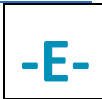


**Sylvant® (siltuximab)**  
**(Intravenous)**



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**I. Length of Authorization <sup>2,6</sup>**

Coverage will be provided for 6 months and may be renewed (unless otherwise specified)

- Management of CAR T-Cell-Related Toxicities: Coverage will be provided for 1 dose only and may NOT be renewed

**II. Dosing Limits**

**Quantity Limit (max daily dose) [NDC Unit]:**

- Sylvant 100 mg single-dose vial: 3 vials per 21-day supply
- Sylvant 400 mg single-dose vial: 3 vials per 21-day supply

**Max Units (per dose and over time) [HCPCS Unit]:**

| Diagnosis                                      | Billable Units | Interval (days)          |
|--|----------------|--------------------------|
| MCD, UCD                                       | 130            | 21                       |
| Management of Immunotherapy-Related Toxicities | 130            | 1 course of therapy only |

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

Coverage is provided in the following conditions:

- Patient is at least 18 years of age; **AND**

**Universal Criteria <sup>1</sup>**

- Patient is human immunodeficiency virus (HIV) negative; **AND**
- Patient is human herpes virus-8 (HHV-8) negative; **AND**
- Patient is currently free of all clinically significant infections; **AND**

- Patient will NOT receive any live vaccines during treatment with siltuximab; **AND**
- Must be used as a single agent; **AND**

**Multicentric Castleman’s Disease (MCD) † Φ<sup>1-4</sup>**

**Unicentric Castleman’s Disease (UCD) ‡<sup>2</sup>**

- Used as second-line therapy for relapsed or refractory disease

**Management of CAR T-Cell-Related Toxicities ‡<sup>2,6</sup>**

- Patient has received or will be receiving chimeric antigen receptor (CAR)-T cell therapy (e.g., axicabtagene ciloleucel, brexucabtagene autoleucel, idecabtagene vicleucel, lisocabtagene maraleucel, tisagenlecleucel, ciltacabtagene autoleucel, etc.); **AND**
  - Used for the management of Grade 4 cytokine release syndrome (CRS); **AND**
    - Patient is refractory to high-dose corticosteroids and anti-interleukin-6 therapy (e.g., tocilizumab); **OR**
  - Used as a replacement for the second dose of tocilizumab when supplies are limited or unavailable; **AND**
    - Used for Grade 1-4 CRS; **OR**
    - Used for Grade 1-4 neurotoxicity as additional therapy if the patient has concurrent CRS

**Preferred therapies and recommendations are determined by review of clinical evidence. NCCN category of recommendation is taken into account as a component of this review. Regimens deemed equally efficacious (i.e., those having the same NCCN categorization) are considered to be therapeutically equivalent.**

† FDA Approved Indication(s); ‡ Compendia Recommended Indication(s); Φ Orphan Drug

#### IV. Renewal Criteria <sup>1,2,6</sup>

Coverage may be renewed based upon the following criteria:

- Patient continues to meet the universal and other indication-specific relevant criteria such as concomitant therapy requirements (not including prerequisite therapy), performance status, etc. identified in section III; **AND**
- Disease response with treated as defined by stabilization of disease or decrease in size of tumor or tumor spread; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: gastrointestinal perforation, severe infusion related reactions and hypersensitivity, etc.

**Management of CAR T-Cell-Related Toxicities**

- May not be renewed

## V. Dosage/Administration <sup>1,3,4,6</sup>

| Indication                                  | Dose  |
|---|---|
| MCD, UCD                                    | Administer 11 mg/kg intravenously every 21 days until treatment failure |
| Management of CAR T-Cell-Related Toxicities | Administer 11 mg/kg intravenously one time only                         |

## VI. Billing Code/Availability Information

### HCPCS Code:

- J2860 – Injection, siltuximab, 10 mg; 10 mg = 1 billable unit

### NDC(s):

- Sylvant 100 mg lyophilized powder in a single-dose vial: 73090-0420-xx
- Sylvant 400 mg lyophilized powder in a single-dose vial: 73090-0421-xx

## VII. References (STANDARD)

1. Sylvant [package insert]. Hemel Hempstead, Hertfordshire, U.K.; EUSA Pharma (UK), Ltd; December 2019. Accessed December 2023.
2. Referenced with permission from the NCCN Drugs & Biologics Compendium (NCCN Compendium<sup>®</sup>) for siltuximab. National Comprehensive Cancer Network, 2024. The NCCN Compendium<sup>®</sup> is a derivative work of the NCCN Guidelines<sup>®</sup>. NATIONAL COMPREHENSIVE CANCER NETWORK<sup>®</sup>, NCCN<sup>®</sup>, and NCCN GUIDELINES<sup>®</sup> are trademarks owned by the National Comprehensive Cancer Network, Inc. To view the most recent and complete version of the Compendium, go online to NCCN.org. Accessed December 2023.
3. van Rhee F, Wong RS, Munshi N, et al. Siltuximab for multicentric Castleman's disease: a randomised, double-blind, placebo-controlled trial. *Lancet Oncol.* 2014 Aug;15(9):966-74. doi: 10.1016/S1470-2045(14)70319-5. Epub 2014 Jul 17.
4. Kurzrock R, Voorhees PM, Casper C, et al. A phase I, open-label study of siltuximab, an anti-IL-6 monoclonal antibody, in patients with B-cell non-Hodgkin lymphoma, multiple myeloma, or Castleman disease. *Clin Cancer Res.* 2013 Jul 1;19(13):3659-70. doi: 10.1158/1078-0432.CCR-12-3349. Epub 2013 May 9.
5. Chen F, Teachey DT, Pequignot E, et al. Measuring IL-6 and sIL-6R in serum from patients treated with tocilizumab and/or siltuximab following CAR T cell therapy. *J Immunol Methods.* 2016 Jul;434:1-8. doi: 10.1016/j.jim.2016.03.005.
6. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines<sup>®</sup>) for Management of Immunotherapy-Related Toxicities Version 1.2024. National

Comprehensive Cancer Network, 2024. NATIONAL COMPREHENSIVE CANCER NETWORK®, NCCN®, and NCCN GUIDELINES® are trademarks owned by the National Comprehensive Cancer Network, Inc. To view the most recent and complete version of the Guidelines, go online to NCCN.org. Accessed December 2023.

## VIII. References (ENHANCED)

- 1e. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines®) B-Cell Lymphomas, Version 6.2023. National Comprehensive Cancer Network, 2024. The NCCN Compendium® is a derivative work of the NCCN Guidelines®. NATIONAL COMPREHENSIVE CANCER NETWORK®, NCCN®, and NCCN GUIDELINES® are trademarks owned by the National Comprehensive Cancer Network, Inc. To view the most recent and complete version of the Compendium, go online to NCCN.org. Accessed December 2023.
- 2e. Gérard L, Bérezné A, Galicier L, et al. Prospective study of rituximab in chemotherapy-dependent human immunodeficiency virus associated multicentric Castleman's disease: ANRS 117 CastlemaB Trial. *J Clin Oncol.* 2007 Aug 1;25(22):3350-6.
- 3e. Zhang L, Zhao AL, Duan MH, et al. Phase 2 study using oral thalidomide-cyclophosphamide-prednisone for idiopathic multicentric Castleman disease. *Blood* 2019;133:1720-1728.
- 4e. Magellan Rx Management. Sylvant Clinical Literature Review Analysis. Last updated December 2023. Accessed December 2023.

## Appendix 1 – Covered Diagnosis Codes

| ICD-10   | ICD-10 Description   |
|----------|--|
| D47.Z2   | Castleman disease  |
| D89.831  | Cytokine release syndrome, grade 1   |
| D89.832  | Cytokine release syndrome, grade 2   |
| D89.833  | Cytokine release syndrome, grade 3   |
| D89.834  | Cytokine release syndrome, grade 4   |
| D89.839  | Cytokine release syndrome, grade unspecified   |
| G92.00   | Immune effector cell-associated neurotoxicity syndrome, grade unspecified                        |
| G92.01   | Immune effector cell-associated neurotoxicity syndrome, grade 1                                  |
| G92.02   | Immune effector cell-associated neurotoxicity syndrome, grade 2                                  |
| G92.03   | Immune effector cell-associated neurotoxicity syndrome, grade 3                                  |
| G92.04   | Immune effector cell-associated neurotoxicity syndrome, grade 4                                  |
| T80.82XA | Complication of immune effector cellular therapy, initial encounter                              |
| T80.82XS | Complication of immune effector cellular therapy, sequela  |
| T80.89XA | Other complications following infusion, transfusion and therapeutic injection, initial encounter |
| T80.89XS | Other complications following infusion, transfusion and therapeutic injection, sequela           |

## 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/LCA/LCD): 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, LLC                                 |
| M (11)  | NC, SC, WV, VA (excluding below)  | Palmetto GBA, LLC                                 |
| L (12)  | DE, MD, PA, NJ, DC (includes Arlington & Fairfax counties and the city of Alexandria in VA) | Novitas Solutions, Inc.                           |
| K (13 & 14)   | NY, CT, MA, RI, VT, ME, NH  | National Government Services, Inc. (NGS)          |
| 15  | KY, OH  | CGS Administrators, LLC                           |