

Opdivo® (nivolumab) (Intravenous)

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I. Length of Authorization ¹

Coverage will be provided for six months and may be renewed (unless otherwise specified).

- Adjuvant use in the treatment of melanoma can be authorized up to a maximum of 12 months of therapy.
- Use in the treatment of NSCLC in combination with ipilimumab can be authorized up to a maximum of 2 years of therapy.
- Use in the treatment of NSCLC in combination with ipilimumab and two (2) cycles of platinum-doublet chemotherapy can be authorized up to a maximum of 2 years of therapy.

II. Dosing Limits

A. Quantity Limit (max daily dose) [NDC Unit]:

- Opdivo 40 mg/4 mL single-use vial: 2 vials per 14 days
- Opdivo 100 mg/10 mL single-use vial: 2 vials per 14 days
- Opdivo 240 mg/24 mL single-use vial: 4 vials per 14 days

B. Max Units (per dose and over time) [HCPCS Unit]:

Indication	Billable Units (BU)	Per unit time (days)
Merkel Cell	340 BU	14 days
Melanoma/HCC (in combination with ipilimumab)	Initial: 140 BU	21 days x 4 doses
	Followed by: 480 BU	28 days
Melanoma/RCC/HCC/NSCLC (as a single agent), cHL, SCCHN, MSI-H/dMMR CRC (as a single agent), Anal Carcinoma, Gestational Trophoblastic Tumor, Esophageal Squamous Cell Carcinoma & Urothelial Carcinoma	480 BU	28 days
Metastatic NSCLC with PD-L1 expressing tumors (in combination with ipilimumab)	340 BU	14 days

Metastatic or recurrent NSCLC (in combination with ipilimumab and platinum-doublet chemotherapy)	380 BU	21 days
SCLC (as a single agent)	240 BU	14 days
MSI-H/dMMR CRC (in combination with ipilimumab)	Initial: 340 BU	21 days x 4 doses
	Followed by: 480 BU	28 days
Small Bowel Adenocarcinoma, CNS Metastases from NSCLC (both as single agents)	340 BU	14 days
Small Bowel Adenocarcinoma (in combination with ipilimumab)	Initial: 340 BU	21 days x 4 doses
	Followed by: 340 BU	14 days
RCC (in combination with ipilimumab)	Initial: 340 BU	21 days x 4 doses
	Followed by: 480 BU	28 days
SCLC (in combination with ipilimumab)	Initial: 340 BU	21 days x 4 doses
	Followed by: 340 BU	14 days
MPM (as a single agent or in combination with ipilimumab)	340 BU	14 days
CNS Metastases from Melanoma & Uveal Melanoma (both in combination with ipilimumab)	Initial: 140 BU	21 days x 4 doses
	Followed by: 340 BU	14 days
CNS Metastases from Melanoma (as a single agent)	340 BU	14 days
Uveal Melanoma (as a single agent)	1140 BU	14 days
Extranodal NK/ T-Cell Lymphoma	40 BU	14 days

III. Initial Approval Criteria ¹

Coverage is provided for the following conditions:

- Patient is at least 18 years of age (unless otherwise specified); **AND**

Universal Criteria

- Patient has not received previous therapy with a programmed death (PD-1/PD-L1)-directed therapy (e.g., cemiplimab, avelumab, pembrolizumab, atezolizumab, durvalumab, etc.) prior to initiation of therapy, unless otherwise specified; **AND**

Cutaneous Melanoma † ⊕ ^{1,2,15-18}

- Patient has unresectable or metastatic disease OR disease with satellite/in-transit metastases or recurrence; **AND**
 - Used as a single agent or in combination with ipilimumab as first line therapy; **OR**
 - Used for retreatment of disease as re-induction as a single agent or in combination with ipilimumab in patients who experienced disease control (*i.e., complete or partial response or stable disease*) from prior checkpoint inhibitor therapy, but subsequently have disease progression/relapse > 3 months after treatment discontinuation; **OR**
 - Used as subsequent therapy after disease progression on first-line therapy or maximum clinical benefit from BRAF-targeted therapy (e.g., dabrafenib/trametinib, vemurafenib/cobimetinib, encorafenib/binimetinib, etc.); **AND**
 - Used as a single agent or in combination with ipilimumab if checkpoint inhibitor immunotherapy was not previously used; **OR**

- Used in combination with ipilimumab for patients who progressed on single agent checkpoint inhibitor immunotherapy; **OR**
- Used as adjuvant treatment as a single agent; **AND**
 - Patient has lymph node involvement and has undergone complete resection, complete lymph node dissection (CLND), therapeutic lymph node dissection (TLND), or nodal basin ultrasound surveillance; **OR**
 - Patient has satellite/in-transit metastases or recurrence and has no evidence of disease after complete excision; **OR**
 - Patient has undergone TLND and/or complete resection of nodal recurrence; **OR**
 - Patient has undergone complete resection of distant metastatic disease

Uveal Melanoma ‡^{2,19,20}

- Patient has distant metastatic disease; **AND**
- Used as a single agent or in combination with ipilimumab

Hepatocellular Carcinoma (HCC) † Φ^{1,2,21}

- Patient has locally advanced, unresectable, inoperable, or metastatic disease; **AND**
- Used as subsequent therapy; **AND**
 - Patient has Child-Pugh Class A or B disease; **AND**
 - Used as a single agent; **OR**
 - Patient has Child-Pugh Class A disease; **AND**
 - Used in combination with ipilimumab

Non-Small Cell Lung Cancer (NSCLC) †^{1,2,22,23}

- Patient has metastatic disease with a high tumor mutational burden (TMB)* (i.e., ≥10 mutations per megabase) ‡; **AND**
 - Used as a single-agent or in combination with ipilimumab as first-line therapy; **OR**
- Used for recurrent, advanced, or metastatic disease (excluding locoregional recurrence or symptomatic local disease without evidence of disseminated disease) or mediastinal lymph node recurrence with prior radiation therapy; **AND**
 - Used as first-line therapy; **AND**
 - Used for one of the following:
 - Used in patients with PS 0-1 who have EGFR, ALK, ROS1, BRAF, MET exon skipping mutation, and RET rearrangement negative** tumors and PD-L1 expression <1%
 - Used in patients with PS 0-1 who are positive for one of the following molecular biomarkers: BRAF V600E mutations, NTRK gene fusions, MET exon 14 skipping mutations, or RET rearrangements
 - Used in patients with PS 0-2 for PD-L1 expression-positive (PD-L1 ≥1%) tumors that are EGFR, ALK, ROS1, BRAF, MET exon skipping mutation, and RET rearrangement negative**;

- Used in combination with ipilimumab; **OR**
- Used in combination with ipilimumab and platinum-doublet chemotherapy; **OR**
- Used as subsequent therapy; **AND**
 - Used as a single agent; **OR**
 - Used for one of the following:
 - Used in patients with PS 0-1 who have EGFR, ALK, ROS1 positive tumors and have received prior targeted therapy§
 - Used in patients with PS 0-1 who are positive for one of the following molecular biomarkers: BRAF V600E mutations, NTRK gene fusions, MET exon 14 skipping mutations, or RET rearrangements; **AND**
- Used in combination with ipilimumab; **OR**
- Used in combination with ipilimumab, pemetrexed, and either carboplatin or cisplatin for nonsquamous cell histology; **OR**
- Used in combination with ipilimumab, paclitaxel, and carboplatin for squamous cell histology

**TMB is an evolving biomarker that may be helpful in selecting patients for immunotherapy. There is no consensus on how to measure TMB.*

*** Note: If there is insufficient tissue to allow testing for all of the EGFR, ALK, ROS1, BRAF, MET, and RET, repeat biopsy and/or plasma testing should be done. If these are not feasible, treatment is guided by available results and, if unknown, these patients are treated as though they do not have driver oncogenes.*

Renal Cell Carcinoma (RCC) † 1,2,25,26

- Used in combination with ipilimumab for clear cell histology; **AND**
 - Used as first-line therapy in patients with advanced, relapsed, or stage IV disease with intermediate or poor risk; **OR**
 - Used as first-line therapy in patients with relapsed or stage IV disease with favorable risk; **OR**
 - Used as subsequent therapy in patients with relapsed or stage IV disease; **OR**
- Used as a single agent; **AND**
 - Used as subsequent therapy in patients with advanced, relapsed, or stage IV disease for clear cell histology; **OR**
 - Patient has relapsed or stage IV disease and non-clear cell histology

Classical Hodgkin Lymphoma (cHL) † ⊕ 1,2,27,28

- Used as a single agent; **AND**
 - Patient is at least 18 years old and has received 2 or more prior lines of therapy OR used as palliative therapy in patients more than 60 years old; **AND**
 - Patient has relapsed or progressive disease after an autologous hematopoietic stem cell transplantation (HSCT) with or without brentuximab vedotin; **OR**

- Patient has relapsed or refractory disease and is either transplant-ineligible based on comorbidities or has failure of second-line chemotherapy; **OR**
- Patient is post-allogeneic stem-cell transplant; **OR**
- Used in combination with brentuximab vedotin as subsequent therapy (if not previously used) for relapsed or refractory disease

Squamous Cell Carcinoma of the Head and Neck (SCCHN) † 1,2,29

- Used as single-agent therapy; **AND**
- Patient has unresectable, recurrent, persistent, or metastatic disease; **AND**
- Disease has progressed on or after platinum-based therapy; **AND**
- Patient does not have nasopharyngeal disease

Urothelial Carcinoma (Bladder Cancer) † 1,2,30

- Used as a single agent; **AND**
- Used as subsequent systemic therapy after previous platinum treatment*; **AND**
- Patient has one of the following diagnoses:
 - Locally advanced or metastatic urothelial carcinoma; **OR**
 - Local bladder cancer recurrence or persistent disease in a preserved bladder ‡; **OR**
 - Local or metastatic bladder cancer recurrence post-cystectomy ‡; **OR**
 - Recurrent or metastatic primary carcinoma of the urethra ‡; **AND**
 - Patient does not have recurrence of stage T3-4 disease or palpable inguinal lymph nodes; **OR**
 - Metastatic upper genitourinary (GU) tract tumors ‡; **OR**
 - Metastatic urothelial carcinoma of the prostate ‡

*** Note:**

- If platinum treatment occurred greater than 12 months ago, the patient should be re-treated with platinum-based therapy if the patient is still platinum eligible (see below for cisplatin- or carboplatin-ineligible comorbidities).
 - Cisplatin-ineligible comorbidities may include the following: GFR < 60 mL/min, PS ≥ 2, hearing loss of ≥ 25 decibels (dB) at two contiguous frequencies, or grades ≥ 2 peripheral neuropathy. Carboplatin may be substituted for cisplatin particularly in those patients with a GFR < 60 mL/min or a PS of 2.
 - Carboplatin-ineligible comorbidities may include the following: GFR < 30 mL/min, PS ≥ 3, grade ≥ 3 peripheral neuropathy, or NYHA class ≥ 3, etc.

Small Cell Lung Cancer (SCLC) † Φ 1,2,24

- Used as subsequent systemic therapy; **AND**
 - Used as a single agent for metastatic disease with progression after platinum-based treatment and at least one other line of therapy †; **OR**
 - Used as a single agent or in combination with ipilimumab ‡; **AND**
 - Used for relapse within 6 months following complete response, partial response, or stable disease with initial treatment; **AND**
 - Patient did not relapse while on maintenance atezolizumab or durvalumab; **OR**
 - Used for primary progressive disease

Colorectal Cancer (CRC) † 1,2,31,32

- Patient is at least 12 years of age; **AND**
- Patient’s disease must be microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR); **AND**
 - Used as a single agent or in combination with ipilimumab as subsequent therapy for advanced or metastatic disease that progressed following treatment with a fluoropyrimidine-, oxaliplatin-, and/or irinotecan-based chemotherapy; **OR**
 - Used as a single agent or in combination with ipilimumab as primary treatment for unresectable or metastatic disease after previous adjuvant FOLFOX (fluorouracil, leucovorin and oxaliplatin) or CapeOX (capecitabine-oxaliplatin) in the past 12 months ‡; **OR**
 - Used as a single agent as primary treatment for unresectable or metastatic disease in patients who are not candidates for intensive therapy ‡; **OR**
 - Used as a single agent for unresectable or metastatic disease that remains unresectable after primary treatment in patients who are not candidates for intensive therapy ‡

Merkel Cell Carcinoma ‡^{2,4,33}

- Used as a single agent; **AND**
- Patient has disseminated metastatic disease

Central Nervous System (CNS) Cancer ‡^{2,5,34,41,42}

- Used in one of the following treatment settings:
 - Used as initial treatment in patients with small asymptomatic brain metastases; **OR**
 - Used for relapsed disease in patients with limited brain metastases and stable systemic disease or reasonable treatment options; **OR**
 - Patient has recurrent limited brain metastases; **OR**
 - Used for recurrent disease in patients with extensive brain metastases and stable systemic disease or reasonable systemic treatment options; **AND**
- Used as a single-agent or in combination with ipilimumab for the treatment of brain metastases in patients with melanoma; **OR**
- Used as a single-agent for the treatment of brain metastases in patients with PD-L1 positive non-small cell lung cancer (NSCLC)

Anal Carcinoma ‡^{2,6,35}

- Patient has metastatic squamous cell disease; **AND**
- Used as a single agent for subsequent therapy

Gestational Trophoblastic Neoplasia ‡^{2,36}

- Used as single-agent therapy for multiagent chemotherapy resistant disease; **AND**
 - Patient has intermediate placental site trophoblastic tumor (PSTT) or epithelioid trophoblastic tumor (ETT); **AND**
 - Patient has recurrent or progressive disease ; **AND**
 - Patient was previously treated with a platinum/etoposide containing regimen; **OR**

- Patient has methotrexate-resistant high risk disease (i.e., FIGO stages II-III and ≥ 7 Prognostic score OR FIGO stage IV disease)

Malignant Pleural Mesothelioma ‡^{2,37,38}

- Used as a single agent or in combination with ipilimumab as subsequent therapy

Small Bowel Adenocarcinoma ‡^{2,31,39}

- Patient has advanced or metastatic disease that is microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR); **AND**
- Used as a single agent or in combination with ipilimumab (in one of the following settings):
 - As subsequent therapy; **OR**
 - As initial therapy in patients with prior oxaliplatin exposure in the adjuvant setting or contraindication

Extranodal NK/ T-Cell Lymphoma ‡^{2,40}

- Used as a single agent for relapsed or refractory nasal type disease; **AND**
- Disease progressed following additional treatment with an alternative asparaginase-based chemotherapy regimen not previously used; **AND**
- Participation in a clinical trial is unavailable

Esophageal Squamous Cell Carcinoma (ESCC) † Φ^{1,44}

- Used as a single agent; **AND**
- Used as subsequent therapy for unresectable advanced (or is not a surgical candidate), recurrent, or metastatic disease

† FDA Approved Indication(s); ‡ Compendia recommended indication(s); Φ Orphan Drug

Genomic Aberration/Mutational Driver Targeted Therapies (Note: not all inclusive, refer to guidelines for appropriate use) §
Sensitizing <i>EGFR</i> mutation-positive tumors <ul style="list-style-type: none"> – Afatinib – Erlotinib – Dacomitinib – Gefitinib – Osimertinib
<i>ALK</i> rearrangement-positive tumors <ul style="list-style-type: none"> – Alectinib – Brigatinib – Ceritinib – Crizotinib – Lorlatinib
<i>ROS1</i> rearrangement-positive tumors <ul style="list-style-type: none"> – Ceritinib – Crizotinib – Entrectinib
<i>BRAF</i> V600E-mutation positive tumors <ul style="list-style-type: none"> – Dabrafenib ± Trametinib – Vemurafenib
<i>NTRK</i> Gene Fusion positive tumors <ul style="list-style-type: none"> – Larotrectinib – Entrectinib

PD-1/PD-L1 expression-positive tumors ($\geq 1\%$) <ul style="list-style-type: none"> - Pembrolizumab - Atezolizumab - Nivolumab \pm ipilimumab
<i>MET</i> Exon-14 skipping mutations <ul style="list-style-type: none"> - Capmatinib - Crizotinib
<i>RET</i> rearrangement-positive tumors <ul style="list-style-type: none"> - Selpercatinib - Cabozantinib - Vandetanib

IV. Renewal Criteria ^{1,2,4-6,15-42}

Authorizations can be renewed based on the following criteria:

- Patient continues to meet universal and other indication-specific relevant criteria such as concomitant therapy requirements (not including prerequisite therapy), performance status, etc. identified in section III; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: severe infusion reactions, complications of allogeneic hematopoietic stem cell transplantation (HSCT), severe immune-mediated adverse reactions (i.e. pneumonitis, colitis, hepatitis, endocrinopathies, nephritis/renal dysfunction, adverse skin reactions/rash, encephalitis), etc.; **AND**
- Disease response with treatment defined as stabilization of disease or decrease in size of tumor or tumor spread; **AND**

Cutaneous Melanoma (adjuvant therapy)

- Patient has not exceeded a maximum of twelve (12) months of therapy

NSCLC (in combination with ipilimumab or in combination with ipilimumab and two (2) cycles of platinum-doublet chemotherapy)

- Patient has not exceeded a maximum of two (2) years of therapy

Cutaneous Melanoma Re-induction ‡

- *Refer to Section III for criteria (see Melanoma – Used for retreatment of disease as re-induction)*

V. Dosage/Administration ^{1,4-6,24,31-42}

Indication	Dose
Merkel Cell	Administer 240 mg intravenously every 2 weeks or 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity
Anal Cancer	Administer 240 mg intravenously every 2 weeks, 480 mg intravenously every 4 weeks, or 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity
Cutaneous Melanoma	<u>Single agent (excluding adjuvant therapy):</u>

	<ul style="list-style-type: none"> Administer 240 mg intravenously every 2 weeks OR 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab (excluding adjuvant therapy):</u></p> <ul style="list-style-type: none"> Administer 1 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then follow with single agent regimen <p><u>Adjuvant treatment:</u></p> <ul style="list-style-type: none"> Administer 240 mg intravenously every 2 weeks or 480 mg intravenously every 4 weeks until disease recurrence or unacceptable toxicity for up to 1 year
Uveal Melanoma	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> Administer up to 10 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> Administer 1 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity
NSCLC	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> Administer 240 mg intravenously every 2 weeks OR 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> Administer 3 mg/kg intravenously every 2 weeks, with ipilimumab 1 mg/kg every 6 weeks, until disease progression or unacceptable toxicity for up to 2 years <p><u>In combination with ipilimumab and platinum-doublet chemotherapy for metastatic or recurrent disease:</u></p> <ul style="list-style-type: none"> Administer 360 mg intravenously every 3 weeks, with ipilimumab 1 mg/kg every 6 weeks and histology-based platinum-doublet chemotherapy every 3 weeks for 2 cycles, until disease progression or unacceptable toxicity for up to 2 years
cHL, SCCHN, Urothelial Carcinoma, Esophageal Squamous Cell Carcinoma, & Gestational Trophoblastic Neoplasia (GTN)	Administer 240 mg intravenously every 2 weeks or 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity
MSI-H/dMMR CRC	<p><u>Adult patients and for pediatric patients ≥ 12 years and ≥ 40 kg</u></p> <ul style="list-style-type: none"> As a single agent: Administer 240 mg intravenously every 2 weeks or 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity In combination with ipilimumab: Administer 3 mg/kg intravenously, with ipilimumab 1 mg/kg on the same day, every 3 weeks for 4 doses, then follow with the single agent regimen <p><u>Pediatric patients ≥ 12 years and < 40 kg</u></p> <ul style="list-style-type: none"> As a single agent: Administer 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity

	<ul style="list-style-type: none"> • In combination with ipilimumab: Administer 3 mg/kg intravenously, with ipilimumab 1 mg/kg on the same day, every 3 weeks for 4 doses, then follow with the single agent regimen
SCLC	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> • Administer 240 mg intravenously every 2 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> • Administer 1 mg/kg to 3 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then 3 mg/kg intravenously every 2 weeks
Renal Cell Carcinoma (RCC)	<p><u>Single-agent:</u></p> <ul style="list-style-type: none"> • Administer 240 mg intravenously every 2 weeks or 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> • Administer 3 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then follow with single-agent regimen
Hepatocellular Carcinoma (HCC)	<p><u>Single-agent:</u></p> <ul style="list-style-type: none"> • Administer 240 mg intravenously every 2 weeks or 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> • Administer 1 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then follow with single-agent regimen
Malignant Pleural Mesothelioma (MPM)	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> • Administer 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> • Administer 3 mg/kg intravenously every 2 weeks, with ipilimumab 1 mg/kg every 6 weeks, until disease progression or unacceptable toxicity OR • Administer 240 mg intravenously every 2 weeks, with ipilimumab 1 mg/kg every 6 weeks (for a total of 4 ipilimumab doses); treatment with nivolumab is continued for up to 2 years or until disease progression or unacceptable toxicity
CNS Metastases from Melanoma	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> • Administer 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity <p><u>In combination with ipilimumab:</u></p> <ul style="list-style-type: none"> • Administer 1 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity
CNS Metastases from NSCLC	Administer 3 mg/kg intravenously every 2 weeks until disease progression or unacceptable toxicity
Small Bowel Adenocarcinoma	<p><u>Single agent:</u></p> <ul style="list-style-type: none"> • Administer 3 mg/kg intravenously every 2 weeks, or 240 mg intravenously every 2 weeks, or 480 mg intravenously every 4 weeks until disease progression or unacceptable toxicity

	<u>In combination with ipilimumab:</u> <ul style="list-style-type: none"> Administer 3 mg/kg intravenously, with ipilimumab on the same day, every 3 weeks for 4 doses, then 3 mg/kg or 240 mg intravenously every 2 weeks until disease progression or unacceptable toxicity
Extranodal NK/ T-Cell Lymphoma	Administer 40 mg intravenously every 2 weeks until disease progression or unacceptable toxicity
<u>Dosing should be calculated using actual body weight and not flat dosing (as applicable) based on the following:</u>	
<u>Weight ≥ 74 kg:</u> <ul style="list-style-type: none"> Standard dose 240 mg IV every 2 weeks OR 480 mg IV every 4 weeks 	
<u>Weight is 67 kg to 73 kg:</u> <ul style="list-style-type: none"> Use 440 mg IV every 4 weeks 	
<u>Weight is < 66kg:</u> <ul style="list-style-type: none"> Use 400 mg IV every 4 weeks 	
<i>Note: This information is not meant to replace clinical decision making when initiating or modifying medication therapy and should only be used as a guide. Patient-specific variables should be taken into account.</i>	

VI. Billing Code/Availability Information

HCPCS Code:

- J9299 - Injection, nivolumab, 1 mg; 1 billable unit = 1 mg

NDC:

- Opdivo 40 mg/4 mL single-use vial: 00003-3772-xx
- Opdivo 100 mg/10 mL single-use vial: 00003-3774-xx
- Opdivo 240 mg/24 mL single-use vial: 00003-3734-xx

VII. References

- Opdivo [package insert]. Princeton, NJ; Bristol-Myers Squibb Company; June 2020. Accessed July 2020.
- Referenced with permission from the NCCN Drugs & Biologics Compendium (NCCN Compendium®) nivolumab. National Comprehensive Cancer Network, 2020. The NCCN Compendium® is a derivative work of the NCCN Guidelines®. NATIONAL COMPREHENSIVE CANCER NETWORK®, NCCN®, and NCCN GUIDELINES® are trademarks owned by the National Comprehensive Cancer Network, Inc. To view the most recent and complete version of the Compendium, go online to NCCN.org. Accessed July 2020.
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Appendix 1 – Covered Diagnosis Codes

ICD-10	ICD-10 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
C06.8	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
C12	Malignant neoplasm of pyriform sinus
C13.0	Malignant neoplasm of postcricoid region

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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
C15.3	Malignant neoplasm of upper third of esophagus
C15.4	Malignant neoplasm of middle third of esophagus
C15.5	Malignant neoplasm of lower third of esophagus
C15.8	Malignant neoplasm of overlapping sites of esophagus
C15.9	Malignant neoplasm of esophagus, unspecified
C16.0	Malignant neoplasm of cardia
C17.0	Malignant neoplasm of duodenum
C17.1	Malignant neoplasm of jejunum
C17.2	Malignant neoplasm of ileum
C17.3	Meckel's diverticulum, malignant
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

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C22.8	Malignant neoplasm of liver, primary, unspecified as to type
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
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

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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
C45.0	Mesothelioma of pleura
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

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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
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
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

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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

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C81.98	Hodgkin lymphoma, unspecified lymph nodes of multiple sites
C81.99	Hodgkin lymphoma, unspecified extranodal and solid organ sites
C84.90	Mature T/NK-cell lymphomas, unspecified, unspecified site
C84.91	Mature T/NK-cell lymphomas, unspecified, lymph nodes of head, face, and neck
C84.92	Mature T/NK-cell lymphomas, unspecified, intrathoracic lymph nodes
C84.93	Mature T/NK-cell lymphomas, unspecified, intra-abdominal lymph nodes
C84.94	Mature T/NK-cell lymphomas, unspecified, lymph nodes of axilla and upper limb
C84.95	Mature T/NK-cell lymphomas, unspecified, lymph nodes of inguinal region and lower limb
C84.96	Mature T/NK-cell lymphomas, unspecified, intrapelvic lymph nodes
C84.97	Mature T/NK-cell lymphomas, unspecified, spleen
C84.98	Mature T/NK-cell lymphomas, unspecified, lymph nodes of multiple sites
C84.99	Mature T/NK-cell lymphomas, unspecified, extranodal and solid organ sites
C84.Z0	Other mature T/NK-cell lymphomas, unspecified site
C84.Z1	Other mature T/NK-cell lymphomas, lymph nodes of head, face, and neck
C84.Z2	Other mature T/NK-cell lymphomas, intrathoracic lymph nodes
C84.Z3	Other mature T/NK-cell lymphomas, intra-abdominal lymph nodes
C84.Z4	Other mature T/NK-cell lymphomas, lymph nodes of axilla and upper limb
C84.Z5	Other mature T/NK-cell lymphomas, lymph nodes of inguinal region and lower limb
C84.Z6	Other mature T/NK-cell lymphomas, intrapelvic lymph nodes
C84.Z7	Other mature T/NK-cell lymphomas, spleen
C84.Z8	Other mature T/NK-cell lymphomas, lymph nodes of multiple sites
C84.Z9	Other mature T/NK-cell lymphomas, extranodal and solid organ sites
C86.0	Extranodal NK/T-cell lymphoma, nasal type
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
D37.8	Neoplasm of uncertain behavior of other specified digestive organs
D37.9	Neoplasm of uncertain behavior of digestive organ, unspecified
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
D39.2	Neoplasm of uncertain behavior of placenta
D39.8	Neoplasm of uncertain behavior of other specified female genital organs
D39.9	Neoplasm of uncertain behavior of female genital organ, unspecified

OPDIVO® (nivolumab) Prior Auth Criteria

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Z85.00	Personal history of malignant neoplasm of unspecified digestive organ
Z85.01	Personal history of malignant neoplasm of esophagus
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
Z85.59	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

Appendix 2 – Centers for Medicare and Medicaid Services (CMS)

Medicare coverage for outpatient (Part B) drugs is outlined in the Medicare Benefit Policy Manual (Pub. 100-2), Chapter 15, §50 Drugs and Biologicals. In addition, National Coverage Determination (NCD), Local Coverage Determinations (LCDs), and Local Coverage Articles (LCAs) may exist and compliance with these policies is required where applicable. They can be found at: <http://www.cms.gov/medicare-coverage-database/search/advanced-search.aspx>. Additional indications may be covered at the discretion of the health plan.

Medicare Part B Covered Diagnosis Codes (applicable to existing NCD/LCD/LCA):

Jurisdiction(s): 6, K	NCD/LCD/Article Document (s): A54862
https://www.cms.gov/medicare-coverage-database/search/article-date-search.aspx?DocID=A54862&bc=gAAAAAAAAAAAAA ==	

Jurisdiction(s): J&M	NCD/LCD/Article Document (s): A56141
https://www.cms.gov/medicare-coverage-database/search/article-date-search.aspx?DocID=A56141&bc=gAAAAAAAAAAAAA	

Medicare Part B Administrative Contractor (MAC) Jurisdictions		
Jurisdiction	Applicable State/US Territory	Contractor
E (1)	CA, HI, NV, AS, GU, CNMI	Noridian Healthcare Solutions, LLC
F (2 & 3)	AK, WA, OR, ID, ND, SD, MT, WY, UT, AZ	Noridian Healthcare Solutions, LLC
5	KS, NE, IA, MO	Wisconsin Physicians Service Insurance Corp (WPS)

Medicare Part B Administrative Contractor (MAC) Jurisdictions		
Jurisdiction	Applicable State/US Territory	Contractor
6	MN, WI, IL	National Government Services, Inc. (NGS)
H (4 & 7)	LA, AR, MS, TX, OK, CO, NM	Novitas Solutions, Inc.
8	MI, IN	Wisconsin Physicians Service Insurance Corp (WPS)
N (9)	FL, PR, VI	First Coast Service Options, Inc.
J (10)	TN, GA, AL	Palmetto GBA, LLC
M (11)	NC, SC, WV, VA (excluding below)	Palmetto GBA, LLC
L (12)	DE, MD, PA, NJ, DC (includes Arlington & Fairfax counties and the city of Alexandria in VA)	Novitas Solutions, Inc.
K (13 & 14)	NY, CT, MA, RI, VT, ME, NH	National Government Services, Inc. (NGS)
15	KY, OH	CGS Administrators, LLC