

**Clinical Policy: Fam-Trastuzumab Deruxtecan-nxki (Enhertu)****Reference Number: LA.PHAR.456****Effective Date:****Last Review Date: 04.22****Line of Business: Medicaid**[Coding  
Implications  
Revision Log](#)

**See [Important Reminder](#) at the end of this policy for important regulatory and legal information.**

**Description**

**Fam-trastuzumab deruxtecan-nxki (Enhertu®) is a human epidermal growth factor receptor 2 (HER2)-directed antibody and topoisomerase inhibitor conjugate.**

**FDA Approved Indication(s)**

**Enhertu is indicated for the treatment of adult patients with:**

- **Unresectable or metastatic HER2-positive breast cancer who have received two or more prior anti-HER2-based regimens in the metastatic setting**
- **Locally advanced or metastatic HER2-positive gastric or gastroesophageal junction (esophagogastric junction; EGJ) adenocarcinoma who have received a prior trastuzumab-based regimen**

Commented [BJ1]: PI updated January 2021 with new indication

Commented [ACE2R1]: PI updated with current status.

**Policy/Criteria**

**Prior authorization is required. Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.**

**It is the policy of Louisiana Healthcare Connections that Enhertu is medically necessary when the following criteria are met:**

**I. Initial Approval Criteria****A. Breast Cancer (must meet all):**

1. **Diagnosis of recurrent, unresectable or metastatic HER2-positive breast cancer;**
2. **Prescribed by or in consultation with an oncologist;**
3. **Age > 18 years;**
4. **Failure of two anti-HER2-based regimens (see Appendix B), unless contraindicated or clinically significant adverse effects are experienced;**  
*\*Prior authorization may be required for anti-HER2-based regimens*
5. **Request meets one of the following (a or b):\***
  - a. **Dose does not exceed 5.4 mg/kg every 3 weeks;**
  - b. **Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (prescriber must submit supporting evidence).**  
*\*Prescribed regimen must be FDA-approved or recommended by NCCN*

**Approval duration:****Medicaid– 6 months**

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#### **B. Gastric and Esophagogastric Junction Cancer (must meet all):**

1. **Diagnosis of HER2-positive gastric or EGJ adenocarcinoma;**
2. **Prescribed by or in consultation with an oncologist;**
3. **Age ≥ 18 years;**
4. **Disease is locally advanced or metastatic;**
5. **Failure of trastuzumab-based regimen (see Appendix B);**
6. **Request meets one of the following (a or b):\***
  - a. **Dose does not exceed 6.4 mg/kg every 3 weeks;**
  - b. **Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (prescriber must submit supporting evidence).**

*\*Prescribed regimen must be FDA-approved or recommended by NCCN*

**Approval duration: Medicaid– 6 months**

#### **C. Other diagnoses/indications**

1. **Refer to the off-label use policy if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): LA.PMN.53 for Medicaid.**

## **II. Continued Therapy**

#### **A. Cancer (must meet all):**

1. **Currently receiving medication via Louisiana Healthcare Connections benefit, or documentation supports that member is currently receiving Enhertu for a covered indication and has received this medication for at least 30 days;**
2. **Member is responding positively to therapy;**
3. **If request is for a dose increase, request meets one of the following (a, b, or c):\***
  - a. **For breast cancer: New dose does not exceed 5.4 mg/kg every 3 weeks;**
  - b. **For gastric or EGJ adenocarcinoma: New dose does not exceed 6.4 mg/kg every 3 weeks;**
  - c. **New dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (prescriber must submit supporting evidence).**

*\*Prescribed regimen must be FDA-approved or recommended by NCCN*

**Approval duration:**

**Medicaid – 12 months**

#### **B. Other diagnoses/indications (must meet 1 or 2):**

1. **Currently receiving medication via Louisiana Healthcare Connections benefit and documentation supports positive response to therapy.**  
**Approval duration: Duration of request or 6 months (whichever is less); or**
2. **Refer to the off-label use policy if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): LA.PMN.53 for Medicaid.**

## **III. Diagnoses/Indications for which coverage is NOT authorized:**

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- A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policy –LA.PMN.53 for Medicaid or evidence of coverage documents.

#### IV. Appendices/General Information

##### Appendix A: Abbreviation/Acronym Key

EGJ: esophagogastric junction

FDA: Food and Drug Administration

HER2: human epidermal growth factor receptor 2

NCCN: National Comprehensive Center Network

##### Appendix B: Therapeutic Alternatives

*This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may require prior authorization.*

<u>Drug Name</u>	<u>Dosing Regimen</u>	<u>Dose Limit/Maximum Dose</u>
<b>Breast Cancer</b> <u>NCCN examples of systemic therapies for HER2-positive recurrent or metastatic disease</u> <ul style="list-style-type: none"> <li>• <u>Aromatase inhibitor ± trastuzumab</u></li> <li>• <u>Aromatase inhibitor ± lapatinib</u></li> <li>• <u>Pertuzumab + trastuzumab + docetaxel</u></li> </ul>	<u>Varies</u>	<u>Varies</u>
<u>Gastric and Esophagogastric Junction Cancer</u> <u>trastuzumab-based regimen</u>	<u>8 mg/kg</u> <u>IV q 3</u> <u>weeks</u>	<u>8 mg/kg</u>

*Therapeutic alternatives are listed as Brand name® (generic) when the drug is available by brand name only and generic (Brand name®) when the drug is available by both brand and generic.*

##### Appendix C: Contraindications/Boxed Warnings

- Contraindication(s): none reported
- Boxed warning(s): interstitial lung disease and pneumonitis; embryo-fetal toxicity

#### V. Dosage and Administration

<u>Indication</u>	<u>Dosing Regimen</u>	<u>Maximum Dose</u>
<u>Breast cancer</u>	<u>5.4 mg/kg IV every 3 weeks</u>	<u>6.4 mg/kg</u>
<u>Gastric cancer</u>	<u>6.4 mg/kg IV every 3 weeks</u>	<u>6.4 mg/kg</u>

#### VI. Product Availability

Single-dose vial: 100 mg lyophilized powder

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#### VII. References

1. Enhertu Prescribing Information. Basking Ridge, NJ: Daiichi Sankyo, Inc.; January 2021. Available at: [www.enhertu.com](http://www.enhertu.com). Accessed September 13, 2021.
2. National Comprehensive Cancer Network Drugs and Biologics Compendium. Available at [www.nccn.org](http://www.nccn.org). Accessed September 13, 2021.
3. National Comprehensive Cancer Network. Breast Cancer Version 8.2021. Available at: [http://www.nccn.org/professionals/physician\\_gls/pdf/breast.pdf](http://www.nccn.org/professionals/physician_gls/pdf/breast.pdf). Accessed September 13, 2021.
4. Modi S, Saura C, Yamashita T, et al. Trastuzumab deruxtecan in previously treated HER2-positive breast cancer. *N Engl J Med*. 2019; doi: 10.1056/NEJMoa1914510.
5. National Comprehensive Cancer Network. Gastric Cancer Version 4.2021. Available at: [https://www.nccn.org/professionals/physician\\_gls/pdf/gastric.pdf](https://www.nccn.org/professionals/physician_gls/pdf/gastric.pdf). Accessed September 13, 2021.

#### Coding Implications

Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

HCPCS Codes	Description
J9358	Injection, fam-trastuzumab deruxtecan-nxki, 1 mg

Reviews, Revisions, and Approvals	Date	LDH Approval Date
Converted corporate to local policy	04.22	

#### Important Reminder

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. LHCC makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results.

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Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable LHCC administrative policies and procedures.

This clinical policy is effective as of the date determined by LHCC. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. LHCC retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

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