Clinical Policy: Siltuximab (Sylvant)
Reference Number: CP.PHAR.329
Effective Date: 03.01.17
Last Review Date: 02.20
Line of Business: HIM, Medicaid

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

Description
Siltuximab (Sylvant®) is an interleukin-6 (IL-6) antagonist.

FDA Approved Indication(s)
Sylvant is indicated for the treatment of patients with multicentric Castleman’s disease (MCD) who are human immunodeficiency virus (HIV) negative and human herpesvirus-8 (HHV-8) negative.

Limitation(s) of use: Sylvant was not studied in patients with MCD who are HIV positive or HHV-8 positive because Sylvant did not bind to virally produced IL-6 in a nonclinical study.

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 Sylvant is medically necessary when the following criteria are met:

I. Initial Approval Criteria
   A. Castleman’s Disease (must meet all):
      1. Diagnosis of Castleman’s disease* (CD, angiofollicular lymph node hyperplasia) confirmed by biopsy of involved tissue (usually a lymph node);
      2. Prescribed by or in consultation with an oncologist;
      3. Age ≥ 18 years;
      4. Meets one of the following (a or b):
         a. FDA-approved use for treatment of multicentric** Castleman’s disease (MCD);
         b. NCCN-recommended use for second-line, single-agent treatment of relapsed or refractory unicentric** Castleman’s disease (UCD);
      5. Meets all of the following parameters prior to treatment (a, b, c, d, and e):
         a. Human immunodeficiency virus (HIV) negative;
         b. Human herpesvirus-8 (HHV-8) negative;
         c. Absolute neutrophil count: ≥ 1.0 x 10^9/L;
         d. Platelet count ≥ 75 x 10^9/L;
         e. Hemoglobin < 17 g/dL;
      6. Request meets one of the following (a or b):
         a. Dose does not exceed 11 mg/kg every 3 weeks.
b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use *(prescriber must submit supporting evidence)*.  

**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): HIM.PHAR.21 for health insurance marketplace and CP.PMN.53 for Medicaid.

**II. Continued Therapy**  
**A. Castleman’s Disease** (must meet all):  
1. Currently receiving medication via Centene benefit, or documentation supports that member is currently receiving Sylvant for a covered indication and has received this medication for at least 30 days;  
2. Member is responding positively to therapy;  
3. Meets the following laboratory parameters:  
   a. Absolute neutrophil count: $\geq 1.0 \times 10^9$/L;  
   b. Platelet count $\geq 50 \times 10^9$/L;  
   c. Hemoglobin < 17 g/dL;  
4. If request is for a dose increase, new dose does not exceed (a or b):  
   a. 11 mg/kg every 3 weeks;  
   b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use *(prescriber must submit supporting evidence)*.  

**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): HIM.PHAR.21 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 – HIM.PHAR.21 for health insurance marketplace and CP.PMN.53 for Medicaid or evidence of coverage documents.

**IV. Appendices/General Information**  
*Appendix A: Abbreviation/Acronym Key*  
CD: Castleman’s disease  
FDA: Food and Drug Administration  
HHV-8: negative and human hperesvirus-8  
HIV: human immunodeficiency virus
MCD: multicentric Castleman’s disease  UCD: unicentric Castleman’s disease

Appendix B: Therapeutic Alternatives
Not applicable

Appendix C: Contraindications/Boxed Warnings
• Contraindication(s): severe hypersensitivity reaction to siltuximab or any of the excipients in Sylvant
• Boxed warning(s): none reported

Appendix D: General Information
*Group of lymphoproliferative disorders (classified under non-Hodgkin B-cell lymphomas) that share common histologic features
**MCD (systemic disease with symptoms that may include generalized peripheral lymphadenopathy, hepatosplenomegaly, frequent fevers, night sweats); UCD (localized disease that generally is asymptomatic)

V. Dosage and Administration

<table>
<thead>
<tr>
<th>Indication</th>
<th>Dosing Regimen</th>
<th>Maximum Dose</th>
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<tr>
<td>Castleman’s disease</td>
<td>11 mg/kg over 1 hour IV every 3 weeks</td>
<td>11 mg/kg</td>
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VI. Product Availability
Lyophilized powder in a single-use vial: 100 mg and 400 mg

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>
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<tbody>
<tr>
<td>J2860</td>
<td>Injection, siltuximab, 10 mg</td>
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Reviews, Revisions, and Approvals

<table>
<thead>
<tr>
<th>Description</th>
<th>Date</th>
<th>P&amp;T Approval Date</th>
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<tbody>
<tr>
<td>Policy split from CP.PHAR.183 Excellus Other Specialty Pharmacy.</td>
<td>02.01.17</td>
<td>03.17</td>
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<tr>
<td>Updated references and template.</td>
<td>08.20.17</td>
<td>11.17</td>
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<tr>
<td>1Q18 annual review: No significant changes; References reviewed and updated.</td>
<td>11.20.17</td>
<td>02.18</td>
</tr>
<tr>
<td>1Q 2019 annual review: added prescriber requirement; allowed COC for continued approval; added option for off-label dosing as supported by guidelines or literature; references reviewed and updated.</td>
<td>11.20.18</td>
<td>02.19</td>
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<tr>
<td>1Q 2020 annual review: added HIM line of business; references reviewed and updated.</td>
<td>11.08.19</td>
<td>02.20</td>
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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.

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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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