

Drug Policy:

Vyloy™ (zolbetuximab-clzb)

POLICY NUMBER UM ONC_1512	SUBJECT Vyloy™ (zolbetuximab-clzb)		DEPT/PROGRAM UM Dept	PAGE 1 OF 3
DATES COMMITTEE REVIEWED 12/12/24	APPROVAL DATE December 12, 2024	EFFECTIVE DATE December 27, 2024	COMMITTEE APPROVAL DATES 12/12/24	
PRIMARY BUSINESS OWNER: UM		COMMITTEE/BOARD APPROVAL Evolut Specialty Services Clinical Guideline Review Committee		
NCQA STANDARDS UM 2		ADDITIONAL AREAS OF IMPACT		
CMS REQUIREMENTS	STATE/FEDERAL REQUIREMENTS		APPLICABLE LINES OF BUSINESS Commercial, Exchange, Medicaid	

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I. PURPOSE

To define and describe the accepted indications for Vyloy (zolbetuximab-clzb) usage in the treatment of cancer, including FDA approved indications, and off-label indications.

Evolut is responsible for processing all medication requests from network ordering providers. Medications not authorized by Evolut may be deemed as not approvable and therefore not reimbursable.

The use of this drug must be supported by one of the following: FDA approved product labeling, CMS-approved compendia, National Comprehensive Cancer Network (NCCN), American Society of Clinical Oncology (ASCO) clinical guidelines, or peer-reviewed literature that meets the requirements of the CMS Medicare Benefit Policy Manual Chapter 15.

II. INDICATIONS FOR USE/INCLUSION CRITERIA

A. Continuation requests for a not-approvable medication shall be exempt from this Evolent policy provided:

1. The member has not experienced disease progression on the requested medication **AND**
2. The requested medication was used within the last year without a lapse of more than 30 days of having an active authorization **AND**
3. Additional medication(s) are not being added to the continuation request.

B. Gastric or gastroesophageal junction (GEJ) adenocarcinoma

1. Vyloy (zolbetuximab-clzb) may be used in combination with fluoropyrimidine- and platinum-containing chemotherapy for the first-line treatment of adult members with locally advanced unresectable or metastatic human epidermal growth factor receptor 2 (HER2)-negative gastric or gastroesophageal junction adenocarcinoma whose tumors are claudin (CLDN) 18.2 positive as determined by an FDA-approved test.

III. EXCLUSION CRITERIA

- A. Disease progression while taking Vyloy (zolbetuximab-clzb).
- B. Members with tumors that are claudin (CLDN) 18.2 negative.
- C. Dosing exceeds single dose limit of 800 mg/m2.
- D. Investigational use of Vyloy (zolbetuximab-clzb) with an off-label indication that is not sufficient in evidence or is not generally accepted by the medical community. Sufficient evidence that is not supported by CMS recognized compendia or acceptable peer reviewed literature is defined as any of the following:
 1. Whether the clinical characteristics of the patient and the cancer are adequately represented in the published evidence.
 2. Whether the administered chemotherapy/biologic therapy/immune therapy/targeted therapy/other oncologic therapy regimen is adequately represented in the published evidence.
 3. Whether the reported study outcomes represent clinically meaningful outcomes experienced by patients. Generally, the definitions of Clinically Meaningful outcomes are those recommended by ASCO, e.g., Hazard Ratio of < 0.80 and the recommended survival benefit for OS and PFS should be at least 3 months.
 4. Whether the experimental design, considering the drugs and conditions under investigation, is appropriate to address the investigative question. (For example, in some clinical studies, it may be unnecessary or not feasible to use randomization, double blind trials, placebos, or crossover).
 5. That non-randomized clinical trials with a significant number of subjects may be a basis for supportive clinical evidence for determining accepted uses of drugs.
 6. That case reports are generally considered uncontrolled and anecdotal information and do not provide adequate supportive clinical evidence for determining accepted uses of drugs.
 7. That abstracts (including meeting abstracts) without the full article from the approved peer-reviewed journals lack supporting clinical evidence for determining accepted uses of drugs.

IV. CODING INFORMATION

HCPCS Code	Description
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C9399	zolbetuximab-clzb
J9999	zolbetuximab-clzb

V. MEDICATION MANAGEMENT

- A. Please refer to the FDA label/package insert for details regarding these topics.

VI. APPROVAL AUTHORITY

- A. Review – Utilization Management Department
- B. Final Approval – Utilization Management Committee

VII. ATTACHMENTS

- A. None

VIII. REFERENCES

- A. Vyloy prescribing information 2024. Astellas Pharma US, Inc., Northbrook, Illinois
- B. Shitara K, et al. Zolbetuximab plus mFOLFOX6 in patients with CLDN18.2-positive, HER2-negative, untreated, locally advanced unresectable or metastatic gastric or gastro-oesophageal junction adenocarcinoma (SPOTLIGHT): a multicentre, randomised, double-blind, phase 3 trial. The Lancet, 2023 May 20, Volume 401, Issue 10389, 1655 – 1668. DOI: [10.1016/S0140-6736\(23\)00620-7](https://doi.org/10.1016/S0140-6736(23)00620-7)
- C. Shah MA, et al. Zolbetuximab plus CAPOX in CLDN18.2-positive gastric or gastroesophageal junction adenocarcinoma: the randomized, phase 3 GLOW trial. Nat Med. 2023 Aug;29(8):2133-2141. doi: 10.1038/s41591-023-02465-7
- D. Clinical Pharmacology Elsevier Gold Standard 2024.
- E. Micromedex® Healthcare Series: Micromedex Drugdex Ann Arbor, Michigan 2024.
- F. National Comprehensive Cancer Network. Cancer Guidelines and Drugs and Biologics Compendium 2024.
- G. Ellis LM, et al. American Society of Clinical Oncology perspective: Raising the bar for clinical trials by defining clinically meaningful outcomes. J Clin Oncol. 2014 Apr 20;32(12):1277-80.
- H. Medicare Benefit Policy Manual Chapter 15 Covered Medical and Other Health Services: <https://www.cms.gov/Regulations-and-Guidance/Guidance/Manuals/Downloads/bp102c15.pdf>.