL the following criteria are met, and documentation is provided:

1. The member will be using Orladeyo for the prevention of hereditary angioedema attacks
2. Orladeyo will not be used in combination with any medication used for the prophylaxis of HAE attacks.
3. Member meets ONE of the following:
  - a. Member has C1 inhibitor deficiency or dysfunction as confirmed by laboratory testing and meets BOTH of the following criteria:
    - i. Member has a C4 level below the lower limit of normal as defined by the laboratory performing the test
    - ii. Member meets ONE of the following criteria:
      - (a) C1 inhibitor (C1-INH) antigenic level below the lower limit of normal as defined by the laboratory performing the test
      - (b) Normal C1-INH antigenic level and a low C1-INH functional level (functional C1-INH less than 50% or C1-INH functional level below the lower limit of normal as defined by the laboratory performing the test).
  - b. Member has normal C1 inhibitor as confirmed by laboratory testing and meets ONE of the following criteria:
    - i. Member has an F12, angiopoietin-1, plasminogen, or kininogen-1 (KNG1) gene mutation as confirmed by genetic testing
    - ii. Member has a documented family history of angioedema and the angioedema was refractory to a trial of high-dose antihistamine (e.g., cetirizine) for at least one month.
4. Member has adverse effect or contraindication to icatibant (generic for Firazyr)

### **Continuation of Therapy**

Reauthorization may be granted for members when ALL of the following are met, and documentation is provided:

1. Member meets all initial approval criteria.
2. Member has experienced a significant reduction in frequency of attacks (e.g.  $\geq 50\%$ ) since starting treatment.
3. Member has reduced the use of medications to treat acute attacks.

### **Limitations**

1. Initial approvals and reauthorizations will be for 6 months.
2. The following quantity limits apply:

|                   |                         |
|-------------------|-------------------------|
| Oraldeyo capsules | 30 capsules per 30 days |
|-------------------|-------------------------|

### **References**

1. Oraldeyo [package insert]. Durham, NC: BioCryst Pharmaceuticals, Inc.; December 2020.
2. Maurer M, Magerl M, Ansotegui I, et al. The international WAO/EAACI guideline for the management of hereditary angioedema – the 2017 revision and update. *Allergy*. 2018;00:1-22.
3. Henao MP, Kraschnewski J, Kelbel T, Craig T. Diagnosis and screening of patients with hereditary angioedema in primary care. *Therapeutics and Clin Risk Management*. 2016; 12: 701-711.
4. Zuraw B, Lumry WR, Johnston DT, et al. Oral once-daily berotralstat for the prevention of hereditary angioedema attacks: A randomized, double-blind, placebo-controlled phase 3 trial. *J Allergy Clin Immunol*. 2020;S0091-6749(20)31484-6.

### **Review History**

05/19/2021 – Created and Reviewed May P&T. Effective 07/01/2021.

9/21/2022 – Reviewed and Updated for Sept P&T; added requirement of adverse effect or contraindication to icatibant (generic for Firazyr). Separated out MH vs. Comm/Exch. Effective 1/1/2023.

