Bevacizumab (Avastin®)
Bevacizumab-adcd (Vegzelma®)
Bevacizumab-awwb (Mvasi®)
Bevacizumab-bvzr (Zirabev®)
Bevacizumab-maly (Alymsys®)

Place of Service
Office Administration
Outpatient Facility Infusion
Administration
Infusion Center Administration

## **HCPCS**

Avastin: **J9035** per 10 mg Mvasi: **Q5107** per 10 mg Zirabev: **Q5118** per 10 mg Alymsys: **Q5126** per 10 mg Vegzelma: **Q5129** per 10 mg

# Condition(s) listed in policy (see criteria for details)

- Ampullary adenocarcinoma
- Appendiceal adenocarcinoma
- Brain cancer
- Breast cancer
- Cervical cancer
- Colorectal cancer
- Cystoid macular degeneration
- Diabetic macular edema or diabetic retinopathy
- Endometrial cancer
- Fallopian tube cancer
- Hepatocellular carcinoma
- Glaucoma associated with vascular disorders
- Macular edema secondary to retinal veinocclusion
- Mesothelioma: peritoneal
- Mesothelioma: pleural
- Neovascular (wet) age-related macular degeneration
- Non-small cell lung cancer
- Ovarian cancer
- Primary peritoneal cancer
- Renal cell carcinoma
- Retinal edema (if macular)
- Retinal neovascularization
- Rubeosis iridis
- Small bowel adenocarcinoma
- Soft tissue sarcoma
- Vulvar cancer, squamous cell carcinoma

AHFS therapeutic class: Antineoplastic agents

**Mechanism of action:** Recombinant humanized monoclonal antibody against the vascular endothelial growth factor (VEGF)

# (1) Special Instructions and pertinent Information

**Covered under the Medical Benefit,** please submit clinical information for prior authorization review.

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(2) Prior Authorization/Medical Review is required for the following condition(s)
All requests for bevacizumab must be sent for clinical review and receive authorization prior to drug administration or claim payment.

#### Ampullary adenocarcinoma

- 1. Intestinal type, AND
- 2. Used in combination with a fluorouracil or capecitabine based regimen, AND
- 3. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

Covered Doses
Up to 15 mg/kg IV every 21 days
Coverage Period
Yearly
ICD-10:
C24.1

## Appendiceal adenocarcinoma or Colorectal cancer

- 1. One of the following:
  - Advanced, unresectable, medically inoperable, metastatic or synchronous metastatic or metachronous metastatic disease, and used in combination with irinotecan, oxaliplatin, or a fluorouracil-based regimen, or
  - b. Advanced or metastatic disease with prior treatment of a fluoropyrimidine, oxaliplatin, and irinotecan, and being used in combination with Lonsurf (trifluridine/tipiracil)

#### AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 10 mg/kg IV infusion every 2 weeks

## Coverage Period

Yearly

### ICD-10:

C17.0-C17.2, C17.8, C17.9, C18.0-C18.9, C19, C20, C21.8, C78.00-C78.02, C78.6, C78.7, Z85.038

## **Brain cancer**

- 1. Either of the following:
  - a. Diagnosis of one of the following brain cancer types:
    - i. Glioblastoma multiforme (also called WHO grade IV glioma/astrocytoma), OR

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- ii. Anaplastic gliomas (also called WHO grade III glioma/astrocytoma), OR
- iii. Ependymomas, OR
- iv. Meningiomas

### OR

b. Short-course single agent therapy for management of symptoms driven by radiation necrosis, poorly controlled vasogenic edema, or mass effect

#### AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 10 mg/kg IV every 2 weeks or up to 15 mg/kg IV every 3 weeks

## Coverage Period

Brain cancer: Yearly

Short-course symptom management: 6 months

#### ICD-10:

C70.0, C70.1, C70.9, C71.0-C71.9, C72.9, D32.0, D32.1, D32.9, D42.0, D42.1, D42.9, D43.0-D43.2, D43.4, Z85.841, Z85.848

### **Cervical cancer**

- 1. Persistent, recurrent, or metastatic disease, AND
- 2. One of the following:
  - a. Being used in combination with an NCCN supported regimen (e.g., paclitaxel and either cisplatin or carboplatin, and in combination with pembrolizumab if PDL-1 positive; paclitaxel and topotecan), OR
  - b. Being used as a single agent for second-line or subsequent therapy

#### AND

3. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

### **Covered Doses**

Up to 15 mg/kg every 3 weeks

## **Coverage Period**

Yearly

### ICD-10:

C53.0, C53.1, C53.8, C53.9, C79.89, C79.9, Z80.49

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## Cystoid macular degeneration

#### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

## Coverage Period

Yearly

#### ICD-10:

H35.351-H35.353, H35.359

## Diabetic macular edema or diabetic retinopathy

### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

## Coverage Period

Yearly

### ICD-10:

(X = 0 - 9)

E08.3XXX, E09.3XXX, E10.3XXX, E11.3XXX, E13.3XXX

### **Endometrial cancer**

- 1. Advanced, recurrent or metastatic disease, AND
- 2. One of the following:
  - a. Used as a single agent and Patient has received prior treatment with systemic chemotherapy, OR
  - b. Used in combination with paclitaxel for adjuvant treatment, OR
  - c. Used in combination with carboplatin and paclitaxel

### AND

3. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

# **Covered Doses**

Up to 15 mg/kg IV infusion every 3 weeks

## Coverage Period

Yearly

#### ICD-10:

C54.0-C54.3, C54.8, C54.9, C55, Z80.49

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# Fallopian tube cancer, Ovarian cancer, OR Primary peritoneal cancer

- 1. One of the following:
  - a. Ovarian cancer
  - b. Fallopian tube cancer
  - c. Primary peritoneal cancer

#### AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 15 mg/kg every 3 weeks, OR Up to 10 mg/kg every 2 weeks

# Coverage Period

Yearly

#### ICD-10:

C48.1, C48.2, C48.8, C56.1, C56.2, C56.9, C57.00-C57.02, C57.10-C57.12, C57.20-C57.22, C57.3, C57.4, C57.7-C57.9, Z85.43

### Glaucoma associated with vascular disorders (e.g, neovascular glaucoma)

## **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

## Coverage Period

Yearly

## ICD-10:

H40.50X0 - H40.50X4 H40.51X0 - H40.51X4 H40.52X0 - H40.52X4 H40.53X0 - H40.53X4

### Hepatocellular carcinoma

- 1. Patient has not received prior systemic drug therapy, AND
- 2. Being used in combination with Tecentriq, AND
- For request for non-preferred product (Avastin): Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

## **Covered Doses**

Up to 15 mg/kg every 3 weeks

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## Coverage Period

Yearly

ICD-10:

C22.0, C22.8, C22.9

### Macular edema secondary to retinal vein occlusion

#### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

# Coverage Period

Yearly

### ICD-10:

H34.8110-8112, H34.8120-8122, H34.8130-8132, H34.8190-8192, H34.8310-8312, H34.8320-8322, H34.8330-8332, H34.8390-8392

## Mesothelioma: peritoneal

- 1. Either of the following:
  - a. First-line therapy and all of the following:
    - i. Unresectable disease, AND
    - ii. One of the following:
      - 1. Used in combination with pemetrexed and cisplatin, OR
      - 2. Used in combination with pemetrexed and carboplatin, OR
      - 3. Used as a single agent for maintenance

## OR

- b. Subsequent therapy and the following:
  - i. One of the following:
    - 1. Used in combination with Tecentriq, OR
    - 2. Used in combination with pemetrexed and cisplatin, OR
    - 3. Used in combination with pemetrexed and carboplatin, OR
    - 4. Used as a single agent for maintenance

## AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 15 mg/kg IV every 3 weeks

## Coverage Period

Yearly

ICD-10:

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## Mesothelioma: pleural

- 1. Meets either of the following:
  - a. First-line therapy and all of the following:
    - i. Patient has unresectable, stage IIIB, sarcomatoid, metastatic or medically inoperable disease, **AND**
    - ii. One of the following:
      - 1. Used in combination with pemetrexed and cisplatin, OR
      - 2. Used in combination with pemetrexed and carboplatin, OR
      - 3. Used as a single agent for maintenance

### OR

- b. Subsequent therapy and all of the following:
  - i. Received immunotherapy as first line therapy, AND
  - ii. Either of the following:
    - 1. Used in combination with pemetrexed and cisplatin, OR
    - 2. Used in combination with pemetrexed and carboplatin

#### AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 15 mg/kg IV every 3 weeks

# Coverage Period

**Initial Authorization:** 

6 cycles

Reauthorization:

Yearly as maintenance if initially used with chemotherapy

Dosing: 15 mg/kg every 3 weeks until disease progression

ICD-10:

C38.4, C45.0, C45.1

## Neovascular (wet) age-related macular degeneration

1. Exudative senile macular degeneration

## **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

# Coverage Period

Yearly

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#### ICD-10:

H35.3210-3213 H35.3220-3223 H35.3230-3233 H35.3290-3293

#### Non-small cell lung cancer

- 1. Unresectable, locally advanced, recurrent, or metastatic disease, AND
- 2. Non-squamous histology, AND
- 3. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 15 mg/kg every 3 weeks

## Coverage Period

Yearly

ICD-10:

C33, C34.00-C34.02, C34.10-C34.12, C34.2, C34.30-C34.32, C34.80-C34.82, C34.90-C34.92, Z85.118

## Renal cell carcinoma

- 1. Either of the following:
  - Diagnosis is <u>non-clear cell</u>, advanced (relapsed, locally advanced, Stage IV, unresectable, or evidence of metastases) renal cell carcinoma,

OR

b. Diagnosis is <u>clear cell</u>, advanced (relapsed, locally advanced, Stage IV, unresectable, or evidence of metastases) renal cell carcinoma, AND being used as first line treatment in combination with interferon,

#### AND

2. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 10 mg/kg every 2 weeks

## Coverage Period

Yearly

ICD-10:

C64.1, C64.2, C64.9, C65.1, C65.2, C65.9, Z85.528

# Retinal edema (if macular)

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#### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

## Coverage Period

Yearly

ICD-10:

H35.81

## Retinal neovascularization NOS (choroidal, subretinal)

- 1. Retinal neovascularization AND
- 2. At least one of the following secondary ICD 10 code describing cause of retinal neovascularization:
  - a. Histoplasma capsulatum infection OR
  - b. Histoplasma duboisii infection OR
  - c. Histoplasmosis retinitis- unspecified OR
  - d. Progressive high (degenerative) myopia

#### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

# Coverage Period

Yearly

ICD-10:

H35.051-H35.053, H35.059, AND

## Secondary code of.

B39.4, B39.5, B39.9, H44.20-H44.23, H44.2A1-H44.2A3, H44.2A9, H44.2B1-H44.2B3, H44.2B9, H44.2C1-H44.2C3, H44.2C9, H44.2D1-H44.2D3, H44.2D9,

H44.2E1-H44.2E3, H44.2E9

## Rubeosis iridis

#### **Covered Doses**

Up to 2.5 mg per injection no sooner than every 4 weeks

## Coverage Period

Yearly

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ICD-10: H21.1X1-H21.1X3, H21.1X9

#### Small bowel adenocarcinoma

- Being used for advanced (unresectable or recurrent) or metastatic disease, AND
- 2. Used in combination with a fluropyrimidine-based (fluorouracil or capecitabine) regimen,

AND

3. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 5 mg/kg IV every 2 weeks, or Up to 7.5 mg/kg IV every 3 weeks

## Coverage period

Yearly

ICD-10:

C17.0, C17.1, C17.2, C17.3, C17.8, C17.9

#### Soft tissue sarcoma

- 1. Given in combination with temozolomide, AND
- For request for non-preferred product (Avastin): Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

#### **Covered Doses**

Up to 5 mg/kg IV on days 8 and 22, repeated at 28-day intervals

## Coverage Period

Yearly

ICD-10:

C48.0, C49.0, C49.3-C49.5, C49.8, C49.9, Z85.831

## Vulvar cancer, squamous cell carcinoma

- 1. Locally advanced unresectable, recurrent, or metastatic disease, AND
- 2. Histology of squamous cell carcinoma, AND
- 3. Used in combination with cisplatin and paclitaxel, AND
- 4. <u>For request for non-preferred product (Avastin)</u>: Intolerable side effect with the preferred bevacizumab products, Alymsys, Mvasi, Vegzelma, and Zirabev, that is not expected with Avastin, or contraindication to all preferred products

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#### Covered Doses

Up to 15 mg/kg every 3 weeks

## **Coverage Period**

Yearly

#### ICD-10:

C51.0-C51.2, C51.8, C51.9

(3) The following condition(s) <u>DO NOT</u> require Prior Authorization/Preservice.

All requests for bevacizumab must be sent for clinical review and receive authorization prior to drug administration or claim payment.

## (4) This Medication is NOT medically necessary for the following condition(s)

Coverage for a Non-FDA approved indication, requires that criteria outlined in Health and Safety
Code § 1367.21, including objective evidence of efficacy and safety are met for the proposed indication.

Please refer to the Doctor Manual and User Guide for more information

## (5) Additional Information

How supplied:

Avastin/Mvasi/Zirabev/Alymsys/Vegzelma:

- 100 mg/4 mL (single-use vial)
- 400 mg/16 mL (single-use vial)

#### (6) References

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- Avastin® (bevacizumab) [Prescribing information], South San Francisco, CA: Genentech; 2020.
- DrugDex<sup>®</sup>. Available by subscription at <a href="http://www.micromedexsolutions.com/home/dispatch">http://www.micromedexsolutions.com/home/dispatch</a>
- Mvasi® (bevacizumab-awwb) [Prescribing information], Thousand Oaks, CA: Amgen; 4/2021.
- National Comprehensive Cancer Network Drugs & Biologics Compendium. Bevacizumab (2023). Available by subscription at: www.nccn.org.
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   Available at <a href="http://www.nccn.org">http://www.nccn.org</a>.
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- National Comprehensive Cancer Network. Hepatobiliary Cancers (Version 1.2022).
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- National Comprehensive Cancer Network. Non-small cell lung cancer (Version 3.2023).
   Available at http://www.nccn.org.
- National Comprehensive Cancer Network. Ovarian Cancer/Fallopian Tube Cancer/Primary Peritoneal Cancer (Version 1.2023). Available at <a href="http://www.nccn.org">http://www.nccn.org</a>.
- National Comprehensive Cancer Network. Pediatric Central Nervous System Cancers (version 2.2023). Available at http://www.nccn.org.
- National Comprehensive Cancer Network. Rectal Cancer (Version 4.2022). Available at http://www.nccn.org.
- National Comprehensive Cancer Network. Small Bowel Adenocarcinoma (Version 1.2023). Available at: http://www.nccn.org.
- National Comprehensive Cancer Network. Soft Tissue Sarcoma (Version 1.2023). Available at http://www.nccn.org.
- National Comprehensive Cancer Network. Uterine Neoplasms (Version 1.2023). Available at http://www.nccn.org.
- National Comprehensive Cancer Network. Vulvar cancer (Version 1.2023). Available at: http://www.nccn.org.
- Zirabev® (bevacizumab-bvzr) [Prescribing information]. New York, NY: Pfizer Inc; 5/2021.

# (7) Policy Update

Date of last revision: 2Q2023 Date of next review: 4Q2023

Changes from previous policy version:

 Added new bevacizumab biosimilar, Vegzelma (bevacizumab-adcd). Vegzelma is a preferred bevacizumab product for all non-ophthalmic indications.

Rationale: Selection of preferred drugs is supported by similar safety and efficacy and are guideline supported agents.

 Section (2): Cervical cancer – Expanded coverage to include single use for second line or subsequent therapy

Rationale: NCCN category 2A support

BSC Drug Coverage Criteria to Determine Medical Necessity Reviewed by P&T Committee

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