PRIOR AUTHORIZATION POLICY

POLICY: Oncology – Braftovi Prior Authorization Policy
- Braftovi® (encorafenib capsules – Array BioPharma)

REVIEW DATE: 08/14/2024; selected revision 01/29/2025

INSTRUCTIONS FOR USE

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CIGNA NATIONAL FORMULARY COVERAGE:

OVERVIEW

Braftovi, a BRAF inhibitor, is indicated for the following:[1]

- **Colorectal cancer**, in combination with Erbitux® (cetuximab intravenous infusion) and mFOLFOX6 (5-FU, leucovorin, and oxaliplatin), for the treatment of metastatic disease with a *BRAF V600E* mutation, as detected by an FDA-approved test, in adults.
- **Colorectal cancer,** in combination with Erbitux, for the treatment of metastatic disease and a *BRAF V600E* mutation, as detected by an FDA-approved test, after prior therapy in adults.
- **Melanoma**, in combination with Mektovi® (binimetinib tablets), for the treatment of unresectable or metastatic disease and a *BRAF V600E* or *V600K* mutation, as detected by an FDA-approved test in adults.
- **Non-small cell lung cancer (NSCLC),** in combination with Mektovi, for the treatment of adult patients with metastatic NSCLC with a *BRAF V600E* mutation, as detected by an FDA-approved test.

The indication for Braftovi in colorectal cancer in the first-line setting is approved under accelerated approval based on response rate and durability of response. Continued approval for this indication may be contingent upon verification and description of clinical benefit in a confirmatory trial(s).

It is a limitation of use that Braftovi is not indicated for treatment of patients with wild-type BRAF melanoma, wild-type BRAF colorectal cancer, or wild-type BRAF NSCLC.

Guidelines

Braftovi is discussed in guidelines from the National Comprehensive Cancer Network (NCCN).[5]

- **Colon** and **Rectal Cancer:** Guidelines for colon cancer (version 6.2024 – January 17, 2025) and rectal cancer (version 5.2024 – January 17, 2025) recommend Braftovi for some situations in patients with *BRAF V600E*-mutated disease.[3] For primary treatment (following adjuvant chemotherapy) or as subsequent use, Braftovi + Erbitux or Vectibix® (panitumumab intravenous infusion) is a recommended treatment option for *BRAF V600E* mutation-positive disease (category 2A). The first-line indication of Braftovi, in combination with Erbitux and mFOLFOX6, is not yet addressed in the guidelines. NCCN Compendium recommends the use of Braftovi for appendiceal adenocarcinoma for BRAF V600E mutation-positive disease, as subsequent therapy, in combination with Erbitux or Vectibix.[5]
- **Melanoma, Cutaneous:** Guidelines (version 2.2024 – April 3, 2024) recommend BRAF/MEK inhibitor combinations among the "Preferred" therapies for first line (category 1) and subsequent treatment (category 2A) of metastatic or unresectable melanoma with a *V600*-activating mutation.[2] The combinations are also recommended for adjuvant treatment (category 2B). While combination BRAF/MEK inhibition is preferred, if a combination is contraindicated, monotherapy with a BRAF inhibitor (Tafinlar® [dabrafenib capsules] or Zelboraf® [vemurafenib tablets]) is a recommended option, especially in patients who are not appropriate candidates for checkpoint immunotherapy.
- **Non-Small Cell Lung Cancer:** Guidelines (version 7.2024 – June 26, 2024) recommend Braftovi + Mektovi and Tafinlar + Mekinist® (trametinib tablets) combinations for first-line "Preferred" regimens and as subsequent therapies (both category 2A) for *BRAF V600E* mutation-positive disease.[6] Zelboraf or Tafinlar monotherapy is also recommended under "Useful in Certain Circumstances" (both category 2A).

POLICY STATEMENT

Prior Authorization is recommended for prescription benefit coverage of Braftovi. All approvals are provided for the duration noted below.

- **Braftovi® (encorafenib capsules - Array BioPharma)**

is(are) covered as medically necessary when the following criteria is(are) met for fda-approved indication(s) or other uses with supportive evidence (if applicable):

FDA-Approved Indications

1. **Colon or Rectal Cancer.** 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 *BRAF V600E* mutation-positive disease; AND
   C) Patient meets ONE of the following (i or ii):
      i. Patient meets BOTH of the following (a and b):
         a. The medication will be used as first-line systemic therapy for metastatic disease; AND

b. The medication will be used in combination with Erbitux (cetuximab intravenous infusion) and mFOLFOX6 (5-FU, leucovorin, and oxaliplatin); OR

ii. Patient meets BOTH of the following (a and b):

a. Patient has previously received a chemotherapy regimen for colon or rectal cancer; AND

Note: Examples of chemotherapy regimens include a fluoropyrimidine such as 5-fluorouracil (5-FU), capecitabine; oxaliplatin, irinotecan, or an adjunctive chemotherapy regimen such as FOLFOX (5-FU, leucovorin, and oxaliplatin) or CapeOX (capecitabine and oxaliplatin).

b. The medication is used in combination with Erbitux or Vectibix (panitumumab intravenous infusion).

2. Melanoma. Approve for 1 year if the patient meets ALL of the following (A, B, and C):

A) Patient is ≥ 18 years of age; AND

B) Patient has unresectable, advanced, or metastatic melanoma; AND

C) Patient has *BRAF V600* mutation-positive disease.

3. Non-Small Cell Lung Cancer. Approve for 1 year if the patient meets ALL of the following (A, B, and C):

A) Patient is ≥ 18 years of age; AND

B) Patient has *BRAF V600E* mutation-positive metastatic disease; AND

C) The medication will be taken in combination with Mektovi (binimetinib tablets).

Other Uses with Supportive Evidence

4. Appendiceal Adenocarcinoma. 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 *BRAF V600E* mutation-positive disease; AND

C) The medication will be used as subsequent therapy for advanced or metastatic disease; AND

D) The medication will be used in combination with Erbitux (cetuximab intravenous infusion) or Vectibix (panitumumab intravenous infusion).

CONDITIONS NOT COVERED

- **Braftovi® (encorafenib capsules - Array BioPharma)**

is(are) considered experimental, investigational, or unproven for ANY other use(s).

REFERENCES

1. Braftovi® capsules [prescribing information]. Boulder, CO: Array BioPharma; December 2024.
2. The NCCN Melanoma Clinical Practice Guidelines in Oncology (version 2.2024 – April 3, 2024). © 2024 National Comprehensive Cancer Network. Available at: http://www.nccn.org/. Accessed on August 9, 2024.
3. The NCCN Colon Cancer Clinical Practice Guidelines in Oncology (version 6.2024 – January 17, 2025). © 2025 National Comprehensive Cancer Network. Available at: http://www.nccn.org/. Accessed on January 27, 2025.

4. The NCCN Rectal Cancer Clinical Practice Guidelines in Oncology (version 5.2024 – January 17, 2025). © 2025 National Comprehensive Cancer Network. Available at: http://www.nccn.org/. Accessed on January 27, 2025.
5. The NCCN Drugs & Biologics Compendium. © 2024 National Comprehensive Cancer Network. Available at: http://www.nccn.org. Accessed on August 9, 2024. Search terms: encorafenib.
6. The NCCN Non-Small Cell Lung Cancer Clinical Practice Guidelines in Oncology (version 7.2024 – June 26, 2024). © 2024 National Comprehensive Cancer Network. Available at: http://www.nccn.org/. Accessed on August 9, 2024.

HISTORY

Type of Revision	Summary of Changes	Review Date
Annual Revision	No criteria changes	07/19/2023
Selected Revision	**Non-Small Cell Lung Cancer:** Added new FDA-approved indication and criteria	10/18/2023
Annual Revision	**Appendiceal Adenocarcinoma:** Added new approval condition and criteria under "Other Uses with Supportive Evidence" based on compendium/guideline recommendations.	8/14/2024
Selected Revision	**Colon or Rectal Cancer:** Added new criterion for Braftovi use in combination with Erbitux and mFOLFOX6 for first-line therapy based on FDA approval. Also, for subsequent therapy use, modified criteria which referred to combination therapy of Braftovi to be more specific. Now the criterion states Braftovi in used in "combination with Erbitux or Vectibix (panitumumab intravenous infusion)". Deleted Note that provided examples of combination therapies.	01/29/2025