

Specialty Guideline Management

Cyramza

Products Referenced by this Document

Drugs that are listed in the following table include both brand and generic and all dosage forms and strengths unless otherwise stated. Over-the-Counter (OTC) products are not included unless otherwise stated.

Brand Name	Generic Name
Cyramza	ramucirumab

Indications

The indications below including FDA-approved indications and compendial uses are considered a covered benefit provided that all the approval criteria are met and the member has no exclusions to the prescribed therapy.

FDA-approved Indications¹

- Gastric Cancer: Cyramza as a single agent, or in combination with paclitaxel, is indicated for the treatment of patients with advanced or metastatic, gastric or gastro-esophageal junction (GEJ) adenocarcinoma with disease progression on or after prior fluoropyrimidine-or platinum-containing chemotherapy.
- Non-Small Cell Lung Cancer (NSCLC):
 - Cyramza, in combination with docetaxel, is indicated for the treatment of patients with metastatic NSCLC with disease progression on or after platinum-based chemotherapy. Patients with epidermal growth factor receptor (EGFR) or anaplastic lymphoma kinase (ALK) genomic tumor aberrations should have disease progression on FDA-approved therapy for these aberrations prior to receiving Cyramza.
 - Cyramza, in combination with erlotinib, is indicated for the first-line treatment of patients with metastatic NSCLC whose tumors have epidermal growth factor receptor (EGFR) exon 19 deletions or exon 21 (L858R) substitution mutations.

- Colorectal Cancer: Cyramza, in combination with FOLFIRI (irinotecan, folinic acid, and fluorouracil), is indicated for the treatment of patients with metastatic colorectal cancer (mCRC) with disease progression on or after prior therapy with bevacizumab, oxaliplatin, and a fluoropyrimidine.
- Hepatocellular Carcinoma: Cyramza as a single agent, is indicated for the treatment of patients with hepatocellular carcinoma (HCC) who have an alpha fetoprotein (AFP) of greater than or equal to 400 ng/mL and have been treated with sorafenib.

Compendial Uses^{2,3}

- Esophageal adenocarcinoma
- Colorectal cancer, advanced, including anal adenocarcinoma and appendiceal adenocarcinoma
- NSCLC, EGFR mutation positive, recurrent, advanced
- Mesothelioma
- Thymic carcinoma

All other indications are considered experimental/investigational and not medically necessary.

Documentation

Submission of the following information is necessary to initiate the prior authorization review: EGFR mutation testing results and alpha fetoprotein (AFP) level results (where applicable).

Coverage Criteria

Gastric, Gastro-esophageal Junction (GEJ), Esophagogastric Junction (EGJ), and Esophageal Adenocarcinoma^{1,2}

Authorization of 12 months may be granted for treatment of gastric, gastro-esophageal junction (GEJ), esophagogastric junction (EGJ), and esophageal adenocarcinoma for members who are not surgical candidates or who have unresectable locally advanced, recurrent or metastatic disease, when used as subsequent therapy as a single agent, in combination with paclitaxel, or in combination with irinotecan with or without fluorouracil.

Non-Small Cell Lung Cancer (NSCLC)^{1,2}

Authorization of 12 months may be granted for treatment of recurrent, advanced or metastatic NSCLC when either of the following criteria is met:

Reference number(s)
1679-A

- Used in combination with docetaxel as subsequent therapy.
- Used in combination with erlotinib for EGFR exon 19 deletion or exon 21 (L858R) substitution mutation positive disease.

Colorectal Cancer (CRC)¹⁻³

Authorization of 12 months may be granted for treatment of advanced or metastatic colorectal cancer, including anal adenocarcinoma and appendiceal adenocarcinoma, in combination with FOLFIRI (irinotecan, folinic acid, and 5-fluorouracil) or irinotecan.

Hepatocellular Carcinoma (HCC)^{1,2}

Authorization of 12 months may be granted for subsequent treatment of progressive hepatocellular carcinoma as a single agent in members who have an alpha fetoprotein (AFP) of greater than or equal to 400 ng/mL.

Mesothelioma²

Authorization of 12 months may be granted for the subsequent treatment of pleural mesothelioma, pericardial mesothelioma, or tunica vaginalis testis mesothelioma when used in combination with gemcitabine.

Thymic Carcinoma²

Authorization of 12 months may be granted for the treatment of thymic carcinoma when any of the following are met:

- Member has recurrent, advanced, or metastatic disease and the requested medication will be used in combination with carboplatin and paclitaxel and continued as a single agent maintenance therapy, or
- Member has had a R1 or R2 resection and the requested medication will be used in combination with carboplatin and paclitaxel as postoperative treatment, or
- Member has surgically resectable disease, R0 resection is considered uncertain, and the requested medication will be used in combination with carboplatin and paclitaxel as preoperative treatment.

Continuation of Therapy

NSCLC

Authorization of 12 months may be granted for continued treatment in members requesting reauthorization for NSCLC when either of the following criteria is met:

Reference number(s)
1679-A

- There is no evidence of unacceptable toxicity or disease progression while on the current regimen, or
- Disease is T790M negative and there is no evidence of unacceptable toxicity

All other indications

Authorization of 12 months may be granted for continued treatment in members requesting reauthorization for an indication listed in the Coverage Criteria section when there is no evidence of unacceptable toxicity or disease progression while on the current regimen.

References

1. Cyramza [package insert]. Indianapolis, IN: Eli Lilly and Company; March 2022.
2. The NCCN Drugs & Biologics Compendium® © 2025 National Comprehensive Cancer Network, Inc. Available at: <https://www.nccn.org>. Accessed January 17, 2025.
3. National Comprehensive Cancer Network. NCCN Clinical Practice Guidelines in Oncology: Anal Carcinoma. Version 1.2025. https://www.nccn.org/professionals/physician_gls/pdf/anal.pdf. Accessed January 17, 2025.