

# **Medical Policy:**

### Cyramza® (ramucirumab) Intravenous

POLICY NUMBER	LAST REVIEW	ORIGIN DATE
MG.MM.PH.48 March 20, 2024		

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The treating physician or primary care provider must submit to EmblemHealth, or ConnectiCare, as applicable (hereinafter jointly referred to as "EmblemHealth"), the clinical evidence that the member meets the criteria for the treatment or surgical procedure. Without this documentation and information, EmblemHealth will not be able to properly review the request preauthorization or post-payment review. The clinical review criteria expressed below reflects how EmblemHealth determines whether certain services or supplies are medically necessary. This clinical policy is not intended to pre-empt the judgment of the reviewing medical director or dictate to health care providers how to practice medicine. Health care providers are expected to exercise their medical judgment in rendering appropriate care.

EmblemHealth established the clinical review criteria based upon a review of currently available clinical information (including clinical outcome studies in the peer reviewed published medical literature, regulatory status of the technology, evidence-based guidelines of public health and health research agencies, evidence-based guidelines and positions of leading national health professional organizations, views of physicians practicing in relevant clinical areas, and other relevant factors). EmblemHealth expressly reserves the right to revise these conclusions as clinical information changes and welcomes further relevant information. Each benefit program defines which services are covered. The conclusion that a particular service or supply is medically necessary does not constitute a representation or warranty that this service or supply is covered and/or paid for by EmblemHealth, as some programs exclude coverage for services or supplies that EmblemHealth considers medically necessary.

If there is a discrepancy between this guideline and a member's benefits program, the benefits program will govern. Identification of selected brand names of devices, tests and procedures in a medical coverage policy is for reference only and is not an endorsement of any one device, test or procedure over another. In addition, coverage may be mandated by applicable legal requirements of a state, the Federal Government or the Centers for Medicare & Medicaid Services (CMS) for Medicare and Medicaid members. All coding and web site links are accurate at time of publication.

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#### **Definition**

Cyramza (ramucirumab) is a vascular endothelial growth factor (VEGF) receptor 2 antagonist that inhibits ligand-stimulated activation of the VEGF receptor 2, ligand-induced proliferation, and migration of human endothelial cells. Unlike all clinically approved angiogenesis inhibitors, the fully human monoclonal antibody ramucirumab, specifically inhibits VEGFR-2.

### **Length of Authorization**

Coverage will be provided for six months and may be renewed.

## **Dosing Limits [Medical Benefit]**

Max Units (per dose and over time):

Gastric/Esophageal/Esophagogastric Junction Cancers, Colorectal Cancer and HCC: 180 billable units every 14 days

NSCLC: 240 billable units every 14 days

#### Guideline

#### I. INITIAL CRITERIA

Cyramza (ramucirumab) may be considered medically necessary for the following diagnosis when all of the following criteria are met:

#### 1. Gastric, Esophageal and Gastro-esophageal Junction Cancers, initiation:

- A. Patient is 18 years of age or older; AND
- B. Patient has advanced or metastatic disease; AND
- C. Used as a single agent OR in combination with paclitaxel OR used in combination with irinotecan; AND
- D. Must be used as subsequent therapy, progressing after treatment with fluoropyrimidine- or platinum-containing chemotherapy

### 2. Non-small cell lung cancer, initiation:

- A. Patient is 18 years of age or older; AND
- B. Patient meets **ONE** of the following criteria (i or ii):
  - i. Cyramza will be used as first-line therapy; AND
    - a. Patient has epidermal growth factor receptor (EGFR) exon 19 deletion or L858R mutation positive disease; AND
    - b. Cyramza will be used in combination with erlotinib; OR
  - ii. Cyramza will be used as subsequent therapy; AND
    - a. Cyramza will be used in combination with docetaxel intravenous infusion; AND
    - b. Patient has received targeted drug therapy if the patient's tumor is positive for a targetable mutation

Note: Examples of targetable mutations include sensitizing epidermal growth factor receptor mutation, anaplastic lymphoma kinase fusions

#### 3. Colorectal cancer, initiation

- A. Patient is 18 years of age or older; AND
- B. Patient's disease is metastatic; AND
- C. Cyramza is being used in combination with FOLFIRI (irinotecan, folinic acid, and 5-fluorouracil) **OR** irinotecan for metastatic disease that progressed on or after therapy with bevacizumab, oxaliplatin and a fluoropyrimidine

#### 4. Hepatocellular carcinoma:

- A. Patient is 18 years of age or older; AND
- B. Cyramza is being used as a single agent; AND
- C. Patient has an alpha fetoprotein of > 400 ng/mL; AND
- D. Patient has Child-Pugh Class A disease; AND
- E. Patient has been previously treated with sorafenib

#### II. Renewal Criteria

Coverage for Cyramza (ramucirumab) may be renewed when the following criteria are met:

- A. Disease response; AND
- B. Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include the following: hemorrhage, arterial thrombotic events, uncontrollable hypertension, infusion-related reactions, severe proteinuria (>3g/24h), gastrointestinal perforation, wound healing complications, etc.

#### **Limitations/Exclusions**

1. Ramucirumab is considered investigational and may not be considered medically necessary for the

following conditions: bladder cancer, breast cancer, chondrosarcoma, genitourinary tumor, fallopian tube cancer, head and neck cancer, malignant pleural diseases, melanoma, ovarian cancer, peritoneal carcinoma, prostate cancer, renal cell carcinoma, soft tissue sarcoma.

### **Dosing/Administration**

Indication	Dose
Colorectal Cancer, Gastric/Esophageal/	Administer 8 mg/kg intravenously every 14 days until disease progression or
Esophagogastric Junction Cancers,	unacceptable toxicity
Hepatocellular Carcinoma	
Non-Small Cell Lung Cancer	In combination with docetaxel:
	Administer 10 mg/kg intravenously every 21 days until disease progression or
	unacceptable toxicity
	In combination with erlotinib:
	Administer 10 mg/kg intravenously every 14 days until disease progression or
	unacceptable toxicity

## **Applicable Procedure Codes**

Code	Description
J9308	Injection, ramucirumab, 5 mg

## **Applicable NDCs**

Code	Description
00002-7669-01	Cyramza 100mg/10mL, 1 vial, single-dose in 1 carton, 10 mL in 1 vial, single-dose
00002-7678-01	Cyramza 500mg/50mL. 1 vial, single-dose in 1 carton, 50 mL in 1 vial, single-dose

## **ICD-10 Diagnoses**

Code	Description	
C15.3	Malignant neoplasm of upper third of esophagus	
C15.4	Malignant neoplasm of middle third of esophagus	
C15.5	Malignant neoplasm of lower third of esophagus	
C15.8	Malignant neoplasm of overlapping sites of esophagus	
C15.9	Malignant neoplasm of esophagus, unspecified	
C16.0	Malignant neoplasm of cardia	
C16.1	Malignant neoplasm of fundus of stomach	
C16.2	Malignant neoplasm of body of stomach	
C16.3	Malignant neoplasm of pyloric antrum	
C16.4	Malignant neoplasm of pylorus	
C16.5	Malignant neoplasm of lesser curvature of stomach, unspecified	
C16.6	Malignant neoplasm of greater curvature of stomach, unspecified	
C16.8	Malignant neoplasm of overlapping sites of stomach	
C16.9	Malignant neoplasm of stomach, unspecified	
C17.0	Malignant neoplasm duodenum	
C17.1	Malignant neoplasm jejunum	
C17.2	Malignant neoplasm ileum	
C17.8	Malignant neoplasm of overlapping sites of small intestines	

C17.9 Malignant neoplasm of small intestine, unspecified C18.1 Malignant neoplasm of accoum C18.1 Malignant neoplasm of accound C18.2 Malignant neoplasm of ascending colon C18.3 Malignant neoplasm of hepatic flexure C18.4 Malignant neoplasm of fransverse colon C18.5 Malignant neoplasm of splenic flexure C18.6 Malignant neoplasm of splenic flexure C18.6 Malignant neoplasm of sigmoid colon C18.7 Malignant neoplasm of sigmoid colon C18.8 Malignant neoplasm of sigmoid colon C18.8 Malignant neoplasm of overlapping sites of large intestines C19.9 Malignant neoplasm of overlapping sites of large intestines C19.9 Malignant neoplasm of overlapping sites of large intestines C19.0 Malignant neoplasm of overlapping sites of rectum, anus and anal canal C3.0 Malignant neoplasm of rectum C21.8 Malignant neoplasm of rectum C3.3 Malignant neoplasm of trachea C3.4.0 Malignant neoplasm of frem overlapping sites of rectum, anus and anal canal C3.4.0 Malignant neoplasm of his pronchus C3.4.0 Malignant neoplasm of left main bronchus C3.4.10 Malignant neoplasm of left main bronchus C3.4.10 Malignant neoplasm of upper lobe, right bronchus or lung C3.4.11 Malignant neoplasm of upper lobe, left bronchus or lung C3.4.12 Malignant neoplasm of upper lobe, left bronchus or lung C3.4.2 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.3 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.3 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.3 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neoplasm of lower lobe, left bronchus or lung C3.4.9 Malignant neop		
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C18.8 Malignant neoplasm of overlapping sites of large intestines  C18.9 Malignant neoplasm of colon, unspecified  C19 Malignant neoplasm of colon, unspecified  C20 Malignant neoplasm of rectosigmoid junction  C20 Malignant neoplasm of rectum  C21.8 Malignant neoplasm of overlapping sites of rectum, anus and anal canal  C33 Malignant neoplasm of trachea  C34.00 Malignant neoplasm of intrachea  C34.01 Malignant neoplasm of right main bronchus  C34.02 Malignant neoplasm of left main bronchus  C34.03 Malignant neoplasm of left main bronchus  C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung  C34.11 Malignant neoplasm of upper lobe, left bronchus or lung  C34.12 Malignant neoplasm of upper lobe, left bronchus or lung  C34.2 Malignant neoplasm of lower lobe, left bronchus or lung  C34.30 Malignant neoplasm of lower lobe, left bronchus or lung  C34.31 Malignant neoplasm of lower lobe, left bronchus or lung  C34.32 Malignant neoplasm of lower lobe, left bronchus or lung  C34.33 Malignant neoplasm of overlapping sites of unspecified bronchus and lung  C34.34 Malignant neoplasm of overlapping sites of inght bronchus and lung  C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung  C34.90 Malignant neoplasm of overlapping sites of left bronchus and lung  C34.91 Malignant neoplasm of unspecified part of inspecified bronchus or lung  C34.92 Malignant neoplasm of unspecified part of inght bronchus or lung  C34.93 Malignant neoplasm of unspecified part of right bronchus or lung  C34.90 Secondary malignant neoplasm of lung  C78.00 Secondary malignant neoplasm of lung  C78.01 Secondary malignant neoplasm of orer of lung and the production of the secondary malignant neoplasm of lung  C78.02 Secondary malignant neoplasm of retroperitoneum and peritoneum  C78.7 Secondary malignant neoplasm of other specified digestive organs  D37.9 Neoplasm of uncertain behavior of other specified digestive organ  D37.9 Neoplasm of uncertain behavior of other specified digestive organ  Z85.01 Personal histor	C18.6	Malignant neoplasm of descending colon
C18.9 Malignant neoplasm of colon, unspecified C19 Malignant neoplasm of rectosigmoid junction C20 Malignant neoplasm of rectosigmoid junction C21.8 Malignant neoplasm of rectosigmoid junction C21.8 Malignant neoplasm of overlapping sites of rectum, anus and anal canal C33 Malignant neoplasm of trachea C34.00 Malignant neoplasm of main bronchus C34.01 Malignant neoplasm of light main bronchus C34.02 Malignant neoplasm of light main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, light bronchus or lung C34.12 Malignant neoplasm of upper lobe, light bronchus or lung C34.13 Malignant neoplasm of lower lobe, bronchus or lung C34.2 Malignant neoplasm of lower lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, bronchus or lung C34.31 Malignant neoplasm of lower lobe, light bronchus or lung C34.32 Malignant neoplasm of lower lobe, light bronchus or lung C34.32 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of iright bronchus and lung C34.82 Malignant neoplasm of overlapping sites of right bronchus and lung C34.90 Malignant neoplasm of overlapping sites of light bronchus or lung C34.91 Malignant neoplasm of unspecified part of inspecified bronchus or lung C34.92 Malignant neoplasm of unspecified part of inspecified bronchus or lung C34.93 Malignant neoplasm of unspecified part of inspecified bronchus or lung C34.90 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of inspecified digestive organs D37.1 Neoplasm of uncertain behavior of of other specified digestive organs D37.9 Neoplasm of uncertain behavior of of other specified digestive organ D37.9 Neoplasm of uncertain behavior of of other specified digestive organ D38.03 Personal history of malignant neoplasm of esophagus D38.03 Personal history of malignant	C18.7	Malignant neoplasm of sigmoid colon
C19 Malignant neoplasm of rectosigmoid junction C20 Malignant neoplasm of rectum C21.8 Malignant neoplasm of rectum C31.8 Malignant neoplasm of verlapping sites of rectum, anus and anal canal C33.0 Malignant neoplasm of trachea C34.00 Malignant neoplasm of right main bronchus C34.01 Malignant neoplasm of left main bronchus C34.02 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, right bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.13 Malignant neoplasm of indidle lobe, bronchus or lung C34.2 Malignant neoplasm of indidle lobe, bronchus or lung C34.3 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.3 Malignant neoplasm of lower lobe, right bronchus or lung C34.3 Malignant neoplasm of lower lobe, left bronchus or lung C34.3 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.8 Malignant neoplasm of overlapping sites of left bronchus and lung C34.8 Malignant neoplasm of overlapping sites of left bronchus and lung C34.9 Malignant neoplasm of overlapping sites of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of right bronchus or lung C34.9 Malignant neoplasm of unspecified part of right bronchus or lung C34.9 Malignant neoplasm of unspecified part of right bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm of unspecified part of left bronchus or lung C34.9 Malignant neoplasm o	C18.8	Malignant neoplasm of overlapping sites of large intestines
C21.8 Malignant neoplasm of overlapping sites of rectum, anus and anal canal C21.8 Malignant neoplasm of overlapping sites of rectum, anus and anal canal C34.00 Malignant neoplasm of trachea C34.01 Malignant neoplasm of right main bronchus C34.02 Malignant neoplasm of left main bronchus C34.02 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, left bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.13 Malignant neoplasm of middle lobe, bronchus or lung C34.14 Malignant neoplasm of lower lobe, left bronchus or lung C34.2 Malignant neoplasm of lower lobe, left bronchus or lung C34.30 Malignant neoplasm of lower lobe, left bronchus or lung C34.31 Malignant neoplasm of lower lobe, left bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.91 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.92 Malignant neoplasm of unspecified part of inght bronchus or lung C34.93 Malignant neoplasm of unspecified part of right bronchus or lung C34.90 Secondary malignant neoplasm of lung C38.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.08 Secondary malignant neoplasm of lung C78.09 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C7	C18.9	Malignant neoplasm of colon, unspecified
C21.8 Malignant neoplasm of overlapping sites of rectum, anus and anal canal C34.00 Malignant neoplasm of trachea C34.00 Malignant neoplasm of main bronchus C34.01 Malignant neoplasm of right main bronchus C34.02 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of left main bronchus C34.11 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.12 Malignant neoplasm of upper lobe, right bronchus or lung C34.13 Malignant neoplasm of inder lobe, bronchus or lung C34.14 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.15 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.16 Malignant neoplasm of lower lobe, left bronchus or lung C34.31 Malignant neoplasm of lower lobe, left bronchus or lung C34.32 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.80 Malignant neoplasm of overlapping sites of left bronchus and lung C34.81 Malignant neoplasm of overlapping sites of left bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus or lung C34.90 Malignant neoplasm of unspecified part of inspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.09 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.09 Secondary ma	C19	Malignant neoplasm of rectosigmoid junction
C33. Malignant neoplasm of trachea C34.00 Malignant neoplasm of main bronchus C34.01 Malignant neoplasm of left main bronchus C34.02 Malignant neoplasm of left main bronchus C34.03 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, left bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of idele lobe, bronchus or lung C34.3. Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.3. Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.3. Malignant neoplasm of lower lobe, left bronchus or lung C34.3. Malignant neoplasm of lower lobe, left bronchus or lung C34.3. Malignant neoplasm of lower lobe, left bronchus or lung C34.8. Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.8. Malignant neoplasm of overlapping sites of right bronchus and lung C34.9. Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.9. Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.9. Malignant neoplasm of unspecified part of left bronchus or lung C34.9. Malignant neoplasm of unspecified part of left bronchus or lung C34.9. Malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.08 Secondary malignant neoplasm of lung C78.09 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 S	C20	Malignant neoplasm of rectum
C34.00 Malignant neoplasm of right main bronchus C34.01 Malignant neoplasm of right main bronchus C34.02 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, left bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of lower lobe, left bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.33 Malignant neoplasm of lower lobe, left bronchus or lung C34.34 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of left bronchus and lung C34.82 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.90 Malignant neoplasm of unspecified part of right bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of of the specified digestive organs D37.9 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organ D37.9 Neoplasm of uncertain behavior of other specified digestive organ D37.9 Personal history of malignant neoplasm of esophagus D37.0 Personal history of malignant neoplasm of stomach D37.0 Personal history of malignant neoplasm of stomach D37.0 Personal history of malignant neoplasm of large intestine	C21.8	Malignant neoplasm of overlapping sites of rectum, anus and anal canal
C34.01 Malignant neoplasm of right main bronchus C34.02 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, ight bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.13 Malignant neoplasm of middle lobe, bronchus or lung C34.20 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.33 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.80 Malignant neoplasm of overlapping sites of right bronchus and lung C34.81 Malignant neoplasm of overlapping sites of left bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of stomach D37.9 Neoplasm of uncertain behavior of of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of unspecified digestive organ Z85.01 Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach	C33	Malignant neoplasm of trachea
C34.02 Malignant neoplasm of left main bronchus C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, right bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.32 Malignant neoplasm of lower lobe, right bronchus or lung C34.33 Malignant neoplasm of lower lobe, left bronchus or lung C34.34 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.80 Malignant neoplasm of overlapping sites of right bronchus and lung C34.81 Malignant neoplasm of overlapping sites of left bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus or lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of left bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of of digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of unspecified digestive organ Z85.01 Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach	C34.00	Malignant neoplasm of main bronchus
C34.10 Malignant neoplasm of upper lobe, unspecified bronchus or lung C34.11 Malignant neoplasm of upper lobe, right bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of lung bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C38.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.08 Secondary malignant neoplasm of lung C78.09 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of lung C78.06 Secondary malignant neoplasm of lung C78.07 Secondary malignant neoplasm of lung C78.09 Secondary malignant neoplasm of lung C78.00 Secondary malignant neoplasm of stomach D37.1 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of stomach D87.01 Personal history of other malignant neoplasm of stomach	C34.01	Malignant neoplasm of right main bronchus
C34.11 Malignant neoplasm of upper lobe, right bronchus or lung C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, left bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.33 Malignant neoplasm of lower lobe, left bronchus or lung C34.480 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.80 Malignant neoplasm of overlapping sites of right bronchus and lung C34.81 Malignant neoplasm of overlapping sites of left bronchus and lung C34.82 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.90 Malignant neoplasm of unspecified part of right bronchus or lung C34.91 Malignant neoplasm of unspecified part of left bronchus or lung C34.92 Malignant neoplasm of lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of uspecified digestive organ Z85.01 Personal history of malignant neoplasm of stomach D87.8 Personal history of malignant neoplasm of stomach D87.9 Personal history of other malignant neoplasm of stomach D87.9 Personal history of other malignant neoplasm of stomach	C34.02	Malignant neoplasm of left main bronchus
C34.12 Malignant neoplasm of upper lobe, left bronchus or lung C34.2 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, left bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of inspecified bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of other specified digestive organ D37.9 Personal history of malignant neoplasm of esophagus D48.02 Personal history of malignant neoplasm of stomach D88.03 Personal history of malignant neoplasm of stomach D89.04 Personal history of malignant neoplasm of stomach D89.05 Personal history of malignant neoplasm of stomach D89.06 Personal history of malignant neoplasm of stomach D89.07 Personal history of malignant neoplasm of stomach D89.08 Personal history of malignant neoplasm of stomach	C34.10	Malignant neoplasm of upper lobe, unspecified bronchus or lung
C34.2 Malignant neoplasm of middle lobe, bronchus or lung C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of left bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach	C34.11	Malignant neoplasm of upper lobe, right bronchus or lung
C34.30 Malignant neoplasm of lower lobe, unspecified bronchus or lung C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of store of the specified duct D37.1 Neoplasm of uncertain behavior of stormach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of large intestine	C34.12	Malignant neoplasm of upper lobe, left bronchus or lung
C34.31 Malignant neoplasm of lower lobe, right bronchus or lung C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.6 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of esophagus Z85.01 Personal history of other malignant neoplasm of stomach D87.8 Personal history of malignant neoplasm of stomach D87.8 Personal history of malignant neoplasm of stomach	C34.2	Malignant neoplasm of middle lobe, bronchus or lung
C34.32 Malignant neoplasm of lower lobe, left bronchus or lung C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.6 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of unspecified digestive organ Z85.01 Personal history of malignant neoplasm of stomach Parsonal history of other malignant neoplasm of stomach Parsonal history of other malignant neoplasm of stomach Parsonal history of malignant neoplasm of large intestine	C34.30	Malignant neoplasm of lower lobe, unspecified bronchus or lung
C34.80 Malignant neoplasm of overlapping sites of unspecified bronchus and lung C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.6 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of esophagus Z85.01 Personal history of other malignant neoplasm of stomach P85.028 Personal history of malignant neoplasm of stomach P85.038 Personal history of malignant neoplasm of large intestine	C34.31	Malignant neoplasm of lower lobe, right bronchus or lung
C34.81 Malignant neoplasm of overlapping sites of right bronchus and lung C34.82 Malignant neoplasm of overlapping sites of left bronchus and lung C34.90 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.91 Malignant neoplasm of unspecified part of right bronchus or lung C34.92 Malignant neoplasm of unspecified part of left bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.6 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.7 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of unspecified digestive organ Z85.01 Personal history of malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of malignant neoplasm of large intestine	C34.32	Malignant neoplasm of lower lobe, left bronchus or lung
C34.92 Malignant neoplasm of overlapping sites of left bronchus and lung C34.91 Malignant neoplasm of unspecified part of unspecified bronchus or lung C34.92 Malignant neoplasm of unspecified part of right bronchus or lung C78.00 Secondary malignant neoplasm of lung C78.01 Secondary malignant neoplasm of lung C78.02 Secondary malignant neoplasm of lung C78.03 Secondary malignant neoplasm of lung C78.04 Secondary malignant neoplasm of lung C78.05 Secondary malignant neoplasm of retroperitoneum and peritoneum C78.76 Secondary malignant neoplasm of liver and intrahepatic bile duct D37.1 Neoplasm of uncertain behavior of stomach D37.8 Neoplasm of uncertain behavior of other specified digestive organs D37.9 Neoplasm of uncertain behavior of digestive organ, unspecified Z85.00 Personal history of malignant neoplasm of unspecified digestive organ Z85.01 Personal history of malignant neoplasm of esophagus Z85.028 Personal history of other malignant neoplasm of stomach Parsonal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of other malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of stomach Personal history of malignant neoplasm of large intestine	C34.80	
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Z85.038 Personal history of malignant neoplasm of large intestine		
Z85.118 Personal history of other malignant neoplasm of bronchus and lung		
	Z85.118	Personal history of other malignant neoplasm of bronchus and lung

# **Revision History**

Company(ies)	DATE	REVISION
EmblemHealth & ConnectiCare	3/20/2024	Annual Review: Removed PI link, added dosing chart, no criteria changes
EmblemHealth & ConnectiCare	7/18/2023	Annual Review:  Gastric, Esophageal and Gastro-esophageal Junction Cancers, initiation: Initial  Criteria: Added" OR used in combination with irinotecan;"  Non-small cell lung cancer, initiation: Initial Criteria: Added "Patient meets of the following criteria (i or ii):  i. Cyramza will be used as first-line therapy; AND  a) Patient has epidermal growth factor receptor (EGFR) exon 19 deletion or L858R mutation positive disease; AND  b) Cyramza will be used in combination with erlotinib; OR ii. Cyramza will be used as subsequent therapy; AND  a) Cyramza will be used in combination with docetaxel intravenous infusion; AND  b) Patient has received targeted drug therapy if the patient's tumor is positive for a targetable mutation Note: Examples of targetable mutations include sensitizing epidermal growth factor receptor mutation, anaplastic lymphoma kinase fusions" Removed "Patient's disease is metastatic; AND In combination with erlotinib, for the first-line treatment of patients whose tumors have epidermal growth factor receptor (EGFR) exon 19 deletions or exon 21 (L858R) substitution mutations; OR Used as subsequent therapy following progression on or after platinum-based chemotherapy; AND Cyramza must be used in combination with docetaxel; AND Patients with EGFR or ALK genomic tumor aberrations should have disease progression on FDA-approved therapy for these aberrations prior to receiving Cyramza" Colorectal cancer, initiation: Initial Criteria Added "OR irinotecan" Hepatocellular carcinoma: Initial Criteria Added "Patient has Child-Pugh Class A disease; AND"
EmblemHealth & ConnectiCare	4/11/2022	Transferred policy to new template
EmblemHealth & ConnectiCare	06/17/2020	Added Criteria for NSCLC indication: In combination with erlotinib, for the first-line treatment of patients whose tumors have epidermal growth factor receptor (EGFR) exon 19 deletions or exon 21 (L858R) substitution mutations
EmblemHealth & ConnectiCare	11/11/2019	Added indication for Hepatocellular carcinoma; removed hepatocellular carcinoma from limitations/exclusions

## References

1. Product Information: CYRAMZA® intravenous injection, ramucirumab intravenous injection. Eli LIIIy and Company (per manufacturer), Indianapolis, IN, 2019.