

PRIOR AUTHORIZATION POLICY

- POLICY:** Oncology (Injectable) – Trastuzumab Products Prior Authorization Policy
- Herceptin[®] (trastuzumab intravenous infusion – Genentech)
 - Hercessi[™] (trastuzumab-strf intravenous infusion – Accord BioPharma)
 - Herzuma[®] (trastuzumab-pkrb intravenous infusion – Celltrion)
 - Kanjinti[™] (trastuzumab-anns intravenous infusion – Amgen)
 - Ogivri[®] (trastuzumab-dkst intravenous infusion – Mylan)
 - Ontruzant[®] (trastuzumab-dttb intravenous infusion – Merck)
 - Trazimera[™] (trastuzumab-qyyp intravenous infusion – Pfizer)

REVIEW DATE: 07/17/2024; selected revision 12/11/2024

OVERVIEW

Trastuzumab products are human epidermal growth factor receptor 2 (HER2)/neu receptor antagonists indicated for the following uses:¹

- **Breast cancer, adjuvant treatment** of HER2-overexpressing node positive or node negative (estrogen receptor [ER]/progesterone receptor [PR] negative or with one high risk feature) 1) as part of a treatment regimen consisting of doxorubicin, cyclophosphamide, and either paclitaxel or docetaxel; 2) as part of treatment regimen with docetaxel and carboplatin; or 3) as a single agent following multi-modality anthracycline based therapy.
- **Breast cancer, metastatic**, HER2-overexpressing, either in combination with paclitaxel for first-line treatment, or as a single agent in patients who have received one or more chemotherapy regimens for metastatic disease.
- **Gastric cancer or gastroesophageal junction adenocarcinoma, metastatic**, HER2-overexpressing, in combination with cisplatin and capecitabine or 5-fluorouracil (5-FU) who have not received prior treatment for metastatic disease.

Herzuma, Hercessi, Kanjinti, Ogivri, Ontruzant, and Trazimera are all approved biosimilars for Herceptin; all of the biosimilars have the same FDA-approved indications as Herceptin. For all indications, patients must be selected for therapy based on an FDA-approved companion diagnostic for trastuzumab. Tests are specific for breast cancer or gastric cancer.

Guidelines

Trastuzumab is discussed in guidelines from the National Comprehensive Cancer Network (NCCN)

- **Breast Cancer:** NCCN guidelines (version 4.2024 – July 3, 2024) recommend trastuzumab in combination with chemotherapy or endocrine therapy for adjuvant treatment of HER2-positive breast cancer (category 2A).^{2,10} Trastuzumab in combination with paclitaxel (category 2A) is a preferred preoperative/adjuvant therapy regimen. The guidelines also list other trastuzumab-containing regimens for preoperative and adjuvant therapy. The preferred first-line agents for HER2-positive recurrent or metastatic disease (either hormone receptor-negative or hormone receptor-positive and refractory to endocrine therapy) include: Perjeta[®] (pertuzumab intravenous infusion) plus trastuzumab plus docetaxel (category 1) or paclitaxel (category 2A). The guidelines list other trastuzumab-containing regimens for HER2-positive metastatic disease.
- **Colon and Rectal Cancer:** NCCN guidelines for colon cancer (version 4.2024 – July 3, 2024) and NCCN guidelines for rectal cancer (version 3.2024 – July 3, 2024) list trastuzumab in combination with Perjeta, Tukysa (tucatinib tablets), or lapatinib tablets in patients with HER2-amplified disease, RAS and BRAF wild-type disease.^{3-4,10}

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- **Gastric Cancer and Esophageal and Esophagogastric Junction Cancers:** NCCN guidelines for Gastric Cancer (version 2.2024 – May 29, 2024) and Esophageal and Esophagogastric Junction Cancers (version 3.2024 – April 26, 2024) state that for metastatic, locally advanced or recurrent disease (where local therapy is not indicated) trastuzumab should be added to first-line systemic chemotherapy for HER2-overexpressing adenocarcinoma.^{5,6,10} The recommended regimens for metastatic or locally advanced HER2-positive gastric, esophageal, or esophagogastric junction adenocarcinoma are trastuzumab in combination with cisplatin or oxaliplatin and a fluoropyrimidine (5-FU or capecitabine) [category 1] or trastuzumab in combination with other chemotherapy agents (category 2A/2B) [various regimens based on individual patient characteristics]. Trastuzumab is not recommended for use in combination with anthracyclines.
- **Head and Neck Cancers:** NCCN guidelines (version 4.2024 – May 1, 2024) recommend trastuzumab as a systemic therapy option for recurrent, unresectable, or metastatic salivary gland tumors, (useful in certain circumstances), for HER2-positive tumors as a single agent or in combination with Perjeta or docetaxel (category 2A).^{7,10}
- **Biliary Tract Cancers:** NCCN guidelines (version 3.2024 – July 2, 2024) recommend trastuzumab + Perjeta and trastuzumab + Tukysa as subsequent-line therapy options for biliary tract cancers for progression on or after systemic treatment for unresectable or metastatic disease that is HER2-positive (both category 2A).^{8,10}
- **Uterine Neoplasms:** NCCN guidelines (version 2.2024 – March 6, 2024) list the combination chemotherapy regimen of carboplatin/paclitaxel/trastuzumab as one of the recommended therapies for patients with HER2-positive endometrial carcinoma for stage III/IV or recurrent uterine serous carcinoma (category 2A).^{9,10}

POLICY STATEMENT

Prior Authorization is recommended for prescription benefit coverage of trastuzumab products. All approvals are provided for the duration noted below. Because of the specialized skills required for evaluation and diagnosis of patients treated with trastuzumab products, as well as the monitoring required for adverse events and long-term efficacy, approval requires trastuzumab products to be prescribed by or in consultation with a physician who specializes in the condition being treated.

Automation: None.

RECOMMENDED AUTHORIZATION CRITERIA

Coverage of trastuzumab products is recommended in those who meet one of the following:

FDA-Approved Indications

1. **Breast Cancer.** Approve for the duration noted if the patient meets ALL of the following (A, B, C, and D):
 - A) Patient is ≥ 18 years of age; AND
 - B) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - C) Patient meets ONE of the following (i or ii):
 - i. Approve for 1 year (total) if trastuzumab is used for neoadjuvant (preoperative)/adjuvant therapy; OR
 - ii. Approve for 1 year if trastuzumab is used for recurrent or metastatic disease; AND
 - D) The medication is prescribed by or in consultation with an oncologist.
2. **Gastric, Esophageal, or Gastroesophageal Junction Cancer.** Approve for 1 year if the patient meets ALL of the following (A, B, C, D, and E):
 - A) Patient is ≥ 18 years of age; AND
 - B) Patient has locally advanced or metastatic disease; AND
 - C) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - D) Patient meets BOTH of the following (i and ii):
 - i. Trastuzumab will be used as first-line therapy; AND
 - ii. Trastuzumab will be used in combination with chemotherapy; AND

Note: Examples of chemotherapy are cisplatin, oxaliplatin, capecitabine, 5-fluorouracil (5-FU).
 - E) The medication is prescribed by or in consultation with an oncologist.

Other Uses with Supportive Evidence

3. **Biliary Tract Cancer.** Approve for 1 year if the patient meets ALL of the following (A, B, C, D, E, and F):
 - A) Patient is ≥ 18 years of age; AND
 - B) Patient has unresectable or metastatic disease; AND
 - C) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - D) The medication will be used in combination with Perjeta (pertuzumab intravenous infusion) or Tukysa (tucatinib tablets); AND
 - E) The patient has tried one systemic regimen; AND

Note: Examples of a systemic regimen include: gemcitabine and cisplatin, 5-fluorouracil and oxaliplatin, capecitabine and oxaliplatin, or gemcitabine and oxaliplatin.

 - F) The medication is prescribed by or in consultation with an oncologist.
4. **Colon or Rectal Cancer.** Approve for 1 year if the patient meets ALL of the following (A, B, C, D, and E):
 - A) Patient is ≥ 18 years of age; AND
 - B) Patient has advanced or metastatic disease; AND
 - C) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - D) The medication is used in combination with Perjeta (pertuzumab intravenous infusion), lapatinib or Tukysa (tucatinib tablets); AND
 - E) The medication is prescribed by or in consultation with an oncologist.

- 5. Endometrial Carcinoma.** Approve for 1 year if the patient meets ALL of the following (A, B, C, D, and E):
- A) Patient is ≥ 18 years of age; AND
 - B) Patient has advanced or recurrent uterine serous carcinoma; AND
 - C) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - D) Trastuzumab will be used in combination with chemotherapy; AND
- Note: Examples of chemotherapy are carboplatin, paclitaxel.
- E) The medication is prescribed by or in consultation with an oncologist.
- 6. Salivary Gland Tumor.** Approve for 1 year if the patient meets ALL of the following (A, B, C, and D):
- A) Patient is ≥ 18 years of age; AND
 - B) Patient has recurrent, unresectable, or metastatic disease; AND
 - C) Patient has human epidermal growth factor receptor 2 (HER2)-positive disease; AND
 - D) The medication is prescribed by or in consultation with an oncologist.

CONDITIONS NOT RECOMMENDED FOR APPROVAL

Coverage of trastuzumab is not recommended in the following situations.

1. Coverage is not recommended for circumstances not listed in the Recommended Authorization Criteria. Criteria will be updated as new published data are available.

REFERENCES

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