Effective Date..... 4/1/2023
Next Review Date..... 4/1/2024

Prior Authorization
Oncology – Kisqali® (ribociclib tablets) and Kisqali® Femara® Co-Pack (ribociclib and letrozole tablets)

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Product Identifier(s)

62093

INSTRUCTIONS FOR USE
The following Coverage Policy applies to health benefit plans administered by Cigna Companies. Certain Cigna Companies and/or lines of business only provide utilization review services to clients and do not make coverage determinations. References to standard benefit plan language and coverage determinations do not apply to those clients. Coverage Policies are intended to provide guidance in interpreting certain standard benefit plans administered by Cigna Companies. Please note, the terms of a customer's particular benefit plan document [Group Service Agreement, Evidence of Coverage, Certificate of Coverage, Summary Plan Description (SPD) or similar plan document] may differ significantly from the standard benefit plans upon which these Coverage Policies are based. For example, a customer's benefit plan document may contain a specific exclusion related to a topic addressed in a Coverage Policy. In the event of a conflict, a customer's benefit plan document always supersedes the information in the Coverage Policies. In the absence of a controlling federal or state coverage mandate, benefits are ultimately determined by the terms of the applicable benefit plan document. Coverage determinations in each specific instance require consideration of 1) the terms of the applicable benefit plan document in effect on the date of service; 2) any applicable laws/regulations; 3) any relevant collateral source materials including Coverage Policies and; 4) the specific facts of the particular situation. Coverage Policies relate exclusively to the administration of health benefit plans. Coverage Policies are not recommendations for treatment and should never be used as treatment guidelines. In certain markets, delegated vendor guidelines may be used to support medical necessity and other coverage determinations.

National Formulary Medical Necessity

Cigna covers ribociclib and ribociclib/ letrozole (Kisqali® and Kisqali® Femara® Co-Pack) as medically necessary when the following criteria are met for FDA Indications or Other Uses with Supportive Evidence:

Prior Authorization is recommended for prescription benefit coverage of Kisqali and Kisqali Femara Co-Pack. All approvals are provided for the duration noted below. In the clinical criteria, as appropriate, an asterisk (*) is noted next to the specified gender. In this context, the specified gender is defined as follows: a woman is defined as an individual with the biological traits of a woman, regardless of the individual's gender identity or gender expression; men are defined as individuals with the biological traits of a man, regardless of the individual's gender identity or

I. Coverage of **Kisqali** is recommended in those who meet one of the following criteria:

FDA Indication(s)

1. **Breast Cancer in Women***. Approve for 1 year if the individual meets the following criteria (A, B, C, D, E, and F):
   - **A)** Individual is ≥ 18 years of age; AND
   - **B)** Individual has recurrent or metastatic disease; AND
   - **C)** Individual has hormone receptor positive (HR+) [i.e., estrogen receptor positive {ER+} and/or progesterone receptor positive {PR+}] disease; AND
   - **D)** Individual has human epidermal growth factor receptor 2 (HER2)-negative breast cancer; AND
   - **E)** Individual meets ONE of the following criteria (i or ii):
     - **i.** Individual is postmenopausal; OR
     - **ii.** Individual is pre/perimenopausal and meets one of the following (a or b):
       - (1) Individual is receiving ovarian suppression/ablation with a gonadotropin-releasing hormone (GnRH) agonist; OR
         Note: Examples of a GnRH agonist include leuprolide acetate, Lupron Depot (leuprolide acetate intramuscular injection), Trelstar (triptorelin pamoate intramuscular injection), Zoladex (goserelin acetate subcutaneous implant).
       - (2) Individual has had surgical bilateral oophorectomy or ovarian irradiation; AND
   - **F)** Individual meets ONE of the following criteria (i or ii):
     - **i.** Kisqali will be used in combination with anastrozole, exemestane, or letrozole; OR
     - **ii.** Kisqali will be used in combination with fulvestrant.

   ***** Refer to the Policy Statement.

2. **Breast Cancer in Men*.** Approve for 1 year if the individual meets the following criteria (A, B, C, D, and E):
   - **A)** Individual is ≥ 18 years of age; AND
   - **B)** Individual has recurrent or metastatic disease; AND
   - **C)** Individual has hormone receptor positive (HR+) [i.e., estrogen receptor positive {ER+} and/or progesterone receptor positive {PR+}] disease; AND
   - **D)** Individual has human epidermal growth factor receptor 2 (HER2)-negative breast cancer; AND
   - **E)** Individual meets ONE of the following criteria (i or ii):
     - **i.** Individual meets BOTH of the following criteria (a and b):
       - **a)** Individual is receiving a gonadotropin-releasing hormone (GnRH) analog; AND
         Note: Examples of GnRH analog include leuprolide acetate, Lupron Depot (leuprolide acetate intramuscular injection), Trelstar (triptorelin pamoate intramuscular injection), Zoladex (goserelin acetate subcutaneous implant), Firmagon (degarelix acetate subcutaneous injection), Orgovyx (relugolix tablet).
       - **b)** Kisqali will be used in combination with anastrozole, exemestane, or letrozole; OR
     - **ii.** Kisqali will be used in combination with fulvestrant.

   ***** Refer to the Policy Statement.

II. Coverage of **Kisqali Femara Co-Pack** is recommended in those who meet one of the following criteria:

FDA Indication(s)

1. **Breast Cancer in Women***. Approve for 1 year if the individual meets the following criteria (A, B, C, D, and E):
   - **A)** Individual is ≥ 18 years of age; AND
   - **B)** Individual has recurrent or metastatic disease; AND
   - **C)** Individual has hormone receptor positive (HR+) [i.e., estrogen receptor positive {ER+} and/or progesterone receptor positive {PR+}] disease; AND
   - **D)** Individual has human epidermal growth factor receptor 2 (HER2)-negative breast cancer; AND
   - **E)** Individual meets ONE of the following criteria (i or ii):
     - **i.** Individual is postmenopausal OR

ii. Individual is pre/perimenopausal and meets one of the following (a or b):
(1)Individual is receiving ovarian suppression/ablation with a gonadotropin-releasing hormone (GnRH) agonist; OR
Note: Examples of a GnRH agonist include leuprolide acetate, Lupron Depot (leuprolide acetate intramuscular injection), Trelstar (triptorelin pamoate intramuscular injection), Zoladex (goserelin acetate subcutaneous implant).
(2)Individual has had surgical bilateral oophorectomy or ovarian irradiation.

***** Refer to the Policy Statement.

2 Breast Cancer in Men*. Approve for 1 year if the individual meets the following criteria (A, B, C, D, and E):
A) Individual is ≥ 18 years of age; AND
B) Individual has recurrent or metastatic disease; AND
C) Individual has hormone receptor positive (HR+) [i.e., estrogen receptor positive {ER+} and/or progesterone receptor positive {PR+}] disease; AND
D) Individual has human epidermal growth factor receptor 2 (HER2)-negative breast cancer; AND
E) Individual is receiving a gonadotropin-releasing hormone (GnRH) analog.
Note: Examples of a GnRH analog include leuprolide acetate, Lupron Depot (leuprolide acetate intramuscular injection), Trelstar (triptorelin pamoate intramuscular injection), Zoladex (goserelin acetate subcutaneous implant), Firmagon (degarelix acetate subcutaneous injection), Orgovyx (relugolix tablet).

***** Refer to the Policy Statement.

Conditions Not Covered

Ribociclib and ribociclib/ letrozole (Kisqali® and Kisqali® Femara® Co-Pack) is considered experimental, investigational or unproven for ANY other use.

Background

Overview
Kisqali, a cyclin-dependent kinase (CDK) 4/6 inhibitor, is indicated for the treatment of adults with hormone receptor-positive (HR+), human epidermal growth factor receptor 2 (HER2)-negative **advanced or metastatic breast cancer** in the following settings:[1-3]

- In combination with an aromatase inhibitor (AI) as initial endocrine-based therapy;
- Kisqali (not Co-Pack) in combination with fulvestrant as initial endocrine based therapy or following disease progression on endocrine therapy in postmenopausal women or in men;
- Kisqali Femara Co-Pack has the same indication with AI, letrozole, included.

Guidelines
The National Comprehensive Cancer Network (NCCN) guidelines on breast cancer (version 2.2023– February 7, 2023) recommends Kisqali + AI or fulvestrant as a first-line "Preferred Regimen" for HR+ and HER2-negative recurrent unresectable (local or regional) or Stage IV disease in postmenopausal women or premenopausal patient receiving ovarian ablation or suppression (category 1).[3,4] The guidelines state in a footnote that in phase III randomized controlled trials, Kisqali + endocrine therapy has shown overall survival benefit in the first-line setting. CDK4/6 inhibitor + fulvestrant is recommended as a "Preferred Regimen" for second- and subsequent-line therapy, if CDK4/6 inhibitor was not previously used (category 1). However, the guidelines also state in a footnote that if there is disease progression on Ibrance® (palbociclib tablets or capsules), there are limited data to support the use of Kisqali in the second-line setting.[3,4] The guidelines state that in phase III randomized controlled trials, fulvestrant in combination with a CDK4/6 inhibitor has shown overall survival benefit in the second-line setting. For men with breast cancer, the compendium recommends they be treated similarly to postmenopausal women, except that the use of an AI is ineffective without concomitant suppression of testicular steroidogenesis.[4]

References

1. Kisqali® tablets [prescribing information]. East Hanover, NJ: Novartis; October 2022.
2. Kisqali® Femara® Co-Pack tablets [prescribing information]. East Hanover, NJ: Novartis; October 2022.
3. The NCCN Breast Cancer Clinical Practice Guidelines in Oncology (version 2.2023 – February 7, 2023). © 2023 National Comprehensive Cancer Network. Available at: http://www.nccn.org. Accessed on February 10, 2023.
4. The NCCN Compendium. © 2023 National Comprehensive Cancer Network. Available at: http://www.nccn.org. Search term: ribociclib. Accessed on February 10, 2023.

Revision History

Type of Revision	Summary of Changes	Approval Date
Annual Revision	No criteria changes.	02/22/2023