Clinical Policy: Ibalizumab-uiyk (Trogarzo)
Reference Number: CP.PHAR.378
Effective Date: 04.17.18
Last Review Date: 05.19
Line of Business: Commercial, HIM-Medical Benefit, Medicaid

See Important Reminder at the end of this policy for important regulatory and legal information.

Description
Ibalizumab-uiyk (Trogarzo™) is a CD4-directed post-attachment human immunodeficiency virus type 1 (HIV-1) inhibitor.

FDA Approved Indication(s)
Trogarzo is indicated for the treatment of HIV-1 infection, in combination with other antiretroviral(s), in heavily treatment-experienced adults with multidrug resistant HIV-1 infection failing their current antiretroviral regimen.

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® that Trogarzo is medically necessary when the following criteria are met:

I. Initial Approval Criteria
   A. HIV-1 Infection (must meet all):
      1. Diagnosis of multidrug resistant HIV-1 infection;
      2. Prescribed by or in consultation with an infectious disease or HIV specialist;
      3. Age ≥ 18 years;
      4. Documentation of resistance to at least 1 antiretroviral agent from each of 4 classes (NRTI, NNRTI, PI, INSTI), unless contraindicated or clinically significant adverse effects are experienced;
      5. Failure of Fuzeon®, unless resistant, contraindicated, or clinically significant adverse effects are experienced;
      6. If CCR5-tropic: Failure of Selzentry®, unless resistant, contraindicated, or clinically significant adverse effects are experienced;
      7. Current (within the past 30 days) HIV ribonucleic acid viral load of ≥ 200 copies/mL;
      8. Prescribed concurrently with additional antiretroviral agents to which member is susceptible, if available;
      9. Dose does not exceed 2,000 mg (10 vials) IV loading dose* and/or 800 mg (4 vials) IV every 14 days.
         *A loading dose may be repeated if the member misses scheduled maintenance dose by 3 days or more.

Approval duration: 6 months
B. Other diagnoses/indications
1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial and CP.PMN.53 for Medicaid and HIM-Medical Benefit.

II. Continued Therapy
A. HIV-1 Infection (must meet all):
   1. Currently receiving medication via Centene benefit, or documentation supports that member is currently receiving Trogarzo for multidrug resistant HIV-1 infection and has received this medication for at least 30 days;
   2. Member is responding positively to therapy;
   3. If request is for a dose increase, new dose does not exceed 2,000 mg (10 vials) IV loading dose* and/or 800 mg (4 vials) IV every 14 days.  
   *A loading dose may be repeated if the member misses scheduled maintenance dose by 3 days or more. 
Approval duration: 12 months

B. Other diagnoses/indications (must meet 1 or 2):
   1. Currently receiving medication via Centene benefit and documentation supports positive response to therapy.  
   Approval duration: Duration of request or 6 months (whichever is less); or
   2. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial and CP.PMN.53 for Medicaid and HIM-Medical Benefit.

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 and CP.PMN.53 for Medicaid and HIM-Medical Benefit or evidence of coverage documents.

IV. Appendices/General Information
   Appendix A: Abbreviation/Acronym Key
   FDA: Food and Drug Administration
   HIV-1: human immunodeficiency virus type 1
   INSTI: integrase strand transfer inhibitors
   NNRTI: non-nucleoside reverse transcriptase inhibitor
   NRTI: nucleos(t)ide reverse transcriptase inhibitor
   PI: protease inhibitor

   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.
<table>
<thead>
<tr>
<th>Drug Name</th>
<th>Dosing Regimen</th>
<th>Dose Limit/ Maximum Dose</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nucleos(t)ide reverse transcriptase inhibitors (NRTIs) (e.g., abacavir, tenofovir disoproxil fumarate, Emtriva®)</td>
<td>Refer to prescribing information</td>
<td>Refer to prescribing information</td>
</tr>
<tr>
<td>Non-nucleoside reverse transcriptase inhibitors (NNRTIs) (e.g., efavirenz, nevirapine, Edurant®)</td>
<td>Refer to prescribing information</td>
<td>Refer to prescribing information</td>
</tr>
<tr>
<td>Integrase strand transfer inhibitors (INSTIs) (e.g., Tivicay®, Isentress®)</td>
<td>Refer to prescribing information</td>
<td>Refer to prescribing information</td>
</tr>
<tr>
<td>Protease inhibitors (PIs) (e.g., atazanavir, fosamprenavir, Invirase®, Viracept®)</td>
<td>Refer to prescribing information</td>
<td>Refer to prescribing information</td>
</tr>
<tr>
<td>Fuzeon® (enfurvirtide, T-20)</td>
<td>Refer to prescribing information</td>
<td>Adults: 180 mg/day Children 6 years and older: 4 mg/kg/day</td>
</tr>
<tr>
<td>Selzentry® (maraviroc, MVC)</td>
<td>Refer to prescribing information</td>
<td>600 mg/day; 1,200 mg/day if taking a potent CYP3A inducer</td>
</tr>
<tr>
<td>Fixed-dose combinations (e.g., Genvoya®, Stribild®, Odefsey®, Descovy®, Truvada®)</td>
<td>Refer to prescribing information</td>
<td>Refer to prescribing information</td>
</tr>
</tbody>
</table>

Therapeutic alternatives are listed as Brand name® (generic) when the drug is available by brand name only and generic (Brand name®) when the drug is available by both brand and generic.

Appendix C: Contraindications/Boxed Warnings
None reported

V. Dosage and Administration

<table>
<thead>
<tr>
<th>Indication</th>
<th>Dosing Regimen</th>
<th>Maximum Dose</th>
</tr>
</thead>
<tbody>
<tr>
<td>HIV-1 infection</td>
<td>A single loading dose of 2,000 mg IV, followed by a maintenance dose of 800 mg every 2 weeks. If a maintenance dose is missed by 3 days or longer beyond the scheduled dosing day, a loading dose of 2,000 mg should be administered as early as possible prior to resuming maintenance dosing of 800 mg every 2 weeks thereafter.</td>
<td>A loading dose of 2,000 mg up to every 17 days* A maintenance dose of 800 mg every 14 days</td>
</tr>
</tbody>
</table>

*Frequency of every 17 days was calculated from frequency of maintenance dose (every 14 days) plus minimum number of days that the dose is missed to qualify for another loading dose (3 days).
VI. Product Availability
Injection in single-dose vial: 200 mg/1.33 mL (150 mg/mL)

VII. References

Coding Implications
Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

<table>
<thead>
<tr>
<th>HCPCS Codes</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>N/A</td>
<td>Injection, ibalizumab, 200 mg</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Reviews, Revisions, and Approvals</th>
<th>Date</th>
<th>P&amp;T Approval Date</th>
</tr>
</thead>
<tbody>
<tr>
<td>Policy created</td>
<td>04.17.18</td>
<td>05.18</td>
</tr>
<tr>
<td>Clarified dosing regimen under section V. No significant changes.</td>
<td>06.19.18</td>
<td></td>
</tr>
<tr>
<td>No significant changes: added HIM-Medical Benefit to included lines of business per SDC.</td>
<td>10.03.18</td>
<td></td>
</tr>
<tr>
<td>2Q 2019 annual review: no significant changes; references reviewed and updated.</td>
<td>01.23.19</td>
<td>05.19</td>
</tr>
</tbody>
</table>

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
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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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Note:
For Medicaid members, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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