

Federal Employee Program.

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5.21.053

Section: Prescription Drugs Effective Date: October 1, 2025

Subsection: Antineoplastic Agents Original Policy Date: January 16, 2015

Subject: Opdivo Page: 1 of 16

Last Review Date: September 19, 2025

### **Opdivo**

#### **Description**

### Opdivo (nivolumab)

#### **Background**

Opdivo (nivolumab) is a monoclonal antibody indicated for the treatment of patients with melanoma, non-small cell lung cancer (NSCLC), malignant pleural mesothelioma, renal cell carcinoma (RCC), hepatocellular carcinoma (HCC), classical Hodgkin lymphoma (cHL), squamous cell carcinoma of the head and neck (SCCHN), urothelial carcinoma, colorectal cancer (CRC), esophageal cancer, gastric cancer, gastroesophageal junction cancer, and esophageal adenocarcinoma. Opdivo works by binding to the programmed cell death-1 (PD-1) receptor, and blocking its interaction with PD-1 ligands, PD-L1 and PD-L2. This interaction releases the inhibitory effects of PD-1 pathway-mediated inhibition of the immune response, including the anti-tumor immune response, resulting in decreased tumor growth (1).

#### **Regulatory Status**

FDA-approved indications: Opdivo is a programmed death receptor-1 (PD-1) blocking antibody indicated for the treatment of patients with: (1)

- 1. Melanoma
  - a. Unresectable or metastatic melanoma, as a single agent or in combination with ipilimumab
  - Adjuvant treatment of patients with completely resected Stage IIB, Stage IIC,
     Stage III, or Stage IV melanoma
- 2. Non-Small Cell Lung Cancer (NSCLC)
  - a. Resectable (tumors ≥4 cm or node positive) NSCLC in the neoadjuvant setting, in combination with platinum-doublet chemotherapy

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b. Resectable (tumors ≥4 cm or node positive) NSCLC and no known EGFR mutations or ALK rearrangements, for neoadjuvant treatment, in combination with platinum-doublet chemotherapy, followed by single-agent Opdivo as adjuvant treatment after surgery

- c. Metastatic non-small cell lung cancer (NSCLC) whose tumors express PD-L1
   (≥1%) as determined by an FDA-approved test, with no EGFR or ALK genomic
   tumor aberrations, as first-line treatment in combination with ipilimumab
- Metastatic or recurrent NSCLC with no EGFR or ALK genomic tumor aberrations as first-line treatment, in combination with ipilimumab and 2 cycles of platinumdoublet chemotherapy
- e. Metastatic NSCLC and progression on or after platinum-based chemotherapy. Patients with EGFR or ALK genomic tumor aberrations should have disease progression on an FDA-approved therapy for these aberrations prior to receiving Opdivo
- 3. Malignant Pleural Mesothelioma
  - a. Unresectable malignant pleural mesothelioma, as first-line treatment in combination with ipilimumab
- 4. Renal Cell Carcinoma (RCC)
  - a. Advanced renal cell carcinoma in patients who have received prior antiangiogenic therapy
  - b. First-line treatment of patients with advanced RCC, in combination with cabozantinib
  - c. Intermediate or poor risk advanced renal cell carcinoma, as a first-line treatment in combination with ipilimumab
- 5. Classical Hodgkin Lymphoma (cHL)
  - a. Classical Hodgkin lymphoma that has relapsed or progressed after:
    - i. Autologous hematopoietic stem cell transplantation (HSCT) and post-transplantation brentuximab vedotin, OR
    - ii. 3 or more lines of systemic therapy that includes autologous HSCT
- 6. Squamous Cell Carcinoma of the Head and Neck (SCCHN)
  - a. Recurrent or metastatic squamous cell carcinoma of the head and neck with disease progression on or after a platinum-based therapy
- 7. Urothelial Carcinoma
  - a. Adjuvant treatment of patients with urothelial carcinoma (UC) who are at high risk of recurrence after undergoing radical resection of UC
  - b. Patients with unresectable or metastatic urothelial carcinoma, as first-line treatment in combination with cisplatin and gemcitabine
  - c. Patients with locally advanced or metastatic urothelial carcinoma who:

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i. Have disease progression during or following platinum-containing chemotherapy

ii. Have disease progression within 12 months of neoadjuvant or adjuvant treatment with platinum-containing chemotherapy

#### 8. Colorectal Cancer

- a. Unresectable or metastatic microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) colorectal cancer (CRC) in combination with ipilimumab
- b. Microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) metastatic colorectal cancer that has progressed following treatment with a fluoropyrimidine, oxaliplatin, and irinotecan
- 9. Hepatocellular Carcinoma (HCC)
  - a. Unresectable or metastatic HCC as a first-line treatment in combination with ipilimumab
  - b. Unresectable or metastatic HCC that has been previously treated with sorafenib, in combination with ipilimumab

#### 10. Esophageal Cancer

- a. Completely resected esophageal or gastroesophageal junction cancer with residual pathologic disease, who have received neoadjuvant chemoradiotherapy (CRT)
- b. Unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC) as first-line treatment in combination with fluoropyrimidine- and platinumcontaining chemotherapy whose tumors express PD-L1 (≥1)
- Unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC) as first-line treatment in combination with ipilimumab whose tumors express PD-L1 (≥1)
- d. Unresectable advanced, recurrent, or metastatic esophageal squamous cell carcinoma (ESCC) after prior fluoropyrimidine- and platinum-based chemotherapy
- 11. Gastric Cancer, Gastroesophageal Junction Cancer, and Esophageal Adenocarcinoma
  - a. Advanced or metastatic gastric cancer, gastroesophageal junction cancer, and esophageal adenocarcinoma whose tumors express PD-L1 (≥1) in combination with fluoropyrimidine- and platinum-containing chemotherapy

#### Off-Label Uses: (2)

- 1. Small cell lung cancer
- 2. Metastatic anal cancer
- 3. Merkel cell carcinoma

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Opdivo carries warnings for immune-mediated adverse reactions, infusion-related reactions, complications of allogeneic hematopoietic stem cell transplantation (HSCT) and embryo-fetal toxicity. Clinically significant immune-mediated adverse reactions may occur with Opdivo therapy including pneumonitis, colitis, hepatitis, nephritis, renal dysfunction, hyperthyroidism, and hypothyroidism. Patients should be monitored for signs and symptoms of adverse reactions and based on the severity, Opdivo should be withheld or discontinued, and corticosteroids administered. Opdivo may cause fetal harm when administered to a pregnant woman. Female patients of reproductive potential should be advised of the potential hazard to a fetus (1).

The safety and effectiveness of Opdivo have not been established in pediatric patients age less than 12 years of age with melanoma or microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) colorectal cancer (CRC) or in pediatric patients less than 18 years of age for the other approved indications (1).

#### **Related Policies**

Bavencio, Keytruda, Loqtorzi, Opdualag, Tecentriq, Yervoy, Zynyz

#### **Policy**

This policy statement applies to clinical review performed for pre-service (Prior Approval, Precertification, Advanced Benefit Determination, etc.) and/or post-service claims.

Opdivo may be considered **medically necessary** if the conditions indicated below are met.

Opdivo may be considered **investigational** for all other indications.

### **Prior-Approval Requirements**

Age 12 years of age or older

#### **Diagnoses**

Patient must have **ONE** of the following:

- 1. Unresectable or metastatic melanoma
  - a. Used as a single agent **OR** in combination with ipilimumab
- 2. Adjuvant treatment of melanoma post resection

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- Stage IIB, Stage IIC, Stage III, or Stage IV melanoma
- 3. Resectable non-small cell lung cancer (NSCLC)
  - a. Tumors ≥4 cm OR node positive
  - b. Used in combination with platinum-doublet chemotherapy in the neoadjuvant setting
- 4. Metastatic non-small cell lung cancer (NSCLC) with **ONE** of the following:
  - a. **NO** EGFR or ALK genomic tumor aberrations with **ONE** of the following:
    - i. Disease progressed on or after platinum-based chemotherapy
    - Tumors express PD-L1 as determined by an FDA-approved test AND used as first-line treatment in combination with ipilimumab
    - iii. Used as first-line treatment in combination with ipilimumab and 2 cycles of platinum-doublet chemotherapy
  - b. Positive for EGFR or ALK genomic tumor aberrations
    - i. Disease must have progressed while on or after platinum-based chemotherapy
    - ii. Patient had disease progression on FDA approved therapy
- 5. Recurrent non-small cell lung cancer (NSCLC)
  - a. NO EGFR or ALK genomic tumor aberrations
  - b. Used as first-line treatment in combination with ipilimumab and 2 cycles of platinum-doublet chemotherapy
- 6. Advanced renal cell carcinoma with **ONE** of the following:
  - a. First-line treatment in combination with cabozantinib
  - b. Prior treatment with anti-angiogenic therapy
  - c. Patient is considered to have an intermediate or poor prognosis
    - i. Used as first-line treatment in combination with ipilimumab
- 7. Relapsed or progressed classical Hodgkin lymphoma with **ONE** of the following:
  - a. Patient has had autologous hematopoietic stem cell transplantation (HSCT) and post-transplantation therapy with brentuximab vedotin
  - Patient has had 3 or more lines of systemic therapy that includes autologous HSCT
- 8. Recurrent or metastatic squamous cell carcinoma of the head and neck
  - a. Disease must have progressed while on or after platinum-based chemotherapy

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- 9. Urothelial carcinoma with **ONE** of the following:
  - a. Patient is at high risk of recurrence after undergoing radical resection
    - i. Used as adjuvant treatment
  - b. Unresectable or metastatic urothelial carcinoma
    - Used as first-line treatment in combination with cisplatin and gemcitabine
  - Locally advanced or metastatic urothelial carcinoma with ONE of the following:
    - Disease must have progressed while on or after platinum-based chemotherapy
    - ii. Disease progression within 12 months of neoadjuvant or adjuvant treatment with platinum-containing chemotherapy
- 10. Unresectable or metastatic hepatocellular carcinoma (HCC)
  - a. Used as first-line treatment **OR** patient has had prior treatment with sorafenib
  - b. Used in combination with ipilimumab
- 11. Completely resected esophageal or gastroesophageal junction cancer with residual pathologic disease
  - a. Patient has received neoadjuvant chemoradiotherapy (CRT)
- 12. Unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC)
  - a. Tumors express PD-L1 as determined by an FDA-approved test
  - b. Used as first-line treatment
  - c. Used in combination with **ONE** of the following:
    - i. Fluoropyrimidine- and platinum-containing chemotherapy
    - ii. Ipilimumab
- 13. Unresectable advanced, recurrent, or metastatic esophageal squamous cell carcinoma (ESCC)
  - a. Prior treatment with fluoropyrimidine- and platinum-based chemotherapy
- 14. Microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) colorectal cancer (CRC) with **ONE** of the following:
  - a. Used as a single agent

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- i. Disease is metastatic
- ii. Progressed following treatment with fluoropyrimidine, oxaliplatin, and irinotecan
- iii. Diagnosis confirmed by PCR-based assay genetic testing
- b. Used in combination with ipilimumab
  - i. Disease is unresectable or metastatic
  - ii. Diagnosis confirmed by PCR-based assay genetic testing
- 15. Unresectable malignant pleural mesothelioma
  - a. Used as first-line treatment in combination with ipilimumab
- Advanced or metastatic gastric cancer, gastroesophageal junction cancer, or esophageal adenocarcinoma
  - a. Tumors express PD-L1 as determined by an FDA-approved test
  - b. Used in combination with fluoropyrimidine- and platinum-containing chemotherapy
- 17. Small cell lung cancer
- 18. Metastatic anal carcinoma
- 19. Merkel cell carcinoma

#### **AND ALL** of the following for **ALL** indications:

- a. Prescriber agrees to discontinue treatment for any immune mediated adverse reaction (encephalitis, nephritis, rash, decreased renal function and endocrinopathies) or disease progression
- Female patients of reproductive potential only: patient will be advised to use
  effective contraception during treatment with Opdivo and for 5 months after the
  last dose

### Prior - Approval Renewal Requirements

Age 12 years of age or older

#### **Diagnoses**

Patient must have **ONE** of the following:

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- 1. Unresectable or metastatic melanoma
- 2. Adjuvant treatment of melanoma post resection: one renewal **only** 
  - a. Stage IIB, Stage IIC, Stage III, or Stage IV melanoma
- 3. Resectable non-small cell lung cancer (NSCLC)
  - a. Used as a single agent after surgery as adjuvant treatment
  - b. **NO** known EGFR mutations or ALK rearrangements
- 4. Metastatic non-small cell lung cancer
  - a. IF used in combination with ipilimumab: one renewal only
- 5. Recurrent non-small cell lung cancer
  - a. Used in combination with ipilimumab: one renewal only
- 6. Advanced renal cell carcinoma
  - a. IF used in combination with cabozantinib: one renewal only
- 7. Relapsed or progressed classical Hodgkin lymphoma
- 8. Recurrent or metastatic squamous cell carcinoma of the head and neck
- 9. Urothelial carcinoma
  - a. **IF** used as adjuvant treatment in patients at high risk of recurrence after radical resection: one renewal **only**
  - b. **IF** used for unresectable or metastatic urothelial carcinoma, as first-line treatment in combination with cisplatin and gemcitabine: one renewal **only**
- 10. Unresectable or metastatic hepatocellular carcinoma (HCC)
- 11. Completely resected esophageal or gastroesophageal junction cancer with residual pathologic disease: one renewal **only**
- 12. Unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC)
  - a. Used in combination with **ONE** of the following:
    - i. Fluoropyrimidine- and platinum-containing chemotherapy: one renewal only

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- ii. Ipilimumab: one renewal only
- 13. Unresectable advanced, recurrent, or metastatic esophageal squamous cell carcinoma (ESCC)
  - a. Prior treatment with fluoropyrimidine- and platinum-based chemotherapy
- 14. Unresectable or metastatic microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) colorectal cancer (CRC)
- 15. Unresectable malignant pleural mesothelioma
  - a. Used in combination with ipilimumab: one renewal only
- 16. Advanced or metastatic gastric cancer, gastroesophageal junction cancer, or esophageal adenocarcinoma
  - a. Used in combination with fluoropyrimidine- and platinum-containing chemotherapy: one renewal **only**
- 17. Small cell lung cancer
- 18. Metastatic anal carcinoma
- 19. Merkel cell carcinoma

#### **AND ALL** of the following for **ALL** indications:

- a. NO disease progression or unacceptable toxicity
- b. Prescriber agrees to discontinue treatment for any immune mediated adverse reaction (encephalitis, nephritis, rash, decreased renal function and endocrinopathies) or disease progression
- Female patients of reproductive potential only: patient will be advised to use
  effective contraception during treatment with Opdivo and for 5 months after the
  last dose

#### **Policy Guidelines**

#### Pre - PA Allowance

None

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# **Prior - Approval Limits**

**Duration** 6 months

### Prior - Approval Renewal Limits

### **Duration\***

Indication	Renewal PA Duration*	Number of Renewals Allowed
Adjuvant treatment of melanoma post resection	6 months	One renewal only
Adjuvant treatment of urothelial carcinoma (patients at high risk of recurrence after radical resection)	6 months	One renewal only
Completely resected esophageal or gastroesophageal junction cancer with residual pathological disease	6 months	One renewal only
Resectable non-small cell lung cancer (NSCLC) as adjuvant treatment after surgery	12 months	One renewal only
Advanced or metastatic gastric cancer, gastroesophageal junction cancer, and esophageal adenocarcinoma	18 months	One renewal only
Unresectable malignant pleural mesothelioma	18 months	One renewal only
Unresectable or metastatic urothelial carcinoma (first-line, in combination with cisplatin and gemcitabine)	18 months	One renewal only
Unresectable or metastatic hepatocellular carcinoma	18 months	One renewal only

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Metastatic non-small cell lung cancer (NSCLC) **	18 months	Used with ipilimumab One renewal only
		As a single agent
		Until disease progression or unacceptable toxicity
Recurrent non-small cell lung cancer (NSCLC)**	18 months	One renewal only
Unresectable or metastatic MSI-H or dMMR colorectal cancer (CRC)	18 months	Used with ipilimumab followed by Opdivo as a single agent:
		One renewal only
Metastatic MSI-H or dMMR colorectal cancer (CRC)	18 months	Prior treatment with fluoropyrimidine, oxaliplatin, and irinotecan:
		Until disease progression or unacceptable toxicity
Esophageal squamous cell carcinoma	18 months	Used with ipilimumab or fluoropyrmidine- and platinum-containing chemotherapy:
		One renewal only
		Prior treatment with fluoropyrimidine- and platinum-based chemotherapy:
		Until disease progression or unacceptable toxicity
Advanced renal cell carcinoma	18 months	Used with cabozantinib:
		One renewal only
		NOT being used with cabozantinib:
		Until disease progression or unacceptable toxicity

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All other indications	18 months	Until disease progression or unacceptable toxicity
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**<sup>\*\*</sup>NO** renewal for Resectable non-small cell lung cancer (NSCLC) used as <u>neoadjuvant</u> treatment

#### Rationale

#### **Summary**

Opdivo (nivolumab) is a monoclonal antibody indicated for the treatment of various types of cancers. Opdivo works by binding to the programmed cell death-1 (PD-1) receptor, and blocking its interaction with PD-1 ligands, PD-L1 and PD-L2. This interaction releases the inhibitory effects of PD-1 pathway-mediated inhibition of the immune response, including the anti-tumor immune response, resulting in decreased tumor growth. Opdivo carries warnings for immune-mediated adverse reactions, infusion-related reactions, complications of allogeneic HSCT and embryo-fetal toxicity. The safety and effectiveness of Opdivo have not been established in pediatric patients age less than 12 years of age with melanoma or microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) colorectal cancer (CRC) or in pediatric patients less than 18 years of age for the other approved indications (1).

Prior authorization is required to ensure the safe, clinically appropriate, and cost-effective use of Opdivo while maintaining optimal therapeutic outcomes.

#### References

- 1. Opdivo [package insert]. Princeton, NJ: Bristol-Myers Squibb Company; June 2025.
- 2. NCCN Drugs & Biologics Compendium<sup>®</sup> Nivolumab 2025. National Comprehensive Cancer Network, Inc. Accessed on July 28, 2025.

Policy History	
Date	Action
January 2015	Addition to PA
March 2015	Annual editorial review and reference update
	Addition of Metastatic squamous non-small cell lung cancer
June 2015	Annual review
October 2015	Addition of BRAF V600 wild-type, the patient must use in combination with ipilimumab, and metastatic non-small cell lung cancer with the squamous cell requirement along with disease must have progressed after FDA-approved therapy if patient has EGFR or ALK tumor expression option.

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December 2015 Annual review

Addition of new indication of renal cell carcinoma after prior treatment with

an anti-angiogenic therapy

March 2016 Annual review

Removal of requirements: disease progression following Yervoy (ipilimumab) if BRAF V600 mutation positive, a BRAF inhibitor, BRAF V600 wild-type the patient must use in combination with ipilimumab

Policy number change from 5.04.53 to 5.21.53

June 2016 Annual review

Addition of relapsed or progressed classical Hodgkin lymphoma in patients who have had autologous hematopoietic stem cell transplantation (HSCT) and post-transplantation therapy with brentuximab vedotin (Adcetris). Addition of Prescriber agrees to discontinue treatment for any immune mediated adverse reaction (encephalitis, nephritis, rash, decreased renal function and endocrinopathies) or disease progression in renewal section

per SME

September 2016 Annual review

December 2016 Addition of recurrent or metastatic squamous cell carcinoma of the head

and neck with progression on or after platinum-based chemotherapy

February 2017 Addition of locally advanced or metastatic urothelial carcinoma with one of

the following: disease progression during or following platinum-containing chemotherapy, or disease progression within 12 months of neoadjuvant or

adjuvant treatment with platinum-containing chemotherapy

June 2017 Annual editorial review

Addition to the relapsed or progressed classical Hodgkin lymphoma: patient has had 3 or more lines systemic therapy that includes autologous

**HSCT** 

August 2017 Addition of microsatellite instability-high (MSI-H) or mismatch repair

deficient (dMMR) metastatic colorectal cancer

September 2017 Annual review

October 2017 Addition of hepatocellular carcinoma

December 2017 Annual review

January 2018 Addition of melanoma with lymph node involvement or metastatic disease

who have undergone complete resection, in the adjuvant setting

March 2018 Annual review

May 2018 Addition of indication: Intermediate or poor risk, previously untreated

advanced renal cell carcinoma, in combination with ipilimumab; malignant pleural mesothelioma, small cell lung cancer, metastatic anal carcinoma, and Merkel cell carcinoma; and changed the age from 18 to 12 yrs of age

June 2018 Annual review

July 2018 Addition of indication: metastatic colorectal cancer as a single agent or in

combination with ipilimumab

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August 2018 Addition of metastatic small cell lung cancer, progression after platinum-

based chemotherapy and at least one other line of therapy

September 2018 Annual editorial review and reference update

November 2018 Annual review

March 2019 Change to indication: unresectable or metastatic melanoma, as a single

agent or in combination with ipilimumab

June 2019 Annual review

April 2020 Revised indication: hepatocellular carcinoma as a single agent or in

combination with ipilimumab

May 2020 Addition of indication: metastatic NSCLC whose tumors express PD-L1, as

first-line treatment used in combination with ipilimumab, with no EGFR or ALK genomic tumor aberrations. Revised metastatic NSCLC indication so

they need to have both disease progression after platinum-based chemotherapy and disease progression after therapy for EGFR or ALK tumor aberration, if present. Addition of indication: metastatic or recurrent NSCLC with no EGFR or ALK tumor aberrations as first-line treatment with ipilimumab and 2 cycles of platinum-doublet chemotherapy. Changed renewal duration from 12 months to 18 months. Added "ONE renewal ONLY for metastatic/recurrent NSCLC when used with ipilimumab and for

adjuvant treatment of melanoma post resection"

June 2020 Annual review. Addition of indication: esophageal squamous cell

carcinoma (ESCC)

September 2020 Annual review

October 2020 Per FEP, revised malignant pleural mesothelioma indication: removed it

from the off-label section, included the requirement that it must be unresectable and used as first-line treatment in combination with ipilimumab. Added "no disease progression or unacceptable toxicity"

renewal requirement

December 2020 Annual review

January 2021 Removed metastatic small cell lung cancer indication per PI. Small cell

lung cancer remains a recommended indication per NCCN

February 2021 Addition of indication: advanced renal cell carcinoma in combination with

cabozantinib as first-line treatment

March 2021 Annual review

May 2021 Addition of indication: advanced or metastatic gastric cancer,

gastroesophageal junction cancer, or esophageal adenocarcinoma

June 2021 Addition of indication: completely resected esophageal or

gastroesophageal junction cancer with residual pathologic disease. Added

renewal duration chart for clarity

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September 2021 Annual editorial review and reference update. Removed indication:

hepatocellular carcinoma as a single agent. Addition of indication:

adjuvant treatment of patients with urothelial carcinoma who are at high

risk of recurrence after undergoing radical resection

March 2022 Annual editorial review and reference update

April 2022 Addition of indication per PI update: neoadjuvant treatment of resectable

**NSCLC** 

June 2022 Annual review and reference update. Addition of indication per PI update:

unresectable advanced or metastatic esophageal squamous cell

carcinoma in combination with fluoropyrimidine- and platinum-containing

chemotherapy or in combination with ipilimumab

September 2022 Annual review and reference update

December 2022 Revised quantity limits chart to separate out metastatic NSCLC when used

with ipilimumab

March 2023 Annual review and reference update September 2023 Annual review and reference update

November 2023 Per PI update, added requirement of Stage IIB, IIC, III, or IV melanoma for

adjuvant treatment of completely resected patients

December 2023 Annual review and reference update
March 2024 Annual review and reference update

April 2024 Per PI update, added indication of unresectable or metastatic urothelial

carcinoma, as first-line treatment in combination with cisplatin and

gemcitabine

June 2024 Annual review and reference update September 2024 Annual review and reference update

October 2024 Per PI update, added indication of resectable NSCLC as adjuvant

treatment as a single agent after surgery. Updated advanced RCC

indication to require use with ipilimumab to be first-line

December 2024 Annual review

March 2025 Annual review and reference update

May 2025 Per PI update, added indication of unresectable colorectal cancer and

added separate indication of metastatic CRC after disease progression following treatment with fluoropyrimidine, oxaliplatin, and irinotecan. Added HCC as first-line treatment and added that HCC must be

unresectable or metastatic. Also added monitoring agreement to initiation criteria and contraception agreement to both initiation and continuation

June 2025 Annual review and reference update

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July 2025 Per PI update, added requirement of tumor expressing PD-L1 for

unresectable advanced or metastatic esophageal squamous cell carcinoma and gastric cancer, gastroesophageal junction cancer, and

esophageal adenocarcinoma

September 2025 Annual review and reference update

Keywords

This policy was approved by the FEP® Pharmacy and Medical Policy Committee on September 19, 2025 and is effective on October 1, 2025.