

## Medical Policy: Opdivo® (nivolumab) Intravenous

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EmblemHealth established the clinical review criteria based upon a review of currently available clinical information (including clinical outcome studies in the peer reviewed published medical literature, regulatory status of the technology, evidence-based guidelines of public health and health research agencies, evidence-based guidelines and positions of leading national health professional organizations, views of physicians practicing in relevant clinical areas, and other relevant factors). EmblemHealth expressly reserves the right to revise these conclusions as clinical information changes and welcomes further relevant information. Each benefit program defines which services are covered. The conclusion that a particular service or supply is medically necessary does not constitute a representation or warranty that this service or supply is covered and/or paid for by EmblemHealth, as some programs exclude coverage for services or supplies that EmblemHealth considers medically necessary.

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### Length of Authorization

Coverage will be provided for six months and may be renewed.

- Adjuvant use in the treatment of melanoma patients can be authorized up to a maximum of 12 months of therapy.

### Dosing Limits [Medical Benefit]

**Max Units (per dose and over time):**

Indication	Billable Units (BU)	Per unit time (days)
Esophageal Cancer, MPM, MSI-H/dMMR CRC, MSI-H/dMMR, Small Bowel Adenocarcinoma	340 BU	14 days
Cutaneous Melanoma, Esophagogastric/Gastroesophageal Junction Cancer, HCC, cHL, NSCLC,	480 BU	28 days

RCC, SCCHN, Urothelial Carcinoma		
Uveal Melanoma	1140 BU	14 days

## Guideline

### I. Initial Approval Criteria

Coverage is provided for the following conditions:

1. Patient must be 18 years of age or older (unless otherwise specified); **AND**
2. Patient has not received previous therapy with a programmed death (PD-1/PD-L1)-directed therapy (e.g., avelumab, pembrolizumab, atezolizumab, durvalumab, etc.) unless otherwise specified; **AND**

#### **Melanoma †**

1. Patient is 12 years of age or older; **AND**
2. Patient's disease is unresectable or metastatic; **AND**
  - a. Used as a single agent or in combination with ipilimumab; **OR**
3. Patient has unresectable or metastatic uveal melanoma; **AND**
  - a. Used as a single agent or in combination with ipilimumab; **OR**
4. Used as adjuvant treatment as a single agent; **AND**
  - a. Patient has lymph node involvement or metastatic disease and has undergone complete resection; **OR**
5. Used for retreatment of disease (*see Renewal Criteria*)

#### **Hepatocellular Carcinoma (HCC) †**

1. Used in combination with ipilimumab; **AND**
2. Patient progressed on or was intolerant to sorafenib; **AND**
3. Patient has a laboratory confirmed diagnosis of hepatocellular carcinoma; **AND**
4. Patient has Child-Pugh Class A

#### **Non-Small Cell Lung Cancer (NSCLC) †**

1. Indicated for the treatment of patients with metastatic NSCLC with progression on or after platinum-based chemotherapy; patients with EGFR or ALK genomic tumor aberrations should have disease progression on FDA-approved therapy for these aberrations prior to receiving nivolumab; **OR**
2. In combination with ipilimumab, it is indicated for the first line treatment of adult patients with metastatic non small cell lung cancer (NSCLC) whose tumors express PD L1 ( $\geq 1\%$ ) as determined by an FDA approved test, with no EGFR or ALK genomic tumor aberrations; **OR**
3. In combination with ipilimumab and 2 cycles of platinum-doublet chemotherapy, it is indicated for the first-line treatment of adult patients with metastatic or recurrent non-small cell lung cancer with no EGFR or ALK genomic tumor aberrations.
4. For the Neoadjuvant Treatment of Resectable Non-Small Cell Lung Cancer:

In combination with platinum-doublet chemotherapy, is indicated as neoadjuvant treatment of adult patients with resectable (tumors  $\geq 4$  cm or node positive) non-small cell lung cancer (NSCLC).

### **Renal Cell Carcinoma (RCC) †**

1. Used in combination with ipilimumab; **AND**
  - a. Used as initial therapy in patients with advanced or metastatic disease with intermediate or poor risk; **OR**
  - b. Used as first-line therapy in patients with relapsed or stage IV disease with favorable risk; **OR**
  - c. Used as subsequent therapy in patients with relapsed or stage IV disease **OR**
2. Used as a single agent; **AND**
  - a. Patient has advanced disease with intermediate or poor risk; **OR**
  - b. Patient has relapsed, unresectable metastatic disease; **AND**
    - i. Used as subsequent therapy for clear cell histology; **OR**
    - ii. Patient has non-clear cell histology
3. Used in combination with cabozantinib,
  - a. Used for the first-line treatment of patients with advanced disease; **OR**
  - b. Used as subsequent therapy in patients with relapsed or stage IV disease

### **Classical Hodgkin Lymphoma (cHL) †**

1. Patient has relapsed or progressive disease; **AND**
  - a. Patient had an autologous hematopoietic stem cell transplantation (HSCT) and post-transplantation brentuximab vedotin; **OR**
  - b. Patient has received 3 or more lines of systemic therapy that includes autologous HSCT

### **Squamous Cell Carcinoma of the Head and Neck (SCCHN) †**

1. Patient has recurrent, or metastatic disease; **AND**
2. Patient has progressed on or after platinum-based therapy; **AND**

### **Colorectal Cancer †**

1. Adult and pediatric (12 years and older) patients with microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR) disease; **AND**
2. Disease progression following treatment with a fluoropyrimidine, oxaliplatin, and irinotecan, as a single agent or in combination with ipilimumab.

### **Urothelial Carcinoma †**

1. Must be used as a single agent; **AND**
  - A. Must be used as subsequent systemic therapy after previous platinum treatment\* \* **OR** as second-line treatment after chemotherapy other than a platinum; **AND**
    - a. Patient has **ONE** of the following:
      - i. Locally advanced or metastatic disease; **OR**

- ii. Muscle invasive bladder cancer with local recurrence or persistent disease in a preserved bladder ‡; **OR**
  - iii. Disease recurrence post-cystectomy ‡; **OR**
  - iv. Recurrent or metastatic Primary Carcinoma of the Urethra ‡; **AND**
    - a) Patient does not have recurrent stage T3-4 disease or palpable inguinal lymph nodes; **OR**
  - v. Metastatic Upper GU Tract Tumors ‡; **OR**
  - vi. Metastatic Urothelial Carcinoma of the Prostate ‡; **OR**
- B. Used as adjuvant therapy †; AND**
- a. Patient has urothelial carcinoma of the bladder, bulbar urethra, prostate with stromal invasion, ureter, or renal pelvis; **AND**
  - b. Patient underwent radical surgical resection or partial cystectomy; **AND**
  - c. Patient is at high risk of disease recurrence\*\*

*\*If platinum treatment occurred greater than 12 months ago, the patient should be re-treated with platinum-based therapy. Patients with comorbidities (e.g., hearing loss, neuropathy, poor PS, renal insufficiency, etc.) may not be eligible for cisplatin. Carboplatin may be substituted for cisplatin particularly in those patients with a GFR <60 mL/min or a PS of 2.*

*\*\* High risk of disease recurrence is defined as:*

*ypT2-ypT4a or ypN+ for patients who received neoadjuvant cisplatin; OR*

*pT3-pT4a or pN+ for patients who did not receive neoadjuvant cisplatin and are also ineligible for or refused adjuvant cisplatin therapy*

### **Esophageal Squamous Cell Carcinoma (ESCC) †**

1. Patient with unresectable advanced, recurrent or metastatic esophageal squamous cell carcinoma after prior fluoropyrimidine- and platinum-based chemotherapy.
  - A. The medication is being used as adjuvant treatment of completely resected esophageal or gastroesophageal junction cancer with residual pathologic disease in adult patients who have received neoadjuvant chemoradiotherapy (CRT); **OR**
  - B. The medication is being used in combination with fluoropyrimidine- and platinum-containing chemotherapy, for the first-line treatment of adult patients with unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC); **OR**
  - C. The medication is being used in combination with ipilimumab, for the first-line treatment of adult patients with unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC); **OR**
  - D. The medication is being used for the treatment of adult patients with unresectable advanced, recurrent or metastatic esophageal squamous cell carcinoma (ESCC) after prior fluoropyrimidine- and platinum-based chemotherapy.

### **Unresectable Malignant Pleural Mesothelioma (MPM) †**

1. Used as a first-line therapy:
  - a. Nivolumab is used in combination with ipilimumab; **OR**

2. Used as a subsequent systemic therapy‡
  - a. As a single agent; **OR**
  - b. Nivolumab is used in combination with ipilimumab (if not administered first-line).

### **Gastric Cancer, Gastroesophageal Junction Cancer and Esophageal Adenocarcinoma†**

1. The patient has advanced or metastatic disease; **AND**
2. Opdivo will be used in combination with fluoropyrimidine- and platinum-containing chemotherapy

## **II. Renewal Criteria**

Authorizations can be renewed based on the following criteria:

1. Patient continues to meet the criteria identified above; **AND**
2. Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include the following: severe infusion reactions, complications of allogeneic HSCT, severe immune-mediated adverse reactions such as pneumonitis, colitis, hepatitis, endocrinopathies, nephritis/renal dysfunction, rash, encephalitis, etc.; **AND**
3. Tumor response with stabilization of disease or decrease in size of tumor or tumor spread; **AND**
4. For the following indication, the patient has not exceeded a maximum of twelve (12) months of therapy:
  - a. Adjuvant treatment of melanoma; **OR**
5. For the following indication, the patient has not exceeded a maximum of twenty four (24) months of therapy:
  - a. Esophageal Cancer (in combination with fluoropyrimidine- and platinum-containing chemotherapy OR ipilimumab)
  - b. Esophagogastric/Gastroesophageal Junction Cancer (in combination with fluoropyrimidine- and platinum-containing chemotherapy)
  - c. Malignant Pleural Mesothelioma
  - d. Non-Small Cell Lung Cancer (in combination with ipilimumab with or without platinum-doublet chemotherapy)
  - e. Renal Cell Carcinoma (in combination with cabozantinib)
6. **Non-Small Cell Lung Cancer (neoadjuvant therapy in combination with platinum-doublet chemotherapy)**
  - Patient has not exceeded a maximum of three (3) cycles
7. **Urothelial Carcinoma (adjuvant therapy)**
  - Patient has not exceeded a maximum of one (1) year of therapy
8. **Esophageal and Esophagogastric/Gastroesophageal Junction Cancer (adjuvant therapy)**
  - Patient has not exceeded a maximum of one (1) year of therapy
9. **Classical Hodgkin Lymphoma (in combination with brentuximab vedotin)**
  - Patient has not exceeded a maximum of twelve (12) weeks of therapy

## 10. Retreatment for Melanoma (metastatic or unresectable disease) ‡

- A. Used for re-treatment of patients who experienced disease control, but subsequently have disease progression/relapse > 3 months after treatment discontinuation; **OR**
- B. Used as subsequent therapy
  - i. in combination with ipilimumab, in patients who experienced disease relapse and/or progression within 3 months after initial monotherapy with an immune checkpoint-inhibitor; **OR**
  - ii. Used as a single agent or in combination with ipilimumab if anti-PD-1 was not previously used

### Limitations/Exclusions

Opdivo is considered investigational when used for any indication not listed above.

## Applicable Procedure Codes

Code	Description
J9299	Injection, nivolumab, 1 mg; 1 billable unit = 1 mg

## Applicable NDCs

Code	Description
00003-3772-xx	Opdivo 40 mg/4 mL single-use vial
00003-3774-xx	Opdivo 100 mg/10 mL single-use vial
00003-3756-xx	Opdivo 120 mg/12 mL single-dose vial
00003-3734-xx	Opdivo 240 mg/24 mL single-use vial

## ICD-10 Diagnoses

Code	Description
C00.0	Malignant neoplasm of external upper lip
C00.1	Malignant neoplasm of external lower lip
C00.2	Malignant neoplasm of external lip, unspecified
C00.3	Malignant neoplasm of upper lip, inner aspect
C00.4	Malignant neoplasm of lower lip, inner aspect
C00.5	Malignant neoplasm of lip, unspecified, inner aspect
C00.6	Malignant neoplasm of commissure of lip, unspecified
C00.8	Malignant neoplasm of overlapping sites of lip
C01	Malignant neoplasm of base of tongue
C02.0	Malignant neoplasm of dorsal surface of tongue
C02.1	Malignant neoplasm of border of tongue
C02.2	Malignant neoplasm of ventral surface of tongue
C02.3	Malignant neoplasm of anterior two-thirds of tongue, part unspecified
C02.4	Malignant neoplasm of lingual tonsil

C02.8	Malignant neoplasm of overlapping sites of tongue
C02.9	Malignant neoplasm of tongue, unspecified
C03.0	Malignant neoplasm of upper gum
C03.1	Malignant neoplasm of lower gum
C03.9	Malignant neoplasm of gum, unspecified
C04.0	Malignant neoplasm of anterior floor of mouth
C04.1	Malignant neoplasm of lateral floor of mouth
C04.8	Malignant neoplasm of overlapping sites of floor of mouth
C04.9	Malignant neoplasm of floor of mouth, unspecified
C05.0	Malignant neoplasm of hard palate
C05.1	Malignant neoplasm of soft palate
C06.0	Malignant neoplasm of cheek mucosa
C06.2	Malignant neoplasm of retromolar area
C06.80	Malignant neoplasm of overlapping sites of unspecified parts of mouth
C06.89	Malignant neoplasm of overlapping sites of other parts of mouth
C06.9	Malignant neoplasm of mouth, unspecified
C09.0	Malignant neoplasm of tonsillar fossa
C09.1	Malignant neoplasm of tonsillar pillar (anterior) (posterior)
C09.8	Malignant neoplasm of overlapping sites of tonsil
C09.9	Malignant neoplasm of tonsil, unspecified
C10.3	Malignant neoplasm of posterior wall of oropharynx
C11.0	Malignant neoplasm of superior wall of nasopharynx
C11.1	Malignant neoplasm of posterior wall of nasopharynx
C11.2	Malignant neoplasm of lateral wall of nasopharynx
C11.3	Malignant neoplasm of anterior wall of nasopharynx
C11.8	Malignant neoplasm of overlapping sites of nasopharynx
C11.9	Malignant neoplasm of nasopharynx, unspecified
C12	Malignant neoplasm of pyriform sinus
C13.0	Malignant neoplasm of postcricoid region
C13.1	Malignant neoplasm of aryepiglottic fold, hypopharyngeal aspect
C13.2	Malignant neoplasm of posterior wall of hypopharynx
C13.8	Malignant neoplasm of overlapping sites of hypopharynx
C13.9	Malignant neoplasm of hypopharynx, unspecified
C14.0	Malignant neoplasm of pharynx, unspecified
C14.2	Malignant neoplasm of Waldeyer's ring
C14.8	Malignant neoplasm of overlapping sites of lip, oral cavity and pharynx
C17.0	Malignant neoplasm of duodenum
C17.1	Malignant neoplasm of jejunum
C17.2	Malignant neoplasm of ileum
C17.8	Malignant neoplasm of overlapping sites of small intestine
C17.9	Malignant neoplasm of small intestine, unspecified
C18.0	Malignant neoplasm of cecum
C18.1	Malignant neoplasm of appendix
C18.2	Malignant neoplasm of ascending colon
C18.3	Malignant neoplasm of hepatic flexure
C18.4	Malignant neoplasm of transverse colon

C18.5	Malignant neoplasm of splenic flexure
C18.6	Malignant neoplasm of descending colon
C18.7	Malignant neoplasm of sigmoid colon
C18.8	Malignant neoplasm of overlapping sites of colon
C18.9	Malignant neoplasm of colon, unspecified
C19	Malignant neoplasm of rectosigmoid junction
C20	Malignant neoplasm of rectum
C21.0	Malignant neoplasm of anus, unspecified
C21.1	Malignant neoplasm of anal canal
C21.2	Malignant neoplasm of cloacogenic zone
C21.8	Malignant neoplasm of overlapping sites of rectum, anus and anal canal
C22.0	Liver cell carcinoma
C22.9	Malignant neoplasm of liver, not specified as primary or secondary
C31.0	Malignant neoplasm of maxillary sinus
C31.1	Malignant neoplasm of ethmoidal sinus
C32.0	Malignant neoplasm of glottis
C32.1	Malignant neoplasm of supraglottis
C32.2	Malignant neoplasm of subglottis
C32.3	Malignant neoplasm of laryngeal cartilage
C32.8	Malignant neoplasm of overlapping sites of larynx
C32.9	Malignant neoplasm of larynx, unspecified
C33	Malignant neoplasm of trachea
C34.00	Malignant neoplasm of unspecified main bronchus
C34.01	Malignant neoplasm of right main bronchus
C34.02	Malignant neoplasm of left main bronchus
C34.10	Malignant neoplasm of upper lobe, unspecified bronchus or lung
C34.11	Malignant neoplasm of upper lobe, right bronchus or lung
C34.12	Malignant neoplasm of upper lobe, left bronchus or lung
C34.2	Malignant neoplasm of middle lobe, bronchus or lung
C34.30	Malignant neoplasm of lower lobe, unspecified bronchus or lung
C34.31	Malignant neoplasm of lower lobe, right bronchus or lung
C34.32	Malignant neoplasm of lower lobe, left bronchus or lung
C34.80	Malignant neoplasm of overlapping sites of unspecified bronchus and lung
C34.81	Malignant neoplasm of overlapping sites of right bronchus and lung
C34.82	Malignant neoplasm of overlapping sites of left bronchus and lung
C34.90	Malignant neoplasm of unspecified part of unspecified bronchus or lung
C34.91	Malignant neoplasm of unspecified part of right bronchus or lung
C34.92	Malignant neoplasm of unspecified part of left bronchus or lung
C45.0	Mesothelioma of pleura
C38.4	Malignant neoplasm of pleura
C43.0	Malignant melanoma of lip
C43.10	Malignant melanoma of unspecified eyelid, including canthus
C43.111	Malignant melanoma of right upper eyelid, including canthus
C43.112	Malignant melanoma of right lower eyelid, including canthus
C43.121	Malignant melanoma of left upper eyelid, including canthus
C43.122	Malignant melanoma of left lower eyelid, including canthus



C43.20	Malignant melanoma of unspecified ear and external auricular canal
C43.21	Malignant melanoma of right ear and external auricular canal
C43.22	Malignant melanoma of left ear and external auricular canal
C43.30	Malignant melanoma of unspecified part of face
C43.31	Malignant melanoma of nose
C43.39	Malignant melanoma of other parts of face
C43.4	Malignant melanoma of scalp and neck
C43.51	Malignant melanoma of anal skin
C43.52	Malignant melanoma of skin of breast
C43.59	Malignant melanoma of other part of trunk
C43.60	Malignant melanoma of unspecified upper limb, including shoulder
C43.61	Malignant melanoma of right upper limb, including shoulder
C43.62	Malignant melanoma of left upper limb, including shoulder
C43.70	Malignant melanoma of unspecified lower limb, including hip
C43.71	Malignant melanoma of right lower limb, including hip
C43.72	Malignant melanoma of left lower limb, including hip
C43.8	Malignant melanoma of overlapping sites of skin
C43.9	Malignant melanoma of skin, unspecified
C44.00	Unspecified malignant neoplasm of skin of lip
C44.02	Squamous cell carcinoma of skin of lip
C44.09	Other specified malignant neoplasm of skin of lip
C4A.0	Merkel cell carcinoma of lip
C4A.10	Merkel cell carcinoma of eyelid, including canthus
C4A.111	Merkel cell carcinoma of right upper eyelid, including canthus
C4A.112	Merkel cell carcinoma of right lower eyelid, including canthus
C4A.121	Merkel cell carcinoma of left upper eyelid, including canthus
C4A.122	Merkel cell carcinoma of left lower eyelid, including canthus
C4A.20	Merkel cell carcinoma of unspecified ear and external auricular canal
C4A.21	Merkel cell carcinoma of right ear and external auricular canal
C4A.22	Merkel cell carcinoma of left ear and external auricular canal
C4A.30	Merkel cell carcinoma of unspecified part of face
C4A.31	Merkel cell carcinoma of nose
C4A.39	Merkel cell carcinoma of other parts of face
C4A.4	Merkel cell carcinoma of scalp and neck
C4A.51	Merkel cell carcinoma of anal skin
C4A.52	Merkel cell carcinoma of skin of breast
C4A.59	Merkel cell carcinoma of other part of trunk
C4A.60	Merkel cell carcinoma of unspecified upper limb, including shoulder
C4A.61	Merkel cell carcinoma of right upper limb, including shoulder
C4A.62	Merkel cell carcinoma of left upper limb, including shoulder
C4A.70	Merkel cell carcinoma of unspecified lower limb, including hip
C4A.71	Merkel cell carcinoma of right lower limb, including hip
C4A.72	Merkel cell carcinoma of left lower limb, including hip
C4A.8	Merkel cell carcinoma of overlapping sites
C4A.9	Merkel cell carcinoma, unspecified
C61	Malignant neoplasm of prostate

C64.1	Malignant neoplasm of right kidney, except renal pelvis
C64.2	Malignant neoplasm of left kidney, except renal pelvis
C64.9	Malignant neoplasm of unspecified kidney, except renal pelvis
C65.1	Malignant neoplasm of right renal pelvis
C65.2	Malignant neoplasm of left renal pelvis
C65.9	Malignant neoplasm of unspecified renal pelvis
C66.1	Malignant neoplasm of right ureter
C66.2	Malignant neoplasm of left ureter
C66.9	Malignant neoplasm of unspecified ureter
C67.0	Malignant neoplasm of trigone of bladder
C67.1	Malignant neoplasm of dome of bladder
C67.2	Malignant neoplasm of lateral wall of bladder
C67.3	Malignant neoplasm of anterior wall of bladder
C67.4	Malignant neoplasm of posterior wall of bladder
C67.5	Malignant neoplasm of bladder neck
C67.6	Malignant neoplasm of ureteric orifice
C67.7	Malignant neoplasm of urachus
C67.8	Malignant neoplasm of overlapping sites of bladder
C67.9	Malignant neoplasm of bladder, unspecified
C68.0	Malignant neoplasm of urethra
C69.30	Malignant neoplasm of unspecified choroid
C69.31	Malignant neoplasm of right choroid
C69.32	Malignant neoplasm of left choroid
C69.40	Malignant neoplasm of unspecified ciliary body
C69.41	Malignant neoplasm of right ciliary body
C69.42	Malignant neoplasm of left ciliary body
C69.60	Malignant neoplasm of unspecified orbit
C69.61	Malignant neoplasm of right orbit
C69.62	Malignant neoplasm of left orbit
C69.90	Malignant neoplasm of unspecified site of unspecified eye
C69.91	Malignant neoplasm of unspecified site of right eye
C69.92	Malignant neoplasm of unspecified site of left eye
C76.0	Malignant neoplasm of head, face and neck
C77.0	Secondary and unspecified malignant neoplasm of lymph nodes of head, face and neck
C78.00	Secondary malignant neoplasm of unspecified lung
C78.01	Secondary malignant neoplasm of right lung
C78.02	Secondary malignant neoplasm of left lung
C78.6	Secondary malignant neoplasm of retroperitoneum and peritoneum
C78.7	Secondary malignant neoplasm of liver and intrahepatic bile duct
C78.89	Secondary malignant neoplasm of other digestive organs
C79.31	Secondary malignant neoplasm of brain
C79.51	Secondary malignant neoplasm of bone
C79.52	Secondary malignant neoplasm of bone marrow
C7A.1	Malignant poorly differentiated neuroendocrine tumors
C7B.1	Secondary Merkel cell carcinoma
C80.0	Disseminated malignant neoplasm, unspecified

C80.1	Malignant (primary) neoplasm, unspecified
C81.10	Nodular sclerosis Hodgkin lymphoma, unspecified site
C81.11	Nodular sclerosis Hodgkin lymphoma, lymph nodes of head, face, and neck
C81.12	Nodular sclerosis Hodgkin lymphoma, intrathoracic lymph nodes
C81.13	Nodular sclerosis Hodgkin lymphoma, intra-abdominal lymph nodes
C81.14	Nodular sclerosis Hodgkin lymphoma, lymph nodes of axilla and upper limb
C81.15	Nodular sclerosis Hodgkin lymphoma, lymph nodes of inguinal region and lower limb
C81.16	Nodular sclerosis Hodgkin lymphoma, intrapelvic lymph nodes
C81.17	Nodular sclerosis Hodgkin lymphoma, spleen
C81.18	Nodular sclerosis Hodgkin lymphoma, lymph nodes of multiple sites
C81.19	Nodular sclerosis Hodgkin lymphoma, extranodal and solid organ sites
C81.20	Mixed cellularity Hodgkin lymphoma, unspecified site
C81.21	Mixed cellularity Hodgkin lymphoma, lymph nodes of head, face, and neck
C81.22	Mixed cellularity Hodgkin lymphoma, intrathoracic lymph nodes
C81.23	Mixed cellularity Hodgkin lymphoma, intra-abdominal lymph nodes
C81.24	Mixed cellularity Hodgkin lymphoma, lymph nodes of axilla and upper limb
C81.25	Mixed cellularity Hodgkin lymphoma, lymph nodes of inguinal region and lower limb
C81.26	Mixed cellularity Hodgkin lymphoma, intrapelvic lymph nodes
C81.27	Mixed cellularity Hodgkin lymphoma, spleen
C81.28	Mixed cellularity Hodgkin lymphoma, lymph nodes of multiple sites
C81.29	Mixed cellularity Hodgkin lymphoma, extranodal and solid organ sites
C81.30	Lymphocyte depleted Hodgkin lymphoma, unspecified site
C81.31	Lymphocyte depleted Hodgkin lymphoma, lymph nodes of head, face, and neck
C81.32	Lymphocyte depleted Hodgkin lymphoma, intrathoracic lymph nodes
C81.33	Lymphocyte depleted Hodgkin lymphoma, intra-abdominal lymph nodes
C81.34	Lymphocyte depleted Hodgkin lymphoma, lymph nodes of axilla and upper limb
C81.35	Lymphocyte depleted Hodgkin lymphoma, lymph nodes of inguinal region and lower limb
C81.36	Lymphocyte depleted Hodgkin lymphoma, intrapelvic lymph nodes
C81.37	Lymphocyte depleted Hodgkin lymphoma, spleen
C81.38	Lymphocyte depleted Hodgkin lymphoma, lymph nodes of multiple sites
C81.39	Lymphocyte depleted Hodgkin lymphoma, extranodal and solid organ sites
C81.40	Lymphocyte-rich Hodgkin lymphoma, unspecified site
C81.41	Lymphocyte-rich Hodgkin lymphoma, lymph nodes of head, face, and neck
C81.42	Lymphocyte-rich Hodgkin lymphoma, intrathoracic lymph nodes
C81.43	Lymphocyte-rich Hodgkin lymphoma, intra-abdominal lymph nodes
C81.44	Lymphocyte-rich Hodgkin lymphoma, lymph nodes of axilla and upper limb
C81.45	Lymphocyte-rich Hodgkin lymphoma, lymph nodes of inguinal region and lower limb
C81.46	Lymphocyte-rich Hodgkin lymphoma, intrapelvic lymph nodes
C81.47	Lymphocyte-rich Hodgkin lymphoma, spleen
C81.48	Lymphocyte-rich Hodgkin lymphoma, lymph nodes of multiple sites
C81.49	Lymphocyte-rich Hodgkin lymphoma, extranodal and solid organ sites
C81.70	Other Hodgkin lymphoma unspecified site
C81.71	Other Hodgkin lymphoma lymph nodes of head, face, and neck
C81.72	Other Hodgkin lymphoma intrathoracic lymph nodes
C81.73	Other Hodgkin lymphoma intra-abdominal lymph nodes
C81.74	Other Hodgkin lymphoma lymph nodes of axilla and upper limb

C81.75	Other Hodgkin lymphoma lymph nodes of inguinal region and lower limb
C81.76	Other Hodgkin lymphoma intrapelvic lymph nodes
C81.77	Other Hodgkin lymphoma spleen
C81.78	Other Hodgkin lymphoma lymph nodes of multiple sites
C81.79	Other Hodgkin lymphoma extranodal and solid organ sites
C81.90	Hodgkin lymphoma, unspecified site
C81.91	Hodgkin lymphoma, unspecified lymph nodes of head, face, and neck
C81.92	Hodgkin lymphoma, unspecified intrathoracic lymph nodes
C81.93	Hodgkin lymphoma, unspecified intra-abdominal lymph nodes
C81.94	Hodgkin lymphoma, unspecified lymph nodes of axilla and upper limb
C81.95	Hodgkin lymphoma, unspecified lymph nodes of inguinal region and lower limb
C81.96	Hodgkin lymphoma, unspecified intrapelvic lymph nodes
C81.97	Hodgkin lymphoma, unspecified spleen
C81.98	Hodgkin lymphoma, unspecified lymph nodes of multiple sites
C81.99	Hodgkin lymphoma, unspecified extranodal and solid organ sites
D09.0	Carcinoma in situ of bladder
D37.01	Neoplasm of uncertain behavior of lip
D37.02	Neoplasm of uncertain behavior of tongue
D37.05	Neoplasm of uncertain behavior of pharynx
D37.09	Neoplasm of uncertain behavior of other specified sites of the oral cavity
D38.0	Neoplasm of uncertain behavior of larynx
D38.5	Neoplasm of uncertain behavior of other respiratory organs
D38.6	Neoplasm of uncertain behavior of respiratory organ, unspecified
Z85.038	Personal history of other malignant neoplasm of large intestine
Z85.068	Personal history of other malignant neoplasm of small intestine
Z85.118	Personal history of other malignant neoplasm of bronchus and lung
Z85.21	Personal history of malignant neoplasm of larynx
Z85.22	Personal history of malignant neoplasm of nasal cavities, middle ear, and accessory sinuses
Z85.51	Personal history of malignant neoplasm of bladder
Z85.528	Personal history of other malignant neoplasm of kidney
Z85.59	Personal history of malignant neoplasm of other urinary tract organ
Z85.71	Personal history of Hodgkin lymphoma
Z85.810	Personal history of malignant neoplasm of tongue
Z85.818	Personal history of malignant neoplasm of other sites of lip, oral cavity and pharynx
Z85.819	Personal history of malignant neoplasm of unspecified site of lip, oral cavity and pharynx
Z85.820	Personal history of malignant melanoma of skin
Z85.821	Personal history of Merkel cell carcinoma

## Revision History

Company(ies)	DATE	REVISION
EmblemHealth & ConnectiCare	1/29/2024	Annual Review: Initial Criteria: Melanoma †: Added: "Patient is 12 years of age or older"

		<p>Hepatocellular Carcinoma (HCC): removed “as a single agent” in the Statement: “Used as a single agent; or in combination with ipilimumab; AND” and removed Patient has Child-Pugh Class B7 disease</p> <p>Renal Cell Carcinoma (RCC) †</p> <p>Added: “Used as first-line therapy in patients with relapsed or stage IV disease with favorable risk; OR”</p> <p>Removed: “unresectable metastatic disease with clear cell histology; “ from the statement: “Used as subsequent therapy in patients with relapsed, unresectable metastatic disease with clear cell histology; or stage IV disease”</p> <p>Added “Used as subsequent therapy in patients with relapsed or stage IV disease” under: Used in combination with cabozantinib,</p> <p>Classical Hodgkin Lymphoma (cHL) † Removed: Must be used as a single agent; Renewal Criteria: Retreatment for Melanoma (metastatic or unresectable disease) Added: “Used as a single agent or in combination with ipilimumab if anti-PD-1 was not previously used”</p> <p>Added NDC 00003-3756-xx</p>
<p>EmblemHealth &amp; ConnectiCare</p>	<p>5/25/2023</p>	<p>Annual Review:</p> <p><u>Urothelial Carcinoma</u>- Initial Criteria, Added “OR as second-line treatment after chemotherapy other than a platinum;” “Muscle invasive bladder cancer with local recurrence or persistent disease in a preserved bladder ‡;” and “Used as adjuvant therapy †; AND</p> <p>a. Patient has urothelial carcinoma of the bladder, bulbar urethra, prostate with stromal invasion, ureter, or renal pelvis; AND</p> <p>b .Patient underwent radical surgical resection or partial cystectomy; AND</p> <p>c. Patient is at high risk of disease recurrence**” AND “** High risk of disease recurrence is defined as:</p> <p>ypT2-ypT4a or ypN+ for patients who received neoadjuvant cisplatin; OR</p> <p>pT3-pT4a or pN+ for patients who did not receive neoadjuvant cisplatin and are also ineligible for or refused adjuvant cisplatin therapy”</p> <p><u>Squamous Cell Carcinoma of the Head and Neck</u>; Initial Criteria: Removed “Patient has unresectable disease” and “Must be used as a single agent”</p> <p>Added “Gastric Cancer, Gastroesophageal Junction Cancer and Esophageal Adenocarcinoma” Indications and initial criteria.</p> <p><u>Esophageal Squamous Cell Carcinoma</u> Initial Criteria:</p> <p>Removed “Patient with unresectable advanced, recurrent or metastatic esophageal squamous cell carcinoma after prior fluoropyrimidine- and platinum-based chemotherapy.” Added “The medication is being used as adjuvant treatment of completely resected esophageal or gastroesophageal junction cancer with residual pathologic disease in adult patients who have received neoadjuvant chemoradiotherapy (CRT); OR</p> <p>B. The medication is being used in combination with fluoropyrimidine- and platinum-containing chemotherapy, for the first-line treatment of adult</p>

		<p>patients with unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC); OR</p> <p>C. The medication is being used in combination with ipilimumab, for the first-line treatment of adult patients with unresectable advanced or metastatic esophageal squamous cell carcinoma (ESCC); OR</p> <p>D. The medication is being used for the treatment of adult patients with unresectable advanced, recurrent or metastatic esophageal squamous cell carcinoma (ESCC) after prior fluoropyrimidine- and platinum-based chemotherapy.”</p> <p><u>RCC</u>: Initial Criteria: Added ” Used in combination with cabozantinib, for the first-line treatment of patients with advanced disease.”</p> <p><u>NSCLC</u>: Initial Criteria: added “Neoadjuvant Treatment of Resectable Non-Small Cell Lung Cancer: in combination with platinum-doublet chemotherapy, is indicated as neoadjuvant treatment of adult patients with resectable (tumors ≥4 cm or node positive) non-small cell lung cancer (NSCLC).”</p> <p><u>Renewal Criteria</u>: added “5. For the following indication, the patient has not exceeded a maximum of twenty four (24) months of therapy:</p> <p>b. Esophageal Cancer (in combination with fluoropyrimidine- and platinum-containing chemotherapy OR ipilimumab)</p> <p>c. Esophagogastric/Gastroesophageal Junction Cancer (in combination with fluoropyrimidine- and platinum-containing chemotherapy)</p> <p>e. Malignant Pleural Mesothelioma</p> <p>g.Non-Small Cell Lung Cancer (in combination with ipilimumab with or without platinum-doublet chemotherapy)</p> <p>h.Renal Cell Carcinoma (in combination with cabozantinib)”</p> <p>added “1 Non-Small Cell Lung Cancer (neoadjuvant therapy in combination with platinum-doublet chemotherapy)</p> <ul style="list-style-type: none"> <li>• Patient has not exceeded a maximum of three (3) cycles</li> </ul> <p>2.Urothelial Carcinoma (adjuvant therapy)</p> <ul style="list-style-type: none"> <li>• Patient has not exceeded a maximum of one (1) year of therapy</li> </ul> <p>3.Esophageal and Esophagogastric/Gastroesophageal Junction Cancer (adjuvant therapy)</p> <ul style="list-style-type: none"> <li>• Patient has not exceeded a maximum of one (1) year of therapy</li> </ul> <p>4.Classical Hodgkin Lymphoma (in combination with brentuximab vedotin)</p> <ul style="list-style-type: none"> <li>• Patient has not exceeded a maximum of twelve (12) weeks of therapy”</li> </ul> <p>Updated dosing limits chart</p>
EmblemHealth & ConnectiCare	09/15/2022	Transferred policy to new template

EmblemHealth & ConnectiCare	2/25/2020	Removed small-cell lung cancer indication from policy as per updated FDA labeling and NCCN guideline recommendation
EmblemHealth & ConnectiCare	11/19/2020	Added newest NSCLC indication for patients with no EGFR or ALK genomic tumor aberrations in combination with ipilimumab and 2 cycles of platinum-doublet chemotherapy
EmblemHealth & ConnectiCare	10/16/2020	Added Malignant Pleural Mesothelioma indication per FDA label update and NCCN guidelines. Added ICD 10 codes
EmblemHealth & ConnectiCare	06/16/2020	Under Covered Indications (added indication and criteria): Esophageal Squamous Cell Carcinoma (ESCC) and dosing per FDA Label
EmblemHealth & ConnectiCare	06/15/2020	Added dosing limits per FDA label for NSCLC
EmblemHealth & ConnectiCare	06/01/2020	Added under Initial Approval Criteria (NSCLC): In combination with ipilimumab, it is indicated for the first line treatment of adult patients with metastatic non-small cell lung cancer (NSCLC) whose tumors express PD L1 ( $\geq 1\%$ ) as determined by an FDA approved test, with no EGFR or ALK genomic tumor aberrations.
EmblemHealth & ConnectiCare	03/30/2020	Added max dosing limits when in combination with ipilimumab for Hepatocellular Carcinoma (HCC)
EmblemHealth & ConnectiCare	01/01/2020	-Under Guideline, added Small Cell Lung Cancer indication, platinum-based chemotherapy and at least one other line of therapy -Under Guideline, added Metastatic Colorectal Cancer indication per FDA label

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