

## **Clinical Policy: Lapatinib (Tykerb)**

Reference Number: CP.PHAR.79

Effective Date: 10.01.11

Last Review Date: 11.18

Line of Business: Commercial, HIM, Medicaid

[Revision Log](#)

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

### **Description**

Lapatinib (Tykerb<sup>®</sup>) is a kinase inhibitor.

### **FDA Approved Indication(s)**

Tykerb is indicated in combination with:

- Capecitabine for the treatment of patients with advanced or metastatic breast cancer whose tumors overexpress human epidermal growth factor receptor 2 (HER2) and who have received prior therapy including an anthracycline, a taxane, and trastuzumab
- Letrozole for the treatment of postmenopausal women with hormone receptor (HR)-positive metastatic breast cancer that overexpresses the HER2 receptor for whom hormonal therapy is indicated

Limitation(s) of use:

- Patients should have disease progression on trastuzumab prior to initiation of treatment with Tykerb in combination with capecitabine.
- Tykerb in combination with an aromatase inhibitor has not been compared to a trastuzumab-containing chemotherapy regimen for the treatment of metastatic breast cancer.

### **Policy/Criteria**

*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 health plans affiliated with Centene Corporation<sup>®</sup> that Tykerb is **medically necessary** when the following criteria are met:

#### **I. Initial Approval Criteria**

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

1. Diagnosis of breast cancer;
2. Prescribed by or in consultation with an oncologist;
3. Age  $\geq$  18 years;
4. Disease is recurrent or metastatic (stage IV), and HER2-positive;
5. Tykerb is prescribed in combination with one of the following (a, b, or c):
  - a. Capecitabine;
  - b. Trastuzumab;
  - c. If HR-positive, an aromatase inhibitor (e.g., anastrozole, letrozole, exemestane), and:

- i. If male, an agent that suppresses testicular steroidogenesis (e.g., gonadotropin-releasing hormone agonists);
6. Request meets one of the following (a or b):
  - a. Dose does not exceed 1500 mg per day;
  - b. Dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

**Approval duration:**

**Medicaid/HIM** – 6 months

**Commercial** – Length of Benefit

**B. Other diagnoses/indications**

1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**II. Continued Therapy**

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

1. Currently receiving medication via Centene benefit or documentation supports that member is currently receiving Tykerb for breast cancer 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, meets one of the following (a or b):
  - a. New dose does not exceed 1500 mg per day;
  - b. New dose is supported by practice guidelines or peer-reviewed literature for the relevant off-label use (*prescriber must submit supporting evidence*).

**Approval duration:**

**Medicaid/HIM** – 12 months

**Commercial** – Length of Benefit

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

1. Currently receiving medication via Centene 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 for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid.

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

- 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 policies – CP.CPA.09 for commercial, HIM.PHAR.21 for health insurance marketplace, and CP.PMN.53 for Medicaid or evidence of coverage documents.

**IV. Appendices/General Information**

*Appendix A: Abbreviation/Acronym Key*

FDA: Food and Drug Administration  
HER2: human epidermal growth factor receptor 2

HR: hormone receptor  
NCCN: National Comprehensive Cancer Network

*Appendix B: Therapeutic Alternatives*

Not applicable

*Appendix C: Contraindications/Boxed Warnings*

- Contraindication(s): known severe hypersensitivity (e.g., anaphylaxis) to this product or any of its components
- Boxed warning(s): hepatotoxicity

*Appendix D: General Information*

- The NCCN recommends that men with HR-positive breast cancer be treated similarly to postmenopausal women, except that the use of an aromatase inhibitor is ineffective without concomitant suppression of testicular steroidogenesis.
- The NCCN supports use of Tykerb in premenopausal women with HR-positive breast cancer when used concomitantly with an aromatase inhibitor. Along with this combination therapy, patients should also be treated with ovarian ablation/suppression. Ovarian ablation can be achieved with surgical oophorectomy or ovarian irradiation. Ovarian suppression can be achieved with luteinizing hormone-releasing hormone agonists (e.g., goserelin, leuprolide).
- The NCCN also recommends use of Tykerb in combination with capecitabine for the treatment of recurrent brain metastases in patients with breast cancer that is responsive to Tykerb.
- HR-positive can be either estrogen receptor (ER)- or progesterone receptor (PR)-positive.

**V. Dosage and Administration**

| Indication    | Dosing Regimen   | Maximum Dose                                   |
|---------------|--|--|
| Breast cancer | Advanced or metastatic: 1250 mg PO QD on Days 1-21 continuously in combination with capecitabine 2,000 mg/m <sup>2</sup> /day (administered PO in 2 doses approximately 12 hours apart) on Days 1-14 in a repeating 21-day cycle | 1500 mg/day                                    |
|               |  | 5500 mg/day if taking a strong CYP3A4 inducer  |
|               | HER2-positive: 1500 mg PO QD continuously in combination with letrozole 2.5 mg PO QD   | 500 mg/day if taking a strong CYP3A4 inhibitor |

**VI. Product Availability**

Tablets: 250 mg

**VII. References**

1. Tykerb Prescribing Information. East Hanover, NJ: Novartis Pharmaceuticals Corporation; April 2017. Available at <https://www.tykerb.com>. Accessed July 5, 2018.

2. National Comprehensive Cancer Network Drugs and Biologics Compendium. Available at: [http://www.nccn.org/professionals/drug\\_compendium](http://www.nccn.org/professionals/drug_compendium). Accessed July 5, 2018.
3. National Comprehensive Cancer Network. Breast Cancer Version 1.2018. Available at [www.NCCN.org](http://www.NCCN.org). Accessed July 5, 2018.

| Reviews, Revisions, and Approvals   | Date     | P&T Approval Date |
|---|----------|-------------------|
| Converted embedded SGM document into Centene policy   | 08.13    |                   |
| Added Limitation of Use to Description<br>Split algorithm into initial and reauthorization<br>Added dosing adjustments to algorithm<br>Added other adverse events to Appendix A<br>Added Appendix B for dosing adjustments<br>Updated Background information  | 12.13    | 12.13             |
| Added question for LFT monitoring in Figure 2<br>Updated background information<br>Updated references<br>Added combination with trastuzumab indication  | 12.14    | 12.14             |
| Policy converted to new template.<br>Dose adjustment criteria removed; max dose limits added to criteria; lab test baselines removed but test result limits included in criteria.<br>Contraindications limited to absolute indications only<br>Appendices limited to abbreviations.   | 11.15    | 12.15             |
| Policy converted to new template.<br>Removed prescriber and age requirements.<br>Hormone receptor-positive and HER2-positive metastatic breast cancer (FDA approved use): Added that Tykerb must be prescribed in combination with letrozole.<br>Normal baseline and follow-up LVEF added.<br>Added max dose with CYP inducers under FDA indications.<br>Added all NCCN recommended uses.<br>Added appendix of breast cancer therapies by drug class. | 11.16    | 12.16             |
| Policy converted to new template. Annual Review.<br>Safety criteria was applied according to the safety guidance discussed at CPAC and endorsed by Centene Medical Affairs.<br>Central nervous system cancer off-label use criteria added per NCCN 2A recommendation.<br>Authorization limits extended from 3 and 6 months to 6 and 12 months for initial and continued approval, respectively.   | 08.17    | 11.17             |
| 4Q 2018 annual review: no significant changes; added commercial and HIM lines of business; summarized NCCN and FDA-approved uses for improved clarity; added specialist involvement in care; added age; added COC; references reviewed and updated.   | 07.05.18 | 11.18             |

**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. The Health Plan 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. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

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. 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 Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. 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. The Health Plan 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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**Note:**

**For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

**For Health Insurance Marketplace members**, when applicable, this policy applies only when the prescribed agent is on your health plan approved formulary. Request for non-formulary drugs must be reviewed using the non-formulary policy; HIM.PA.103.

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