

Medical Policy: AVASTIN® (bevacizumab)

| POLICY NUMBER | LAST REVIEW | ORIGIN DATE |
|---------------|--------------------|---------------|
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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. 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.

EmblemHealth may also use tools developed by third parties, such as the MCG™ Care Guidelines, to assist us in administering health benefits. The MCG™ Care Guidelines are intended to be used in connection with the independent professional medical judgment of a qualified health care provider and do not constitute the practice of medicine or medical advice. EmblemHealth Services Company, LLC, has adopted this policy in providing management, administrative and other services to EmblemHealth Plan, Inc., EmblemHealth Insurance Company, EmblemHealth Services Company, LLC, and Health Insurance Plan of Greater New York (HIP) related to health benefit plans offered by these entities. ConnectiCare, an EmblemHealth company, has also adopted this policy. All of the aforementioned entities are affiliated companies under common control of EmblemHealth Inc.

Definitions

Avastin is a recombinant humanized monoclonal IgG1 antibody that binds to vascular endothelial growth factor (VEGF) and inhibits the proliferation of endothelial cells and the formation of new blood vessels.

Length of Authorization

- Coverage will be provided for **6 months** and may be renewed.
- For **CNS cancers** (Symptom management), coverage will be provided for 12 weeks and may **NOT** be renewed.

Dosing Limits [Medical Benefit]

Max Units (per dose and over time):

Oncology indications (J9035):

- 170 billable units per 21 days
- 120 billable units per 14 days

Guideline

I. INITIAL APPROVAL CRITERIA

For Commercial, Medicaid, and Medicare members:

- Non-preferred agent: Avastin, Alymsys, Vegzelma
- Preferred agents: Mvasi, Zirabev.

Coverage is provided for the following conditions (in addition to use supported by the National Comprehensive Cancer Network [NCCN] Clinical Practice Guidelines [NCCN Guidelines®] and/or NCCN Drugs & Biologics Compendium [NCCN Compendium®] with a recommendation of category level 1 or 2A*):

1. Patient is 18 years of age or older; **AND**
2. Must be prescribed by or in consultation with an oncologist; **AND**
3. Patient does not have recent history of hemorrhage or hemoptysis (the presence of blood in sputum); **AND**
4. Patient must not have had a surgical procedure within the preceding 28 days or have a surgical wound that has not fully healed; **AND**
5. For newly started Avastin, Alymsys, or Vegzelma therapy, for Commercial, Medicaid, and Medicare members:

Coverage may be considered medically necessary when:

- Patient has experienced a therapeutic failure or intolerance with the plan-preferred medications (Mvasi AND Zirabev); **OR**
- Avastin or Alymsys is requested for an indication for which the plan-preferred biosimilar agents (Mvasi or Zirabev) have not been FDA-approved OR are not supported by NCCN Guidelines® or NCCN Compendium® with a recommendation of category level 1 or 2A; **AND**

**Please note: Coverage for an appropriate biosimilar substitution will be allowed where NCCN Guidelines or Compendium state that an FDA-approved biosimilar is an appropriate substitution for bevacizumab.*

Hepatocellular Carcinoma

1. The medication is used in combination with Tecentriq (atezolizumab intravenous infusion); **AND**
2. Patient has not received prior systemic therapy

Colorectal Cancer (CRC)

1. Patient's disease is metastatic, unresectable, or advanced; **AND**
2. Medication is not used as adjuvant treatment **AND**
3. Used in combination with a fluoropyrimidine- (e.g., 5-fluorouracil/5-FU or capecitabine) or irinotecan-based regimen as first-line or subsequent therapy for metastatic, unresectable (or medically inoperable), or advanced disease; **OR**
4. Used in combination with a fluoropyrimidine-irinotecan- or fluoropyrimidine-oxaliplatin based regimen (not used first line) as second-line therapy for metastatic disease that has progressed on a first-line bevacizumab-containing regimen; **OR**
5. Used in combination with trifluridine and tipiracil as subsequent therapy for advanced or metastatic disease after progression on all available regimens

Non-squamous non-small cell lung cancer (NSCLC)

1. Patient's disease must be recurrent, unresectable, locally advanced, or metastatic; **AND**
2. Used as first-line treatment in combination with carboplatin and paclitaxel **OR**

3. The Patient meets **ONE** of the following criteria (a, b, c, **or** d):
 - a. The tumor is positive for epidermal growth factor receptor (*EGFR*) exon 19 deletion or L858R mutations and bevacizumab is used in combination with erlotinib; **OR**
 - b. The tumor is positive for one of the following mutations and bevacizumab is used in combination with other systemic therapies (i, ii,iii, iv, v, **or** vi):

Note: Examples include carboplatin plus paclitaxel or Alimta (pemetrexed intravenous infusion); cisplatin plus Alimta; and Tecentriq (atezolizumab intravenous infusion) plus carboplatin and paclitaxel.

 - i. Epidermal growth factor receptor (*EGFR*) exon 20 mutation; **OR**
 - ii. *KRAS G12C* mutation; **OR**
 - iii. *BRAF V600E*; **OR**
 - iv. *NTRK1/2/3* gene fusion; **OR**
 - v. *MET* exon 14 skipping mutation; **OR**
 - vi. *RET* rearrangement positive; **OR**
 - c. Patient has previously received targeted drug therapy for an actionable mutation; **OR**

*Note: Examples of actionable mutations include sensitizing epidermal growth factor receptor (*EGFR*) mutation, anaplastic lymphoma kinase (*ALK*) fusions, *RET* rearrangement positive, *MET* exon 14 skipping, *NTRK* gene fusion positive, *BRAF V600E* mutation positive, and *ROS* proto-oncogene 1 (*ROS1*) rearrangement positive.*
 - d. The NSCLC tumor is negative or unknown for actionable mutations and the patient meets **ONE** of the following criteria (i **or** ii):

*Note: Examples of actionable mutations include sensitizing epidermal growth factor receptor (*EGFR*) mutation, anaplastic lymphoma kinase (*ALK*) fusions, *RET* rearrangement positive, *MET* exon 14 skipping, *NTRK* gene fusion positive, *BRAF V600E* mutation positive, and *ROS* proto-oncogene 1 (*ROS1*) rearrangement positive.*

 - i. Bevacizumab is used as initial therapy in combination with other systemic therapies; **OR**

Note: Examples of systemic therapies are cisplatin, carboplatin, Tecentriq (atezolizumab intravenous infusion), Alimta (pemetrexed intravenous infusion), paclitaxel.
 - ii. Bevacizumab is used as subsequent therapy

Note: Bevacizumab can be used either as a single agent or in combination with other agents.

Cervical Cancer

1. Patient's disease must be persistent, recurrent, or metastatic; **AND**
2. Used in combination with paclitaxel **AND** either cisplatin/carboplatin, or topotecan

Renal cell carcinoma (RCC)

1. Patient has metastatic or relapsed disease; **AND**
 - a. Must be used as a single agent for predominantly non-clear cell histology; **OR**
 - b. Must be used in combination with interferon alfa; **OR**
 - c. Used in combination with everolimus or erlotinib in patients with papillary or hereditary leiomyomatosis disease

Central nervous system (CNS) cancer

1. Patient has tried at least one previous therapy; **AND**

Note: Examples are temozolomide capsules or injection, etoposide, carmustine, radiotherapy.
2. Patient has **ONE** of the following (a, b, c, d, e, f, g, h, **or** i):
 - a. Anaplastic gliomas; **OR**
 - b. Glioblastoma; **OR**
 - c. Intracranial and spinal ependymoma (excluding subependymoma) in patient \geq 18 years of age; **OR**

- d. Meningiomas; **OR**
- e. Brain, Spine, or Leptomeningeal metastases; **OR**
- f. Primary CNS lymphoma **OR**
- g. Medulloblastoma; **OR**
- h. Supratentorial Astrocytoma/Oligodendroglioma (Infiltrative, WHO Grade II); **OR**
- i. Symptoms due to one of the following (i, ii, **or** iii):
 - i. Radiation necrosis; **OR**
 - ii. Poorly controlled vasogenic edema; **OR**
 - iii. Mass effect

Ovarian cancer

1. Patient has Stage II-IV ovarian cancer after primary surgery; **AND**
 - a. Medication is used in combination with carboplatin and paclitaxel followed by Avastin as a single agent; **OR**
2. Patient has persistent or recurrent disease; **AND** (a **or** b)
 - a. If patient is platinum sensitive, medication is used in combination with carboplatin **AND** one of the following: gemcitabine or paclitaxel; **OR**
 - b. If patient is platinum resistant, medication is used in combination with one of the following: PEGylated liposomal doxorubicin, paclitaxel, or topotecan; **OR**
3. Medication is used as single agent maintenance therapy if used previously as part of combination therapy in patients with a partial or complete remission following primary therapy or therapy for platinum-sensitive recurrence; **OR**
4. Medication is used as neoadjuvant therapy in combination with paclitaxel and carboplatin; **AND**
 - a. Patient has bulky stage III or IV disease or is a poor surgical candidate; **OR**
5. Medication is used as adjuvant therapy in combination with paclitaxel and carboplatin; **AND**
 - a. Patient has stage II-IV disease; **OR**
 - b. Patient has stage I-IV carcinosarcoma histologic disease

Soft tissue Sarcoma ‡

1. Used as a single agent for Angiosarcoma; **OR**
2. Used in combination with temozolomide for Solitary Fibrous Tumor or Hemangiopericytoma

Endometrial Carcinoma ‡

1. Used as a single agent therapy for disease that has progressed on prior cytotoxic therapy; **OR**
2. Used in combination with carboplatin and paclitaxel for advanced or recurrent disease

Malignant Pleural Mesothelioma ‡

1. Patient has unresectable or metastatic disease; **AND**
2. One of the following applies (a, b, **or** c):
 - a. Bevacizumab will be used in combination with a chemotherapy regimen; **OR**
Note: Examples of chemotherapy are Alimta (pemetrexed intravenous infusion), cisplatin, carboplatin.
 - b. Bevacizumab will be used in combination with Tecentriq (atezolizumab intravenous infusion); **OR**
 - c. Bevacizumab is being used as a single agent for maintenance therapy after the patient has received combination chemotherapy regimen
Note: Examples of chemotherapy are Alimta (pemetrexed intravenous infusion), cisplatin, carboplatin.

Breast Cancer ‡

1. Patient must have recurrent or metastatic disease; **AND**
2. Patient has a high tumor burden or rapidly progressive disease; **AND**
3. Must be used in combination with paclitaxel; **AND**
4. Patient must be human epidermal growth factor receptor 2 (HER2)-negative; **AND**
 - a. Disease is hormone receptor-negative; **OR**
 - b. Disease is hormone receptor-positive and refractory to endocrine therapy; **OR**
 - c. Patient has symptomatic visceral disease or visceral crisis

AIDS-Related Kaposi Sarcoma ‡

1. Patient has relapsed or refractory disease; **AND**
2. Patient has advanced cutaneous, oral, visceral or nodal disease; **AND**
3. Used as subsequent therapy in combination with antiretroviral therapy (ART) after failure to two lines of systemic therapy

‡ Compendia recommended indication(s)

| Genomic Aberration Targeted Therapies (<i>not all inclusive</i>) § |
|---|
| Sensitizing EGFR mutation-positive tumors <ul style="list-style-type: none"> • Erlotinib • Afatinib • Gefitinib • Osimertinib |
| ALK rearrangement-positive tumors <ul style="list-style-type: none"> • Crizotinib • Ceritinib • Brigatinib • Alectinib |
| ROS1 rearrangement-positive tumors <ul style="list-style-type: none"> • Crizotinib • Ceritinib |
| BRAF V600E-mutation positive tumors <ul style="list-style-type: none"> • Dabrafenib/Trametinib |
| PD-L1 expression-positive tumors (>50%) <ul style="list-style-type: none"> • Pembrolizumab |

II. RENEWAL CRITERIA

Coverage can be renewed based upon the following criteria:

1. **Continuation of documented current and/or successful therapy with a non-preferred agent (Avastin and Alymsys); AND**
2. Tumor response with stabilization of disease or decrease in size of tumor or tumor spread; **AND**
3. Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: gastrointestinal perforation, surgical/wound healing complications, hemorrhage, arterial and venous thromboembolic events (ATE & VTE), uncontrolled hypertension, posterior reversible encephalopathy syndrome (PRES), nephrotic syndrome, severe infusion reactions, ovarian failure, congestive heart failure (CHF), etc.; **AND**
4. **CNS Cancers – symptom management (short-course therapy):** May NOT be renewed

5. **Ovarian cancer - Platinum sensitive disease or recurrence:** Must be used as a single agent for maintenance therapy; **OR** Used in combination with chemotherapy, for completion of initial therapy, up to 10 cycles total

Dosing/Administration

| Indication | Dose |
|--------------------------------|--|
| CRC | 5 to 10 mg/kg every 2 weeks or 7.5 mg/kg every 3 weeks |
| NSCLC & Cervical Cancer | 15 mg/kg every 3 weeks until disease progression or unacceptable toxicity. |
| CNS Cancers | <ul style="list-style-type: none"> • For disease treatment: 10 mg/kg every 2 weeks until disease progression or unacceptable toxicity. • For symptom management: 5-10 mg/kg every 2 weeks up to 12 weeks duration |
| RCC | 10 mg/kg every 2 weeks until disease progression or unacceptable toxicity. |
| MPM | 15 mg/kg every 3 weeks in combination with chemotherapy for up to 6 cycles followed by single agent use, at the same dose/frequency, until disease progression or unacceptable toxicity. |
| Ovarian Cancer | <p><u>Platinum-sensitive:</u> 15 mg/kg every 3 weeks for up to 8 cycles when used with paclitaxel or up to 10 cycles when used with gemcitabine; followed by single-agent bevacizumab 15 mg/kg IV every 3 weeks until disease progression or unacceptable toxicity</p> <p><u>Platinum-resistant:</u> 10 mg/kg every 2 weeks or 15 mg/kg every 3 weeks until disease progression or unacceptable toxicity</p> |
| All Other Oncology Indications | 5-10 mg/kg every 2 weeks OR 7.5-15 mg/kg every 3 weeks |

Applicable Procedure Codes

| Code | Description |
|-------|--|
| J9035 | Injection, bevacizumab, 10 mg; 1 billable unit = 10 mg |
| J9999 | Not otherwise classified, antineoplastic drugs (Alymsys only) |
| J3590 | Unclassified biologics (Vegzelma only) |

Applicable NDCs

| Code | Description |
|---------------|--|
| 50242-0060-xx | Avastin single-use vial, 100 mg/4 mL solution for injection |
| 50242-0061-xx | Avastin single-use vial, 400 mg/16 mL solution for injection |
| 70121-1754-xx | Alymsys single-dose vial, 100 mg/4 mL solution for injection |
| 70121-1755-xx | Alymsys single-dose vial, 400 mg/16 mL solution for injection |
| 32228-0011-xx | Vegzelma single-dose vial, 100 mg/4 mL solution for injection |
| 32228-0011-xx | Vegzelma single-dose vial, 400 mg/16 mL solution for injection |

ICD-10 Diagnoses

| Code | Description |
|--------|---|
| 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 |
| C18.0 | Malignant neoplasm of cecum |
| C18.1 | Malignant neoplasm of appendix |
| C18.2 | Malignant neoplasm of ascending colon |
| C18.3 | Malignant neoplasm of hepatic flexure |
| C18.4 | Malignant neoplasm of transverse colon |
| C18.5 | Malignant neoplasm of splenic flexure |
| C18.6 | Malignant neoplasm of descending colon |
| C18.7 | Malignant neoplasm of sigmoid colon |
| C18.8 | Malignant neoplasm of overlapping sites of large intestines |
| C18.9 | Malignant neoplasm of colon, unspecified |
| C19 | 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 unspecified 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, 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 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 |
| C38.4 | Malignant neoplasm of pleura |
| C45.0 | Mesothelioma of pleura |
| C46.0 | Kaposi's sarcoma of skin |
| C46.1 | Kaposi's sarcoma of soft tissue |
| C46.2 | Kaposi's sarcoma of palate |
| C46.3 | Kaposi's sarcoma of lymph nodes |
| C46.4 | Kaposi's sarcoma of gastrointestinal sites |
| C46.50 | Kaposi's sarcoma of unspecified lung |

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| C46.51 | Kaposi's sarcoma of right lung |
| C46.52 | Kaposi's sarcoma of left lung |
| C46.7 | Kaposi's sarcoma of other sites |
| C46.9 | Kaposi's sarcoma, unspecified |
| C48.0 | Malignant neoplasm of retroperitoneum |
| C48.1 | Malignant neoplasm of specified parts of peritoneum |
| C48.2 | Malignant neoplasm of peritoneum, unspecified |
| C48.8 | Malignant neoplasm of overlapping sites of retroperitoneum and peritoneum |
| C49.0 | Malignant neoplasm of connective and soft tissue of head, face and neck |
| C49.10 | Malignant neoplasm of connective and soft tissue of unspecified upper limb, including shoulder |
| C49.11 | Malignant neoplasm of connective and soft tissue of right upper limb including shoulder |
| C49.12 | Malignant neoplasm of connective and soft tissue of left upper limb, including shoulder |
| C49.20 | Malignant neoplasm of connective and soft tissue of unspecified lower limb, including hip |
| C49.21 | Malignant neoplasm of connective and soft tissue of right lower limb, including hip |
| C49.22 | Malignant neoplasm of connective and soft tissue of left lower limb, including hip |
| C49.3 | Malignant neoplasm of connective and soft tissue of thorax |
| C49.4 | Malignant neoplasm of connective and soft tissue of abdomen |
| C49.5 | Malignant neoplasm of connective and soft tissue of pelvis |
| C49.6 | Malignant neoplasm of connective and soft tissue of trunk, unspecified |
| C49.8 | Malignant neoplasm of overlapping sites of connective and soft tissue |
| C49.9 | Malignant neoplasm of connective and soft tissue, unspecified |
| C50.011 | Malignant neoplasm of nipple and areola, right female breast |
| C50.012 | Malignant neoplasm of nipple and areola, left female breast |
| C50.019 | Malignant neoplasm of nipple and areola, unspecified female breast |
| C50.021 | Malignant neoplasm of nipple and areola, right male breast |
| C50.022 | Malignant neoplasm of nipple and areola, left male breast |
| C50.029 | Malignant neoplasm of nipple and areola , unspecified male breast |
| C50.111 | Malignant neoplasm of central portion of right female breast |
| C50.112 | Malignant neoplasm of central portion of left female breast |
| C50.119 | Malignant neoplasm of central portion of unspecified female breast |
| C50.121 | Malignant neoplasm of central portion of right male breast |
| C50.122 | Malignant neoplasm of central portion of left male breast |
| C50.129 | Malignant neoplasm of central portion of unspecified male breast |
| C50.211 | Malignant neoplasm of upper-inner quadrant of right female breast |
| C50.212 | Malignant neoplasm of upper-inner quadrant of left female breast |
| C50.219 | Malignant neoplasm of upper-inner quadrant of unspecified female breast |
| C50.221 | Malignant neoplasm of upper-inner quadrant of right male breast |
| C50.222 | Malignant neoplasm of upper-inner quadrant of left male breast |
| C50.229 | Malignant neoplasm of upper-inner quadrant of unspecified male breast |
| C50.311 | Malignant neoplasm of lower-inner quadrant of right female breast |
| C50.312 | Malignant neoplasm of lower-inner quadrant of left female breast |
| C50.319 | Malignant neoplasm of lower-inner quadrant of unspecified female breast |
| C50.321 | Malignant neoplasm of lower-inner quadrant of right male breast |
| C50.322 | Malignant neoplasm of lower-inner quadrant of left male breast |
| C50.329 | Malignant neoplasm of lower-inner quadrant of unspecified male breast |
| C50.411 | Malignant neoplasm of upper-outer quadrant of right female breast |

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| C50.412 | Malignant neoplasm of upper-outer quadrant of left female breast |
| C50.419 | Malignant neoplasm of upper-outer quadrant of unspecified female breast |
| C50.421 | Malignant neoplasm of upper-outer quadrant of right male breast |
| C50.422 | Malignant neoplasm of upper-outer quadrant of left male breast |
| C50.429 | Malignant neoplasm of upper-outer quadrant of unspecified male breast |
| C50.511 | Malignant neoplasm of lower-outer quadrant of right female breast |
| C50.512 | Malignant neoplasm of lower-outer quadrant of left female breast |
| C50.519 | Malignant neoplasm of lower-outer quadrant of unspecified female breast |
| C50.521 | Malignant neoplasm of lower-outer quadrant of right male breast |
| C50.522 | Malignant neoplasm of lower-outer quadrant of left male breast |
| C50.529 | Malignant neoplasm of lower-outer quadrant of unspecified male breast |
| C50.611 | Malignant neoplasm of axillary tail of right female breast |
| C50.612 | Malignant neoplasm of axillary tail of left female breast |
| C50.619 | Malignant neoplasm of axillary tail of unspecified female breast |
| C50.621 | Malignant neoplasm of axillary tail of right male breast |
| C50.622 | Malignant neoplasm of axillary tail of left male breast |
| C50.629 | Malignant neoplasm of axillary tail of unspecified male breast |
| C50.811 | Malignant neoplasm of overlapping sites of right female breast |
| C50.812 | Malignant neoplasm of overlapping sites of left female breast |
| C50.819 | Malignant neoplasm of overlapping sites of unspecified female breast |
| C50.821 | Malignant neoplasm of overlapping sites of right male breast |
| C50.822 | Malignant neoplasm of overlapping sites of left male breast |
| C50.829 | Malignant neoplasm of overlapping sites of unspecified male breast |
| C50.911 | Malignant neoplasm of unspecified site of right female breast |
| C50.912 | Malignant neoplasm of unspecified site of left female breast |
| C50.919 | Malignant neoplasm of unspecified site of unspecified female breast |
| C50.921 | Malignant neoplasm of unspecified site of right male breast |
| C50.922 | Malignant neoplasm of unspecified site of left male breast |
| C50.929 | Malignant neoplasm of unspecified site of unspecified male breast |
| C51.0 | Malignant neoplasm of labium majus |
| C51.1 | Malignant neoplasm of labium minus |
| C51.2 | Malignant neoplasm of clitoris |
| C51.8 | Malignant neoplasm of overlapping sites of vulva |
| C53.0 | Malignant neoplasm of endocervix |
| C53.1 | Malignant neoplasm of exocervix |
| C53.8 | Malignant neoplasm of overlapping sites of cervix uteri |
| C53.9 | Malignant neoplasm of cervix uteri, unspecified |
| C54.0 | Malignant neoplasm of isthmus uteri |
| C54.1 | Malignant neoplasm of endometrium |
| C54.2 | Malignant neoplasm of myometrium |
| C54.3 | Malignant neoplasm of fundus uteri |
| C54.8 | Malignant neoplasm of overlapping sites of corpus uteri |
| C54.9 | Malignant neoplasm of corpus uteri, unspecified |
| C55 | Malignant neoplasm of uterus, part unspecified |
| C56.1 | Malignant neoplasm of right ovary |
| C56.2 | Malignant neoplasm of left ovary |

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| C56.9 | Malignant neoplasm of unspecified ovary |
| C57.00 | Malignant neoplasm of unspecified fallopian tube |
| C57.01 | Malignant neoplasm of right fallopian tube |
| C57.02 | Malignant neoplasm of left fallopian tube |
| C57.10 | Malignant neoplasm of unspecified broad ligament |
| C57.11 | Malignant neoplasm of right broad ligament |
| C57.12 | Malignant neoplasm of left broad ligament |
| C57.20 | Malignant neoplasm of unspecified round ligament |
| C57.21 | Malignant neoplasm of right round ligament |
| C57.22 | Malignant neoplasm of left round ligament |
| C57.3 | Malignant neoplasm of parametrium |
| C57.4 | Malignant neoplasm of uterine adnexa, unspecified |
| C57.7 | Malignant neoplasm of other specified female genital organs |
| C57.8 | Malignant neoplasm of overlapping sites of female genital organs |
| C57.9 | Malignant neoplasm of female genital organ, unspecified |
| C64.1 | Malignant neoplasm of right kidney, except renal pelvis |
| C64.2 | Malignant neoplasm of left kidney, except renal pelvis |
| C64.9 | Malignant neoplasm of unspecified kidney, except renal pelvis |
| C65.1 | Malignant neoplasm of right renal pelvis |
| C65.2 | Malignant neoplasm of left renal pelvis |
| C65.9 | Malignant neoplasm of unspecified renal pelvis |
| C70.0 | Malignant neoplasm of cerebral meninges |
| C70.1 | Malignant neoplasm of spinal meninges |
| C70.9 | Malignant neoplasm of meninges, unspecified |
| C71.0 | Malignant neoplasm of cerebrum, except lobes and ventricles |
| C71.1 | Malignant neoplasm of frontal lobe |
| C71.2 | Malignant neoplasm of temporal lobe |
| C71.3 | Malignant neoplasm of parietal lobe |
| C71.4 | Malignant neoplasm of occipital lobe |
| C71.5 | Malignant neoplasm of cerebral ventricle |
| C71.6 | Malignant neoplasm of cerebellum |
| C71.7 | Malignant neoplasm of brain stem |
| C71.8 | Malignant neoplasm of overlapping sites of brain |
| C71.9 | Malignant neoplasm of brain, unspecified |
| C72.0 | Malignant neoplasm of spinal cord |
| C72.9 | Malignant neoplasm of central nervous system, unspecified |
| C78.00 | Secondary malignant neoplasm of unspecified lung |
| C78.01 | Secondary malignant neoplasm of right lung |
| C78.02 | Secondary malignant neoplasm of left lung |
| C78.6 | Secondary malignant neoplasm of retroperitoneum and peritoneum |
| C78.7 | Secondary malignant neoplasm of liver and intrahepatic bile duct |
| C79.31 | Secondary malignant neoplasm of brain |
| C79.32 | Secondary malignant neoplasm of cerebral meninges |
| C79.89 | Secondary malignant neoplasm of other specified sites |
| C79.9 | Secondary malignant neoplasm of unspecified site |
| C83.30 | Diffuse large B-cell lymphoma unspecified site |

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| C83.31 | Diffuse large B-cell lymphoma lymph nodes of head, face, and neck |
| C83.39 | Diffuse large B-cell lymphoma extranodal and solid organ sites |
| C83.80 | Other non-follicular lymphoma unspecified site |
| C83.81 | Other non-follicular lymphoma lymph nodes of head, face, and neck |
| C83.89 | Other non-follicular lymphoma extranodal and solid organ sites |
| D32.0 | Benign neoplasm of cerebral meninges |
| D32.1 | Benign neoplasm of spinal meninges |
| D32.9 | Benign neoplasm of meninges, unspecified |
| D42.0 | Neoplasm of uncertain behavior of cerebral meninges |
| D42.1 | Neoplasm of uncertain behavior of spinal meninges |
| D42.9 | Neoplasm of uncertain behavior of meninges, unspecified |
| D43.0 | Neoplasm of uncertain behavior of brain, supratentorial |
| D43.1 | Neoplasm of uncertain behavior of brain, infratentorial |
| D43.2 | Neoplasm of uncertain behavior of brain, unspecified |
| D43.4 | Neoplasm of uncertain behavior of spinal cord |
| I67.89 | Other cerebrovascular disease |
| Z85.038 | Personal history of other malignant neoplasm of large intestine |
| Z85.068 | Personal history of other malignant neoplasm of small intestine |
| Z85.118 | Personal history of other malignant neoplasm of bronchus and lung |
| Z85.3 | Personal history of malignant neoplasm of breast |
| Z85.43 | Personal history of malignant neoplasm of ovary |
| Z80.49 | Family history of malignant neoplasm of other genital organs |
| Z85.528 | Personal history of other malignant neoplasm of kidney |
| Z85.831 | Personal history of malignant neoplasm of soft tissue |
| Z85.841 | Personal history of malignant neoplasm of brain |
| Z85.848 | Personal history of malignant neoplasm of other parts of nervous tissue |

Revision History

| Company(ies) | DATE | REVISION |
|-----------------------------|------------|---|
| EmblemHealth & ConnectiCare | 04/21/2023 | Added Vegzelma as non-preferred agent to criteria |
| EmblemHealth & ConnectiCare | 9/14/2022 | Under CNS Cancer – Removed. Used as a single agent OR in combination with one of the following: irinotecan, carmustine, lomustine, or temozolomide in patients with recurrent Glioblastomas † or Anaplastic Gliomas; OR 4. Medication is used as a single agent for progressive or recurrent Intracranial or Spinal Ependymoma (excluding subependymoma) after prior radiation therapy; OR 5. Medication is used as a single agent for patients with surgically inaccessible recurrent or progressive Meningioma when radiation is not possible |
| EmblemHealth & ConnectiCare | 08/11/2022 | Added Alymsys as non-preferred agent to Criteria |

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|-----------------------------|------------|--|
| EmblemHealth & ConnectiCare | 8/02/2022 | <p>Added Hepatocellular Carcinoma indication</p> <p>Under Colorectal Cancer – Added additional criteria “Used in combination with trifluridine and tipiracil as subsequent therapy for advanced or metastatic disease after progression on all available regimens”</p> <p>Under NSCLC criteria- removal of “Patient must have an ECOG performance status 0-2” and “Patient does not have locoregional recurrence without evidence of disseminated disease” Added – “The tumor is positive for epidermal growth factor receptor (EGFR) exon 19 deletion or L858R mutations and bevacizumab is used in combination with erlotinib”</p> <p>For Malignant Pleural Mesothelioma – added examples of chemotherapy regimens</p> |
| EmblemHealth & ConnectiCare | 07/28/2022 | <p>Updated Initial approval criteria: Must be prescribed by, or in consultation with an oncologist</p> <p>Updated Colorectal cancer, Cervical Cancer, RCC, CNS, Ovarian carcinoma Cancer to match FDA Label</p> |
| EmblemHealth & ConnectiCare | 3/24/2022 | Transferred policy to new template |
| EmblemHealth & ConnectiCare | 12/20/2020 | <p>Clarifications:</p> <ul style="list-style-type: none"> • Step therapy will apply to NEW starts only • NCCN-supported use (with 1 or 2A recommendation) will be covered <p>Renewal criteria updated:</p> <ul style="list-style-type: none"> • Removed: “Patient continues to meet criteria identified above” <p>Added coverage: “Continuation of documented current and/or successful therapy with a non-preferred agent (Avastin)”</p> |
| EmblemHealth & ConnectiCare | 11/2/2020 | <p>Effective 01/01/2021 Member must fail trial of Mvasi AND Zirabev, prior to using Avastin (Medicare members are subject to this step therapy).</p> |
| EmblemHealth & ConnectiCare | 03/31/2020 | <p>Added to the Initial Criteria: Effective 07/01/2020, Mvasi and Zirabev are the preferred agents for Commercial and Medicaid members. Member must fail trial of Mvasi AND Zirabev, prior to using Avastin (Only Commercial and Medicaid members are subject to this step therapy).</p> <p>Initial Criteria: Added Patient is 18 years of age or older.</p> |
| EmblemHealth & ConnectiCare | 2/12/2019 | Added Diagnosis Codes C51.0, C51.1, C51.2, C51.8 |

References

1. Avastin [package insert]. South San Francisco, CA; Genentech; June 2018. Accessed September 2019.

2. Referenced with permission from the NCCN Drugs & Biologics Compendium (NCCN Compendium®) bevacizumab. National Comprehensive Cancer Network, 2018. 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 July 2018.
3. Ceresoli GL, Zucali PA, Mencoboni M, et al. Phase II study of pemetrexed and carboplatin plus bevacizumab as first-line therapy in malignant pleural mesothelioma. *Br J Cancer*. 2013 Aug 6; 109(3): 552–558
4. Delishaj D, Ursino S, Pasqualetti F, et al. Bevacizumab for the Treatment of Radiation-Induced Cerebral Necrosis: A Systematic Review of the Literature. *J Clin Med Res*. 2017 Apr; 9(4): 273–280.
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