

# Keytruda Qlex™ (pembrolizumab and berahyaluronidase alfa-pmph) (Subcutaneous)

-E-

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## I. Length of Authorization <sup>Δ 1</sup>

- Initial: Prior authorization validity will be provided initially for 6 months (180 days), unless otherwise specified.
  - Substitution/Switch-Therapy for Intravenous Pembrolizumab: Prior authorization validity will follow the same parameters for intravenous pembrolizumab based on the indication requested [see Keytruda IV-E policy Document Number: IC-0523]
  - Neoadjuvant therapy for TNBC: Prior authorization validity may be provided for up to a maximum of 24 weeks of therapy.\*
  - Neoadjuvant therapy for Head and Neck Squamous Cell Cancer (HNSCC): Prior authorization validity may be provided for up to a maximum of 6 weeks of therapy.
  - Neoadjuvant therapy for resectable NSCLC: Prior authorization validity may be provided for up to a maximum of 12 weeks of therapy
  - Neoadjuvant therapy for Urothelial Carcinoma: Prior authorization validity may be provided for up to a maximum of 9 weeks of therapy (3 doses).
- Renewal: Prior authorization validity may be renewed every 6 months (180 days) thereafter, unless otherwise specified.
  - Substitution/Switch-Therapy for Intravenous Pembrolizumab: Prior authorization validity will follow the same parameters for intravenous pembrolizumab based on the indication requested [see Keytruda IV-E policy Document Number: IC-0523]
  - Biliary Tract Cancer, Urothelial Carcinoma (excluding neoadjuvant/adjvant therapy), Cervical Cancer, cSCC, Endometrial Carcinoma, Esophageal Cancer, Gastric Cancer, HCC, MCC, MSI-H/dMMR Cancer, NSCLC (excluding neoadjuvant/adjvant therapy), RCC (excluding adjuvant therapy), HNSCC (excluding neoadjuvant/adjvant therapy), TMB-H Cancer, TNBC (excluding neoadjuvant/adjvant therapy), Ovarian Cancer, and Malignant Pleural Mesothelioma: Prior authorization validity may be renewed up to a maximum of 24 months of therapy.\*

- Neoadjuvant therapy for all the following: Urothelial Carcinoma, TNBC, NSCLC, Head and Neck Squamous Cell Cancer: Prior authorization validity may not be renewed.
- Adjuvant therapy for NSCLC (no prior neoadjuvant therapy), RCC, Head and Neck Squamous Cell Cancer, and Cutaneous Melanoma: Prior authorization validity may be renewed up to a maximum of 12 months of therapy.\*
- Adjuvant therapy for NSCLC (following neoadjuvant therapy): Prior authorization validity may be renewed for up to a maximum of 39 weeks of therapy.\*
- Adjuvant therapy for TNBC: Prior authorization validity may be renewed up to a maximum of 27 weeks of therapy.\*
- Adjuvant therapy in Urothelial Carcinoma: Prior authorization validity may be renewed for up to a maximum of 42 weeks of therapy\*

| <b><i>*Note: The maximum number of doses is dependent on the dosing frequency and duration of therapy. Refer to Section V for exact dosage.</i></b> |                                  |                                |
|-----------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------|--------------------------------|
| <b>Dosing Frequency</b>                                                                                                                             | <b>Maximum length of therapy</b> | <b>Maximum number of doses</b> |
| 3 weeks                                                                                                                                             | 24 weeks                         | 8 doses                        |
|                                                                                                                                                     | 27 weeks                         | 9 doses                        |
|                                                                                                                                                     | 42 weeks                         | 14 doses                       |
|                                                                                                                                                     | 1 year                           | 18 doses                       |
|                                                                                                                                                     | 2 years                          | 35 doses                       |
| 6 weeks                                                                                                                                             | 24 weeks                         | 4 doses                        |
|                                                                                                                                                     | 27 weeks                         | 5 doses                        |
|                                                                                                                                                     | 42 weeks                         | 7 doses                        |
|                                                                                                                                                     | 1 year                           | 9 doses                        |
|                                                                                                                                                     | 2 years                          | 18 doses                       |

## II. Dosing Limits

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

- 790 billable units every 6 weeks

## III. Initial Approval Criteria <sup>1</sup>

Prior authorization validity is provided in the following conditions:

- Member is at least 18 years of age, unless otherwise specified; **AND**

### Universal Criteria

- Member has not received previous therapy with a programmed death (PD-1/PD-L1)-directed therapy unless otherwise specified <sup>Δ</sup> (*Note: Not applicable when used as switch-therapy from intravenous pembrolizumab*); **AND**
- Therapy will not be used concomitantly with intravenous pembrolizumab; **AND**
- Therapy will not be used concurrently with intravenous chemotherapy agents (*not applicable when used for FDA approved combination therapy indicated with †*); **AND**
- Intravenous pembrolizumab must be used for members weighing <55 kg; **AND**

#### **Substitution/Switch-Therapy for Intravenous Pembrolizumab <sup>Δ</sup> ‡**

- Used as substitution for OR switch-therapy from intravenous pembrolizumab; **AND**
  - Member has previously met criteria for use of intravenous pembrolizumab [*see Keytruda IV-E policy Document Number: IC-0523*]; **AND**
    - Member has been receiving treatment with intravenous pembrolizumab and has shown a beneficial disease response and absence of unacceptable toxicity while on therapy; **OR**
  - Member currently meets criteria for use of intravenous pembrolizumab [*see Keytruda IV-E policy Document Number: IC-0523*]

#### **Biliary Tract Cancer † <sup>1</sup>**

- Used in combination with gemcitabine and cisplatin; **AND**
- Member has locally advanced unresectable or metastatic disease; **AND**
- Used as primary treatment

#### **Urothelial Carcinoma † <sup>1</sup>**

- Used in combination with enfortumab vedotin; **AND**
  - Member has locally advanced or metastatic urothelial carcinoma; **AND**
    - Used as first-line therapy; **OR**
  - Member has muscle invasive bladder cancer (MIBC); **AND**
    - Member is ineligible for cisplatin-containing chemotherapy\*; **AND**
    - Used as neoadjuvant treatment, and then continued after cystectomy as adjuvant treatment\*\*\*; **AND**
    - Member has stage II or IIIA disease; **OR**
- Used as a single agent; **AND**
  - Member has Bacillus Calmette-Guerin (BCG)-unresponsive\*\*, high-risk, non-muscle invasive bladder cancer (NMIBC) defined as one of the following:

- Persistent disease despite adequate BCG therapy
- Disease recurrence after an initial tumor free state following an adequate BCG course of therapy
- T1 disease following a single induction course of BCG therapy; **AND**
- Member has carcinoma in situ (CIS); **AND**
- Member is ineligible for or has elected not to undergo cystectomy; **OR**
- Member has locally advanced or metastatic urothelial carcinoma; **AND**
- Used for disease that progressed during or following platinum-containing chemotherapy or within 12 months of neoadjuvant or adjuvant treatment with a platinum containing chemotherapy; **OR**
- Used as first-line therapy for members who are not eligible for any platinum-containing chemotherapy (i.e., both cisplatin and carboplatin-ineligible\*)

**\* Note:** 10,71,79

- *Cisplatin-ineligible comorbidities may include the following: CrCl < 60 mL/min, ECOG PS ≥ 2 or KPS ≤ 70%, hearing loss of ≥ 25 decibels (dB) at two contiguous frequencies, grade ≥ 2 peripheral neuropathy, or NYHA Heart Failure class ≥ 3. Carboplatin may be substituted for cisplatin in the metastatic setting for cisplatin-ineligible members such as those with a GFR less than 60 mL/min.*
- *Platinum-ineligible comorbidities may include the following: CrCl < 30 mL/min, ECOG PS ≥ 3, grade ≥ 2 peripheral neuropathy, or NYHA Heart Failure class > 3, etc.*

**\*\* Adequate BCG therapy is defined as administration of at least five of six doses of an initial induction course AND at least two of three doses of maintenance therapy or at least two of six doses of a second induction course.**

**\*\*\* When used as adjuvant treatment for MIBC, enfortumab vedotin will be given in combination for six doses, and then the member will continue pembrolizumab for the remainder of the adjuvant treatment course.**

### Triple-Negative Breast Cancer (TNBC) † Ψ<sup>1</sup>

- Used as first-line therapy for locally recurrent unresectable or metastatic disease; **AND**
  - Used in combination with albumin-bound paclitaxel, paclitaxel, OR gemcitabine with carboplatin; **AND**
  - Tumor expresses PD-L1 (combined positive score [CPS] ≥10) as determined by an FDA-approved or Clinical Laboratory Improvement Amendments (CLIA)-compliant test❖; **OR**
- Member has high-risk early-stage disease; **AND**
  - Used as neoadjuvant therapy in combination with carboplatin and docetaxel; **AND**

- Use of pembrolizumab is restricted to members with a contraindication or intolerance to a neoadjuvant regimen including doxorubicin and/or cyclophosphamide; **OR**

- Used as adjuvant therapy as a single agent following use as neoadjuvant therapy in combination with chemotherapy

### **Cervical Cancer †<sup>1</sup>**

- Member has FIGO 2014 Stage III-IVA disease\*; **AND**
  - Used in combination with platinum-containing chemoradiotherapy (CRT); **OR**
- Tumor expresses PD-L1 (CPS ≥1) as determined by an FDA-approved or CLIA-compliant test❖; **AND**
  - Used as a single agent; **AND**
    - Member has recurrent or metastatic disease with disease progression on or after chemotherapy; **OR**
  - Used in combination with cisplatin or carboplatin AND paclitaxel (with or without bevacizumab); **AND**
    - Member has persistent, recurrent, or metastatic disease; **AND**
    - Disease is not amenable to curative treatment (i.e., surgery and/or radiation); **AND**
    - Used as first-line therapy

*\*FIGO 2014 Stage III-IVA disease is locally advanced cervical cancer involving the lower third of the vagina, with or without extension to pelvic sidewall, or hydronephrosis/non-functioning kidney, or spread to adjacent pelvic organs*

### **Esophageal Cancer †<sup>1</sup>**

- Member has locally advanced or metastatic esophageal or gastroesophageal junction (GEJ) (tumors with epicenter 1 to 5 cm above the GEJ) carcinoma; **AND**
  - Used as first-line therapy and tumor expresses PD-L1 (CPS ≥ 1) as determined by an FDA-approved or CLIA compliant test❖; **AND**
    - Disease is not amenable to surgical resection or definitive chemoradiation; **AND**
      - Member has HER2-positive adenocarcinoma; **AND**
        - Used in combination with trastuzumab AND fluorouracil or capecitabine AND oxaliplatin or cisplatin; **OR**
      - Member has HER2-negative adenocarcinoma; **AND**
        - Used in combination with oxaliplatin or cisplatin AND either fluorouracil or capecitabine; **OR**
      - Member has squamous cell carcinoma; **AND**

- Used in combination with oxaliplatin or cisplatin AND either fluorouracil or capecitabine; **OR**
- Tumor expresses PD-L1 (CPS  $\geq 10$ ) as determined by an FDA-approved or CLIA compliant test❖; **AND**
  - Used as a single agent after one or more prior lines of systemic therapy; **AND**
  - Used for tumors of squamous cell histology; **AND**
  - Members with HER2-positive disease must have previously received HER2-directed therapy (e.g., trastuzumab, etc.), unless contraindicated

### **Gastric Cancer †<sup>1</sup>**

- Member has locally advanced unresectable or metastatic gastric or gastroesophageal junction (GEJ) adenocarcinoma; **AND**
- Tumor expresses PD-L1 (CPS  $\geq 1$ ) as determined by an FDA-approved or CLIA compliant test❖; **AND**
- Used as first-line therapy; **AND**
  - Member has HER2-positive disease; **AND**
    - Used in combination with trastuzumab AND fluorouracil or capecitabine AND oxaliplatin or cisplatin; **OR**
  - Member has HER2-negative disease; **AND**
    - Used in combination with oxaliplatin or cisplatin AND either fluorouracil or capecitabine

### **Head and Neck Squamous Cell Cancer (HNSCC) †<sup>1</sup>**

- Used as neoadjuvant therapy, followed by adjuvant therapy for resectable locally advanced disease; **AND**
  - Tumor expresses PD-L1 (CPS  $\geq 1$ ) as determined by an FDA-approved or CLIA-compliant test❖; **AND**
    - Used as a single agent for neoadjuvant treatment; **OR**
    - Used as adjuvant treatment in combination with radiotherapy (with or without cisplatin) following neoadjuvant treatment, and then continued as a single agent; **OR**
- Used as first line therapy for metastatic or unresectable, recurrent disease; **AND**
  - Used in combination with platinum and fluorouracil-containing chemotherapy; **OR**
  - Used as a single agent; **AND**
    - Tumor expresses PD-L1 (CPS  $\geq 1$ ) as determined by an FDA-approved or CLIA-compliant test❖; **OR**
- Used as subsequent therapy for recurrent or metastatic disease; **AND**

- Used as a single agent; **AND**
- Member had disease progression on or after platinum-containing therapy; **AND**
- Tumor expresses PD-L1 (CPS  $\geq 1$ ) as determined by an FDA-approved or CLIA-compliant test❖

### **Hepatocellular Carcinoma (HCC) † <sup>1</sup>**

- Used as a single agent; **AND**
- Member does not have Child-Turcotte-Pugh (CTP) Class C liver disease; **AND**
- Disease is secondary to hepatitis B; **AND**
- Member has received prior systemic therapy other than a PD-1/PD-L1- containing regimen

### **Renal Cell Carcinoma (RCC) † <sup>1</sup>**

- Used as first line therapy for advanced disease; **AND**
  - Used in combination with axitinib; **AND**

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to pembrolizumab/lenvatinib; **OR**
  - Used in combination with lenvatinib; **OR**
- Used as adjuvant therapy; **AND**
  - Used as a single agent; **AND**
    - Member has intermediate-high or high risk of recurrence following nephrectomy; **OR**
    - Member has undergone nephrectomy and resection of metastatic lesions

### **Malignant Pleural Mesothelioma (MPM) † <sup>1</sup>**

- Used in combination with pemetrexed and either cisplatin or carboplatin; **AND**
- Used as first line therapy for unresectable advanced or metastatic disease

### **Cutaneous Melanoma † <sup>1</sup>**

- Used as a single agent for unresectable or metastatic disease; **OR**
- Used as a single agent for adjuvant treatment; **AND**
  - Member has stage IIB, IIC, or III melanoma following complete resection; **AND**
  - Member is at least 12 years of age

### **Merkel Cell Carcinoma (MCC) † <sup>1</sup>**

- Member is at least 12 years of age; **AND**

- Used as a single agent; **AND**
- Used as first-line therapy; **AND**
- Used for one of the following:
  - Member has recurrent locally advanced disease; **AND**
    - Both curative surgery and curative radiation therapy are not feasible; **OR**
  - Member has M1 disseminated disease; **AND**

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to retifanlimab

### Non-Small Cell Lung Cancer (NSCLC) †<sup>1</sup>

- Used for stage III disease; **AND**
  - Used as first-line therapy as a single agent in members who are not candidates for surgical resection or definitive chemoradiation; **AND**
  - Used in members with tumors expressing PD-L1 (TPS ≥1%) as determined by an FDA-approved or CLIA compliant test❖ and with no EGFR or ALK genomic tumor aberrations; **AND**

#### PD-L1 expression ≥50%:

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to cemiplimab; **OR**
- Used as neoadjuvant therapy, followed by adjuvant therapy; **AND**
    - Member has resectable disease (tumors ≥4 cm or node positive); **AND**
    - Used in combination with platinum-containing chemotherapy and then continued as a single agent after surgery; **OR**
  - Used as adjuvant therapy; **AND**
    - Used as a single agent; **AND**
    - Used following resection and previous adjuvant platinum-based chemotherapy; **AND**
    - Member has stage IB (T2a ≥4 cm), II, or IIIA disease; **OR**
  - Used for metastatic disease; **AND**
    - Used as first-line therapy; **AND**
      - Used in combination with pemetrexed AND either carboplatin or cisplatin for non-squamous cell histology with no EGFR or ALK genomic tumor aberrations; **AND**

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to cemiplimab/pemetrexed/(carboplatin or cisplatin); **OR**

- Used in combination with carboplatin AND either paclitaxel or albumin-bound paclitaxel for squamous cell histology; **AND**

➤ Use of pembrolizumab will be restricted to members with a contraindication or intolerance to cemiplimab/paclitaxel/(carboplatin or cisplatin); **OR**

- Used as a single agent; **AND**
  - Used in members with tumors expressing PD-L1 (TPS  $\geq 1\%$ ) as determined by an FDA-approved or CLIA compliant test❖; **AND**
  - Member has no EGFR or ALK genomic tumor aberrations; **AND**

PD-L1 expression  $\geq 50\%$ :

➤ Use of pembrolizumab will be restricted to members with a contraindication or intolerance to cemiplimab; **OR**

- Used as subsequent therapy as a single agent; **AND**
  - Used in members with tumors expressing PD-L1 (TPS  $\geq 1\%$ ) as determined by an FDA-approved or CLIA compliant test❖; **AND**
  - Member has disease progression on or after platinum-containing chemotherapy (*Note: Members with EGFR or ALK genomic tumor aberrations should have disease progression on FDA-approved therapy for these aberrations*)

### Cutaneous Squamous Cell Carcinoma (cSCC) † <sup>1</sup>

- Used as a single agent; **AND**
- Member has locally advanced, recurrent or metastatic disease that is not curable by surgery or radiation; **AND**

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to cemiplimab

### Endometrial Carcinoma † <sup>1</sup>

- Used in combination with lenvatinib; **AND**
  - Disease is mismatch repair proficient (pMMR) or NOT microsatellite instability-high (MSI-H) as determined by an FDA-approved or CLIA-compliant test❖; **AND**
  - Used as subsequent therapy for advanced disease; **AND**
  - Member has disease progression following prior platinum-based therapy in any setting and is not a candidate for curative surgery or radiation; **OR**
- Used in combination with carboplatin and paclitaxel, followed by single agent therapy; **AND**
  - Used for one of the following:

- Primary advanced disease (*excluding use in members with carcinosarcoma*)
- First-line therapy for recurrent disease (*excluding use in members with carcinosarcoma*);  
**AND**

Members with dMMR tumors ONLY:

- Use of pembrolizumab will be restricted to members with a contraindication or intolerance to durvalumab/carboplatin/paclitaxel

### **Ovarian Cancer †<sup>1</sup>**

- Member has epithelial ovarian, fallopian tube, or primary peritoneal carcinoma; **AND**
- Tumor expresses PD-L1 (CPS  $\geq 1$ ) as determined by an FDA-approved or CLIA compliant test❖; **AND**
- Member has platinum-resistant disease; **AND**
- Member has received one or two prior lines of systemic therapy; **AND**
- Used in combination with paclitaxel with or without bevacizumab

### **Microsatellite Instability-High (MSI-H) or Mismatch Repair Deficient (dMMR) Cancer †<sup>1</sup>**

- Used as a single agent; **AND**
- Member has microsatellite instability-high (MSI-H) or mismatch repair deficient (dMMR), as determined by an FDA-approved or CLIA compliant test❖; **AND**
  - Member has unresectable or metastatic solid tumors; **AND**
    - Used for disease progression following prior treatment; **AND**
    - Member is at least 12 years of age; **AND**
      - Member has Colorectal Cancer and was previously treated with a fluoropyrimidine AND either oxaliplatin or irinotecan, unless contraindicated; **OR**
      - Member has no satisfactory alternative treatment options; **AND**
  - Used as initial therapy for unresectable or metastatic colorectal cancer; **OR**
  - Used as subsequent therapy for advanced endometrial carcinoma; **AND**
    - Member has disease progression following prior systemic therapy in any setting and are not candidates for curative surgery or radiation

### **Tumor Mutational Burden-High (TMB-H) Cancer †<sup>1</sup>**

- Member is at least 12 years of age; **AND**
- Member has tumor mutational burden-high (TMB-H) [ $\geq 10$  mutations/megabase (mut/Mb)] disease, as determined by an FDA-approved or CLIA-compliant test❖; **AND**

- Used as a single agent; **AND**
- Pediatric members must not have a diagnosis of TMB-H central nervous system cancer; **AND**
- Member has unresectable or metastatic solid tumors; **AND**
- Used for disease progression following prior treatment and member has no satisfactory alternative treatment options

| <b>Ψ ER Scoring Interpretation</b> (following ER testing by validated IHC assay) <sup>116</sup>                                                                                                                                                                                                                                                                  |                       |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------|
| <b>Results</b>                                                                                                                                                                                                                                                                                                                                                   | <b>Interpretation</b> |
| – 0% – <1% of nuclei stain                                                                                                                                                                                                                                                                                                                                       | – ER-negative         |
| – 1%–10% of nuclei stain                                                                                                                                                                                                                                                                                                                                         | – ER-low–positive*    |
| – >10% of nuclei stain                                                                                                                                                                                                                                                                                                                                           | – ER-positive         |
| <p><i>*Note: Invasive cancers with between 1%–10% ER positivity are considered ER-low–positive. However, this group is noted to be heterogeneous and the biologic behavior of ER-low–positive cancers may be more similar to ER-negative cancers. This should be considered in decision making for other adjuvant therapy and overall treatment pathway.</i></p> |                       |

**Preferred therapies and recommendations are determined by review of clinical evidence. NCCN category of recommendation is taken into account as a component of this review. Regimens deemed equally efficacious (i.e., those having the same NCCN categorization) are considered to be therapeutically equivalent.**

#### **Enhanced Oncology Value (EOV) Program – Redacted indications**

Uses not listed above have inadequate data to support efficacy and are excluded from prior authorization validity.

Other treatment options including, but are not limited to, the following may be appropriate: radiation therapy, surgery, traditional chemotherapy (e.g., platinum, taxane), compassionate use/expanded access programs, clinical trials, supportive care, integrative and complementary therapies.

❖ If confirmed using an FDA approved assay – <http://www.fda.gov/companiondiagnostics>

† FDA Approved Indication(s); ‡ Compendia Recommended Indication(s); Ⓢ Orphan Drug

## IV. Renewal Criteria <sup>Δ 1</sup>

Prior authorization validity can be renewed based upon the following criteria:

- Member 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**
- Duration of authorization has not been exceeded (*refer to Section I*); **AND**
- Disease response with treatment as defined by stabilization of disease or decrease in size of tumor or tumor spread; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: immune-mediated adverse reactions (e.g., pneumonitis, hepatitis, colitis, endocrinopathies, nephritis/renal dysfunction, rash/dermatitis [including Stevens-Johnson syndrome (SJS), drug rash with eosinophilia and systemic symptoms (DRESS), and toxic epidermal necrolysis (TEN)], myocarditis, pericarditis, vasculitis, solid organ transplant rejection, etc.), severe administration-related reactions, complications of allogeneic hematopoietic stem cell transplantation (HSCT), etc.

### <sup>Δ</sup> Notes:

- Members responding to therapy who relapse  $\geq 6$  months after discontinuation due to duration (i.e., receipt of 24 months of therapy) are eligible to re-initiate PD-directed therapy.
- Members previously presenting with aggressive disease who are exhibiting stable disease on treatment as their best response (or if therapy improved performance status) may be eligible for continued therapy beyond the 24-month limit without interruption or discontinuation.
- Members who complete adjuvant therapy and progress  $\geq 6$  months after discontinuation are eligible to re-initiate PD-directed therapy for metastatic disease.
- Members whose tumors, upon re-biopsy, demonstrate a change in actionable mutation (e.g., MSS initial biopsy; MSI-H subsequent biopsy) may be eligible to re-initiate PD-directed therapy and will be evaluated on a case-by-case basis.

## V. Dosage/Administration <sup>Δ 1</sup>

| Indication                                                  | Dose                                                                                                                                                                                                                                                                                                                                                                                                           |
|-------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Substitution/Switch-Therapy for Intravenous Pembrolizumab ‡ | <ul style="list-style-type: none"><li>• Dose will be provided as one of the following:<ul style="list-style-type: none"><li>○ Every 3-week dosing: 395 mg/4,800 units</li><li>○ Every 6-week dosing: 790 mg/9,600 units; <b>AND</b></li></ul></li><li>• Dosing duration will follow the same indication-specific parameters for intravenous pembrolizumab [see Keytruda IV policy – Document Number:</li></ul> |

|                                      |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
|--------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
|                                      | <i>IC-0209 or Keytruda IV-E policy Document Number: IC-0523, as applicable]</i>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
| Urothelial Carcinoma †               | <p><b><u>Neoadjuvant therapy:</u></b><br/>395 mg/4,800 units every 3 weeks for 3 doses or until disease progression that precludes curative-intent cystectomy or unacceptable toxicity</p> <p><b><u>Adjuvant therapy:</u></b><br/>395 mg/4,800 units every 3 weeks for 14 doses or 790 mg/9,600 units every 6 weeks for 7 doses or until disease recurrence or unacceptable toxicity</p> <p><b><u>All other settings:</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p>                                                                                                                                                                                                                                                                                                                                                                           |
| Head and Neck Squamous Cell Cancer † | <p><b><u>Neoadjuvant therapy:</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks for 6 weeks or until disease progression that precludes definitive surgery or unacceptable toxicity</p> <p><b><u>Adjuvant therapy:</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 12 months in members without disease recurrence or unacceptable toxicity</p> <p><b><u>All other settings:</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p>                                                                                                                                                                                                                                                                                                                                 |
| NSCLC †                              | <p><b><u>Neoadjuvant followed by adjuvant treatment: †</u></b></p> <ul style="list-style-type: none"> <li>• <b><u>Neoadjuvant therapy:</u></b> 395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks for 12 weeks or until disease progression that precludes definitive surgery or unacceptable toxicity</li> <li>• <b><u>Adjuvant therapy:</u></b> 395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks for 39 weeks or until disease recurrence or unacceptable toxicity</li> </ul> <p><b><u>Adjuvant treatment (no prior neoadjuvant therapy):</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 12 months in members without disease recurrence or unacceptable toxicity</p> <p><b><u>All other settings:</u></b><br/>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p> |
| TNBC †                               | <b><u>Neoadjuvant therapy:</u></b>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |

|                         |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             |
|-------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
|                         | <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 24 weeks in members without disease progression or unacceptable toxicity (up to 8 doses of 395 mg/4,800 units every 3 weeks or 4 doses of 790 mg/9,600 units every 6 weeks)</p> <p><b><u>Adjuvant therapy*:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 27 weeks in members without disease recurrence or unacceptable toxicity (up to 9 doses of 395 mg/4,800 units every 3 weeks or 5 doses of 790 mg/9,600 units every 6 weeks)</p> <p><i>* Members who experience disease progression or unacceptable toxicity related to pembrolizumab with neoadjuvant treatment in combination with chemotherapy should not receive adjuvant single agent pembrolizumab.</i></p> <p><b><u>Locally recurrent unresectable or metastatic disease:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p> |
| Cutaneous Melanoma †    | <p><b><u>Adjuvant therapy:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 12 months in members without disease recurrence or unacceptable toxicity</p> <p><b><u>Unresectable or metastatic disease:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity</p>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      |
| Renal Cell Carcinoma †  | <p><b><u>Adjuvant therapy:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks up to a maximum of 12 months in members without disease recurrence or unacceptable toxicity</p> <p><b><u>First line therapy:</u></b></p> <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  |
| All other indications † | <p>395 mg/4,800 units every 3 weeks or 790 mg/9,600 units every 6 weeks until disease progression or unacceptable toxicity for up to 24 months</p>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          |

## VI. Billing Code/Availability Information

### HCPSC Code(s):

- J9277 – Injection, pembrolizumab, 1 mg and berahyaluronidase alfa-pmph; 1 billable unit = 1 mg

### NDC(s):

- Keytruda Qlex single-dose vial providing 395 mg pembrolizumab and 4,800 units berahyaluronidase alfa per 2.4 mL (165 mg/2,000 units per mL): 00006-3083-xx

### Medical Necessity Criteria

- Keytruda Qlex single-dose vial providing 790 mg pembrolizumab and 9,600 units berahyaluronidase alfa per 4.8 mL (165 mg/2,000 units per mL): 00006-5083-xx

## VII. References (STANDARD)

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## Appendix A – Non-Quantitative Treatment Limitations (NQTL) Factor Checklist

Non-quantitative treatment limitations (NQTLs) refer to the methods, guidelines, standards of evidence, or other conditions that can restrict how long or to what extent benefits are provided under a health plan. These may include things like utilization review or prior authorization. The utilization management NQTL applies comparably, and not more stringently, to mental health/substance use disorder (MH/SUD) Medical Benefit Prescription Drugs and medical/surgical (M/S) Medical Benefit Prescription Drugs. The table below lists the factors that were considered in designing and applying prior authorization to this drug/drug group, and a summary of the conclusions that Prime’s assessment led to for each.

| Factor                     | Conclusion            |
|----------------------------|-----------------------|
| Indication                 | Yes: Consider for PA  |
| Safety and efficacy        | No: PA not a priority |
| Potential for misuse/abuse | No: PA not a priority |

|              |                      |
|--------------|----------------------|
| Cost of drug | Yes: Consider for PA |
|--------------|----------------------|

## Appendix 1 – Covered Diagnosis Codes

| ICD-10 | ICD-10 Description                                                    |
|--------|-----------------------------------------------------------------------|
| 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                        |
| C00.9  | Malignant neoplasm of lip, unspecified                                |
| 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                                     |
| C05.2  | Malignant neoplasm of uvula                                           |
| C05.8  | Malignant neoplasm of overlapping sites of palate                     |
| C05.9  | Malignant neoplasm of palate, unspecified                             |
| 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       |

### Medical Necessity Criteria

| ICD-10 | ICD-10 Description                                                      |
|--------|-------------------------------------------------------------------------|
| C06.9  | Malignant neoplasm of mouth, unspecified                                |
| C07    | Malignant neoplasm of parotid gland                                     |
| C08.0  | Malignant neoplasm of submandibular gland                               |
| C08.1  | Malignant neoplasm of sublingual gland                                  |
| C08.9  | Malignant neoplasm of major salivary gland, 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                               |
| C10.0  | Malignant neoplasm of vallecula                                         |
| C10.1  | Malignant neoplasm of anterior surface of epiglottis                    |
| C10.2  | Malignant neoplasm of lateral wall of oropharynx                        |
| C10.3  | Malignant neoplasm of posterior wall of oropharynx                      |
| C10.4  | Malignant neoplasm of branchial cleft                                   |
| C10.8  | Malignant neoplasm of overlapping sites of oropharynx                   |
| C10.9  | Malignant neoplasm of oropharynx, 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 |
| 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                                            |
| C16.1  | Malignant neoplasm of fundus of stomach                                 |
| C16.2  | Malignant neoplasm of body of stomach                                   |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                     |
|--------|------------------------------------------------------------------------|
| C16.3  | Malignant neoplasm of pyloric antrum                                   |
| C16.4  | Malignant neoplasm of pylorus                                          |
| C16.5  | Malignant neoplasm of lesser curvature of stomach, unspecified         |
| C16.6  | Malignant neoplasm of greater curvature of stomach, unspecified        |
| C16.8  | Malignant neoplasm of overlapping sites of stomach                     |
| C16.9  | Malignant neoplasm of stomach, unspecified                             |
| 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                                                   |
| C22.1  | Intrahepatic bile duct carcinoma                                       |
| C22.8  | Malignant neoplasm of liver, primary, unspecified as to type           |
| C22.9  | Malignant neoplasm of liver, not specified as primary or secondary     |
| C23    | Malignant neoplasm of gallbladder                                      |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                       |
|--------|--------------------------------------------------------------------------|
| C24.0  | Malignant neoplasm of extrahepatic bile duct                             |
| C24.1  | Malignant neoplasm of ampulla of Vater                                   |
| C24.8  | Malignant neoplasm of overlapping sites of biliary tract                 |
| C24.9  | Malignant neoplasm of biliary tract, unspecified                         |
| C25.0  | Malignant neoplasm of head of pancreas                                   |
| C25.1  | Malignant neoplasm of body of the pancreas                               |
| C25.2  | Malignant neoplasm of tail of pancreas                                   |
| C25.3  | Malignant neoplasm of pancreatic duct                                    |
| C25.7  | Malignant neoplasm of other parts of pancreas                            |
| C25.8  | Malignant neoplasm of overlapping sites of pancreas                      |
| C25.9  | Malignant neoplasm of pancreas, unspecified                              |
| 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        |

#### Medical Necessity Criteria

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| ICD-10  | ICD-10 Description                                                                          |
|---------|---------------------------------------------------------------------------------------------|
| 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                             |
| C37     | Malignant neoplasm of thymus                                                                |
| C40.00  | Malignant neoplasm of scapula and long bones of unspecified upper limb                      |
| C40.01  | Malignant neoplasm of scapula and long bones of right upper limb                            |
| C40.02  | Malignant neoplasm of scapula and long bones of left upper limb                             |
| C40.10  | Malignant neoplasm of short bones of unspecified upper limb                                 |
| C40.11  | Malignant neoplasm of short bones of right upper limb                                       |
| C40.12  | Malignant neoplasm of short bones of left upper limb                                        |
| C40.20  | Malignant neoplasm of long bones of unspecified lower limb                                  |
| C40.21  | Malignant neoplasm of long bones of right lower limb                                        |
| C40.22  | Malignant neoplasm of long bones of left lower limb                                         |
| C40.30  | Malignant neoplasm of short bones of unspecified lower limb                                 |
| C40.31  | Malignant neoplasm of short bones of right lower limb                                       |
| C40.32  | Malignant neoplasm of short bones of left lower limb                                        |
| C40.80  | Malignant neoplasm of overlapping sites of bone and articular cartilage of unspecified limb |
| C40.81  | Malignant neoplasm of overlapping sites of bone and articular cartilage of right limb       |
| C40.82  | Malignant neoplasm of overlapping sites of bone and articular cartilage of left limb        |
| C40.90  | Malignant neoplasm of unspecified bones and articular cartilage of unspecified limb         |
| C40.91  | Malignant neoplasm of unspecified bones and articular cartilage of right limb               |
| C40.92  | Malignant neoplasm of unspecified bones and articular cartilage of left limb                |
| C41.0   | Malignant neoplasm of bones of skull and face                                               |
| C41.1   | Malignant neoplasm of mandible                                                              |
| C41.2   | Malignant neoplasm of vertebral column                                                      |
| C41.3   | Malignant neoplasm of ribs, sternum and clavicle                                            |
| C41.4   | Malignant neoplasm of pelvic bones, sacrum and coccyx                                       |
| C41.9   | Malignant neoplasm of bone and articular cartilage, unspecified                             |
| 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                                  |

#### Medical Necessity Criteria

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| ICD-10   | ICD-10 Description                                                              |
|----------|---------------------------------------------------------------------------------|
| 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                                         |
| C44.02   | Squamous cell carcinoma of skin of lip                                          |
| C44.121  | Squamous cell carcinoma of skin of unspecified eyelid, including canthus        |
| C44.1221 | Squamous cell carcinoma of skin of right upper eyelid, including canthus        |
| C44.1222 | Squamous cell carcinoma of skin of right lower eyelid, including canthus        |
| C44.1291 | Squamous cell carcinoma of skin of left upper eyelid, including canthus         |
| C44.1292 | Squamous cell carcinoma of skin of left lower eyelid, including canthus         |
| C44.221  | Squamous cell carcinoma of skin of unspecified ear and external auricular canal |
| C44.222  | Squamous cell carcinoma of skin of right ear and external auricular canal       |
| C44.229  | Squamous cell carcinoma of skin of left ear and external auricular canal        |
| C44.320  | Squamous cell carcinoma of skin of unspecified parts of face                    |
| C44.321  | Squamous cell carcinoma of skin of nose                                         |
| C44.329  | Squamous cell carcinoma of skin of other parts of face                          |
| C44.42   | Squamous cell carcinoma of skin of scalp and neck                               |
| C44.520  | Squamous cell carcinoma of anal skin                                            |

#### Medical Necessity Criteria

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| ICD-10  | ICD-10 Description                                                                    |
|---------|---------------------------------------------------------------------------------------|
| C44.521 | Squamous cell carcinoma of skin of breast                                             |
| C44.529 | Squamous cell carcinoma of skin of other part of trunk                                |
| C44.621 | Squamous cell carcinoma of skin of unspecified upper limb, including shoulder         |
| C44.622 | Squamous cell carcinoma of skin of right upper limb, including shoulder               |
| C44.629 | Squamous cell carcinoma of skin of left upper limb, including shoulder                |
| C44.721 | Squamous cell carcinoma of skin of unspecified lower limb, including hip              |
| C44.722 | Squamous cell carcinoma of skin of right lower limb, including hip                    |
| C44.729 | Squamous cell carcinoma of skin of left lower limb, including hip                     |
| C44.82  | Squamous cell carcinoma of overlapping sites of skin                                  |
| C44.92  | Squamous cell carcinoma of skin, unspecified                                          |
| C45.0   | Mesothelioma of pleura                                                                |
| C45.1   | Mesothelioma of peritoneum                                                            |
| C45.2   | Mesothelioma of pericardium                                                           |
| C45.7   | Mesothelioma of other sites                                                           |
| C45.9   | Mesothelioma, unspecified                                                             |
| C46.0   | Kaposi's sarcoma of skin                                                              |
| C46.1   | Kaposi's sarcoma of soft tissue                                                       |
| C46.2   | Kaposi's sarcoma of palate                                                            |
| C46.3   | Kaposi's sarcoma of lymph nodes                                                       |
| C46.4   | Kaposi's sarcoma of gastrointestinal sites                                            |
| C46.50  | Kaposi's sarcoma of unspecified lung                                                  |
| C46.51  | Kaposi's sarcoma of right lung                                                        |
| C46.52  | Kaposi's sarcoma of left lung                                                         |
| C46.7   | Kaposi's sarcoma of other sites                                                       |
| C46.9   | Kaposi's sarcoma, unspecified                                                         |
| C47.0   | Malignant neoplasm of peripheral nerves of head, face and neck                        |
| C47.10  | Malignant neoplasm of peripheral nerves of unspecified upper limb, including shoulder |
| C47.11  | Malignant neoplasm of peripheral nerves of right upper limb, including shoulder       |
| C47.12  | Malignant neoplasm of peripheral nerves of left upper limb, including shoulder        |
| C47.20  | Malignant neoplasm of peripheral nerves of unspecified lower limb, including hip      |
| C47.21  | Malignant neoplasm of peripheral nerves of right lower limb, including hip            |
| C47.22  | Malignant neoplasm of peripheral nerves of left lower limb, including hip             |
| C47.3   | Malignant neoplasm of peripheral nerves of thorax                                     |

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| ICD-10  | ICD-10 Description                                                                             |
|---------|------------------------------------------------------------------------------------------------|
| C47.4   | Malignant neoplasm of peripheral nerves of abdomen                                             |
| C47.5   | Malignant neoplasm of peripheral nerves of pelvis                                              |
| C47.6   | Malignant neoplasm of peripheral nerves of trunk, unspecified                                  |
| C47.8   | Malignant neoplasm of overlapping sites of peripheral nerves and autonomic nervous system      |
| C47.9   | Malignant neoplasm of peripheral nerves and autonomic nervous system, unspecified              |
| C48.0   | Malignant neoplasm of retroperitoneum                                                          |
| C48.1   | Malignant neoplasm of specified parts of peritoneum                                            |
| C48.2   | Malignant neoplasm of peritoneum, unspecified                                                  |
| C48.8   | Malignant neoplasm of overlapping sites of retroperitoneum and peritoneum                      |
| C49.0   | Malignant neoplasm of connective and soft tissue of head, face and neck                        |
| C49.10  | Malignant neoplasm of connective and soft tissue of unspecified upper limb, including shoulder |
| C49.11  | Malignant neoplasm of connective and soft tissue of right upper limb, including shoulder       |
| C49.12  | Malignant neoplasm of connective and soft tissue of left upper limb, including shoulder        |
| C49.20  | Malignant neoplasm of connective and soft tissue of unspecified lower limb, including hip      |
| C49.21  | Malignant neoplasm of connective and soft tissue of right lower limb, including hip            |
| C49.22  | Malignant neoplasm of connective and soft tissue of left lower limb, including hip             |
| C49.3   | Malignant neoplasm of connective and soft tissue of thorax                                     |
| C49.4   | Malignant neoplasm of connective and soft tissue of abdomen                                    |
| C49.5   | Malignant neoplasm of connective and soft tissue of pelvis                                     |
| C49.6   | Malignant neoplasm of connective and soft tissue of trunk, unspecified                         |
| C49.8   | Malignant neoplasm of overlapping sites of connective and soft tissue                          |
| C49.9   | Malignant neoplasm of connective and soft tissue, unspecified                                  |
| 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                                                                  |

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| ICD-10  | ICD-10 Description                                                      |
|---------|-------------------------------------------------------------------------|
| 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                                      |
| C50.011 | Malignant neoplasm of nipple and areola, right female breast            |
| C50.012 | Malignant neoplasm of nipple and areola, left female breast             |
| C50.019 | Malignant neoplasm of nipple and areola, unspecified female breast      |
| C50.021 | Malignant neoplasm of nipple and areola, right male breast              |
| C50.022 | Malignant neoplasm of nipple and areola, left male breast               |
| C50.029 | Malignant neoplasm of nipple and areola, unspecified male breast        |
| C50.111 | Malignant neoplasm of central portion of right female breast            |
| C50.112 | Malignant neoplasm of central portion of left female breast             |
| C50.119 | Malignant neoplasm of central portion of unspecified female breast      |
| C50.121 | Malignant neoplasm of central portion of right male breast              |
| C50.122 | Malignant neoplasm of central portion of left male breast               |
| C50.129 | Malignant neoplasm of central portion of unspecified male breast        |
| C50.211 | Malignant neoplasm of upper-inner quadrant of right female breast       |
| C50.212 | Malignant neoplasm of upper-inner quadrant of left female breast        |
| C50.219 | Malignant neoplasm of upper-inner quadrant of unspecified female breast |
| C50.221 | Malignant neoplasm of upper-inner quadrant of right male breast         |
| C50.222 | Malignant neoplasm of upper-inner quadrant of left male breast          |
| C50.229 | Malignant neoplasm of upper-inner quadrant of unspecified male breast   |
| C50.311 | Malignant neoplasm of lower-inner quadrant of right female breast       |
| C50.312 | Malignant neoplasm of lower-inner quadrant of left female breast        |

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| ICD-10  | ICD-10 Description                                                      |
|---------|-------------------------------------------------------------------------|
| C50.319 | Malignant neoplasm of lower-inner quadrant of unspecified female breast |
| C50.321 | Malignant neoplasm of lower-inner quadrant of right male breast         |
| C50.322 | Malignant neoplasm of lower-inner quadrant of left male breast          |
| C50.329 | Malignant neoplasm of lower-inner quadrant of unspecified male breast   |
| C50.411 | Malignant neoplasm of upper-outer quadrant of right female breast       |
| C50.412 | Malignant neoplasm of upper-outer quadrant of left female breast        |
| C50.419 | Malignant neoplasm of upper-outer quadrant of unspecified female breast |
| C50.421 | Malignant neoplasm of upper-outer quadrant of right male breast         |
| C50.422 | Malignant neoplasm of upper-outer quadrant of left male breast          |
| C50.429 | Malignant neoplasm of upper-outer quadrant of unspecified male breast   |
| C50.511 | Malignant neoplasm of lower-outer quadrant of right female breast       |
| C50.512 | Malignant neoplasm of lower-outer quadrant of left female breast        |
| C50.519 | Malignant neoplasm of lower-outer quadrant of unspecified female breast |
| C50.521 | Malignant neoplasm of lower-outer quadrant of right male breast         |
| C50.522 | Malignant neoplasm of lower-outer quadrant of left male breast          |
| C50.529 | Malignant neoplasm of lower-outer quadrant of unspecified male breast   |
| C50.611 | Malignant neoplasm of axillary tail of right female breast              |
| C50.612 | Malignant neoplasm of axillary tail of left female breast               |
| C50.619 | Malignant neoplasm of axillary tail of unspecified female breast        |
| C50.621 | Malignant neoplasm of axillary tail of right male breast                |
| C50.622 | Malignant neoplasm of axillary tail of left male breast                 |
| C50.629 | Malignant neoplasm of axillary tail of unspecified male breast          |
| C50.811 | Malignant neoplasm of overlapping sites of right female breast          |
| C50.812 | Malignant neoplasm of overlapping sites of left female breast           |
| C50.819 | Malignant neoplasm of overlapping sites of unspecified female breast    |
| C50.821 | Malignant neoplasm of overlapping sites of right male breast            |
| C50.822 | Malignant neoplasm of overlapping sites of left male breast             |
| C50.829 | Malignant neoplasm of overlapping sites of unspecified male breast      |
| C50.911 | Malignant neoplasm of unspecified site of right female breast           |
| C50.912 | Malignant neoplasm of unspecified site of left female breast            |
| C50.919 | Malignant neoplasm of unspecified site of unspecified female breast     |
| C50.921 | Malignant neoplasm of unspecified site of right male breast             |
| C50.922 | Malignant neoplasm of unspecified site of left male breast              |

#### Medical Necessity Criteria

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| ICD-10  | ICD-10 Description                                                |
|---------|-------------------------------------------------------------------|
| C50.929 | Malignant neoplasm of unspecified site of unspecified male breast |
| C50.A0  | Malignant inflammatory neoplasm of unspecified breast             |
| C50.A1  | Malignant inflammatory neoplasm of right breast                   |
| C50.A2  | Malignant inflammatory neoplasm of left breast                    |
| C51.0   | Malignant neoplasm of labium majus                                |
| C51.1   | Malignant neoplasm of labium minus                                |
| C51.2   | Malignant neoplasm of clitoris                                    |
| C51.8   | Malignant neoplasm of overlapping sites of vulva                  |
| C51.9   | Malignant neoplasm of vulva, unspecified                          |
| C52     | Malignant neoplasm of vagina                                      |
| C53.0   | Malignant neoplasm of endocervix                                  |
| C53.1   | Malignant neoplasm of exocervix                                   |
| C53.8   | Malignant neoplasm of overlapping sites of cervix uteri           |
| C53.9   | Malignant neoplasm of cervix uteri, unspecified                   |
| C54.0   | Malignant neoplasm of isthmus uteri                               |
| C54.1   | Malignant neoplasm of endometrium                                 |
| C54.2   | Malignant neoplasm of myometrium                                  |
| C54.3   | Malignant neoplasm of fundus uteri                                |
| C54.8   | Malignant neoplasm of overlapping sites of corpus uteri           |
| C54.9   | Malignant neoplasm of corpus uteri, unspecified                   |
| C55     | Malignant neoplasm of uterus, part unspecified                    |
| C56.1   | Malignant neoplasm of right ovary                                 |
| C56.2   | Malignant neoplasm of left ovary                                  |
| C56.3   | Malignant neoplasm of bilateral ovaries                           |
| C56.9   | Malignant neoplasm of unspecified ovary                           |
| C57.00  | Malignant neoplasm of unspecified fallopian tube                  |
| C57.01  | Malignant neoplasm of right fallopian tube                        |
| C57.02  | Malignant neoplasm of left fallopian tube                         |
| C57.10  | Malignant neoplasm of unspecified broad ligament                  |
| C57.11  | Malignant neoplasm of right broad ligament                        |
| C57.12  | Malignant neoplasm of left broad ligament                         |
| C57.20  | Malignant neoplasm of unspecified round ligament                  |
| C57.21  | Malignant neoplasm of right round ligament                        |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                               |
|--------|------------------------------------------------------------------|
| C57.22 | Malignant neoplasm of left round ligament                        |
| C57.3  | Malignant neoplasm of parametrium                                |
| C57.4  | Malignant neoplasm of uterine adnexa, unspecified                |
| C57.7  | Malignant neoplasm of other specified female genital organs      |
| C57.8  | Malignant neoplasm of overlapping sites of female genital organs |
| C57.9  | Malignant neoplasm of female genital organ, unspecified          |
| C60.0  | Malignant neoplasm of prepuce                                    |
| C60.1  | Malignant neoplasm of glans penis                                |
| C60.2  | Malignant neoplasm of body of penis                              |
| C60.8  | Malignant neoplasm of overlapping sites of penis                 |
| C60.9  | Malignant neoplasm of penis, unspecified                         |
| C61    | Malignant neoplasm of prostate                                   |
| C63.7  | Malignant neoplasm of other specified male genital organs        |
| C63.8  | Malignant neoplasm of overlapping sites of male genital organs   |
| 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                       |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                  |
|--------|---------------------------------------------------------------------|
| 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                                    |
| C71.0  | Malignant neoplasm of cerebrum, except lobes and ventricles         |
| C71.1  | Malignant neoplasm of frontal lobe                                  |
| C71.2  | Malignant neoplasm of temporal lobe                                 |
| C71.3  | Malignant neoplasm of parietal lobe                                 |
| C71.4  | Malignant neoplasm of occipital lobe                                |
| C71.5  | Malignant neoplasm of cerebral ventricle                            |
| C71.6  | Malignant neoplasm of cerebellum                                    |
| C71.7  | Malignant neoplasm of brain stem                                    |
| C71.8  | Malignant neoplasm of overlapping sites of brain                    |
| C71.9  | Malignant neoplasm of brain, unspecified                            |
| C72.0  | Malignant neoplasm of spinal cord                                   |
| C72.1  | Malignant neoplasm of cauda equina                                  |
| C72.9  | Malignant neoplasm of central nervous system, unspecified           |
| C73    | Malignant neoplasm of thyroid gland                                 |
| C74.00 | Malignant neoplasm of cortex of unspecified adrenal gland           |
| C74.01 | Malignant neoplasm of cortex of right adrenal gland                 |
| C74.02 | Malignant neoplasm of cortex of left adrenal gland                  |
| C74.90 | Malignant neoplasm of unspecified part of unspecified adrenal gland |
| C74.91 | Malignant neoplasm of unspecified part of right adrenal gland       |
| C74.92 | Malignant neoplasm of unspecified part of left adrenal gland        |
| C76.0  | Malignant neoplasm of head, face and neck                           |
| C78.00 | Secondary malignant neoplasm of unspecified lung                    |
| C78.01 | Secondary malignant neoplasm of right lung                          |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                                |
|--------|-----------------------------------------------------------------------------------|
| 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                  |
| C79.31 | Secondary malignant neoplasm of brain                                             |
| C79.70 | Secondary malignant neoplasm of unspecified adrenal gland                         |
| C79.71 | Secondary malignant neoplasm of right adrenal gland                               |
| C79.72 | Secondary malignant neoplasm of left adrenal gland                                |
| C7A.1  | Malignant poorly differentiated neuroendocrine tumors                             |
| C7A.8  | Other malignant neuroendocrine tumors                                             |
| C7B.00 | Secondary carcinoid tumors unspecified site                                       |
| C7B.01 | Secondary carcinoid tumors of distant lymph nodes                                 |
| C7B.02 | Secondary carcinoid tumors of liver                                               |
| C7B.03 | Secondary carcinoid tumors of bone                                                |
| C7B.04 | Secondary carcinoid tumors of peritoneum                                          |
| C7B.1  | Secondary Merkel cell carcinoma                                                   |
| C7B.8  | Other secondary neuroendocrine tumors                                             |
| 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          |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                                  |
|--------|-------------------------------------------------------------------------------------|
| 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                                                       |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                           |
|--------|------------------------------------------------------------------------------|
| 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, 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              |
| C83.00 | Small cell B-cell lymphoma, unspecified site                                 |
| C83.01 | Small cell B-cell lymphoma, lymph nodes of head, face, and neck              |
| C83.02 | Small cell B-cell lymphoma, intrathoracic lymph nodes                        |
| C83.03 | Small cell B-cell lymphoma, intra-abdominal lymph nodes                      |
| C83.04 | Small cell B-cell lymphoma, lymph nodes of axilla and upper limb             |
| C83.05 | Small cell B-cell lymphoma, lymph nodes of inguinal region and lower limb    |
| C83.06 | Small cell B-cell lymphoma, intrapelvic lymph nodes                          |
| C83.07 | Small cell B-cell lymphoma, spleen                                           |
| C83.08 | Small cell B-cell lymphoma, lymph nodes of multiple sites                    |
| C83.09 | Small cell B-cell lymphoma, extranodal and solid organ sites                 |
| C83.30 | Diffuse large B-cell lymphoma, unspecified site                              |
| C83.31 | Diffuse large B-cell lymphoma, lymph nodes of head, face, and neck           |
| C83.32 | Diffuse large B-cell lymphoma, intrathoracic lymph nodes                     |
| C83.33 | Diffuse large B-cell lymphoma, intra-abdominal lymph nodes                   |
| C83.34 | Diffuse large B-cell lymphoma, lymph nodes of axilla and upper limb          |
| C83.35 | Diffuse large B-cell lymphoma, lymph nodes of inguinal region and lower limb |
| C83.36 | Diffuse large B-cell lymphoma, intrapelvic lymph nodes                       |
| C83.37 | Diffuse large B-cell lymphoma, spleen                                        |
| C83.38 | Diffuse large B-cell lymphoma, lymph nodes of multiple sites                 |
| C83.39 | Diffuse large B-cell lymphoma, extranodal and solid organ sites              |
| C83.90 | Non-follicular (diffuse) lymphoma, unspecified, unspecified site             |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                                            |
|--------|-----------------------------------------------------------------------------------------------|
| C83.91 | Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of head, face, and neck           |
| C83.92 | Non-follicular (diffuse) lymphoma, unspecified, intrathoracic lymph nodes                     |
| C83.93 | Non-follicular (diffuse) lymphoma, unspecified, intra-abdominal lymph nodes                   |
| C83.94 | Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of axilla and upper limb          |
| C83.95 | Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of inguinal region and lower limb |
| C83.96 | Non-follicular (diffuse) lymphoma, unspecified, intrapelvic lymph nodes                       |
| C83.97 | Non-follicular (diffuse) lymphoma, unspecified, spleen                                        |
| C83.98 | Non-follicular (diffuse) lymphoma, unspecified, lymph nodes of multiple sites                 |
| C83.99 | Non-follicular (diffuse) lymphoma, unspecified, extranodal and solid organ sites              |
| C84.00 | Mycosis fungoides, unspecified site                                                           |
| C84.01 | Mycosis fungoides, lymph nodes of head, face, and neck                                        |
| C84.02 | Mycosis fungoides, intrathoracic lymph nodes                                                  |
| C84.03 | Mycosis fungoides, intra-abdominal lymph nodes                                                |
| C84.04 | Mycosis fungoides, lymph nodes of axilla and upper limb                                       |
| C84.05 | Mycosis fungoides, lymph nodes of inguinal region and lower limb                              |
| C84.06 | Mycosis fungoides, intrapelvic lymph nodes                                                    |
| C84.07 | Mycosis fungoides, spleen                                                                     |
| C84.08 | Mycosis fungoides, lymph nodes of multiple sites                                              |
| C84.09 | Mycosis fungoides, extranodal and solid organ sites                                           |
| C84.10 | Sézary disease, unspecified site                                                              |
| C84.11 | Sézary disease, lymph nodes of head, face, and neck                                           |
| C84.12 | Sézary disease, intrathoracic lymph nodes                                                     |
| C84.13 | Sézary disease, intra-abdominal lymph nodes                                                   |
| C84.14 | Sézary disease, lymph nodes of axilla and upper limb                                          |
| C84.15 | Sézary disease, lymph nodes of inguinal region and lower limb                                 |
| C84.16 | Sézary disease, intrapelvic lymph nodes                                                       |
| C84.17 | Sézary disease, spleen                                                                        |
| C84.18 | Sézary disease, lymph nodes of multiple sites                                                 |
| C84.19 | Sézary disease, extranodal and solid organ sites                                              |
| C84.90 | Mature T/NK-cell lymphomas, unspecified site                                                  |
| C84.91 | Mature T/NK-cell lymphomas, lymph nodes of head, face, and neck                               |
| C84.92 | Mature T/NK-cell lymphomas, intrathoracic lymph nodes                                         |
| C84.93 | Mature T/NK-cell lymphomas, intra-abdominal lymph nodes                                       |

#### Medical Necessity Criteria

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| ICD-10 | ICD-10 Description                                                                        |
|--------|-------------------------------------------------------------------------------------------|
| C84.94 | Mature T/NK-cell lymphomas, lymph nodes of axilla and upper limb                          |
| C84.95 | Mature T/NK-cell lymphomas, lymph nodes of inguinal region and lower limb                 |
| C84.96 | Mature T/NK-cell lymphomas, intrapelvic lymph nodes                                       |
| C84.97 | Mature T/NK-cell lymphomas, spleen                                                        |
| C84.98 | Mature T/NK-cell lymphomas, lymph nodes of multiple sites                                 |
| C84.99 | Mature T/NK-cell lymphomas, 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                        |
| C85.20 | Mediastinal (thymic) large B-cell lymphoma, unspecified site                              |
| C85.21 | Mediastinal (thymic) large B-cell lymphoma, lymph nodes of head, face and neck            |
| C85.22 | Mediastinal (thymic) large B-cell lymphoma, intrathoracic lymph nodes                     |
| C85.23 | Mediastinal (thymic) large B-cell lymphoma, intra-abdominal lymph nodes                   |
| C85.24 | Mediastinal (thymic) large B-cell lymphoma, lymph nodes of axilla and upper limb          |
| C85.25 | Mediastinal (thymic) large B-cell lymphoma, lymph nodes of inguinal region and lower limb |
| C85.26 | Mediastinal (thymic) large B-cell lymphoma, intrapelvic lymph nodes                       |
| C85.27 | Mediastinal (thymic) large B-cell lymphoma, spleen                                        |
| C85.28 | Mediastinal (thymic) large B-cell lymphoma, lymph nodes of multiple sites                 |
| C85.29 | Mediastinal (thymic) large B-cell lymphoma, extranodal and solid organ sites              |
| C22.3  | Extranodal NK/T-cell lymphoma, nasal type not having achieved remission                   |
| C86.60 | Primary cutaneous CD30-positive T-cell proliferations not having achieved remission       |
| C91.10 | Chronic lymphocytic leukemia of B-cell type not having achieved remission                 |
| C91.12 | Chronic lymphocytic leukemia of B-cell type in relapse                                    |
| D09.0  | Carcinoma in situ of bladder                                                              |
| D15.0  | Benign neoplasm of other and unspecified intrathoracic organs                             |
| D37.02 | Neoplasm of uncertain behavior of tongue                                                  |

#### Medical Necessity Criteria

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| ICD-10  | ICD-10 Description                                                                          |
|---------|---------------------------------------------------------------------------------------------|
| D37.031 | Neoplasm of uncertain behavior of the sublingual salivary glands                            |
| D37.032 | Neoplasm of uncertain behavior of the submandibular salivary glands                         |
| D37.039 | Neoplasm of uncertain behavior of the major salivary glands, unspecified                    |
| D37.04  | Neoplasm of uncertain behavior of the minor salivary glands                                 |
| D37.05  | Neoplasm of uncertain behavior of pharynx                                                   |
| D37.09  | Neoplasm of uncertain behavior of other specified sites of the oral cavity                  |
| D37.1   | Neoplasm of uncertain behavior of stomach                                                   |
| D37.3   | Neoplasm of uncertain behavior of appendix                                                  |
| 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                                                    |
| D38.4   | Neoplasm of uncertain behavior of thymus                                                    |
| D38.5   | Neoplasm of uncertain behavior of other respiratory organs                                  |
| D38.6   | Neoplasm of uncertain behavior of respiratory organ, unspecified                            |
| D48.60  | Neoplasm of uncertain behavior of unspecified breast                                        |
| D48.61  | Neoplasm of uncertain behavior of right breast                                              |
| D48.62  | Neoplasm of uncertain behavior of left breast                                               |
| Z85.00  | Personal history of malignant neoplasm of unspecified digestive organ                       |
| Z85.01  | Personal history of malignant neoplasm of esophagus                                         |
| Z85.028 | Personal history of other malignant neoplasm of stomach                                     |
| Z85.038 | Personal history of other malignant neoplasm of large intestine                             |
| Z85.068 | Personal history of other malignant neoplasm of small intestine                             |
| Z85.07  | Personal history of malignant neoplasm of pancreas                                          |
| Z85.09  | Personal history of malignant neoplasm of other digestive organs                            |
| Z85.118 | Personal history of other malignant neoplasm of bronchus and lung                           |
| Z85.12  | Personal history of malignant neoplasm of trachea                                           |
| Z85.22  | Personal history of malignant neoplasm of nasal cavities, middle ear, and accessory sinuses |
| Z85.238 | Personal history of other malignant neoplasm of thymus                                      |
| Z85.3   | Personal history of malignant neoplasm of breast                                            |
| Z85.42  | Personal history of malignant neoplasm of other parts of uterus                             |
| Z85.43  | Personal history of malignant neoplasm of ovary                                             |
| Z85.46  | Personal history of malignant neoplasm of prostate                                          |
| Z85.51  | Personal history of malignant neoplasm of bladder                                           |

#### Medical Necessity Criteria

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| ICD-10  | ICD-10 Description                                                                     |
|---------|----------------------------------------------------------------------------------------|
| 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 |
| Z85.820 | Personal history of malignant melanoma of skin                                         |
| Z85.821 | Personal history of Merkel cell carcinoma                                              |
| Z85.830 | Personal history of malignant neoplasm of bone                                         |
| Z85.831 | Personal history of malignant neoplasm of soft tissue                                  |
| Z85.841 | Personal history of malignant neoplasm of brain                                        |
| Z85.848 | Personal history of malignant neoplasm of other parts of nervous tissue                |
| Z85.850 | Personal history of malignant neoplasm of thyroid                                      |
| Z85.858 | Personal history of malignant neoplasm of other endocrine glands                       |

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

The preceding information is intended for non-Medicare coverage determinations. 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 Determinations (NCDs) and/or Local Coverage Determinations (LCDs) may exist and compliance with these policies is required where applicable. Local Coverage Articles (LCAs) may also exist for claims payment purposes or to clarify benefit eligibility under Part B for drugs which may be self-administered. The following link may be used to search for NCD, LCD, or LCA documents: <https://www.cms.gov/medicare-coverage-database/search.aspx>. Additional indications, including any preceding information, may be applied at the discretion of the health plan.

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

| 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) |
| 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) |

| Medicare Part B Administrative Contractor (MAC) Jurisdictions |                                                                                             |                                          |
|---------------------------------------------------------------|---------------------------------------------------------------------------------------------|------------------------------------------|
| Jurisdiction                                                  | Applicable State/US Territory                                                               | Contractor                               |
| N (9)                                                         | FL, PR, VI                                                                                  | First Coast Service Options, Inc.        |
| J (10)                                                        | TN, GA, AL                                                                                  | Palmetto GBA                             |
| M (11)                                                        | NC, SC, WV, VA (excluding below)                                                            | Palmetto GBA                             |
| 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                  |