Cigna Medical Coverage Policies – Radiology Oncology Imaging Guidelines

Effective February 01, 2024





Instructions for use

The following coverage policy applies to health benefit plans administered by Cigna. Coverage policies are intended to provide guidance in interpreting certain standard Cigna benefit plans and are used by medical directors and other health care professionals in making medical necessity and other coverage determinations. Please note the terms of a customer's particular benefit plan document may differ significantly from the standard benefit plans upon which these coverage policies are based. For example, a customer's benefit plan document may contain a specific exclusion related to a topic addressed in a coverage policy.

In the event of a conflict, a customer's benefit plan document always supersedes the information in the coverage policy. In the absence of federal or state coverage mandates, benefits are ultimately determined by the terms of the applicable benefit plan document. Coverage determinations in each specific instance require consideration of:

- 1. The terms of the applicable benefit plan document in effect on the date of service
- 2. Any applicable laws and regulations
- 3. Any relevant collateral source materials including coverage policies
- 4. The specific facts of the particular situation

Coverage policies relate exclusively to the administration of health benefit plans. Coverage policies are not recommendations for treatment and should never be used as treatment guidelines.

This evidence-based medical coverage policy has been developed by eviCore, Inc. Some information in this coverage policy may not apply to all benefit plans administered by Cigna.

These guidelines include procedures eviCore does not review for Cigna. Please refer to the <u>Cigna CPT</u> <u>code list</u> for the current list of high-tech imaging procedures that eviCore reviews for Cigna.

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Abbreviations for Oncology Imaging Guidelines

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Abbreviations for Oncology Imaging Guidelines		
ACTH	adrenocorticotropic hormone	
AFP	alpha-fetoprotein	
ALKP	alkaline phosphatase	
AP	anteroposterior	
betaHCG	beta human chorionic gonadotropin	
CA 125	cancer antigen 125 test	
CA 19-9	cancer antigen 19-9	
CA 15-3	cancer antigen 15-3	
CA 27-29	cancer antigen 27-29	
CBC	complete blood count	
CEA	carcinoembryonic antigen	
CNS	central nervous system	
CR	complete response	
СТА	computed tomography angiography	
DCIS	ductal carcinoma in situ	
DLBCL	diffuse large B cell lymphomas	
DRE	digital rectal exam	
EGD	esophagogastroduodenoscopy	
ENT	ear, nose, throat	
EOT	end of therapy	
ERCP	endoscopic retrograde cholangiopancreatography	
ESR	erythrocyte sedimentation rate	
EUA	exam under anesthesia	
EUS	endoscopic ultrasound	
FDG	fluorodeoxyglucose	
FNA	fine needle aspiration	

Abbreviations for Oncology Imaging Guidelines		
FUO	fever of unknown origin	
GE	gastroesophageal	
GI	gastrointestinal	
GU	genitourinary	
GTR	gross total resection	
HG	high-grade	
HIV	human immunodeficiency disease	
HRPC	hormone refractory prostate cancer	
hypermet	hypermetabolic	
IFRT	involved field radiation therapy	
inv	invasive	
LAR	low anterior resection	
LCIS	lobular carcinoma in situ	
LDH	lactate dehydrogenase	
LFT	liver function tests	
LND	lymph node dissection	
MALT	mucosa associated lymphoid tissue	
maint	maintenance	
MEN	multiple endocrine neoplasia	
MG	myasthenia gravis	
MGUS	monoclonal gammopathy of unknown significance	
MIBG	I-123 metaiodobenzylguanidine scintigraphy	
MRA	magnetic resonance angiography	
MRI	magnetic resonance imaging	
MUGA	'multiple gated acquisition' cardiac nuclear scan	
MWA	microwave ablation	
NaF	sodium fluoride	
NET	neuroendocrine tumor	
NCCN®	National Comprehensive Cancer Network	

Abbreviations for Oncology Imaging Guidelines		
NHL	non-Hodgkin's lymphoma	
NPC	nasopharyngeal carcinoma	
NSABP	National Surgical Adjuvant Breast and Bowel Project	
NSAIDS	nonsteroidal anti-inflammatory drugs	
NSCLC	non-small cell lung cancer	
NSGCT	non-seminomatous germ cell tumor	
PA	posteroanterior	
PCI	prophylactic cranial irradiation	
PET	positron emission tomography	
COG	Children's Oncology Group	
PSA	prostate specific antigen	
RFA	radiofrequency ablation	
RPLND	retroperitoneal lymph node dissection	
SqCCa	squamous cell carcinoma	
SCLC	small cell lung cancer	
SIADH	syndrome of inappropriate secretion of antidiuretic hormone	
TCC	transitional cell carcinoma	
TLH	total laparoscopic hysterectomy	
TNM	tumor node metastasis staging system	
TSH	thyroid-stimulating hormone	
TURBT	trans-urethral resection of bladder tumor	
VIPoma	vasoactive intestinal polypeptide	
WLE	wide local incision	
WB-MRI	whole body MRI	
WM	Waldenstrom's macroglobulinemia	
WBXRT	whole brain radiation therapy	

General Guidelines (ONC-1.0)

ON.GG.0001.0.C

- A recent clinical evaluation (within 60 days) or meaningful contact (telephone call, electronic mail or messaging) should be performed prior to considering advanced imaging, unless the individual is undergoing guideline-supported scheduled off therapy surveillance evaluation or cancer screening. The clinical evaluation may include a relevant history and physical examination, including biopsy, appropriate laboratory studies, and results of non-advanced or advanced imaging modalities.
- Unless otherwise stated in the disease-specific guideline, a histological confirmation of malignancy (or recurrence) and the stage of disease is required to perform a medical necessity review of the requested imaging.
- Generally, the studies listed in the disease-specific sections reflect the studies supported by current literature and research for that condition. If a study is not listed, then it is not supported.
- Routine imaging of brain, spine, neck, chest, abdomen, pelvis, bones, or other body areas is not indicated except where explicitly stated in a diagnosis-specific guideline section, or if one of the following applies:
 - o Known prior disease involving the requested body area
 - New or worsening symptoms or physical exam findings involving the requested body area (including non-specific findings such as ascites or pleural effusion)
 - o New finding on basic imaging study such as plain x-ray or ultrasound
 - New finding on adjacent body area CT/MRI study (i.e., pleural effusion observed on CT abdomen)
- Unless otherwise stated in the disease-specific guideline, advanced imaging of asymptomatic individuals is not routinely supported without signs or symptoms of systemic involvement of cancer.
- Repeat imaging studies are not generally necessary unless there is evidence of disease progression, recurrence of disease, and/or the repeat imaging will affect an individual's clinical management.
- Conventional imaging performed prior to diagnosis should not be repeated unless there is a delay of at least 6 weeks since previous imaging and treatment initiation or there are new or significantly worsening clinical signs or symptoms

Phase	Imaging Timeframe	
After definitive local therapy of primary tumor (surgery or radiation therapy)	Follow surveillance guidelines	
During adjuvant chemotherapy	Follow surveillance guidelines	
After ablative therapy	See disease-specific guidelines	
During chemotherapy or immunotherapy for measurable disease	Every 2 cycles (generally every 6 to 8 weeks)	

Phase	Imaging Timeframe
During endocrine/hormonal therapy for measurable disease	Every 3 months (12 weeks)
Measurable metastatic disease being monitored off therapy	Every 3 months (12 weeks)
Minimal metastatic disease on maintenance therapy	Every 3 months (12 weeks)
Surveillance for history of metastatic disease with complete response and being observed off-therapy	Imaging typically not indicated beyond 5 years from completion of treatment for metastatic disease

- Advanced imaging is not indicated for evaluation of in situ or non-invasive cancers or cancer surveillance after complete surgical removal of primary disease unless otherwise stated in the cancer-specific guidelines.
- Advanced imaging is not indicated for monitoring disease in individuals who choose
 to not receive standard oncologic therapy, but may be receiving alternative
 therapies or palliative care and/or hospice. All advanced imaging indicated for initial
 staging of the specific cancer type can be approved once when the individual is
 considering initiation of a standard therapeutic approach (surgery, chemotherapy, or
 radiation therapy).
- Brain imaging is performed for signs or symptoms of brain disease
 - MRI Brain without and with contrast (CPT® 70553) is the recommended study for evaluation of suspected or known brain metastases. If a non-contrast CT head shows suspicious lesion, MRI brain may be obtained to further characterize the lesion
 - CT without and with contrast (CPT® 70470) can be approved when MRI is contraindicated or not available, or if there is skull bone involvement
 - Certain malignancies including, but not limited to melanoma and lung cancer have indications for brain imaging for asymptomatic individuals
 - If stage IV disease is demonstrated elsewhere or if systemic disease progression is noted, refer to disease specific guidelines
 - Initiation of angiogenesis therapy is not an indication for advanced imaging of the brain in asymptomatic individuals (Avastin/Bevacizumab; < 3% risk of bleeding and < 1% risk of serious bleeding)

Bone Scan:

- Primarily used for evaluation of bone metastases in individuals with solid malignancies.
- Indications for bone scan in individuals with history of malignancy include bone pain, rising tumor markers, elevated alkaline phosphatase or in individuals with primary bone tumor.
- For evaluation of suspected or known bony metastases, CPT[®] 78306 (Nuclear bone scan whole body), may be approved.

- Radiopharmaceutical Localization scan SPECT (CPT® 78803 or CPT® 78831) or SPECT/CT (CPT® 78830 or CPT® 78832) may be approved as an add-on test for further evaluation of a specific area of interest.
- CPT® codes 78300 (Nuclear bone scan limited), 78305 (Nuclear bone scan multiple areas) or 78315 do not have any indications in oncology nuclear medicine imaging.
- Bone scan supplemented by plain x-rays are the initial imaging modalities for suspected malignant bone pain. For specific imaging indications, see also:
 - o Bone (including Vertebral) Metastases (ONC-31.5)
 - o Spinal Cord Compression (ONC-31.6)
 - Carcinoma of Unknown Primary Site (ONC-31.7)
- Advanced imaging used for radiation therapy treatment planning should not be authorized using any of the diagnostic imaging codes for CT, MRI, or PET
 - In the absence of written payor guidelines, advanced imaging performed in support of radiation therapy treatment planning should be reported with CPT[®] 76498 for Unlisted MRI or CPT[®] 76497 for Unlisted CT scan
- Delay PET/CT for at least 12 weeks after completion of radiation treatment, unless required sooner for imminent surgical resection.
- PET/CT may be considered prior to biopsy in order to determine a more favorable site for biopsy when a prior biopsy was nondiagnostic or a relatively inaccessible site is contemplated which would require invasive surgical intervention for biopsy attempt.
- PET/CT may be indicated if:
 - Conventional imaging (CT, MRI or bone scan) reveals findings that are inconclusive or negative, with continued suspicion for recurrence
- Unless specified in diagnosis-specific guideline section PET/CT Imaging is NOT indicated for:
 - Infection, inflammation, trauma, post-operative healing, granulomatous disease, rheumatological conditions
 - Concomitantly with separate diagnostic CT studies
 - Conclusive evidence of distant or diffuse metastatic disease on recent conventional imaging studies
 - o Metastatic disease in the central nervous system (CNS)
 - Lesions less than 8 mm in size
 - Follow up after localized therapy (i.e. radiofrequency ablation, embolization, stereotactic radiation, etc.)
 - Rare malignancies, due to lack of available evidence regarding the diagnostic accuracy of PET in rare cancers
 - Surveillance
 - Serial monitoring of individuals who are not currently receiving anti-tumor treatment or are receiving maintenance treatment

- Serial monitoring of FDG avidity until resolution.
- PET/CT avidity in a residual mass at the end of planned therapy is not an indication for PET/CT imaging during surveillance.
- Residual mass that has not changed in size since the last conventional imaging does not justify PET imaging
- Unless otherwise specified for a specific cancer type, once PET has been documented to be negative for a given individual's cancer or all PET-avid disease has been surgically resected, PET should not be used for continued disease monitoring or surveillance.
- PET/MRI is generally not supported by eviCore for imaging adults with a vast majority of oncologic conditions due to lack of standardization in imaging technique and interpretation.
- However, it may be indicated in certain pediatric oncologic conditions. See: <u>PET Imaging in Pediatric Oncology (PEDONC-1.4)</u> for indications.
- The specific radiotracer planned to be used with PET/CT imaging is required to perform a medical necessity review. Indications for PET/CT imaging using non-FDG radiotracers are listed in diagnosis-specific guidelines.
 - Supported radiotracers:
 - ¹⁸F-FDG
 - 68Ga-DOTATATE (NETSPOT®) for low-grade neuroendocrine tumors and medullary thyroid cancer
 - 64Cu-DOTATATE (DETECTNET®) for low-grade neuroendocrine tumors
 - 68Ga-DOTA-TOC for low-grade neuroendocrine tumors
 - ¹¹C Choline for prostate cancer
 - 18F-Fluciclovine (AXUMIN®) for prostate cancer
 - ⁶⁸Ga PSMA-11 for prostate cancer
 - ¹⁸F Piflufolastat (Pylarify®) for prostate cancer
 - ⁶⁸Ga Gozetotide (Illuccix® and Locametz®) for prostate cancer
 - 18F Flotufolastat (Posluma®) for prostate cancer
 - ¹⁸F-Na Fluoride PET bone scan
 - Unsupported radiotracers:
 - ¹⁸F Fluoroestradiol
 - PET/CT imaging using isotopes other than those specified above
- Octreotide scan:
 - Specific for low and intermediate grade neuroendocrine tumors which express specific cell surface somatostatin receptors. See cancer specific guidelines for recommended use.
 - One of the following codes may be approved when Octreotide scan is requested:
 - CPT® 78802 (Radiopharmaceutical localization of tumor whole-body single day study)

- CPT® 78804 (Radiopharmaceutical localization of tumor whole-body two or more days)
- In addition to one of the above CPT codes, CPT® 78803 (Radiopharmaceutical localization of tumor SPECT), SPECT CPT® 78831, or hybrid SPECT/CT (CPT® 78830 or 78832) may be approved as an add-on test for further evaluation of a specific area of interest.

Clinical Trials

- Similar to investigational and experimental studies, clinical trial imaging requests will be considered to determine whether they meet Health Plan coverage and eviCore's evidence-based guidelines.
- Imaging studies which are inconsistent with established clinical standards, or are requested for data collection and not used in direct clinical management are not supported.

Key Principles (ONC-1.1)

ON.GG.001.1.C

AGE APPROPRIATE GUIDELINES		
Age of Individual	Appropriate Imaging Guidelines	
≥18 years old at initial diagnosis	General Oncology Imaging Guidelines, except where directed otherwise by a specific guideline section	
<18 years old at initial diagnosis	Pediatric and Special Populations Oncology Imaging Guidelines, except where directed otherwise by a specific guideline section	
15 to 39 years old at initial diagnosis (defined as Adolescent and Young Adult (AYA) oncology individuals)	 When unique guidelines for a specific cancer type exist only in either General Oncology or Pediatric and Special Populations Oncology, AYA individuals should be imaged according to the guideline section for their specific cancer type, regardless of the individual's age When unique guidelines for a specific cancer type exist in both General Oncology and Pediatric and Special Populations Oncology, AYA individuals should be imaged according to the age rule in the previous bullet 	

- Brain imaging is performed for signs or symptoms of brain disease:
 - MRI Brain without and with contrast (CPT® 70553) is the recommended study for evaluation of suspected or known brain metastases.
 - MRI Brain without and with contrast (CPT® 70553) may be obtained if a noncontrast CT Head shows suspicious lesion.
 - o CT Head without and with contrast (CPT® 70470) can be approved when MRI is contraindicated or not available, or if there is skull bone involvement.
 - Initiation of angiogenesis therapy is not an indication for advanced imaging of the brain in asymptomatic individuals (Avastin/bevacizumab; <3% risk of bleeding and <1% risk of serious bleeding).
- Individuals receiving cardiotoxic chemotherapy (such as doxorubicin, trastuzumab, pertuzumab, mitoxantrone, etc.) may undergo cardiac evaluation - at baseline and for monitoring while on active therapy.

- Echocardiography (CPT® 93306, CPT® 93307, or CPT® 93308) rather than MUGA scan for determination of LVEF and/or wall motion
 - MUGA Scan may be performed instead of ECHO in individuals who have a low LV ejection fraction of <50% on a prior ECHO or MUGA, pre-existing left ventricular wall motion abnormalities from ischemic or non-ischemic cardiomyopathies, congestive heart failure or when ECHO is technically limited and prevents accurate assessment of LV function.
 - A prior MUGA is not a reason to approve another MUGA (it is not necessary to compare LVEF by the same modality)
- The timeframe for monitoring the ejection fraction should be determined by the provider but no more often than baseline and at every 6 weeks.
- May repeat every 4 weeks if cardiotoxic chemotherapeutic drug is withheld for significant left ventricular cardiac dysfunction.
- See: <u>Oncologic Indications for Cancer Therapeutics-Related Cardiac</u>
 <u>Dysfunction (CTRCD) (CD-12.1)</u> in the Cardiology Imaging Guidelines
- CTA or MRA of a specific anatomic region is indicated when requested for surgical planning when there is suspected vascular proximity to proposed resection margin
- Adults (≥18 years) with a diagnosis of Li-Fraumeni Syndrome (LFS) may be screened for malignancy with a Whole-Body MRI (CPT® 76498) on an annual basis. Annual Brain MRI (CPT® 70553) may be performed as part of Whole-Body MRI or as a separate exam. Due to lack of standardization of technique, interpretation, and availability of Whole-Body MRI, individuals with LFS are encouraged to participate in clinical trials.

Use of Contrast

- CT imaging should be performed with contrast for known or suspected body regions, unless contraindicated.
 - Shellfish allergy is not a contraindication to contrast
 - For iodinated contrast dye allergy, either CT scans without contrast or MRI scan without and with contrast are indicated.
 - If CT scanning is considered strongly indicated in an individual with known contrast allergy, CT with contrast may be considered to be safely performed following prednisone premedication over a 24-hour period prior to the study.
- Severe renal insufficiency, i.e. an eGFR less than 30, is a contraindication for an MRI using a gadolinium-based contrast agent (GBCA). In individuals with eGFR greater than 40, GBCA administration can be safely performed. GBCA administered to individuals with acute kidney injury or severe chronic kidney disease can result in a syndrome of nephrogenic systemic fibrosis (NSF), but GBCAs are not considered nephrotoxic at dosages approved for MRI.
- Gadolinium deposition has been found in individuals with normal renal function following the use of gadolinium based contrast agents (GBCAs).
 - The U.S. Food and Drug Administration (FDA) is investigating the risk of brain deposits following repeated use of GBCAs.

- The FDA has noted that, "It is unknown whether these gadolinium deposits are harmful or can lead to adverse health effects." and have recommended:
 - To reduce the potential for gadolinium accumulation, health care professionals should consider limiting GBCA use to clinical circumstances in which the additional information provided by the contrast is necessary.
 - Health care professionals are also urged to reassess the necessity of repetitive GBCA MRIs in established treatment protocols

Radiation Exposure

The use of MRI in place of CT scans to reduce risk of secondary malignancy from radiation exposure during CT is not supported by the peer-reviewed literature. Unless otherwise specified in the Guidelines, MRI in place of CT scans for this purpose alone is not indicated. In some instances (i.e., testicular cancer surveillance), MRI may be considered inferior to CT scans.

Phases of Oncology Imaging and General Phase-Related Considerations (ONC-1.2)

ON.GG.0001.2.A

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Phases of Oncology Imaging	Definition	
Screening	Imaging requested for individuals at increased risk for a particular cancer in the absence of known clinical signs or symptoms	
Suspected Diagnosis	Imaging requested to evaluate a suspicion of cancer, prior to histological confirmation	
Initial work-up and Staging	Imaging requested after biopsy confirmation and prior to starting specific treatment	
Treatment response or Interim Restaging	Imaging performed during active treatment with chemotherapy, targeted therapy, immunotherapy, or endocrine therapy	
Restaging of locally treated lesions	Imaging performed to evaluate primary or metastatic lesions with ablation using cryoablation, radiofrequency, radioactive isotope, microwave or chemotherapy	
Restaging / Suspected Recurrence	Imaging requested when there is suspicion for progression or recurrence of known cancer based on clinical signs/symptoms, laboratory tests or basic imaging studies	
Surveillance	 Imaging performed in individuals who: Are asymptomatic or have chronic stable symptoms, and Have no clinical suspicion of change in disease status, and Are not receiving active anti-tumor treatment or are receiving maintenance treatment 	

General phase-related considerations

 Conventional imaging performed prior to diagnosis should not be repeated unless there is a delay of at least 6 weeks since previous imaging and treatment initiation or there are new or significantly worsening clinical signs or symptoms

Phase	Imaging Timeframe	
After definitive local therapy of primary tumor (surgery or radiation therapy)	Follow surveillance guidelines	

Phase	Imaging Timeframe
During adjuvant chemotherapy or endocrine therapy	Follow surveillance guidelines
After ablative therapy	See disease-specific guidelines
During chemotherapy or immunotherapy for measurable disease	Every 2 cycles (generally every 6 to 8 weeks)
During endocrine/hormonal therapy for measurable disease	Every 3 months (12 weeks)
Metastatic disease on maintenance therapy	Every 3 months (12 weeks)
Measurable metastatic disease being monitored off therapy	Every 3 months (12 weeks) for up to 5 years after completion of treatment for metastatic disease

PET Imaging in Oncology (ONC-1.4)

ON.GG.0001.4.C

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CPT codes:

- PET Imaging in oncology should use PET/CT fusion (CPT® 78815 or CPT® 78816). Unbundling PET/CT imaging into separate PET and diagnostic CT codes is otherwise not supported
- "Limited area" protocol is done infrequently, but may be considered, and is reported with PET (CPT® 78811) or for PET/CT (CPT® 78814).

Radiotracers:

- Unless specified otherwise, the term "PET" refers to ¹⁸F-FDG-PET and PET/CT fusion studies
- Indications for PET/CT imaging using non-FDG radiotracers are listed in diagnosis-specific guidelines. The indications may be as follows:

Supported radiotracers:

- o ¹⁸F-FDG
- 68Ga-DOTATATE (NETSPOT®) for low-grade neuroendocrine tumors and medullary thyroid cancer
- o 64Cu-DOTATATE (DETECTNET®) for low-grade neuroendocrine tumors
- o 68Ga-DOTA-TOC for low-grade neuroendocrine tumors
- o ¹¹C Choline for prostate cancer
- o ¹⁸F-Fluciclovine (AXUMIN®) for prostate cancer
- o 68Ga PSMA-11 for prostate cancer
- o ¹⁸F Piflufolastat (Pylarify®) for prostate cancer
- o 68Ga Gozetotide (Illuccix® and Locametz®) for prostate cancer
- o ¹⁸F Flotufolastat (Posluma®) for prostate cancer
- o ¹⁸F-Na Fluoride PET bone scan

· Unsupported radiotracers:

- o ¹⁸F Fluoroestradiol
- o PET/CT imaging using isotopes other than those specified above

CPT/ HCPCS Code	Code Description	Brand or common name	Guideline Section and Cancer Type
A9552	¹⁸ F Fluoro deoxyglucose	FDG	Various guideline sections where PET is indicated
A9580	¹⁸ F Sodium fluoride	N/A	ONC-1, ONC-11: Breast Cancer

CPT/ HCPCS Code	Code Description	Brand or common name	Guideline Section and Cancer Type
A9587	⁶⁸ Ga-68 Dotatate	NETSPOT®	ONC-15: Low-grade neuroendocrine tumors, ONC-6: Medullary thyroid cancer
A9515	¹¹ C Choline	N/A	ONC-19, Prostate Cancer
A9588	¹⁸ F-Fluciclovine	AXUMIN®	ONC-19, Prostate Cancer
A9593	⁶⁸ Ga PSMA-11	N/A	ONC-19, Prostate Cancer
A9594			
A9595	¹⁸ F Piflufolastat	Pylarify®	ONC-19, Prostate Cancer
A9596	⁶⁸ Ga Gozetotide	Illuccix®	ONC-19, Prostate Cancer
A9800	⁶⁸ Ga Gozetotide	Locametz®	ONC-19, Prostate Cancer
A9597	¹⁸ F Flotufolastat	Posluma®	ONC-19, Prostate Cancer
A9591	¹⁸ F Fluoroestradiol	Cerianna®	ONC-1
A9592	⁶⁴ Cu Copper dotatate	Detectnet®	ONC-15, Low-grade neuroendocrine tumors
C9067	⁶⁸ Ga Gallium-DOTA-TOC	N/A	ONC-15, Low-grade neuroendocrine tumors

- Unless specified in diagnosis-specific guideline section PET/CT Imaging is not indicated for:
 - Infection, inflammation, trauma, post-operative healing, granulomatous disease, rheumatologic conditions
 - o Concomitantly with separate diagnostic CT studies
 - Conclusive evidence of distant or diffuse metastatic disease on recent conventional imaging studies
 - Metastatic disease in the central nervous system (CNS)
 - Lesions less than 8 mm in size
 - Follow up after localized therapy (i.e. radiofrequency ablation, embolization, stereotactic radiation, etc.)
 - Rare malignancies, due to lack of available evidence regarding the diagnostic accuracy of PET in rare cancers
 - o Surveillance:
 - Serial monitoring of individuals who are not currently receiving anti-tumor treatment or are receiving maintenance treatment
 - Serial monitoring of FDG avidity until resolution.

- PET/CT avidity in a residual mass at the end of planned therapy is not an indication for PET/CT imaging during surveillance.
- Residual mass that has not changed in size since the last conventional imaging does not justify PET imaging
- Unless otherwise specified for a specific cancer type, once PET has been documented to be negative for a given individual's cancer or all PET-avid disease has been surgically resected, PET should not be used for continued disease monitoring or surveillance.
- PET/CT may be indicated if:
 - Conventional imaging (CT, MRI or bone scan) reveals findings that are inconclusive or negative, with continued suspicion for recurrence
 - The individual is undergoing salvage treatment for a recurrent solid tumor with residual measurable disease on conventional imaging and confirmed repeat negative PET imaging will allow the individual to transition from active treatment to surveillance
 - PET/CT may be considered prior to biopsy in order to determine a more favorable site for biopsy when a prior biopsy was nondiagnostic or a relatively inaccessible site is contemplated which would require invasive surgical intervention for biopsy attempt
- PET/CT for rare malignancies is not covered by eviCore guidelines due to lack of available evidence regarding diagnostic accuracy of PET/CT in the majority of rare cancers. Conventional imaging studies should be used for initial staging and treatment response for these diagnoses. PET/CT can be approved if all of the following apply:
 - Conventional imaging (CT, MRI or bone scan) reveals equivocal or suspicious findings
 - No other specific metabolic imaging (MIBG, octreotide, technetium, etc.) is appropriate for the disease type
 - The submitted clinical information describes a specific decision regarding the individual's care that will be made based on the PET/CT results
- Delay PET/CT for at least 12 weeks after completion of radiation treatment, unless required sooner for imminent surgical resection.
- PET mammography (PEM, generally reported with CPT® 78811) is considered experimental and investigational at this time

Unlisted Procedure Codes in Oncology (ONC-1.5)

ON.GG.0001.5.C

- eviCore does not routinely authorize requests for PET associated with imagedirected biopsy or radiation therapy treatment planning.
- There is often no unique procedure code for a service performed solely for treatment planning purposes. AMA instructions in the CPT state that if no specific code exists for a particular service, the service is reported with an unlisted code.
- Advanced imaging being used for radiation therapy treatment planning should not be authorized using any of the diagnostic imaging codes for CT, MRI or PET:
 - of radiation therapy to be delivered ONLY to the brain, prostate and cervix. The use of this code for radiation treatment planning of any other cancers/body parts not listed above may be reviewed on a case-by-case basis.
 - <u>CPT® 76497 for Unlisted CT</u> may NOT be used for radiation treatment planning. CT imaging performed in support of radiation therapy treatment planning is bundled in with the concurrent radiation treatment authorization codes and a separate authorization for treatment planning is not required.
 - CPT® 78999 for Unlisted procedure, nuclear medicine (PET) eviCore does not perform prior authorization for this CPT code. This code may not be reviewed or offered as an alternative recommendation to the provider.
 - Imaging associated with image-directed biopsy should be reported with the corresponding interventional codes.
 - For advanced imaging used solely for the purpose of Surgical planning, see:
 <u>Unlisted Procedures/Therapy treatment planning (Preface-4.3)</u> in the Preface Imaging Guidelines.

Predisposition Syndromes (ONC-1.6)

ON.GG.0001.6.A

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 For predisposition syndrome screening in adult individuals, see: <u>Screening</u> <u>Imaging in Cancer Predisposition Syndromes (PEDONC-2)</u> in the Pediatric Oncology Imaging Guidelines

References (ONC-1)

- 1. ACR Committee on Drugs and Contrast Media. ACR Manual on Contrast Media, version 10.3. Reston, VA: American College of Radiology; 2018.
- 2. The American College of Radiology. *Practice parameter for the performance of skeletal scintigraphy (bone scan)*. Rev. 2017.
- The American College of Radiology. Practice parameter for performing FDG-PET/CT in oncology. Rev. 2016
- 4. The American College of Radiology. *Practice parameter for the performance of tumor scintigraphy with gamma cameras*). Rev. 2015.
- 5. Erdi YE. Limits of tumor detectability in nuclear medicine and PET. *Mol Imaging Radionucl Ther.* 2012;21(1):23-28. doi:10.4274/Mirt.128.
- 6. Hapani S, Sher A, Chu D, Wu S. Increased risk of serious hemorrhage with bevacizumab in cancer patients: a meta-analysis. *Oncology*. 2010;79(1):27-38. doi:10.1159/000314980.
- 7. ACR Appropriateness Criteria. Pretreatment planning of Invasive cancer of Cervix. Rev. 2015.
- 8. ACR Appropriateness Criteria. External Beam Radiation therapy treatment planning for clinically localized prostate cancer. Rev. 2016.
- 9. Metcalfe P, Liney GP, Holloway L, et al. The potential for an enhanced role for MRI in radiation-therapy treatment planning. *Technol Cancer Res Treat*. 2013;12(5):429-46. doi:10.7785/tcrt.2012.500342.
- 10. Daly MB, Pal T, AlHilli Z, et. al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 February 13, 2023, Genetic/Familial High Risk Assessment: Breast and Ovarian, available at: http s://www.nccn.org/professionals/physician_gls/pdf/genetics_bop.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Genetic/Familial High Risk Assessment: Breast and Ovarian V3.2023 February 13, 2023 [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 11. Coverage of Clinical Trials under the Patient Protection and Affordable Care Act; 42 U.S.C.A. § 300gg-8.
- 12. Shah MH, Goldner WS, Benson III AB, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2022 December 21, 2022. Neuroendocrine tumors, available at: https://www.nccn.org/professionals/physician_gls/pdf/neuroendocrine.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Neuroendocrine tumors V2.2022 December 21, 2022. [©]2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.

Primary Central Nervous System Tumors (ONC-2)

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Primary Central Nervous System Tumors – General Considerations (ONC-2.1)

ON.CN.0002.1.C

- This guideline section applies to primary CNS tumors only. For imaging guidelines
 in metastatic brain cancer, see the appropriate diagnosis-specific section or <u>Brain</u>
 <u>Metastases (ONC-31.3)</u> for imaging guidelines.
- Primary brain tumors presenting only with uncomplicated headache are very uncommon. Most primary brain tumors present with specific CNS symptoms.
- Histologic confirmation is critical. Therapeutic decisions should not be made on radiographic findings alone, except for ANY of the following:
 - Medically fragile individuals for whom attempted biopsy carries excess medical risk, as stated in writing by both the attending physician and surgeon.
 - Brain stem tumors or other sites where the imaging findings are pathognomonic and the risk of permanent neurological damage is excessive with even a limited biopsy attempt
- For evaluation of known or suspected spinal cord compromise, see: <u>Spinal Cord</u> <u>Compression (ONC-31.6)</u>
- For suspected brain tumors in neurofibromatosis, see: <u>Screening Imaging in Cancer Predisposition Syndromes (PEDONC-2)</u> in the Pediatric Oncology Imaging Guidelines
- Rare tumors occurring more commonly in the pediatric population should be imaged according to the imaging guidelines in: <u>Pediatric Central Nervous System</u> <u>Tumors (PEDONC-4)</u> in the Pediatric Oncology Imaging Guidelines

Indication	lmaging Study	
Characterization and follow up of all brain tumors	 MRI Brain without and with contrast (CPT® 70553) CT Head without and with contrast (CPT® 70470) can be approved when MRI is contraindicated or not available, or there is skull bone involvement CT Head (contrast as requested) can be approved for preoperative planning when requested by the operating surgeon 	
Preoperative planning or to clarify inconclusive findings on MRI or CT	 MRA Head (CPT® 70544) or CTA Head (CPT® 70496) 	
Within 24 to 72 hours following brain tumor surgery	MRI Brain without and with contrast (CPT® 70553)	

Indication	Imaging Study
Clinical deterioration or development of new neurological features	 MRI Brain without and with contrast (CPT® 70553) MRI Spine without and with contrast (Cervical-CPT® 72156, Thoracic-CPT® 72157, Lumbar-CPT® 72158) for signs/symptoms of spinal involvement or if spinal involvement is suspected

MR Spectroscopy in Brain Tumors (MRS, CPT® 76390)

- MRS is only supported for use in brain tumors of specified histologies where diagnostic accuracy has been established in peer-reviewed literature.
 - See diagnosis-specific guidelines for MRS indications
- MRS is considered investigational/experimental for all other histologies and indications not listed in a diagnosis-specific guideline section.

PET Brain Imaging (CPT® 78608 and CPT® 78609)

- PET Brain Metabolic Imaging (CPT® 78608) is considered experimental, investigational, or unproven for all other histologies and indications not listed in a diagnosis-specific guideline section
- PET Brain Perfusion Imaging (CPT® 78609) is considered experimental, investigational, or unproven in the evaluation or management of primary CNS tumors, and is nationally non-covered by Medicare per NCD 220.6.17
- Body PET studies (CPT® 78811, CPT® 78812, and CPT® 78813) and fusion PET/CT studies (CPT® 78814, CPT® 78815, or CPT® 78816) are considered experimental, investigational, or unproven in the evaluation or management of primary CNS tumors
- See: Other Imaging Studies (HD-24) in the Head Imaging Guidelines for details on other advanced neuro-imaging studies

Low Grade Gliomas (ONC-2.2)

ON.CN.0002.2.A

- These tumors are defined as having a WHO histologic grade of I or II (out of IV), can occur anywhere in the CNS, and includes the following tumors:
 - Pilocytic Astrocytoma
 - Fibrillary (or Diffuse) Astrocytoma
 - Optic Pathway Gliomas
 - Pilomyxoid Astrocytoma
 - Oligodendroglioma
 - Oligoastrocytoma
 - Oligodendrocytoma
 - Subependymal Giant Cell Astrocytoma (SEGA)
 - Ganglioglioma
 - Gangliocytoma
 - Dysembryoplastic infantile astrocytoma (DIA)
 - Dysembryoplastic infantile ganglioglioma (DIG)
 - Dysembryoplastic neuroepithelial tumor (DNT)
 - Tectal plate gliomas
 - Cervicomedullary gliomas
 - Pleomorphic xanthoastrocytoma (PXA)
 - Any other glial tumor with a WHO grade of I or II

Indication	Imaging Study
Initial Staging	 MRI Brain without and with contrast (CPT® 70553) if not already done MRI Spine without and with contrast (Cervical-CPT® 72156, Thoracic-CPT® 72157, Lumbar-CPT® 72158)
	 MRI Spine with contrast only (Cervical-CPT® 72142, Thoracic- CPT® 72147, Lumbar-CPT® 72149) can be approved if being performed immediately following a contrast- enhanced MRI Brain
After initial resection or other treatment (radiation therapy, etc.)	MRI Brain without and with contrast (CPT® 70553)

Indication	Imaging Study
For individuals undergoing chemotherapy treatment	 MRI Brain without and with contrast (CPT® 70553) every 2 cycles Individuals with spinal cord involvement at diagnosis can have MRI without and with contrast of the involved spinal region on the same schedule as MRI brain
 ONE of the following: Determine need for biopsy when transformation to high-grade glioma is suspected based on clinical symptoms or recent MRI findings Evaluate a brain lesion of indeterminate nature when the PET findings will be used to determine whether biopsy/resection can be safely postponed 	 ANY of the following: PET Brain Metabolic Imaging (CPT® 78608) MRI Perfusion imaging (CPT® 70553)
 ONE of the following: Distinguish low-grade from high-grade gliomas Evaluate a brain lesion of indeterminate nature when the MRS findings will be used to determine whether biopsy/resection can be safely postponed Distinguish radiation-induced tumor necrosis from progressive disease within 18 months of completing radiotherapy 	ANY of the following: • MR Spectroscopy (CPT® 76390) • MRI Perfusion imaging (CPT® 70553)
Suspected intracranial or intraspinal recurrence	All imaging supported for initial staging may be repeated
Surveillance	 MRI Brain without and with contrast (CPT® 70553) every 3 months for 2 years, then every 6 months thereafter Individuals with spinal cord involvement at diagnosis can have MRI Spine without and with contrast (Cervical-CPT® 72156, Thoracic-CPT® 72157, Lumbar-CPT® 72158) on the same schedule as MRI Brain

High Grade Gliomas (ONC-2.3)

ON.CN.0002.3.A

- These tumors are defined as having a WHO histologic grade of III or IV (out of IV can occur anywhere in the CNS (though the majority occur in the brain), and include the following tumors:
 - o Anaplastic astrocytoma
 - Glioblastoma multiforme
 - Diffuse intrinsic pontine glioma (DIPG, or "brainstem glioma")
 - o Gliomatosis cerebri
 - o Gliosarcoma
 - o Anaplastic oligodendroglioma
 - Anaplastic ganglioglioma
 - Anaplastic mixed glioma
 - o Anaplastic mixed ganglioneuronal tumors
 - o Any other glial tumor with a WHO grade of III or IV

Indication	Imaging Study
Initial Staging	 MRI Brain without and with contrast (CPT® 70553) if not already done MRI Spine without and with contrast (Cervical-CPT® 72156, Thoracic-CPT® 72157, Lumbar-CPT® 72158)
	 MRI Spine with contrast only (Cervical- CPT® 72142, Thoracic-CPT® 72147, Lumbar-CPT® 72149) can be approved if being performed immediately following a contrast-enhanced MRI Brain
Immediately following partial or complete resection	MRI Brain without and with contrast (CPT® 70553)
Immediately following radiation therapy (XRT)	MRI Brain without and with contrast (CPT® 70553) once within 2 to 6 weeks following completion of treatment, and then go to surveillance imaging

Indication	Imaging Study
For individuals undergoing chemotherapy treatment	 MRI Brain without and with contrast (CPT® 70553) every 2 cycles Individuals with spinal cord involvement at diagnosis can have MRI without and with contrast of the involved spinal region on the same schedule as MRI Brain
 ONE of the following: Distinguish low-grade from high-grade gliomas Evaluate a brain lesion of indeterminate nature when the MRS findings will be used to determine whether biopsy/resection can be safely postponed Distinguish radiation-induced tumor necrosis from progressive disease within 18 months of completing radiotherapy 	 ANY of the following: MR Spectroscopy (CPT® 76390) MRI Perfusion imaging (CPT® 70553)
 ONE of the following: Distinguish radiation-induced tumor necrosis from progressive disease Evaluate inconclusive MRI findings when the PET findings will be used to determine need for biopsy or change in therapy, including a change from active therapy to surveillance Evaluate a brain lesion of indeterminate nature when the PET findings will be used to determine whether biopsy/resection can be safely postponed 	 ANY of the following: MRI Perfusion imaging (CPT® 70553) PET Brain metabolic imaging (CPT® 78608) PET Brain is not indicated in gliomas occurring in the brain stem due to poor uptake and lack of impact on individual outcomes
Suspected intracranial or intraspinal recurrence	All imaging supported for initial staging may be repeated

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Indication	Imaging Study
Surveillance	 MRI Brain without and with contrast (CPT® 70553) every 3 months for 3 years and every 6 months thereafter Individuals with spinal cord involvement at diagnosis can have MRI Spine without and with contrast (Cervical-CPT® 72156,
	Thoracic-CPT® 72157, Lumbar-CPT® 72158) on the same schedule as MRI Brain

Medulloblastoma and Supratentorial Primitive Neuroectodermal Tumors (sPNET) (ONC-2.4)

ON.CN.0002.4.A

v1.0.2024

 Medulloblastoma and sPNET imaging indications in adult individuals are identical to those for pediatric individuals. See: <u>Medulloblastoma (MDB), Supratentorial</u> <u>Primitive Neuroectodermal Tumors (sPNET), and Pineoblastoma (PEDONC-4.4)</u> in the Pediatric Oncology Imaging Guidelines.

Ependymoma (ONC-2.5)

ON.CN.0002.5.A

v1.0.2024

 Ependymoma imaging indications in adult individuals are identical to those for pediatric individuals. See: <u>Ependymoma (PEDONC-4.8)</u> in the Pediatric Oncology Imaging Guidelines.

Central Nervous System Germ Cell Tumors (ONC-2.6)

ON.CN.0002.6.A

v1.0.2024

 Central nervous system germ cell tumor imaging indications in adult individuals are identical to those for pediatric individuals. See: <u>CNS Germinomas and Non-</u> <u>Germinomatous Germ Cell Tumors (NGGCT) (PEDONC-4.7)</u> in the Pediatric Oncology Imaging Guidelines.

CNS Lymphoma (also known as Microglioma) (ONC-2.7)

ON.CN.0002.7.A

V1.0.2024			
Indication	Imaging Study		
Initial Staging	 ALL of the following are indicated: MRI Brain without and with contrast (CPT® 70553) MRI Cervical Spine without and with contrast (CPT® 72156) MRI Thoracic Spine without and with contrast (CPT® 72157) MRI Lumbar Spine without and with contrast (CPT® 72158) 		
Extra-neural evaluation to confirm CNS primary *Individuals with CNS Lymphoma that is metastatic should be imaged according to: • Non-Hodgkin Lymphomas (ONC-27) for individuals age ≥18 years • Pediatric Aggressive Mature B-Cell Non-Hodgkin Lymphomas (NHL) (PEDONC-5.3) in the Pediatric Oncology Imaging Guidelines for individuals age ≤17 years	 ANY or ALL of the following are indicated: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) PET/CT (CPT® 78815) can be approved for evaluation of inconclusive findings on CT imaging 		
Treatment Response	MRI without and with contrast of all positive disease sites every 2 cycles		
Suspected intracranial or intraspinal recurrence	All imaging supported for initial staging may be repeated		
Surveillance	MRI without and with contrast of all positive disease sites every 3 months for 2 years, then every 6 months for 3 years, then annually thereafter		

Meningiomas (Intracranial and Intraspinal) (ONC-2.8)

ON.CN.0002.8.A

Indication	Imaging Study
Initial Staging of Intracranial Meningioma	 ANY or ALL of the following are indicated: MRI Brain without and with contrast (CPT® 70553) CT Head (contrast as requested)
Initial staging of Intraspinal Meningioma	 ONE of the following: MRI without and with contrast of appropriate spinal region (Cervical CPT® 72156, Thoracic CPT® 72157, and Lumbar CPT® 72158) OR CT without and with contrast of the appropriate spinal region (Cervical CPT® 72127, Thoracic CPT® 72130, and Lumbar CPT® 72133)
Treatment Response	MRI without and with contrast of all positive disease sites every 2 cycles
Suspected recurrence of intracranial or intraspinal recurrence	All imaging supported for initial staging may be repeated
Suspected recurrence with inconclusive findings on MRI	 Any ONE of the following studies: Octreotide SPECT Brain (CPT® 78803) Octreotide SPECT/CT Brain (CPT® 78830) Dotatate PET/CT Brain (CPT® 78814)
Surveillance for Grade I (low- grade) and Grade II (atypical) intracranial meningioma	 MRI Brain without and with contrast (CPT® 70553) at 3, 6, and 12 months, then annually for 5 years
(completely resected, partially resected, and unresected)	 Imaging beyond 5 years is only indicated for evaluation of new signs or symptoms

Indication	Imaging Study
Surveillance for Grade I (low- grade) and Grade II (atypical) intraspinal meningioma (completely resected, partially resected, and unresected)	 ONE of the following at 3, 6, and 12 months, and then annually for 5 years: MRI without and with contrast (CPT® 72156 [Cervical spine], CPT® 72157 [Thoracic spine], CPT® 72158 [Lumbar spine]) of the involved spinal level
	OR
	 CT without and with contrast (CPT® 72127 [Cervical spine], CPT® 72130 [Thoracic spine], CPT® 72133 [Lumbar spine]) of the involved spinal level
	 Imaging beyond 5 years is only indicated for evaluation of new signs or symptoms
Surveillance for Grade III (malignant or anaplastic) Meningioma	 Intracranial Meningioma: MRI Brain without and with contrast (CPT® 70553) every 3 months for 3 years, and then every 6 months thereafter
	 Intraspinal Meningioma: MRI or CT without and with contrast of the involved spinal region every 3 months for 3 years and then every 6 months thereafter

Spinal Cord Tumors (Benign and Malignant) (ONC-2.9)

ON.CN.0002.9.A

- See: <u>Low Grade Gliomas (ONC-2.2)</u> and <u>High Grade Gliomas (ONC-2.3)</u> for imaging guidelines of low-grade and high-grade gliomas of the spinal cord
- See: Malignant Tumors of the Spinal Cord (PEDONC-4.9) in the Pediatric Oncology Imaging Guidelines for other malignant spinal cord tumors
- See: <u>Neurofibromatosis 1 and 2 (NF1 and NF2) (PEDONC-2.3)</u> in the Pediatric Oncology Imaging Guidelines for spinal tumors in individuals with Neurofibromatosis 1 or 2
- See: **Spinal Cord Compression (ONC-31.6)** for known secondary malignancy involving the spine/spinal canal/spinal cord

Choroid Plexus Tumors (ONC-2.10)

ON.CN.0002.10.A

v1.0.2024

 Choroid Plexus Tumor imaging indications in adult individuals are identical to those for pediatric individuals. See: <u>Choroid Plexus Tumors (PEDONC-4.13)</u> in the Pediatric Oncology Imaging Guidelines.

References (ONC-2)

- 1. Nabors LB, Portnow J, Baehring J, et. al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 March 24, 2023 Central Nervous System Cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/cns.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Central Nervous System Tumors Cancer V1.2023. March 24, 2023 [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Brandão LA, Castillo M. Adult brain tumors: clinical applications of magnetic resonance spectroscopy. Magn Reson Imaging Clin N Am. 2016;24(4):781-809. doi:10.1016/j.mric.2016.07.005.
- 3. Pasquier D, Bijmolt S, Veninga T, et al. Atypical and malignant meningioma: outcome and prognostic factors in 119 irradiated patients. A multicenter, retrospective study of the Rare Cancer Network. *Int J Radiat Oncol Biol Phys.* 2008;71(5):1388. doi:10.1016.j.ijrobp.2007.12.020.
- 4. Modha A, Gutin PH. Diagnosis and treatment of atypical and anaplastic meningiomas: a review. *Neurosurgery*. 2005;57(3):538-550.
- 5. Horská A, Barker PB. Imaging of brain tumors: MR spectroscopy and metabolic imaging. *Neuroimaging Clin N Am.* 2010;20(3):293-310. doi:10.1016/j.nic.2010.04.003.
- Sundgren PC. MR Spectroscopy in radiation Injury. Am J Neuroradiol. 2009;30(8):1469-1476. doi:10.3174/ ajnr.A1580.
- 7. American College of Radiology. ACR–ASNR-SPR practice parameter for the performance of intracranial magnetic resonance perfusion imaging. 2017; Available at: https://www.acr.org/-/media/ACR/Files/Practice-Parameters/MR-Perfusion.pdf?la=en.

Squamous Cell Carcinomas of the Head and Neck (ONC-3)

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Squamous Cell Carcinomas of the Head and Neck – General Considerations (ONC-3.0)

ON.HN.0003.0.A

- Individuals with esthesioneuroblastoma should be imaged according to this guideline section
- Stage III/IV disease encompasses any primary tumor larger than 4 cm or documented lymph node positive disease

Squamous Cell Carcinomas of the Head and Neck – Suspected/Diagnosis (ONC-3.1)

ON.HN.0003.1.A

- See: <u>Neck Masses Imaging (NECK-5.1)</u> in the Neck Imaging Guidelines for evaluation of suspected malignancy in the neck
- PET may be considered prior to biopsy in order to determine a more favorable site for biopsy when:
 - o A prior biopsy was nondiagnostic or
 - A relatively inaccessible site is contemplated which would require invasive surgical intervention for biopsy attempt

Squamous Cell Carcinomas of the Head and Neck – Initial Work-up/Staging (ONC-3.2)

ON.HN.0003.2.C

Indication	Imaging Study
All Stages of Disease	 CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck (OFN) without and with contrast (CPT® 70543) CT Chest with contrast (CPT® 71260)
For sentinel lymph node evaluation when nodes are not clinically positive	 Lymph system imaging (lymphoscintigraphy, CPT® 78195) SPECT/CT (CPT® 78830) is indicated as an add on code if requested
Nasal cavity and paranasal sinuses (bony erosion or skull base and intracranial involvement)	 ONE of the following studies is indicated: CT Maxillofacial with contrast (CPT® 70487) CT Neck with contrast (CPT® 70491) MRI Orbits/Face/Neck without and with contrast (CPT® 70543)
Nasopharyngeal Cancer (NPC)	 MRI Orbits/Face/Neck without and with contrast (CPT® 70543) is the preferred study CT Neck (CPT® 70491) and/or CT Maxillofacial (CPT® 70487) with contrast can be approved if contraindication to MRI Chest x-ray or CT Chest with contrast (CPT® 71260)

Indication	Imaging Study
 For ANY of the following: Known stage III or IV disease Prior to start of primary chemoradiotherapy and have not undergone definitive surgical resection Nasopharyngeal primary site Inconclusive findings on conventional imaging (CT, MRI) In order to direct laryngoscopy/exam under anesthesia for biopsy Pulmonary nodule(s) ≥8 mm in size Cervical lymph node biopsy positive for squamous cell carcinoma and no primary site identified on CT or MRI Neck and Chest 	• PET/CT (CPT® 78815)
Signs or symptoms of abdominal metastatic disease, including elevated liver function tests	CT Abdomen with contrast (CPT® 74160)
Any head and neck cancer with neurological findings or suspicion of skull base invasion	MRI Brain without and with contrast (CPT® 70553)

Squamous Cell Carcinomas of the Head and Neck – Restaging/Recurrence (ONC-3.3)

ON.HN.0003.3.A

Indication	Imaging Study
Following complete resection and/or radical neck dissection	See: Surveillance/Follow-up (ONC-3.4)
Following primary chemoradiotherapy or radiation therapy in individuals who have not undergone surgical resection of primary tumor or neck dissection	 ONE of the following: CT Neck with contrast (CPT® 70491); or MRI Orbits/Face/Neck without and with contrast (CPT® 70543); or PET/CT (CPT® 78815) no sooner than 12 weeks (3 months) post completion of radiation therapy If post-treatment PET/CT scan is negative, further surveillance imaging is not routinely indicated.
Induction chemotherapy response	 CT Neck with contrast (CPT® 70491) or MRI Orbits/ Face/Neck without and with contrast (CPT® 70543) PET not indicated to assess response to induction chemotherapy
Measurable or metastatic disease undergoing active treatment	 Every 2 cycles (6-8 weeks): CT Neck with contrast (CPT® 70491) OR MRI Orbits/ Face/Neck without and with contrast (CPT® 70543) AND CT with contrast of involved body sites
Suspected local recurrence	 CT Neck with contrast (CPT® 70491) or MRI Orbits/ Face/Neck without and with contrast (CPT® 70543) CT Chest with contrast (CPT® 71260)

Indication	Imaging Study
Biopsy proven local recurrence	ONE of the following: • PET/CT (CPT® 78815)
	 or CT Neck with contrast (CPT® 70491) or MRI Orbits/ Face/Neck without and with contrast (CPT® 70543) and CT Chest with contrast (CPT® 71260)
Inconclusive conventional imaging (CT or MRI)	• PET/CT (CPT® 78815)
If new pulmonary symptoms or chest previously involved	CT Chest with contrast (CPT® 71260)

Squamous Cell Carcinomas of the Head and Neck – Surveillance/Follow-up (ONC-3.4)

ON.HN.0003.4.A

v1.0.2024

Indications	Imaging Study
Individuals treated with surgical resection of primary site and/or neck dissection (with or without postoperative radiation therapy)	Once within 6 months of completing all treatment: CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck without and with contrast (CPT® 70543) CT with contrast of any other involved body area
Individuals treated with definitive radiation therapy or combined chemoradiation, and post-treatment imaging is negative	Further surveillance imaging is not routinely indicated
If post-treatment imaging shows residual abnormalities	ONE of the following, once within 6 months of prior imaging: CT Neck with contrast (CPT® 70491) OR MRI Orbits/Face/Neck without and with contrast (CPT® 70543)
After initial post-treatment study, for ANY of the following: Nasopharyngeal primary site Physical exam unable to visualize deep-seated primary site	Annually for 3 years: • CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck without and with contrast (CPT® 70543)
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 CT Chest is not indicated for surveillance. Individuals with smoking history may undergo annual low dose CT cancer screening if criteria are met (See: <u>Lung</u> <u>Cancer Screening (CH-33)</u> in the Chest Imaging Guidelines)

References (ONC-3)

- 1. Pfister DG, Spencer S, Adkins D, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 May 15, 2023 Head and Neck Cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/head-and-neck.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Head and Neck Cancer V2.2023 May 15, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Goel R, Moore W, Sumer B, Khan S, Sher D, Subramaniam RM. Clinical practice in PET/CT for the management of head and neck squamous cell cancer. AJR Am J Roentgenol. 2017;209(2):289-303. doi:10.2214/AJR.17.18301.
- 3. Moncrieff M, Pywell S, Snelling A, et. al. Effectiveness of SPECT/CT imaging for sentinel node biopsy staging of primary cutaneous melanoma and patient outcomes. *Ann Surg Oncol.* 2022;29(2):767-775. doi:10.1245/s10434-021-10911-4.
- Quartuccio N, Garau LM, Arnone A, et. al. Comparsion of 99mTc-labeled colloid SPECT/CT and planar lymphoscintigraphy in sentinel lymph node detection in patients with melanoma: a meta-analysis. *J Clin Med*. 2020;9(6):1680. doi:10.3390/jcm9061680.
- 5. Bennie G, Vorster M, Buscombe J, Sathekge M. The added value of a single-photon emission computed tomography-computed in sentinel lymph node mapping in patients with breast cancer and malignant melanoma. *World J Nucl Med*. 2015;14(01):41-46. doi:10.4103/1450-1147.150543.

Salivary Gland Cancers (ONC-4)

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Salivary Gland Cancers – General Considerations (ONC-4.0)

ON.SG.0004.0.A

- Salivary gland tumors may originate within the parotid, submandibular, sublingual or minor salivary glands in the mouth.
- Histological subtypes include:
 - Mucoepidermoid
 - Acinic
 - Adenocarcinoma
 - Adenoid cystic carcinoma
 - Malignant myoepithelial tumors
 - o Squamous cell carcinoma
 - Lymphoma and metastatic squamous carcinoma can occur in the parotid gland
- Over 80% of parotid gland tumors are benign. A bilateral parotid tumor is most likely Warthin's tumor.
- The role of PET in salivary gland tumors is considered experimental, investigational, or unproven.

Salivary Gland Cancers – Suspected/Diagnosis (ONC-4.1)

ON.SG.0004.1.A

v1.0.2024

• See: <u>Salivary Gland Disorders (NECK-11)</u> and <u>Neck Masses – Imaging (NECK-5.1)</u> in the Neck Imaging Guidelines for evaluation of salivary gland masses, salivary gland stones and neck masses.

Salivary Gland Cancers – Initial Workup/Staging (ONC-4.2)

ON.SG.0004.2.A

Indication	Imaging Study
Biopsy-proven malignancy	 ONE of the following can be approved: MRI Orbits/Face/Neck without and with contrast (CPT® 70543) CT Neck with contrast (CPT® 70491) CT Neck without contrast (CPT® 70490)
Skull base invasion	MRI Brain without and with contrast (CPT® 70553)
 Adenoid cystic carcinoma Lymphadenopathy in the neck Pulmonary signs or symptoms Abnormal chest x-ray 	CT Chest with contrast (CPT® 71260)

Salivary Gland Cancers – Restaging/Recurrence (ONC-4.3)

ON.SG.0004.3.A

Indication	Imaging Study
After complete surgical resection	See: Salivary Gland Cancers - Surveillance (ONC-4.4)
Individuals with unresected disease receiving systemic therapy (chemotherapy)	 ONE of the following may be approved every 2 cycles: CT Neck with contrast (CPT® 70491) and any other sites of disease MRI Orbits/Face/Neck without and with contrast (CPT® 70543) and any other sites of disease
Recurrence or progression suspected based on new or worsening signs or symptoms	 ONE of the following may be approved: CT Neck with contrast (CPT® 70491) MRI Orbits/Face/Neck without and with contrast (CPT® 70543) In addition, for all individuals: CT Chest with contrast (CPT® 71260)
All other individuals	No routine advanced imaging indicated

Salivary Gland Cancers – Surveillance/ Follow-up (ONC-4.4)

ON.SG.0004.4.A

Indication	Imaging Study
Total surgical resection	No routine advanced imaging indicated
Unresectable or partially resected disease, including those treated with radiation therapy	Either CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck without and with contrast (CPT® 70543) once within 6 months of completion of treatment
Adenoid cystic carcinoma	 ANY of the following, annually for up to 10 years: CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck without and with contrast (CPT® 70543) CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250)

References (ONC-4)

- 1. Pfister DG, Spencer S, Adkins D et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 May 15, 2023 Head and Neck Cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/head-and-neck.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Head and Neck Cancer V2.2023 May 15, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Palacios E, Ellis M, Lam EC, Neitzschman H, Haile M. Pitfalls in imaging the submandibular glands with PET/CT. *Ear Nose Throat* J. 2015;94(10-11):E37-E39.
- 3. Seo YL, Yoon DY, Baek S, et al. Incidental focal FDG uptake in the parotid glands on PET/CT in patients with head and neck malignancy. *Eur Radiol*. 2015;25(1):171-177. doi:10.1007/s00330-0140339701.
- 4. Park HL, Yoo le R, Lee N, et al. The value of F-18 FDG PET for planning treatment and detecting recurrence in malignant salivary gland tumors: comparison with conventional imaging studies. *Nucl Med Mol Imaging*. 2013;47(4):242-248. doi:10.1007/s13139-013-0222-8.
- 5. Bertagna F, Nicolai P, Maroldi R. Diagnostic role of 18F-FDG-PET or PET/CT in salivary gland tumors: a systematic review. *Rev Esp Med Nucl Imagen Mol.* 2015;34(5):295-302.
- Garg M, Tudor-Green B, Bisase B. Current thinking in the management of adenoid cystic carcinoma of the head and neck. *British Journal of Oral and Maxillofacial Surgery*. 2019;57(8):716-721. doi:10.1016/j.bjoms.2019.07.021.
- 7. Geiger JL, Ismaila N, Beadle B, et al. Management of salivary gland malignancy: ASCO guideline. *Journal of Clinical Oncology*. 2021;39(17):1909-1941. doi:10.1200/JCO.21.00449.

Melanomas and Other Skin Cancers (ONC-5)

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Melanoma – General Considerations (ONC-5.0)

ON.SC.0005.0.A

v1.0.2024

• Melanomas can metastasize in an unpredictable fashion.

Melanoma – Suspected/Diagnosis (ONC-5.1)

ON.SC.0005.1.A

Indication	Imaging Study
All	Imaging is not indicated until histologic diagnosis is confirmed

Melanoma – Initial Work-Up/Staging (ONC-5.2)

ON.SC.0005.2.C

Indication	Imaging Study
Stage 0 or IA (in situ or disease <1 mm)	Routine advanced imaging is not indicated
 Stage IB (<0.8 mm with ulceration or 0.8-1 mm without or with ulceration) Stage II (lesions >1 mm thick, but node negative) 	CT with contrast or MRI without and with contrast of specific areas, only if signs or symptoms indicate need for further evaluation
For sentinel lymph node evaluation in stages IB and II	 Lymph system imaging (lymphoscintigraphy, CPT® 78195) SPECT/CT (CPT® 78830) is indicated as an add on code if requested
 ANY of the following: Stage III (sentinel node positive, palpable regional nodes) Stage IV (metastatic) 	 PET/CT (CPT® 78815 or CPT® 78816) OR CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Brain without and with contrast (CPT® 70553)
 Head or neck primary site Palpable lymphadenopathy in the neck Mucosal melanoma of the head or neck region 	In addition to above initial staging imaging, if PET/CT not performed: • CT Neck with contrast (CPT® 70491)
Primary site of melanoma is unknown and CT Chest, Abdomen, and Pelvis are negative	• PET/CT (CPT® 78815 or CPT® 78816)

Melanoma – Restaging/Recurrence (ONC-5.3)

ON.SC.0005.3.C

v1.0.2024

 All recurrences should be confirmed histologically, except when excessive morbidity from a biopsy may occur, such as a biopsy requiring craniotomy

Indication	Imaging Study
Individuals receiving chemotherapy, with measurable disease, after every 2 cycles	 CT Chest with contrast (CPT® 71260); and CT Abdomen and Pelvis with contrast (CPT® 74177) In addition, for individuals receiving systemic treatment for brain metastases: MRI Brain without and with contrast (CPT® 70553)
All in situ recurrences	Restaging imaging is not needed after adequate aggressive local therapy (See Surveillance)
Documented or clinically suspected (See top of page regarding biopsy morbidity) recurrence at: Primary site In-transit disease Regional lymph nodes Metastatic site ANY of the following: Inconclusive findings on conventional imaging Isolated metastatic site found on conventional imaging	 CT Chest with contrast (CPT® 71260); and CT Abdomen and Pelvis with contrast (CPT® 74177) In addition, for all individuals: MRI Brain without and with contrast (CPT® 70553) PET/CT (CPT® 78815 or CPT® 78816)
 Brain imaging is indicated for: New discovery of metastatic disease or progression of metastatic disease Signs or symptoms of CNS disease If considering Interleukin (IL-2) therapy 	MRI Brain without and with contrast (CPT® 70553)

Melanoma – Surveillance/Follow-up (ONC-5.4)

ON.SC.0005.4.C

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Indication	Imaging Study
Stage 0, IA, IB and IIA Melanomas	No advanced imaging indicated
Stage IIB, IIC, IIIAb and IIIB Melanomas	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast every 6 months for 2 years, then annually for 3 years For melanoma arising from extremities, advanced imaging of the primary site is not routinely indicated for surveillance in asymptomatic individuals.
Stage IIIC and IV Melanomas	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast every 3 months for 2 years, then every 6 months for 3 years MRI Brain without and with contrast (CPT® 70553) annually for 3 years For melanoma arising from extremities, advanced imaging of the primary site is not routinely indicated for surveillance in asymptomatic individuals.
Mucosal Melanoma of the head or neck region	In addition to above stage-based surveillance imaging, the following may be obtained ONCE within 6 months of completing all treatment: • CT Neck with contrast (CPT® 70491) or MRI Orbits/Face/Neck without and with contrast (CPT® 70543) • CT with contrast of any other involved body area
Liver metastases treated with focal therapy	See: Liver Metastases (ONC-31.2)

Non-Melanoma Skin Cancers – General Considerations (ONC-5.5)

ON.SC.0005.5.A

- Advanced imaging is generally not indicated for basal cell and squamous cell skin cancers
- PET/CT scan is not indicated for evaluation of non-melanoma skin cancers unless specified within the guidelines below (e.g. Merkel cell carcinoma)
- Merkel cell carcinoma is an unusual skin cancer with neuroendocrine-like histologic features, which has a high propensity (25% to 33%) for regional lymph node spread and occasionally, metastatic spread to lungs.
- Merkel cell carcinoma may present as a primary cancer or as a skin metastasis
 from a non-cutaneous primary neuroendocrine carcinoma (i.e., small cell lung
 cancer), therefore conventional imaging is indicated initially to confirm the absence
 of metastasis prior to considering PET scan.

Non-Melanoma Skin Cancers – Initial Work-up/Staging (ONC-5.6)

ON.SC.0005.6.A

Indication	Imaging Study
Body area with unexplained signs or symptoms	CT with contrast of that body area
Perineural invasion or local regional extension (i.e. bone; deep soft tissue) involvement	ONE of the following may be approved of the primary site: • MRI without contrast or without and with contrast • CT (contrast as requested)
Skin lesion may be a dermal metastasis from distant primary	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast PET/CT (CPT® 78815 or CPT® 78816) is indicated if conventional imaging (CT or MRI) is unable to identify a primary site
Squamous cell carcinoma head or neck skin with regional lymphadenopathy	CT Neck (CPT® 70491) and CT Chest (CPT® 71260) with contrast
Merkel Cell carcinoma	 CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of other involved body area(s) PET/CT (CPT® 78815 or CPT® 78816) if inconclusive conventional imaging Lymph system imaging (lymphoscintigraphy, CPT® 78195) for sentinel lymph node evaluation SPECT/CT (CPT® 78830) is indicated as an add on code if requested
Signs or symptoms of CNS involvement	MRI Brain with and without contrast (CPT® 70553)

Non-Melanoma Skin Cancers – Restaging/Recurrence (ONC-5.7)

ON.SC.0005.7.A

v1.0.2024

 All recurrences should be confirmed histologically, except when excessive morbidity from a biopsy may occur, such as a biopsy requiring craniotomy.

Indication	Imaging Study
Recurrence where planned therapy is more extensive than simple wide local excision	CT with contrast of the primary and recurrent site(s)
Suspected or biopsy- proven recurrence of Merkel cell carcinoma	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast CT with contrast of other symptomatic body area(s)
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815 or 78816)
Signs or symptoms of CNS involvement	MRI Brain without and with contrast (CPT® 70553)

Non-Melanoma Skin Cancers – Surveillance/Follow-up (ONC-5.8)

ON.SC.0005.8.A

Indication	Imaging Study
Merkel cell cancer – only if node positive	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast every 6 months for 5 years Add CT Neck with contrast (CPT® 70491) if known prior neck disease or scalp/facial/neck disease
All others	 Routine advanced imaging for surveillance is not indicated Imaging indicated only for signs and symptoms of recurrent disease

Ocular Melanoma (ONC-5.9)

ON.SC.0005.9.A

v1.0.2024

General Considerations

- Approximately 95% of ocular melanomas arise from the uvea (iris, ciliary body and choroid) and 5% arise from the conjunctiva or orbit.
- Biopsy is usually not necessary for initial diagnosis of uveal melanoma but may be useful in cases when diagnosis is uncertain (e.g. amelanotic tumors, retinal detachment) or for prognostic analysis and risk stratification.
- Treatment is directed to the affected eye with systemic therapy reserved only for known metastatic disease.
- The most common site of metastatic disease is the liver.
- Surveillance of the affected eye is with clinical examination only; advanced imaging
 is supported for surveillance of systemic metastatic disease based on individual risk
 factors. See risk categories below for surveillance recommendations.

Ocular Melanoma Risk Categories

Low Risk	Medium Risk	High-Risk
T1	T2 and T3	T4
Class IA	Class IB	Class 2
Spindle cell histology	Mixed Spindle and Epitheloid cells	Epitheloid cell histology
No extraoccular extension	No extraoccular extension	Extraoccular extension present
No ciliary body involvement	No ciliary body involvement	Ciliary body involvement present
Chromosome mutations: Disomy 3 EIF1AX mutation Gain of chromosome 6p	Chromosome mutations: • SF3B1 mutation	Chromosome mutations: BAP1 mutation PRAME mutation Monosomy 3 Gain of chromosome 8q

Indication	Imaging Study
Initial staging of suspected or biopsy- proven uveal melanoma	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Orbits/Face/Neck without and with contrast (CPT® 70543)
Neurological signs/symptoms	MRI Brain without and with contrast (CPT® 70553)
Restaging/Suspected Recurrence	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Orbits/Face/Neck without and with contrast (CPT® 70543) MRI Brain without and with contrast (CPT® 70553)
Surveillance for Low Risk disease	Annually for 10 years: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or MRI Abdomen without and with contrast (CPT® 74183)
Surveillance for Medium Risk disease	 Every 6 months for 2 years and then annually up to year 10: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or MRI Abdomen without and with contrast (CPT® 74183)
Surveillance for High Risk disease	 Every 3 months for 2 years, every 6 months for 3 years, then annually up to year 10: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or MRI Abdomen without and with contrast (CPT® 74183)

References (ONC-5)

- 1. Swetter SM, Johnson, D, Thompson JA, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 March 10, 2023 Cutaneous Melanoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/cutaneous_melanoma.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Cutaneous Melanoma V2.2023 March 10, 2023 ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Swetter S, Johnson D, Thompson JA, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 May 4, 2023 Melanoma: Uveal, available at: https://www.nccn.org/professionals/physician_gls/pdf/uveal.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Melanoma: Uveal V1.2023 May 4, 2023 ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Schmults CD, Blitzblau R, Aasi SZ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 April 10, 2023 Merkel Cell Carcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/mcc.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Merkel Cell Carcinoma V1.2023 April 10, 2023 ®2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 4. Schmults CD, Blitzblau R, Aasi SZ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 March 10, 2023 Basal Cell Skin Cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/nmsc.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Basal Cell Skin Cancer 1.2023 March 10, 2023 [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org .
- 5. Schmults CD, Blitzblau R, Aasi SZ, et al, National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 May 10, 2023 Squamous Cell Skin Cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/squamous.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Squamous Cell Skin Cancer V1.2023 May 10, 2023 [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org .
- 6. Schröer-Günther MA, Wolff RF, Westwood ME, et al. F-18-fluoro-2-deoxyglucose positron emission tomography (PET) and PET/computed tomography imaging in primary staging of patients with malignant melanoma: a systematic review. *Syst Rev.* 2012;1:62. doi:10.1186/2046-4053-1-62.
- Xing Y, Bronstein Y, Ross MI, et al. Contemporary diagnostic imaging modalities for the staging and surveillance of melanoma patients: a meta-analysis. *J Natl Cancer Inst*. 2011;103(2):129-142. doi:10.1093/inci/dig455.
- 8. Rodriguez Rivera AM, Alabbas H, Ramjuan A, Meguerditchian AN. Value of positron emission tomography scan in stage III cutaneous melanoma: a systematic review and meta-analysis. *Surg Oncol*. 2014;23(1):11-16. doi: 10.1016/j.suronc.2014.01.002.
- 9. Nathan P, Cohen V, Coupland S, et al. Uveal melanoma UK national guidelines. *European Journal of Cancer*. 2015;51(16):2404-2412. doi:10.1016/j.ejca/2015.07.013.
- Moncrieff M, Pywell S, Snelling A, et. al. Effectiveness of SPECT/CT imaging for sentinel node biopsy staging of primary cutaneous melanoma and patient outcomes. *Ann Surg Oncol*. 2022;29(2):767-775. doi:10.1245/s10434-021-10911-4.
- 11. Bennie G, Vorster M, Buscombe J, Sathekge M. The added value of a single-photon emission computed tomography-computed in sentinel lymph node mapping in patients with breast cancer and malignant melanoma. *World J Nucl Med.* 2015;14(01):41-46. doi:10.4103/1450-1147.150543

12. Quartuccio N, Garau LM, Arnone A, et. al. Comparsion of 99mTc-labeled colloid SPECT/CT and planar lymphoscintigraphy in sentinel lymph node detection in patients with melanoma: a meta-analysis. *J Clin Med*. 2020;9(6):1680. doi:10.3390/jcm9061680.

Thyroid Cancer (ONC-6)

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Thyroid Cancer – General Considerations (ONC-6.0)

ON.TC.0006.0.A

- Individuals of all ages with thyroid cancer are imaged according to this guideline.
- Whole-Body Thyroid Nuclear scan (also known as whole-body radioiodine scan) is coded with CPT[®] 78018. If CPT[®] 78018 is obtained and found to be positive, CPT[®] 78020 may be approved as an add-on test to evaluate the degree of iodine uptake.
- Single photon emission computed tomography (SPECT) imaging –
 Radiopharmaceutical Localization of Tumor SPECT (CPT® 78803 or CPT® 78831)
 or SPECT/CT Hybrid study (CPT® 78830 or CPT® 78832) may complement planar
 and pinhole imaging and can be approved as an add-on wherever radioiodine (RAI)
 scans are indicated.
- Whole-Body Thyroid Nuclear scan (also known as whole-body RAI scan) is the
 imaging modality of choice for differentiated thyroid cancers, as these are usually
 not well visualized on FDG-PET/CT scans. Individuals who have RAI-diagnostic
 scan negative and PET-positive disease will generally not respond to RAI
 treatment, whereas individuals who have PET-negative and RAI-diagnostic scan
 negative disease may still be candidates for empiric RAI treatment.
- Radioiodine (RAI) refractory disease is defined as: (i) the malignant/metastatic
 tissue does not ever concentrate RAI (no uptake outside the thyroid bed at the first
 therapeutic WBS), (ii)the tumor tissue loses the ability to concentrate RAI after
 previous evidence of RAI-avid disease (in the absence of stable iodine
 contamination), (iii) RAI is concentrated in some lesions but not in others, and (iv)
 metastatic disease progresses despite significant concentration of RAI⁶.

Thyroid Cancer – Suspected/Diagnosis (ONC-6.1)

ON.TC.0006.1.A

v1.0.2024

• See: <u>Thyroid Nodule (NECK-8.1)</u> in the Neck Imaging Guidelines for suspected thyroid malignancies

Thyroid Cancer – Initial Work-Up/Staging (ONC-6.2)

ON.TC.006.2.C

	V1.U.2U24
Follicular, Papillary and Hürthle Cell Carcinomas	Imaging Study
 ONE of the following: Locally advanced disease or fixation suggested by clinical exam and/or ultrasound Substernal or bulky disease Disease precluding full ultrasound examination Vocal cord paresis 	 ONE of the following: MRI Neck without contrast (CPT® 70540) MRI Neck without and with contrast (CPT® 70543) CT Neck without contrast (CPT® 70490) CT Neck with contrast (CPT® 70491) can be approved if contrast study is necessary for complete pre-operative assessment and use of IV contrast will not delay post-operative use of RAI therapy.
Post-thyroidectomy to assess thyroid remnant and to look for iodine-avid metastases for ONE of the following: • Extent of thyroid remnant cannot be accurately ascertained from the surgical report or neck ultrasound • When the results may alter the decision to treat • Prior to administration of RAI therapy	 Whole-Body Thyroid Nuclear scan (CPT® 78018) The following may be approved as an add-on test: CPT® 78020 to evaluate the degree of iodine uptake AND/OR SPECT (CPT® 78803, or CPT® 78831), OR SPECT/CT Hybrid study (CPT® 78830, or CPT® 78832)
Skeletal pain	 Bone scan Whole-Body Thyroid Nuclear scan (CPT® 78018) The following may be approved as an add-on test: CPT® 78020 to evaluate the degree of iodine uptake AND/OR SPECT (CPT® 78803, or CPT® 78831), OR SPECT/CT Hybrid study (CPT® 78830, or CPT® 78832)
Suspicious findings on Chest X-ray, US, or substernal extension of mass	CT Chest without contrast (CPT® 71250)

Follicular, Papillary and Hürthle Cell Carcinomas	Imaging Study
All other individuals	Routine preoperative advanced imaging is not indicated

Medullary Thyroid Carcinomas	Imaging Study
ANY of the following: • Elevated CEA levels • Calcitonin level >400pg/mL • Positive lymph nodes	 ANY or ALL of the following: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or CT Abdomen without and with contrast (CPT® 74170) Bone scan
Skeletal pain	Bone scan
Inconclusive findings on conventional imaging	⁶⁸ Gallium-labeled PET/CT (CPT® 78815)

Anaplastic Thyroid Carcinomas	Imaging Study
All	 ONE of the following combinations, not both: CT Neck with contrast (CPT® 70491), CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177) OR FDG PET/CT (CPT® 78815) In addition to one of the above studies: MRI Brain without and with contrast (CPT® 70553)
Skeletal pain	Bone scan

Thyroid Cancer – Restaging/Recurrence (ONC-6.3)

ON.TC.0006.3.C

Follicular, Papillary and Hürthle Cell Carcinomas	Imaging Study
Gross residual disease found in the neck post-thyroidectomy	 ANY one of the following: CT Neck with contrast (CPT® 70491) MRI Neck without and with contrast (CPT® 70543)
Within 2 weeks (ideally 7 to 10 days) following the administration of Radioactive lodine therapy	 Whole-body Thyroid Nuclear Scan (CPT[®] 78018) The following may be approved as an add-on test: CPT[®] 78020 to evaluate the degree of iodine uptake SPECT (CPT[®] 78803, or CPT[®] 78831) or SPECT/CT Hybrid study (CPT[®] 78830, or CPT[®] 78832)
 ANY of the following: Recurrence documented by biopsy Increasing thyroglobulin level without Thyrogen® stimulation Thyroglobulin level >2 ng/mL or higher than previous after Thyrogen® stimulation Anti-thyroglobulin antibody present Evidence of residual thyroid tissue on ultrasound or physical exam after thyroidectomy or ablation 	 ALL of the following: CT Neck with contrast (CPT® 70491) or MRI Neck without and with contrast (CPT® 70543) CT Chest with contrast (CPT® 71260) CT with contrast of any symptomatic body area Whole-body Thyroid Nuclear Scan (CPT® 78018) The following may be approved as an add-on test: CPT® 78020 to evaluate the degree of iodine uptake SPECT (CPT® 78803 or CPT® 78831), or SPECT/CT Hybrid study (CPT® 78830, or CPT® 78832)

Follicular, Papillary and Hürthle Cell Carcinomas	Imaging Study
 ANY of the following: Rising thyroglobulin level with negative CT scans AND radioiodine scan Inconclusive findings on conventional imaging (CT scans and radioiodine scan) Known radioiodine-refractory disease and CT scans are negative or inconclusive 	• FDG PET/CT (CPT® 78815)
Measurable metastatic disease on systemic therapy (no more often than every 2 cycles)	 CT Chest with contrast (CPT® 71260) CT with contrast of affected or symptomatic body area

Medullary Thyroid Carcinoma	Imaging Study
 ANY of the following: Elevated CEA levels Calcitonin level ≥150 pg/mL Signs or symptoms of recurrence 	 ANY or ALL of the following: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or CT Abdomen without and with contrast (CPT® 74170) Bone scan
Inconclusive conventional imaging with calcitonin ≥150 pg per mL	⁶⁸ Gallium-labeled DOTATATE PET/CT (CPT® 78815)

Anaplastic Thyroid Carcinoma	Imaging Study
Measurable metastatic disease on systemic treatment	 Any of the following every 2 cycles (usually every 6-8 weeks): CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT of any other involved/symptomatic sites

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Anaplastic Thyroid Carcinoma	Imaging Study
Signs or symptoms of recurrence	 ONE of the following combinations, not both: CT Neck with contrast (CPT® 70491), CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177) OR FDG PET/CT (CPT® 78815) In addition to one of the above studies: MRI Brain without and with contrast (CPT® 70553)

Thyroid Cancer – Surveillance/Follow-up (ONC-6.4)

ON.TC.0006.4.A

Follicular, Papillary and Hürthle Cell Carcinomas	Imaging Study
Individuals being monitored on active surveillance	 Neck ultrasound (CPT® 76536) every 6 months for 2 years, and then annually thereafter
All other individuals post- treatment	Neck ultrasound (CPT® 76536) once at 6-12 months post-treatment, and then annually thereafter
For individuals with ANY of the following: Node positive disease RAI-avid metastases	 Whole-body Thyroid Nuclear Scan annually (CPT® 78018) The following may be approved as an add-on test: CPT® 78020 to evaluate the degree of iodine uptake SPECT (CPT® 78803, or CPT® 78831), OR SPECT/CT Hybrid study (CPT® 78830, or CPT® 78832)

Medullary Carcinomas	Imaging Study
All individuals	 CEA and calcitonin are required for monitoring medullary carcinomas Routine surveillance imaging is not indicated

Anaplastic Thyroid Carcinomas	Imaging Study
All individuals	 Every 3 months for 2 years: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Brain without and with contrast (CPT® 70553)

References (ONC-6)

- 1. Haddad RH, Bischoff L, Ball D, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 July 27, 2023 Thyroid carcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/thyroid.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Thyroid carcinoma V3.2023 Juy 27, 2023 ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Slough CM, Randolph GW. Workup of well-differentiated thyroid carcinoma. *Cancer Control.* 2006;13(2):99-105. doi:10.1177/107327480601300203.
- 3. Smallridge RC, Ain KB, Asa SL, et al. American Thyroid Association guidelines for management of patients with anaplastic thyroid cancer. *Thyroid*. 2012;22(11):1104-1139. doi:10.1089/thy.2012.0302.
- 4. Wells SA Jr, Asa SL, Dralle H, et al. American Thyroid Association guidelines for the management of medullary thyroid carcinoma. *Thyroid*. 2015;25(6):567-610. doi:10.1089/thy.2014.0335.
- Yeh MW, Bauer AJ, Bernet VA, et al. American Thyroid Association statement on preoperative imaging for thyroid cancer surgery. *Thyroid*. 2015;25:3-14. doi:10.1089/thy.2014.0096.
- Haugen BR, Alexander EK, Bible KB, et al. 2015 American Thyroid Association management guidelines for adult patients with thyroid nodules and differentiated thyroid cancer: the American Thyroid Association guidelines task force on thyroid nodules and differentiated thyroid cancer. *Thyroid*. 2016;26(1):1-133. doi:10.1089/thy.2015.0020.
- 7. Silberstein EB, Alavi A, Balon HR, et al. The SNMMI Practice Guideline for therapy of thyroid disease with ¹³¹I3.0. *J Nucl Med*. 2012;53(10):1633-1651. doi:10.2967/jnumed.112.105148.
- 8. Avram AM, Fig LM, Frey KA, Gross MD, Wong KK. Preablation 131-I scans with SPECT/CT in postoperative thyroid cancer patients: what is the impact on staging? *J Clin Endocrinol Metab*. February 21, 2013 [Epub ahead of print].

Small Cell Lung Cancer (ONC-7)

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Small Cell Lung Cancer – General Considerations (ONC-7.0)

ON.SL.0007.0.A

- Combined histologies of Small and Non-Small cell are considered Small cell lung cancer. Use this guideline for imaging recommendations for small and large cell high-grade (poorly differentiated) neuroendocrine tumors of the lung.
- Imaging is presently guided by traditional staging of limited or extensive disease.
 - Extensive stage is either metastatic disease or an extent which cannot be encompassed by a single radiotherapy portal.
 - Limited staging is confined to one side of the chest.
- Individuals treated curatively for SCLC are at increased risk for developing a second lung cancer. If new lung nodule is seen on imaging without any evidence of other systemic disease, follow <u>Lung Metastases (ONC-31.1)</u> for work-up of nodule.
- For carcinoid (low-grade neuroendocrine tumors) of the lung, see: **Neuroendocrine Cancers and Adrenal Tumors (ONC-15)**

Small Cell Lung Cancer – Suspected/Diagnosis (ONC-7.1)

ON.SL.0007.1.A

Indication	Imaging Study
Abnormal chest x-ray or clinical suspicion remains high despite a normal chest x-ray in symptomatic individual	 CT Chest without contrast (CPT® 71250) or CT Chest with contrast (CPT® 71260)
Pulmonary nodule <8 mm in size noted on CT Chest	See: Incidental Pulmonary Nodules Detected on CT Images (CH-16.2) in the Chest Imaging Guidelines
Pulmonary nodule 8 mm (0.8 cm) to 30 mm (3 cm) seen on CT Chest or MRI Chest	 See: PET (CH-16.4) in the Chest Imaging Guidelines If PET is Positive: Qualifies as initial staging PET/CT Biopsy is indicated prior to PET imaging for pulmonary masses ≥31 mm (3.1 cm) in size
Mediastinal/Hilar Mass	See: Lymphadenopathy (CH-2) in the Chest Imaging Guidelines
Paraneoplastic syndrome suspected	See: Paraneoplastic Syndromes (ONC-30.3)

Small Cell Lung Cancer – Initial Workup/Staging (ONC-7.2)

ON.SL.0007.2.C

Indication	Imaging Study
Initial staging	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Brain without and with contrast (CPT® 70553) Bone scan, if PET/CT not being done
To confirm the extent of disease when initial CT and MRI indicate limited stage disease (confined to one side of the chest)	• PET/CT (CPT® 78815)

Small Cell Lung Cancer – Restaging/Recurrence (ONC-7.3)

ON.SL.0007.3.C

Indication	Imaging Study
 Treatment Response: After every 2 cycles of chemotherapy Following completion of chemoradiation 	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Brain without and with contrast (CPT® 70553) for measurable brain metastases being treated with systemic therapy Bone scan PET is not indicated for evaluation of treatment response in SCLC, but can be considered on a case-by-case basis.
Restaging (suspected recurrence)	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Brain without and with contrast (CPT® 70553) Bone scan PET is not indicated for evaluation of recurrent SCLC but can be considered on a case-by-case basis.
Complete or partial response to initial treatment, if prophylactic cranial irradiation (PCI) is planned.	MRI Brain without and with contrast (CPT® 70553)

Small Cell Lung Cancer – Surveillance/ Follow-up ONC-7.4

ON.PC.0007.4.A

Indication	Imaging Study
Limited stage SCLC	 Every 3 months for one year, every 6 months for two years, and then annually: CT Chest without (CPT® 71250) or CT Chest with (CPT® 71260) contrast CT Abdomen and Pelvis with contrast (CPT® 74177) For new nodules, see: Lung Metastases (ONC-31.1)
Extensive stage SCLC	 Every 2 months for one year, every 4 months for two years, every 6 months for two years, and then annually: CT Chest without (CPT® 71250) or CT Chest with (CPT® 71260) contrast CT Abdomen and Pelvis with contrast (CPT® 74177) For new nodules, see: Lung Metastases (ONC-31.1)
Screening for brain metastases, regardless of PCI status	MRI Brain without and with contrast (CPT® 70553) every 4 months for 1 year and then every 6 months for 1 year

References (ONC-7)

- 1. Ganti AKP, Loo Jr. BW, Bassetti M, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 December 21, 2022 Small Cell Lung Cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/sclc.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Small Cell Lung Cancer V3.2023 December 21, 2022 ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Lu YY, Chen JH, Liang JA, Chu S, Lin WY, Kao CH. 18F-FDG PET or PET/CT for detecting extensive disease in small-cell lung cancer: a systematic review and meta-analysis. *Nucl Med Commun*. 2014;35(7):697-703. doi:10.1097/MNM.000000000000122.
- 3. Carter BW, Glisson BS, Truong MT, Erasmus JJ. Small cell lung carcinoma: staging, imaging, and treatment considerations. *Radiographics*. 2014;34(6):1707-1721. doi:10.1148/rg.346140178.
- 4. Kalemkerian G. Staging and imaging of small cell lung cancer. *Cancer Imag.* 2011;11(1):253-258.doi:10.1102/1470-7330.2011.0036.

Non-Small Cell Lung Cancer (ONC-8)

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Non-Small Cell Lung Cancer – General Considerations (ONC-8.0)

ON.NL.0008.0.C

- Non-small cell lung cancer includes adenocarcinoma, squamous cell carcinoma, adenosquamous, and large cell tumors.
- See: <u>Bronchopulmonary or Thymic Carcinoid Initial Staging (ONC-15.6)</u> for evaluation of low-grade neuroendocrine tumors (carcinoid) of the lung.
- See: **Small Cell Lung Cancer (ONC-7)** for evaluation of high-grade small cell and large cell neuroendocrine tumors of the lung.
- PET/CT may be considered to confirm solitary focus of extra-pulmonary metastatic disease (i.e., brain or adrenal) if the individual is being considered for an aggressive treatment for oligometastatic disease.

Non-Small Cell Lung Cancer – Asymptomatic Screening (ONC-8.1)

ON.NL.0008.1.A

v1.0.2024

• See: <u>Lung Cancer Screening (CH-33)</u> in the Chest Imaging Guidelines for criteria for Low-dose CT Chest for lung cancer screening.

Non-Small Cell Lung Cancer – Suspected/Diagnosis (ONC-8.2)

ON.NL.0008.2.A

Indication	Imaging Study
Abnormal Chest X-ray or clinical suspicion remains high despite a normal Chest X-ray in symptomatic individual	 CT Chest without contrast (CPT® 71250) or CT Chest with contrast (CPT® 71260)
Pulmonary nodule <8 mm in size noted on CT Chest	See: Incidental Pulmonary Nodules Detected on CT Images (CH-16.2) in the Chest Imaging Guidelines
Pulmonary nodule 8 mm (0.8 cm) to 30 mm (3 cm) seen on CT Chest or MRI Chest	 PET/CT (CPT® 78815) See: PET (CH-16.4) in the Chest Imaging Guidelines If PET is Positive: Qualifies as initial staging PET/CT
Pulmonary mass 31 mm (3.1 cm) or greater seen on CT or MRI	 PET/CT (CPT® 78815) can be approved prior to biopsy if ONE or MORE of the following applies: Definitive treatment with resection or radiation will be utilized instead of biopsy if PET confirms limited disease Multiple possible biopsy options are present within the chest and PET findings will be used to determine the most favorable biopsy site Biopsy is indicated prior to PET imaging for all other indications in pulmonary masses ≥31 mm (3.1 cm) in size
Mediastinal/Hilar Lymphadenopathy	See: Mediastinal Lymphadenopathy (CH-2.3) in the Chest Imaging Guidelines
Mediastinal/Hilar Mass	See: Mediastinal Mass (CH-20) in the Chest Imaging Guidelines
Paraneoplastic syndrome suspected	See: Paraneoplastic Syndromes (ONC-30.3)

Non-Small Cell Lung Cancer – Initial Work-Up/Staging (ONC-8.3)

ON.NL.0008.3.C

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Indication	Imaging Study
All individuals	 ANY or ALL of the following: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) CT Abdomen may be omitted if CT Chest report clearly documents upper abdomen through level of adrenals Bone scan, if PET/CT not being done
 ANY of the following: Stage I-IIIB Stage IV confined to the chest region (including pleural/pericardial effusion) Stage IV with oligometastatic disease on conventional imaging and individual is a candidate for aggressive surgical resection or other localized treatment of metastases with a curative intent Conventional imaging is inconclusive 	PET/CT (CPT® 78815) (If not already completed prior to histological diagnosis)
 ANY of the following: All Stage II-IV disease Stage I disease and considering surgical resection as primary therapy 	MRI Brain without and with contrast (CPT® 70553)
Superior sulcus (Pancoast) tumor suspected	 ANY or ALL of the following: MRI Chest without and with contrast (CPT® 71552) MRI Cervical Spine without and with contrast (CPT® 72156) MRI Thoracic Spine without and with contrast (CPT® 72157)

Non-Small Cell Lung Cancer – Restaging/Recurrence (ONC-8.4)

ON.NL.0008.4.C

Indication	Imaging Study
Stage I or II individuals who undergo definitive local treatment with surgery, radiation, or radiosurgery	Restaging imaging is not indicated. See: <u>Surveillance/Follow-Up (ONC-8.5)</u>
 ANY of the following: After neoadjuvant treatment for evaluation of surgical resectability Prior to starting adjuvant therapy Inadequately resected disease 	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250)
Measurable metastatic disease, undergoing active treatment, after every 2 cycles of chemotherapy	 ANY or ALL of the following studies: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) CT Abdomen and Pelvis with contrast (CPT® 74177) may be substituted for known pelvic disease or pelvic symptoms CT with contrast of other involved body areas In addition to the above studies, for individuals receiving systemic treatment for brain metastases: MRI Brain without and with contrast (CPT® 70553) In addition to the above studies, for individuals receiving systemic treatment for bone metastases: Bone scan PET/CT is not indicated for routine evaluation of NSCLC that is metastatic outside the chest cavity

Indication	Imaging Study
Suspected recurrence	 CT Chest with contrast (CPT® 71260); and CT Abdomen with contrast (CPT® 74160) CT Abdomen and Pelvis with contrast (CPT® 74177) may be substituted for known pelvic disease or pelvic symptoms For individuals with prior history of brain metastases or current signs or symptoms of brain metastasis: MRI Brain without and with contrast (CPT® 70553)
 ANY of the following: Biopsy proven recurrence localized to the chest cavity Inconclusive findings conventional imaging To differentiate tumor from radiation scar/fibrosis Stage IV with oligometastatic disease on conventional imaging and individual is a candidate for aggressive surgical resection or other localized treatment of metastases with a curative intent 	• PET/CT (CPT® 78815)
 ANY of the following: Following a demonstrated adequate response to neoadjuvant therapy if intracranial disease will preclude surgery Documented recurrence/progression New or worsening neurological signs or symptoms 	MRI Brain without and with contrast (CPT® 70553)

Non-Small Cell Lung Cancer – Surveillance/Follow-up (ONC-8.5)

ON.NL.0008.5.A

Indication	Study
Stage I-II	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) every 6 months for 3 years and then annually
	***Individuals treated with radiation therapy and residual abnormality on imaging may undergo CT Chest every 3 months for the first year after therapy, every 6 months for 2 years, and then annually thereafter
Stage III-IV (metastatic sites treated with definitive intent)	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) every 3 months for 2 years, every 6 months for 3 years and then annually
New lung nodule	See: Lung Metastases (ONC-31.1)

References (ONC-8)

- 1. Ettinger DS, Wood DE, Aisner DL, et al, National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 April 13, 2023. Non small cell lung cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/nscl.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Non-small cell lung cancer V3.2023 April 13, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Scheider BJ, Ismaila N, Aerts J, et al. Lung cancer surveillance after definitive curative-intent therapy: ASCO guideline. *J Clin Oncol*. 2020;38(7):753-766. doi:10.1200/JCO.19.02748.
- MacMahon H, Naidich DP, Goo JM, et al. Guidelines for management of incidental pulmonary nodules detected on CT images: from the Fleischner Society 2017. *Radiology*. 2017;284(1):228-243. doi:10.1148/radiol.2017161659.
- 4. Calman L, Beaver K, Hind D, Lorigan P, Roberts C, Lloyd-Jones M. Survival benefits from follow-up of patients with lung cancer: a systematic review and meta-analysis. *J Thorac Oncol*. 2011;6(12):1993-2004. doi:10.1097/JTO.0b013e31822b01a1.
- 5. Lou F, Huang J, Sima CS et al. Patterns of recurrence and second primary lung cancer in early-stage lung cancer survivors followed with routine computed tomography surveillance. *J Thorac Cardiovasc Surg*. 2013;145:75-81. https://www.ncbi.nlm.nih.gov/pubmed/23127371.
- Colt HG, Murgu SD, Korst RJ, et al. Follow-up and surveillance of the patient with lung cancer after curative-intent therapy: diagnosis and management of lung cancer, 3rd ed: American College of Chest Physicians evidence-based clinical practice guidelines. *Chest.* 2013;143:e437S-454S. https://www. ncbi.nlm.nih.gov/pubmed/23649451
- 7. Dane B, Grechushkin V, Plank A, et al. PET/CT vs. non-contrast CT alone for surveillance 1-year post lobectomy for stage I non-small cell lung cancer. *Am J Nucl Med Mol Imaging*. 2013; 3:408-416. https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3784804/.
- Zhao L, He ZY, Zhong XN, et al. (18)FDG-PET/CT for detection of mediastinal nodal metastasis in non-small cell lung cancer: a meta-analysis. Surg Oncol. 2012;21(3):230-236. https://www.ncbi.nlm.nih.gov/pubmedhealth/PMH0049561/.
- 9. Li J, Xu W, Kong F, et al. Meta-analysis: accuracy of 18FDG PET-CT for distant metastasis in lung cancer patients. *Surg Oncol.* 2013;22(3):151-155. https://www.ncbi.nlm.nih.gov/pubmed/23664848.
- Ravenel JG. Evidence-based imaging in lung cancer: a systematic review. J Thorac Imaging. 2012; 27(5):315-324. http://journals.lww.com/thoracicimaging/Abstract/2012/09000/ Evidence_based_Imaging_in_Lung_Cancer__A.8.aspx.
- 11. Bille A, Pelosi E, Skanjeti A, et al. Preoperative intrathoracic lymph node staging in patients with non-small-cell lung cancer: accuracy of integrated positron emission tomography and computed tomography. *Eur J Cardiothorac Surg.* 2009;36(3):440-445. https://academic.oup.com/ejcts/article-lookup/doi/10.1016/j.ejcts.2009.04.003.

Esophageal and GE Junction Cancer (ONC-9)

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Esophageal and GE Junction Cancer – General Considerations (ONC-9.0)

ON.EJ.0009.0.C

- Imaging for esophageal cancer is determined by the cell type and in which third of the esophagus it occurs.
- These guidelines may be used for imaging of esophageal and gastroesophageal (GE) junction cancers.

Esophageal and GE Junction Cancer – Suspected/Diagnosis (ONC-9.1)

ON.EJ.0009.1.A

v1.0.2024

• See: **Dysphagia and Esophageal Disorders (NECK-3.1)** in the Neck Imaging Guidelines for evaluation of suspected esophageal malignancy.

Esophageal and GE Junction Cancer – Initial Work-up/Staging (ONC-9.2)

ON.EJ.0009.2.A

Indication	Imaging Study
Biopsy proven	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
	 CT Abdomen and Pelvis with contrast (CPT® 74177) may be approved instead of CT Abdomen if there are pelvic signs or symptoms
Upper 1/3 or neck mass	CT Neck with contrast (CPT® 70491)
If no evidence of metastatic disease on conventional imaging	• PET/CT (CPT® 78815)

Esophageal and GE Junction Cancer – Restaging/Recurrence (ONC-9.3)

ON.EJ.0009.3.A

Indication	Imaging Study
After primary chemoradiation therapy prior to surgery	 Any ONE of the following, not both: CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast OR PET/CT (CPT® 78815) no sooner than 8 weeks post completion of radiation therapy
Post-surgical resection	See: Surveillance/Follow-up (ONC-9.4)
Monitoring response to chemotherapy for stage IV/metastatic disease	 Every 2 cycles of treatment (~every 6-8 weeks): CT Abdomen with contrast (CPT® 74160) CT Chest with contrast (CPT® 71260)
 If conventional imaging is inconclusive or Salvage surgical candidate with recurrence and no metastatic disease documented by conventional imaging 	• PET/CT (CPT® 78815)
 For ANY of the following: Signs or symptoms of recurrence Biopsy proven on follow-up endoscopy Recurrence suggested by other imaging (i.e. Chest x-ray or barium swallow) 	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
If previously involved or new signs or symptoms	CT Pelvis with contrast (CPT® 72193) and/or CT Neck with contrast (CPT® 70491)

Esophageal and GE Junction Cancer – Surveillance/Follow-up (ONC-9.4)

ON.EJ.0009.4.A

Indication	Imaging Study
Stage 0-IA (Tis, T1a) disease	No routine advanced imaging indicated
Stage IB (T1b) disease	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast annually for 3 years
Stage II-III disease	 CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast every 6 months for 2 years and then annually for 3 more years
Stage IV disease	See: Phases of Oncology Imaging and General Phase-Related Considerations (ONC-1.2)

References (ONC-9)

- 1. Ajani JA, D'Amico TA, Bentrem DJ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 March 10, 2023. Esophageal and esophagogastric junction cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/esophageal.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Esophageal and esophagogastric junction cancers V2.2023 March 10, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Klaeser B, Nitzsche E, Schuller JC, et al. Limited predictive value of FDG-PET for response assessment in the preoperative treatment of esophageal cancer: results of a prospective multi-center trial (SAKK 75/02). Onkologie. 2009;32(12):724-730. doi:10.1159/000251842.
- 3. Malik V, Lucey JA, Duffy GJ, et al. Early repeated 18F-FDG PET scans during neoadjuvant chemoradiation fail to predict histopathologic response or survival benefit in adenocarcinoma of the esophagus. *J Nucl Med*. 2010;51(12):1863-1869. doi:10.2967/jnumed.110.079566.
- Stiekema J, Vermeulen D, Vegt E, et al. Detecting interval metastases and response assessment using 18F-FDG PET/CT after neoadjuvant chemoradiotherapy for esophageal cancer. *Clin Nucl Med*. 2014;39(10):862-867. doi:10.1097/RLU.000000000000517.
- Sudo K, Xiao L, Wadhwa R, et al. Importance of surveillance and success of salvage strategies after definitive chemoradiation in patients with esophageal cancer. *J Clin Oncol*. 2014;32(30):3400-3405. doi:10.1200/JCO.2014.56.7156.
- Lou F, Sima CS, Adusumilli PS, et al. Esophageal cancer recurrence patterns and implications for surveillance. J Thorac Oncol. 2013;8(12):1558–1562. doi:10.1097/01.JTO.0000437420.38972.fb.
- Goense L, van Rossum PS, Reitsma JB, et al. Diagnostic performance of 18F-FDG PET and PET/CT for the detection of recurrent esophageal cancer after treatment with curative intent: a systematic review and meta-analysis. J Nucl Med. 2015;56(7):995-1002. doi:10.2967/jnumed.115.155580.

Other Thoracic Tumors (ONC-10)

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Malignant Pleural Mesothelioma – Suspected/Diagnosis (ONC-10.1)

ON.OT.0010.1.A

v1.0.2024

• See: <u>Asbestos Exposure (CH-9.1)</u> in the Chest Imaging Guidelines for evaluation of suspected mesothelioma.

Malignant Pleural Mesothelioma – Initial Work-up/Staging (ONC-10.2)

ON.OT.0010.2.A

Indication	Imaging Study
Cytologically or pathologically proven	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
	 CT Abdomen and Pelvis with contrast (CPT® 74177) may be approved instead of CT Abdomen if there are pelvic signs or symptoms PET/CT (CPT® 78815) if no evidence of metastatic disease or inconclusive conventional imaging
Preoperative planning	MRI Chest without and with contrast (CPT® 71552)

Malignant Pleural Mesothelioma – Restaging (ONC-10.3)

ON.OT.0010.3.A

Indication	Imaging Study
Signs or symptoms of recurrence	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
	 CT Abdomen and Pelvis with contrast (CPT® 74177) may be approved instead of CT Abdomen if there are pelvic signs or symptoms
Treatment with	Every 2 cycles:
chemotherapy	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
	 CT Abdomen and Pelvis with contrast (CPT® 74177) may be approved instead of CT Abdomen if there are pelvic signs or symptoms
Following induction chemotherapy prior to surgical resection	CT Chest (CPT® 71260) and CT Abdomen (CPT® 74160) with contrast
	 CT Abdomen and Pelvis with contrast (CPT® 74177) may be approved instead of CT Abdomen if there are pelvic signs or symptoms PET/CT (CPT® 78815) if no evidence of metastatic disease
Inconclusive CT Chest	MRI Chest without and with contrast (CPT® 71552)

Malignant Pleural Mesothelioma – Surveillance (ONC-10.4)

ON.OT.0010.4.A

Indication	Imaging Study
All	 CT Chest with contrast (CPT® 71260) and previously involved regions every 3 months for 2 years, then annually thereafter

Thymoma and Thymic Carcinoma – Suspected/Diagnosis (ONC-10.5)

ON.OT.0010.5.A

- See: <u>Mediastinal Mass (CH-20.1)</u> in the Chest Imaging Guidelines for evaluation of suspected thymic malignancies.
- See: <u>Bronchopulmonary or Thymic Carcinoid Initial Staging (ONC-15.6)</u> for imaging guidelines for thymic carcinoid.

Thymoma and Thymic Carcinoma – Initial Work-up/Staging (ONC-10.6)

ON.OT.0010.6.A

Indication	Imaging Study
Encapsulated or invasive limited disease	CT Chest with contrast (CPT® 71260)
Extensive mediastinal involvement on CT Chest	 CT Abdomen with contrast (CPT® 74160) CT Neck with contrast (CPT® 70491)
Inconclusive finding on CT	ONE of the following: • PET/CT (CPT® 78815) • MRI Chest without and with contrast (CPT® 71552)
Preoperative planning	MRI Chest without and with contrast (CPT® 71552)
Thymic Carcinomas	Image according to Non-Small Cell Lung Cancer - Initial Work-up/Staging (ONC-8.3)

Thymoma and Thymic Carcinoma – Restaging (ONC-10.7)

ON.OT.0010.7.A

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Indication	Study	
Adjuvant therapy following surgical resection	Follow surveillance imaging	
Following induction chemotherapy prior to surgical resection, if no evidence of metastatic disease	• PET/CT (CPT® 78815)	
For suspected recurrence	CT Chest with contrast (CPT® 71260)	
Recurrence with extensive mediastinal involvement on CT Chest	 CT Abdomen with contrast (CPT® 74160) CT Neck with contrast (CPT® 70491) 	
Inconclusive finding on CT	ONE of the following: • PET/CT (CPT® 78815) • MRI Chest without and with contrast (CPT® 71552)	
Metastatic disease on chemotherapy	CT Neck (CPT® 70491), CT Chest (CPT® 71260), and CT Abdomen (CPT® 74160) with contrast, every 2 cycles of therapy	
Thymic carcinomas	See: Non-Small Cell Lung Cancer Restaging/Recurrence (ONC-8.4)	

Thymoma and Thymic Carcinoma – Surveillance (ONC-10.8)

ON.OT.0010.8.A

Indication	Study	
Thymoma	 CT Chest with contrast (CPT® 71260) and previously involved regions every 6 months for 2 years, then annually for next 10 years 	
Thymic carcinomas	CT Chest with contrast (CPT® 71260) every 6 months for 2 years and then annually for next 5 years	

References (ONC-10)

- 1. Ettinger DS, Wood DE, Stevenson J, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 December 15, 2022. Mesothelioma: Pleural, available at: https://www.nccn.org/professionals/physician_gls/pdf/meso_pleural.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Mesothelioma: Pleural V1.2023 December 15, 2022. [©]2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Ettinger DS, Wood DE, Stevenson J, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 July 20, 2023. Mesothelioma: Peritoneal, available at: https://www.nccn.org/professionals/physician_gls/pdf/meso_peritoneal.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Mesothelioma: Peritoneal V2.2023 July 20, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Ettinger DS, Wood DE, Riely GJ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 December 15, 2022. Thymoma and Thymic carcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/thymic.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Thymoma and Thymic carcinoma, V1.2023 December 15, 2022. ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 4. Sørensen JB, Ravn J, Loft A, Brenøe J, Berthelsen AK, Nordic Mesothelioma Group. Preoperative staging of mesothelioma by 18F-fluoro-2-deoxy-D-glucose positron emission tomography/computer tomography fused imaging and mediastinoscopy compared to pathological findings after extrapleural pneumonectomy. *Eur J Cardiothorac Surg.* 2008;34:1090-1096. doi:10.1016/j.ejcts.2008.07.050.
- 5. Wilcox BE, Subramaniam RM, Peller PJ, et al. Utility of computed tomography-positron emission tomography for selection of operable malignant pleural mesothelioma. *Clin lung cancer*. 2009;10:244-248. doi: 10.3816/CLC.2009.n.033.
- Marom EM. Imaging thymoma. J Thorac Oncol. 2010;5(10 Suppl 4):S296-S303. doi:10.1097/JTO.0b013e3181f209ca.
- Marom EM. Advances in thymoma imaging. *J Thorac Imaging*. 2013;28(2):69-80. doi:10.1097/RTI.0b013e31828609a0.
- 8. Hayes SA, Huang J, Plodkowski AJ, et al. Preoperative computed tomography findings predict surgical resectability of thymoma. *J Thorac Oncol*. 2014;9(7):1023-1030. doi:10.1097/JTO.0000000000000204.
- 9. Mineo TC, Ambrogi V. Malignant pleural mesothelioma: factors influencing the prognosis. *Oncology*. 2012;26(12):1164-75.

Breast Cancer (ONC-11)

Guideline	Page
Breast Cancer – General Considerations (ONC-11.0)	114
Breast Cancer – Suspected/Diagnosis (ONC-11.1)	115
Breast Cancer – Initial Work-Up/Staging (ONC-11.2)	
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Breast Cancer – General Considerations (ONC-11.0)

ON.BC.0011.0.C

- Advanced imaging to evaluate for distant metastases is not indicated for asymptomatic individuals with invasive or pre-invasive or in-situ breast cancer (histologies such as DCIS and LCIS).
- Bone scan has a high concordance rate with PET for detecting bone metastases.
- Scintimammography and Breast Specific Gamma Imaging (BSGI) are considered experimental, investigational, or unproven.

Breast Cancer – Suspected/Diagnosis (ONC-11.1)

ON.BC.0011.1.A

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 See: <u>Breast MRI Indications (BR-5)</u> in the Breast Imaging Guidelines for evaluation of suspected breast cancer.

Breast Cancer – Initial Work-Up/Staging (ONC-11.2)

ON.BC.0011.2.C

Indication	Imaging Study
All newly diagnosed breast cancer or carcinoma in situ	 Diagnostic bilateral mammogram and/or Ultrasound Breast (CPT® 76641 or CPT® 76642) are imaging modalities of choice MRI Breast bilateral without and with contrast (CPT® 77049) (per Cigna Medical Coverage Policy - Oncology imaging Amendment (DV002))
ANY of the following:Ductal carcinoma in situStage I and II	 For planned sentinel lymph node (SLN) biopsy: Lymph system imaging (lymphoscintigraphy, CPT® 78195 SPECT/CT (CPT® 78830) is indicated as an add on code if requested
Stages I, II, and III	Routine systemic imaging is not indicated for initial staging of non-metastatic breast cancer in the absence of signs or symptoms
ANY of the following:	ANY or ALL of the following:
 Clinically suspected or biopsy proven metastatic/Stage IV disease Signs or symptoms of systemic disease Elevated liver function tests or tumor markers Inflammatory breast cancer (stage T4d) 4 or more axillary lymph nodes positive for cancer involvement 	
Inconclusive CT and/or bone scan	• PET/CT (CPT® 78815)

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Indication	Imaging Study
Bone pain	 Bone scan PET/CT (CPT® 78815) with Sodium Fluoride radiotracer may be obtained if CT, MRI, Bone scan and FDG PET/CT scan are inconclusive for bone metastases See: Bone (including Vertebral) Metastases (ONC-31.5) See: Spinal Cord Compression (ONC-31.6)

Breast Cancer – Restaging/Recurrence (ONC-11.3)

ON.BC.0011.3.C

Indication	Imaging Study
 ANY of the following: End of planned neoadjuvant chemotherapy to determine resectability Biopsy proven local recurrence Suspicion of recurrence with inconclusive mammogram and/or ultrasound (BIRADS 0) Mammogram and ultrasound conflicts with physical exam 	MRI Breast Bilateral without and with contrast (CPT® 77049)
After neoadjuvant chemotherapy, if sentinel node evaluation is planned	 Lymph system imaging (lymphoscintigraphy, CPT® 78195) SPECT/CT (CPT® 78830) is indicated as an add on code if requested
 ANY of the following: Assessing for residual disease after surgery Assessing response to neoadjuvant chemotherapy After lumpectomy or mastectomy, prior to adjuvant therapy 	Neither PET nor CT are indicated for systemic restaging after neoadjuvant chemotherapy or after surgery

Indication		Imaging Study
with metas measurabl • For ind chemot indicate • For ind or endo	response in individuals tatic disease and e disease on imaging: viduals receiving herapy, imaging is ed after every 2 cycles viduals receiving hormonal ecrine therapy, imaging is ed every 3 months	 CT Chest with contrast (CPT® 71260); and CT Abdomen and Pelvis with contrast (CPT® 74177) Bone scan PET/CT (CPT® 78815) with Sodium Fluoride radiotracer may be obtained if CT, MRI, Bone scan and FDG PET/CT scan are inconclusive for bone metastases In addition to the above options, for individuals receiving systemic treatment for brain metastases: MRI Brain without and with contrast (CPT® 70553)
Signs or sy	•	 CT Chest with contrast (CPT® 71260); and CT Abdomen and Pelvis with contrast (CPT® 74177) Bone scan PET/CT (CPT® 78815) with Sodium Fluoride radiotracer may be obtained if CT, MRI, Bone scan and FDG PET/CT scan are inconclusive for bone metastases For individuals with prior history of brain metastases or current signs or symptoms of brain metastasis: MRI Brain without and with contrast (CPT® 70553)
stage IV di metastase has not be compariso	stasis as the only site of sease (excluding brain s) and a prior bone scan en performed for serial n ve findings on CT/MRI scan	• PET/CT (CPT® 78815)

Breast Cancer – Surveillance/Follow- Up (ONC-11.4)

ON.BC.0011.4.C

Indication	Imaging Study
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 ANY or ALL of the following, every 3 months for up to 5 years after completion of active treatment: CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast Bone scan
 Asymptomatic non-metastatic disease Individuals receiving post-operative adjuvant therapy 	No advanced imaging indicated

Indication **Imaging Study** Breast surveillance in an individual Bilateral MRI Breast without and with contrast with prior history of breast cancer (CPT® 77049) annually (not treated with bilateral See also: **Breast MRI Indications (BR-5)** mastectomy) AND any one of the following: Known high risk genetic mutations: Li-Fraumeni Syndrome/TP53 Syndrome BRCA1 BRCA2 Peutz-Jehgers Syndrome (STK11/LKB1 gene variations) PTEN Mutation/Cowden Syndrome CDH1 NF1 PALB2 ATM CHEK2 NBN BARD1 RAD51C RAD51D Clinical lifetime risk estimated to be ≥20% using genetic or clinical risk estimator, calculated prior to the initial diagnosis of breast cancer Extremely dense breast tissue (breast density category D) on mammography

Age at diagnosis ≤50 years Individuals with a history of: Atypical ductal hyperplasia (ADH), atypical lobular hyperplasia (ALH), lobular carcinoma in situ (LCIS), or invasive lobular carcinoma

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Indication	Imaging Study
 Individuals treated with bilateral mastectomy All other individuals with no high risk features (as stated above) 	Breast MRI is not indicated for routine surveillance of asymptomatic individuals

References (ONC-11)

- 1. Gradishar WJ, Moran MS, Abraham J, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 4.2023 March 23, 2023. Breast cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/breast.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Breast Cancer V4.2023 March 23, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Cardoso F, Costa A, Norton L, et al. ESO-ESMO 2nd international consensus guidelines for advanced breast cancer (ABC2). Ann Oncol. 2014;25(10):1871-1888. doi:10.1093/annonc/mdu385.
- Khatcheressian JL, Hurley P, Bantug E, et al. Breast cancer follow-up and management after primary treatment: American Society of Clinical Oncology clinical practice guideline update. *J Clin Oncol*. 2013;31:961-965. doi:10.1200/JCO.2012.45.9859.
- 4. Puglisi F, Follador A, Minisini AM, et al. Baseline staging tests after a new diagnosis of breast cancer: further evidence of their limited indications. *Ann Oncol*. 2005;16(2):263-266. doi:10.1093/annonc/mdi063.
- Rong J, Wang S, Ding Q, Yun M, Zheng Z, Ye S. Comparison of 18 FDG PET-CT and bone scintigraphy for detection of bone metastases in breast cancer patients. A meta-analysis. *Surg Oncol.* 2013;22(2):86-91. doi:10.1016/j.suronc.2013.01.002.
- Hong S, Li J, Wang S. 18FDG PET-CT for diagnosis of distant metastases in breast cancer patients. A meta-analysis. Surg Oncol. 2013;22(2):139-143. doi:10.1016/j.suronc.2013.03.001.
- 7. Cheng X, Li Y, Liu B, Xu Z, Bao L, Wang J. 18F-FDG PET/CT and PET for evaluation of pathological response to neoadjuvant chemotherapy in breast cancer: a meta-analysis. Acta Radiol. 2012;53(6):615-627. doi:10.1258/ar.2012.110603.
- 8. Simos D, Catley C, van Walraven C, et al. Imaging for distant metastases in women with early-stage breast cancer: a population-based cohort study. *CMAJ*. 2015;187(12):E387-E397. doi:10.1503/cmaj.150003.
- Crivello ML, Ruth K, Sigurdson ER, et al. Advanced imaging modalities in early stage breast cancer: preoperative use in the United States Medicare population. *Ann Surg Oncol*. 2013;20(1):102-110. doi:10.1245/s10434-012-2571-4.
- Heller SL, Lourenco AP, Niell BL, et. al. ACR Appropriateness Criteria[®] Imaging after Mastectomy and Breast Reconstruction. Available at https://acsearch.acr.org/docs/3155410/Narrative/. American College of Radiology.
- 11. Landercasper J, Bailey L, Berry TS, et. al. Don't routinely order breast MRI in new breast cancer patients. American Society of Breast Surgeons. https://www.choosingwisely.org/clinician-lists/breast-surgeons-mris-in-new-breast-cancer-patients/
- 12. Chagpar AB, Howard-McNatt M, Chiba A, et. al. Factors affecting time to surgery in breast cancer patients. *Am Surg.* 2022;88(4):648-652. doi:10.1177/00031348211054714.
- 13. Peters NHGM, van Esser S, van den Bosch MAAJ, et. al. Preoperative MRI and surgical management in patients with nonpalpable breast cancer: the MONET randomized controlled trial. *Eur J Cancer*. 2011;47(6):879-886. doi:10.1016/j.ejca.2010.11.035
- 14. Turnbull L, Brown S, Harvey I, et. al. Comparative effectiveness of MRI in breast cancer (COMICE) trial: a randomized controlled trial. *Lancet*. 2010;375(9714):563-571. doi:10.1016/S0140-6736(09)62070-5.
- 15. Daly MB, Pal T, AlHilli Z, et. al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 February 13, 2023. Genetic/Familial High-Risk Assessment: Breast, Ovarian, and Pancreatic, available at: https://www.nccn.org/professionals/physician_gls/pdf/genetics_bop.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Genetic/Familial High-Risk Assessment: Breast, Ovarian, and Pancreatic V3.2023 February 13, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 16. Moncrieff M, Pywell S, Snelling A, et. al. Effectiveness of SPECT/CT imaging for sentinel node biopsy staging of primary cutaneous melanoma and patient outcomes. *Ann Surg Oncol*. 2022;29(2):767-775. doi:10.1245/s10434-021-10911-4.
- 17. Quartuccio N, Garau LM, Arnone A, et. al. Comparsion of 99mTc-labeled colloid SPECT/CT and planar lymphoscintigraphy in sentinel lymph node detection in patients with melanoma: a meta-analysis. *J Clin Med.* 2020;9(6):1680. doi:10.3390/jcm9061680.

18. Bennie G, Vorster M, Buscombe J, Sathekge M. The added value of a single-photon emission computed tomography-computed in sentinel lymph node mapping in patients with breast cancer and malignant melanoma. *World J Nucl Med.* 2015;14(01):41-46. doi:10.4103/1450-1147.150543

Sarcomas – Bone, Soft Tissue, and GIST (ONC-12)

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Bone and Soft Tissue Sarcomas – General Considerations (ONC-12.1)

ON.SS.0012.1.A

- Sarcomas are tumors of mesenchymal origin, classified as high-, intermediate-, and low-grade (G) tumors (sometimes described as "spindle cell" cancers). They can arise in any bony, cartilaginous, smooth muscle, skeletal muscle, or cardiac muscle tissue.
- Malignant nerve sheath tumor cell types should be imaged as high-grade sarcoma.
- Sarcomas occur in both adult and pediatric individuals, but some are more common in one age group than the other. Unless specified below, individuals age ≥18 years old should be imaged according to this guideline section.
- Exceptions include:
 - Rhabdomyosarcoma in individuals of all ages should be imaged according to guidelines in <u>Rhabdomyosarcoma (RMS) (PEDONC-8.2)</u> in the Pediatric Oncology Imaging Guidelines
 - Osteogenic sarcoma (Osteosarcoma) in individuals of all ages should be imaged according to guidelines in <u>Osteogenic Sarcoma (OS) (PEDONC-9.3)</u> in the Pediatric Oncology Imaging Guidelines
 - Ewing sarcoma and Primitive Neuroectodermal Tumor in individuals of all ages should be imaged according to guidelines in <u>Ewing Sarcoma and Primitive</u> <u>Neuroectodermal Tumors (ESFT) (PEDONC-9.4)</u> in the Pediatric Oncology Imaging Guidelines
 - Kaposi's sarcoma in individuals of all ages should be imaged according to guidelines in <u>Kaposi's Sarcoma (ONC-31.10)</u>
 - See: <u>Uterine Cancer (ONC-22)</u> for imaging recommendations for uterine sarcoma
 - Desmoplastic small round cell tumor in individuals of all ages should be imaged according to guidelines in <u>Non-Rhabdomyosarcoma Soft Tissue Sarcomas</u> (NRSTS) (PEDONC-8.3)

Soft Tissue Sarcomas – Initial Workup/Staging (ONC-12.2)

ON.SS.0012.2.A

	v1.0.2024
Indication	Imaging Study
Retroperitoneal or intra-abdominal primary site	 ANY or ALL of the following: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast Either CT Abdomen and Pelvis with contrast (CPT® 74177) or MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
ANY of the following:Extremity or trunk primary siteHead or neck primary site	 ANY or ALL of the following: MRI without and with contrast of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast
 ANY of the following: Angiosarcoma Alveolar soft part sarcoma Clear cell sarcoma Epithelioid sarcoma Hemangiopericytoma Leiomyosarcoma Other histologies documented to have propensity for lymphatic spread and deep-seated tumors 	 ANY or ALL of the following: MRI without and with contrast of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast Either CT Abdomen and Pelvis with contrast (CPT® 74177) or MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Myxoid round cell liposarcoma	 ANY or ALL of the following: MRI without and with contrast of involved area CT Chest with (CPT® 71260) or without contrast (CPT® 71250) Either CT Abdomen and Pelvis with contrast (CPT® 74177) or MRI Abdomen (CPT® 74183) and Pelvis (CPT® 72197) without and with contrast MRI Cervical/Thoracic/Lumbar Spine without and with contrast (CPT® 72156, CPT® 72157, and CPT® 72158)

Indication	Imaging Study
 ANY of the following: Angiosarcoma Alveolar soft part sarcoma All individuals with signs/symptoms of brain metastases 	MRI Brain without and with contrast (CPT® 70553)
 ANY of the following: Grade of tumor in doubt following biopsy Conventional imaging suggests solitary metastasis amenable to surgical resection 	PET/CT (CPT® 78815 or CPT® 78816)
Desmoid Tumors	 ONE of the following: CT without contrast or with contrast of the affected body part MRI without contrast or without and with contrast of the affected body part Imaging of lung, lymph node, and metastatic site for these tumors is not indicated
Dermatofibrosarcoma Protuberans (DFSP)	 ONE of the following: CT without contrast or with contrast of the affected body part MRI without contrast or without and with contrast of the affected body part CT Chest with (CPT® 71260) or without (CPT® 71250) contrast for: Pulmonary symptoms Abnormal Chest X-ray Sarcomatous differentiation

Soft Tissue Sarcomas – Restaging/Recurrence (ONC-12.3)

ON.SS.0012.3.A

	v1.0.2024
Indication	Imaging Study
 ANY of the following: After preoperative radiotherapy After surgical resection After adjuvant radiotherapy 	 MRI without and with contrast or CT with contrast of affected body area Chest or lymph node imaging is not indicated if no abnormality on previous imaging
 ANY of the following: Differentiate tumor from radiation or surgical fibrosis Determine response to neoadjuvant therapy Confirm oligometastatic disease prior to curative intent surgical resection 	• PET/CT (CPT® 78815 or CPT® 78816)
Chemotherapy response for individuals with measurable disease	CT with contrast or MRI without and with contrast of affected body area every 2 cycles
Local recurrence suspected	Repeat all imaging for initial workup of specific histology and/or primary site
Preoperative planning prior to resection	 ANY or ALL of the following: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area
Dermatofibrosarcoma Protuberans (DFSP)	 CT without contrast or with contrast of the affected body part or MRI without contrast or without and with contrast of the affected body part CT Chest with (CPT® 71260) or without (CPT® 71250) contrast for: Pulmonary symptoms Abnormal Chest X-ray Sarcomatous differentiation

Soft Tissue Sarcomas Surveillance/Follow-up (ONC-12.4)

ON.SS.0012.4.A

Indication	Imaging Study
Retroperitoneal/intra- abdominal primary site	 ANY or ALL of the following every 3 months for 2 years, then every 6 months for 2 more years, then annually: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast or MRI without and with contrast of any other involved body areas
Extremity, trunk, or Head/Neck primary site, low- grade Stage I disease	ANY or ALL of the following every 6 months for 5 years, then annually thereafter: • Chest x-ray
	CT Chest with (CPT® 71260) or without (CPT® 71250) contrast is indicated for new findings on Chest X-ray or new/worsening pulmonary signs/ symptoms
	CT with contrast, MRI without contrast, or MRI without and with contrast of primary site if primary site not easily evaluated by physical exam
 ANY of the following: Extremity/trunk primary site - grade II/stage II or higher Head/neck primary site 	 ANY or ALL of the following every 3 months for 2 years, then every 6 months for 2 more years, then annually: CT with contrast, MRI without contrast, or MRI without and with contrast of primary site CT Chest with (CPT® 71260) or without (CPT® 71250) contrast
	CT with contrast or MRI without and with contrast of any other involved body areas
Desmoid tumors	 ONE of the following every 6 months for 3 years, then annually: CT without contrast or with contrast of the affected body part MRI without contrast or without and with contrast of the affected body part
Dermatofibrosarcoma Protuberans	No routine imaging unless clinical signs/symptoms of recurrence

Gastrointestinal Stromal Tumor (GIST) (ONC-12.5)

ON.SS.0012.5.A

v1.0.2024

General Considerations

GISTs are mesenchymal neoplasms of the gastrointestinal (GI) tract, mostly found
in the stomach and upper small bowel, commonly metastasizing to the liver and
abdominal cavity and primarily treated with surgery.

Indication	Imaging Study
Suspected/Diagnosis	CT Abdomen and Pelvis with contrast (CPT® 74177)
Initial Work-up/Staging	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast MRI Abdomen without and with contrast (CPT® 74183) is indicated for evaluation of liver lesions that are equivocal on CT imaging or for preoperative assessment of liver PET (CPT® 78815) is indicated for evaluation of inconclusive findings on conventional imaging
Restaging/Recurrence	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) if prior evidence of chest disease or signs or symptoms of chest disease PET (CPT® 78815) is indicated for evaluation of inconclusive findings on conventional imaging
Monitoring response to treatment (every 8 to 12 weeks) in either of the following: Unresectable primary disease Metastatic disease	CT Abdomen and Pelvis with contrast (CPT® 74177)
 Prior evidence of chest disease Signs or symptoms of chest disease 	CT Chest with contrast (CPT® 71260)
Evaluation of inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

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Indication	Imaging Study
Surveillance/Follow-up	 CT Abdomen and Pelvis with contrast (CPT[®] 74177) every 6 months for 5 years, then annually

Bone Sarcomas - Initial Work-Up/Staging (ONC-12.6)

ON.SS.0012.6.C

Indication	Imaging Study
 Chondrosarcoma Low-grade intracompartmental High-grade (grade II or grade III) Clear cell Extracompartmental 	 ANY or ALL of the following: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast
Dedifferentiated chondrosarcoma	See: Osteogenic Sarcoma (OS) (PEDONC-9.3) for imaging recommendations
Mesenchymal chondrosarcoma	See: Ewing's Sarcoma Family of Tumors (PEDONC-9.4) for imaging recommendations
Chordoma	 ANY or ALL of the following: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), and Lumbar (CPT® 72158) Spine without and with contrast Bone scan PET may be approved for inconclusive conventional imaging

Bone Sarcomas - Restaging/Recurrence (ONC-12.7)

ON.SS.0012.7.C

Indication	Imaging Study
 Chondrosarcoma Low-grade intracompartmental High-grade (grade II or grade III) Clear cell Extracompartmental 	 ANY or ALL of the following, after completion of radiotherapy or every 2 cycles of chemotherapy: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast
Dedifferentiated chondrosarcoma	See: Osteogenic Sarcoma (OS) (PEDONC-9.3) for imaging recommendations
Mesenchymal chondrosarcoma	See: Ewing's Sarcoma Family of Tumors (PEDONC-9.4) for imaging recommendations
Chordoma	 ANY or ALL of the following, after completion of radiotherapy or every 2 cycles of chemotherapy: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area Bone scan PET may be approved for inconclusive conventional imaging

Bone Sarcomas – Surveillance/Followup (ONC-12.8)

ON.SS.0012.8.A

	V1.0.2024
Indication	Imaging Study
 Grade I Chondrosarcoma Intra-compartmental Chondrosarcoma 	 ANY or ALL of the following every 6 months for 2 years, then annually for 10 years: Plain x-ray of primary site MRI without and with contrast is indicated for new findings on plain x-ray or new/worsening clinical symptoms. Chest x-ray CT Chest with (CPT® 71260) or without (CPT® 71250) contrast for new findings on chest x-ray, or new/worsening signs/symptoms
 Grade II or III Chondrosarcoma Clear Cell Chondrosarcoma Extra-compartmental Chondrosarcoma 	 ANY or ALL of the following every 6 months for 5 years, then annually for 10 years: Plain x-ray of primary site MRI without and with contrast is indicated for new findings on plain x-ray or new/worsening clinical symptoms. Chest x-ray or CT Chest with (CPT® 71260) or CT Chest without (CPT® 71250) contrast
Dedifferentiated chondrosarcoma	See: Osteogenic Sarcoma (OS) (PEDONC-9.3) for imaging recommendations
Mesenchymal chondrosarcoma	See: Ewing's Sarcoma Family of Tumors (PEDONC-9.4) for imaging recommendations
Chordoma	 Plain x-ray of primary site every 6 months for 5 years and then annually until year 10 MRI without and with contrast is indicated for new findings on plain x-ray or new/worsening clinical symptoms. Chest x-ray every 6 months for 5 years and then annually until year 10 CT Chest with (CPT® 71260) or without (CPT® 71250) contrast may be obtained annually or for evaluation of any new findings on chest x-ray or new/worsening signs/symptoms

Benign Bone Tumors – General Considerations (ONC-12.9)

ON.SS.0012.9.A

- Variety of diagnoses, including osteoid osteochondroma, chondroblastoma, desmoplastic fibroma, Paget's disease, osteoid osteoma and others.
- Plain x-ray appearance is diagnostic for many benign bone tumors and advanced imaging is generally unnecessary except for preoperative planning.
- MRI without and with contrast is the primary modality for advanced imaging of bone tumors, and can be approved to help narrow differential diagnoses and determine whether biopsy is indicated.
- Some benign bone tumor types carry a risk of malignant degeneration over time, but routine advanced imaging surveillance has not been shown to improve outcomes for these individuals.
- MRI without and with contrast can be approved to evaluate new findings on Plain x-ray new/worsening clinical symptoms not explained by a recent Plain x-ray.
- There are no data to support the use of PET/CT in the evaluation of benign bone tumors, and PET requests should not be approved without biopsy confirmation of a malignancy.
- Other benign bone tumors should be imaged according to guidelines in <u>Lesion of Bone (MS-10.1)</u> in the General Musculoskeletal Imaging Guidelines or <u>Mass Involving Bone (including Lytic and Blastic Metastatic Disease) (PEDMS-3.4)</u> in the Pediatric Musculoskeletal Imaging Guidelines.

Benign Bone Tumors - Initial Work-Up/ Staging (ONC-12.10)

ON.SS.0012.10.C

Indication	Imaging Study
Giant Cell Tumor of Bone (GCTB)	 ANY or ALL of the following: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area CT Chest with (CPT® 71260) or without (CPT® 71250) contrast Bone scan
Enchondroma	MRI without contrast or without and with contrast of primary site

Benign Bone Tumors - Restaging/Recurrence (ONC-12.11)

ON.SS.0012.11.C

Indication	Imaging Study
Giant Cell Tumor of Bone (GCTB)	 ANY or ALL of the following, after completion of radiotherapy or every 2 cycles of chemotherapy: MRI without contrast or without and with contrast of involved area CT (contrast as requested) of involved area Bone scan
Enchondroma	Plain films of primary site

Benign Bone Tumors – Surveillance/Follow-up (ONC-12.12)

ON.SS.0012.12.A

Indication	Imaging Study	
Giant Cell Tumor of Bone (GCTB)	ANY or ALL of the following every 6 months for 4 years, then annually thereafter: Plain x-ray of primary site	
	 MRI without and with contrast is indicated for new findings on plain x-ray or new/worsening clinical symptoms. Chest x-ray 	
	 CT Chest with (CPT® 71260) or without (CPT® 71250) contrast for new findings on chest x-ray, or new/worsening signs/symptoms. 	
Enchondroma	Plain films of primary site	

References (ONC-12)

- 1. Mehren MV, Kane III JM, Armstrong SA, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 April 25, 2023. Soft Tissue Sarcoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/sarcoma.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Soft Tissue Sarcoma V2.2023 April 25, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Biermann JS, Hirbe A, Agulnik M, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 April 4, 2023. Bone cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/bone.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Bone cancer V3.2023 April 4, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Von Mehren M, Kane III JM, Armstrong SA, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 March 13, 2023. Gastrointestinal Stromal Tumors (GISTs), available at: https://www.nccn.org/professionals/physician_gls/pdf/gist.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Gastrointestinal Stromal Tumors (GISTs) V1.2023 March 13, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN™ Guidelines, go online to NCCN.org
- Nishiguchi T, Mochizuki K, Ohsawa M, et al, Differentiating benign notochordal cell tumors from chordomas: radiographic features on MRI, CT, and tomography. *Am Jour Roentgenol*. 2011;196(3):644-650. doi:10.2214/AJR.10.4460.
- 5. Van den Abbeele AD. The lessons of GIST-PET and PET/CT: a new paradigm for imaging. *Oncologist*. 2008;13:8-13. doi:10.1634/theoncologist.13-S2-8.
- Demetri GD, von Mehren M, Antonescu CR, et al. NCCN Task Force report: update on the management of
 patients with gastrointestinal stromal tumors. J Natl Compr Canc Netw. 2010;8(Suppl 2):S42-44.
- Peng PD, Hyder O, Mavros MN, et al. Management and recurrence patterns of desmoids tumors: a multi-institutional analysis of 211 patients. *Ann Surg Oncol*. 2012;19(13):4036-4042. doi:10.1245/s10434-012-2634-6
- 8. Tseng WW, Amini B, Madewell JE. Follow-up of the soft tissue sarcoma patient. *J Surg Oncol.* 2015;111(5):641-645. doi:10.1002/jso.23814.
- 9. Grotz TE, Donohue JH. Surveillance strategies for gastrointestinal stromal tumors. *J Surg Oncol*. 2011;104(8):921-927. doi:10.1002/jso.21862.
- Akram J, Wooler G, Lock-Andersen J. Dermatofibrosarcoma protuberans: clinical series, national Danish incidence data and suggested guidelines. *J Plast Surg Hand Surg*. 2014;48(1):67-73. doi:10.3109/2000656X.2013.812969.
- 11. Puri A, Gulia A, Hawaldar R, Ranganathan P, Badwe RA. Does intensity of surveillance affect survival after surgery for sarcomas? Results of a randomized noninferiority trial. *Clin Orthop Relat Res*. 2014;472(5):1568-1575. doi:10.1007/s11999-013-3385-9.
- 12. Biermann JS, Adkins DR, Aqulnik M, et al. Bone cancer. J Natl Compr Canc Netw. 2013;11(6):688-723.

Pancreatic Cancer (ONC-13)

Guideline	Page
Pancreatic Cancer – General Considerations (ONC-13.0) Pancreatic Cancer – Screening Studies for Pancreatic Cancer (ONC 143	
Pancreatic Cancer – Suspected/Diagnosis (ONC-13.2) Pancreatic Cancer – Initial Work-up/Staging (ONC-13.3)	
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Pancreatic Cancer – General Considerations (ONC-13.0)

ON.PC.0013.0.A

- This guideline refers only to adenocarcinoma of the exocrine pancreas, which accounts for over 90% of pancreatic malignancies. This guideline may also be used for cancer of the Ampulla of Vater.
- Neuroendocrine and carcinoid tumors of the pancreas are not included in this guideline, see: <u>Neuroendocrine Cancers and Adrenal Tumors (ONC-15)</u>.

Pancreatic Cancer – Screening Studies for Pancreatic Cancer (ONC-13.1)

ON.PC.0013.1.A

- Detailed history of any known inherited syndrome in the individual and detailed family history in first- and second-degree relatives, including the age and lineage, is essential to guide screening recommendations. See table below for age- and riskspecific screening recommendations
- New onset of diabetes in individuals older than 50 has been recognized as a potential indicator of the development of pancreatic cancer. Approximately 1% of individuals in this category are diagnosed with cancer within 3 years. A prediction model has been established which identifies those individuals at greatest risk for pancreatic malignancy. The scoring system, known as ENDPAC (Enriching New-Onset Diabetes for Pancreatic Cancer) is based on 3 discriminatory factors, including change in blood glucose, change in weight, and age of onset at the time of the new diagnosis of diabetes. A score of >3 imparts an elevated risk of pancreatic cancer (3.6%), and these individuals should be screened. Screening is not indicated at this time for scores of 0-2.

not indicated at time time for coords of a 2.			
Indications	Imaging Study		
 Individuals who meet BOTH of the following criteria: One or more first- or second-degree relative affected with pancreatic adenocarcinoma AND Known mutation carrier of ONE of the following genes: Lynch Syndrome (MLH1, MSH2, or MSH6 gene mutations) BRCA1, BRCA2 (Familial Breast and Ovarian syndrome) PALB2 mutation ATM (Ataxia-Telangiectasia) 	MRI Abdomen without and with contrast (CPT® 74183) starting at age 50 or 10 years earlier than the youngest affected family member, repeat annually		
 Individuals with family history of pancreatic cancer but no known genetic mutation: Individuals with 2 relatives with pancreatic adenocarcinoma where one is a first-degree relative Individuals with 3 or more relatives with pancreatic adenocarcinoma 	 MRI Abdomen without and with contrast (CPT® 74183) starting at age 45 or 10 years earlier than the youngest affected family member, repeat annually 		

Pancreatic Cancer Kindred (individuals who have at least one first-degree relative with pancreatic adenocarcinoma who in turn also has a first-degree relative with pancreatic adenocarcinoma) and NO known genetic germline mutations Hereditary Pancreatitis (PRSS1, CPA1, and CTRC gene mutations) Hereditary Pancreatitis (PRSS1, CPA1, and CTRC gene mutations) Peutz-Jeghers Syndrome (LKB1/STK11 gene mutation) CDKN2A mutation (also known as p16, p16INK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole Syndrome) Screening MRI reveals cystic lesion of the pancreas Screening MRI reveals indeterminate solid lesion Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without and with contrast (CPT® 74183) in 3 months after the CT scan New onset diabetes in adults with ENDPAC score of ≥3 MRI Abdomen without and with contrast (CPT® 74183) beginning at age 40, repeat annually. MRI Abdomen without and with contrast or MRCP (CPT® 74183) beginning at age 40, repeat annually. **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan		
who have at least one first-degree relative with pancreatic adenocarcinoma who in turn also has a first-degree relative with pancreatic adenocarcinoma) and NO known genetic germline mutations Hereditary Pancreatitis (PRSS1, CPA1, and CTRC gene mutations) Peutz-Jeghers Syndrome (LKB1/STK11 gene mutation) CDKN2A mutation (also known as p16, p16INK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole Syndrome) Screening MRI reveals cystic lesion of the pancreas Screening MRI reveals indeterminate solid lesion Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without a mass We onset diabetes in adults with ENDPAC score of ≥3 We Also starting at age 50 or 10 years earlier than the youngest affected family member, repeat annually (CPT* 74183) starting at age 40 or 20 years after the first pancreatitis attack, repeat annually. MRI Abdomen without and with contrast or MRCP (CPT* 74183) beginning at age 30, repeat annually. MRI Abdomen without and with contrast or MRCP (CPT* 74183) beginning at age 40, repeat annually. **OTAbdomen with contrast - pancreatic protocol (CPT* 74160) **May repeat MRI Abdomen without and with contrast (CPT* 74160) **May repeat MRI Abdomen without and with contrast (CPT* 74160) **May repeat MRI Abdomen without and with contrast (CPT* 74183) in 3 months after the CT scan **CT Abdomen without and with contrast (CPT* 74183) in 3 months after the CT scan **New onset diabetes in adults with ENDPAC score of ≥3 **CT Abdomen without and with contrast (CPT* 74183) at baseline; if negative, can be repeated	Indications	Imaging Study
(CPT® 74183) beginning at age 40 or 20 years after the first pancreatitis attack, repeat annually. Peutz-Jeghers Syndrome (LKB1/STK11 gene mutation) CDKN2A mutation (also known as p16, p16lNK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole Syndrome) Screening MRI reveals cystic lesion of the pancreas Screening MRI reveals indeterminate solid lesion CT Abdomen without and with contrast or MRCP (CPT® 74183) beginning at age 40, repeat annually. Repeat MRI Abdomen without and with contrast (CPT® 74183) in 6 months CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without and with contrast (CPT® 74183) in 3 months after the CT scan New onset diabetes in adults with ENDPAC score of ≥3 CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan	who have at least one first-degree relative with pancreatic adenocarcinoma who in turn also has a first-degree relative with pancreatic adenocarcinoma) and NO known	(CPT® 74183) starting at age 50 or 10 years earlier than the youngest affected
(LKB1/STK11 gene mutation) (CPT® 74183) starting at age 30, repeat annually CDKN2A mutation (also known as p16, p16INK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole Syndrome) Screening MRI reveals cystic lesion of the pancreas Screening MRI reveals indeterminate solid lesion CT Abdomen with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without a mass New onset diabetes in adults with ENDPAC score of ≥3 (CPT® 74183) starting at age 30, repeat annually MRI Abdomen without and with contrast or MRCP (CPT® 74183) beginning at age 40, repeat annually. CT Abdomen without and with contrast — pancreatic protocol (CPT® 74183) in 3 months after the CT scan CT Abdomen with contrast — pancreatic protocol (CPT® 74180) May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan		(CPT® 74183) beginning at age 40 or 20 years after the first pancreatitis attack,
(also known as p16, p16INK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole Syndrome) Screening MRI reveals cystic lesion of the pancreas Screening MRI reveals indeterminate solid lesion **CT Abdomen with contrast − pancreatic protocol (CPT® 74183) in 3 months after the CT scan **Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without a mass **NRCP (CPT® 74183) beginning at age 40, repeat annually. **Repeat MRI Abdomen without and with contrast (CPT® 74183) in 6 months **CT Abdomen with contrast − pancreatic protocol (CPT® 74183) in 3 months after the CT scan **CT Abdomen with contrast − pancreatic protocol (CPT® 74160) **May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan **NRCP (CPT® 74183) beginning at age 40, repeat annually. **CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan **NRCP (CPT® 74183) beginning at age 40, repeat annually. **CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan **NRCP (CPT® 74183) beginning at age 40, repeat annually. **CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan **NRCP (CPT® 74183) beginning at age 40, repeat annually.	,	(CPT® 74183) starting at age 30, repeat
the pancreas Screening MRI reveals indeterminate solid lesion • CT Abdomen with contrast – pancreatic protocol (CPT® 74183) in 3 months after the CT scan Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without a mass • CT Abdomen with contrast – pancreatic protocol (CPT® 74183) in 3 months after the CT scan • CT Abdomen with contrast – pancreatic protocol (CPT® 74160) • May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan New onset diabetes in adults with ENDPAC score of ≥3 • CT Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan	(also known as p16, p16INK4a, and MTS1, FAMM-Familial Atypical Multiple Melanoma and Mole	MRCP (CPT® 74183) beginning at age 40,
solid lesion protocol (CPT® 74160) May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan Screening MRI reveals pancreatic stricture and/or dilation ≥6 mm without a mass CT Abdomen with contrast – pancreatic protocol (CPT® 74160) May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan New onset diabetes in adults with ENDPAC score of ≥3 CT Abdomen without and with contrast (CPT® 74183) or MRI Abdomen without and with contrast (CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183) at baseline; if negative, can be repeated		·
stricture and/or dilation ≥6 mm without a mass • May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months after the CT scan New onset diabetes in adults with ENDPAC score of ≥3 • CT Abdomen without and with contrast (CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183) at baseline; if negative, can be repeated		 protocol (CPT® 74160) May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months
ENDPAC score of ≥3 (CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183) at baseline; if negative, can be repeated	stricture and/or dilation ≥6 mm without	 protocol (CPT® 74160) May repeat MRI Abdomen without and with contrast (CPT® 74183) in 3 months
		(CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183) at baseline; if negative, can be repeated

Pancreatic Cancer – Suspected/Diagnosis (ONC-13.2)

ON.PC.0013.2.A

V 1.0.20	
Indication	Imaging Study
For any suspected symptoms only (e.g. epigastric pain, weight loss, pain radiating to back, etc.)	 Ultrasound (CPT® 76700 or CPT® 76705) Also see: Epigastric Pain and Dyspepsia (AB-2.5)
Symptoms suspicious for pancreatic cancer AND any one of the following: • Abnormal labs (e.g. elevated CA 19-9, ALKP, bilirubin, or GGTP) • Abnormal physical exam findings (e.g. abdominal mass) • Abnormal or non-diagnostic ultrasound/ERCP	 Any ONE of the following: CT Pancreatic Protocol (CT Abdomen with contrast with dual phase imaging, CPT® 74160) MRI Abdomen without and with contrast (CPT® 74183)
Preoperative studies for potentially resectable tumors without confirmed histologic diagnosis	See: Pancreatic Cancer – Initial Work-up/Staging (ONC-13.3)

Pancreatic Cancer – Initial Work-up/Staging (ONC-13.3)

ON.PC.0013.3.A

Indication	Imaging Study
All individuals	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with (CPT® 74177) or CT Abdomen and Pelvis without and with contrast (CPT® 74178) EUS
 For any of the following: Preoperative planning CT insufficient to determine resectability Evaluation of indeterminate liver lesions 	MRI Abdomen without and with contrast (CPT® 74183)
No evidence of metastatic disease on CT or MRI AND any of the following high-risk features: • Borderline resectable disease • Markedly elevated CA 19-9 • Large primary tumor(s) • Enlarged regional lymph nodes	• PET/CT (CPT® 78815)

Pancreatic Cancer – Restaging/Recurrence (ONC-13.4)

ON.PC.0013.4.A

V1.0.202-	
Indication	Imaging Study
 For ANY of the following: After neoadjuvant chemoradiation Post-operative baseline Suspected recurrence 	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with (CPT® 74177) or CT Abdomen and Pelvis without and with contrast (CPT® 74178) CT with contrast of other involved or symptomatic areas
Unresectable disease or metastatic disease on chemotherapy	 Every 2 cycles of treatment (commonly every 6 to 8 weeks): CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with (CPT® 74177) or CT Abdomen and Pelvis without and with contrast (CPT® 74178) CT with contrast of other involved or symptomatic areas
Unexplained elevated liver enzymes or inconclusive recent CT abnormality	MRI Abdomen without and with contrast (CPT® 74183)
If complete surgical resection was initial therapy	See: Pancreatic Cancer – Surveillance/Follow-up for surveillance imaging (ONC-13.5)

Pancreatic Cancer – Surveillance/Follow-up (ONC-13.5)

ON.PC.0013.5.A

Indication	Imaging Study
All individuals	 Every 3 months for 2 years, then annually: CT Chest with contrast (CPT® 71260) And ANY ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen without and with contrast (CPT® 74183) and MRI Pelvis without and with contrast (CPT® 72197)
Measurable metastatic disease on maintenance therapy or being monitored off therapy	Every 3 months for up to 5 years after completion of definitive treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)

References (ONC-13)

- 1. Tempero MA, Malafa MP, Al-Hawary M, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 June 19, 2023. Pancreatic Adenocarcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/pancreatic.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Pancreatic Adenocarcinoma V2.2023 June 19, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Syngal S, Brand RE, Church JM, et al. ACG clinical guideline: genetic testing and management of hereditary gastrointestinal cancer syndromes. *Am. J. Gastroenterol.* 2015;110(2):223-262. doi:10.1038/ajg.2014.435.
- 3. Canto MI, Harinck F, Hruban RH, et al. International Cancer of the Pancreas Screening (CAPS) consortium summit on the management of patients with increased risk for familial pancreatic cancer. *Gut.* 2013;62(3):339-347. doi:10.1136/gutjnl-2012-303108.
- 4. U.S. Preventive Services Task Force. *Screening for pancreatic cancer: recommendation statement.* Rockville, Maryland: Agency for Healthcare Research and Quality (AHRQ); 2004.
- Heinrich S, Goerres GW, Schafer M, et al. Positron emission tomography/computed tomography influences on the management of resectable pancreatic cancer and its cost-effectiveness. *Ann Surg*. 2005;242(2):235-243.
- 6. Gemmel C, Eickhoff A, Helmstädter L, Riemann JF. Pancreatic cancer screening: state of the art. *Expert Rev Gastroenterol Hepatol*. 2009;3(1):89-96. doi:10.1586/17474124.3.1.89.
- 7. Al-Hawary MM, Francis IR, Chari ST, et al. Pancreatic ductal adenocarcinoma radiology reporting template: consensus statement of the Society of Abdominal Radiology and the American Pancreatic Association. *Gastroenterology*. 2014;146(1):291-304. doi:10.1053/j.gastro.2013.11.004.
- 8. Tersmette AC, Petersen GM, Offerhaus GJ. Increased risk of incident pancreatic cancer among first-degree relatives of patients with familial pancreatic cancer. *Clin Cancer Res.* 2001;7(3):738-44.
- Tzeng CW, Abbott DE, Cantor SB et al. Frequency and intensity of postoperative surveillance after curative treatment of pancreatic cancer: a cost-effectiveness analysis. *Ann Surg Oncol*. 2013;20(7):2197-2203. doi:10.1245/s10434-013-2889-6.
- 10. Furman MJ, Lambert LA, Sullivan ME, Whalen GF. Rational follow-up after curative cancer resection. *Journal of Clinical Oncology*. 2013;31(9):1130-1133. doi:10.1200/JCO.2012.46.4438.
- 11. Tzeng C, Fleming J, Lee J, et al. Yield of clinical and radiographic surveillance in patients with resected pancreatic adenocarcinoma following multimodal therapy. *HPB*. 2012;14(6):365-372. doi:10.1111/j.1477-2574.2012.00445.x.
- 12. Sharma, A, Kandlakunta H, Nagpal SJS, et.al. Model to determine risk of pancreatic cancer in patients with new-onset diabetes. *Gastroenterology*. 2018;155(3):730-739.
- 13. Goggins M, Overbeek KA, Brand R, et. al. Management of patients with increased risk for familial pancreatic cancer: updated recommendations from the International Cancer of the Pancreas Screening Consortium. *Gut.* 2020;69(1):7-17. doi:10.1136/gutjnl-2019-319352.

Upper GI Cancers (ONC-14)

Guideline	је
Hepatocellular Carcinoma (HCC) – General Considerations (ONC-14.1)	
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Hepatocellular Carcinoma (HCC) – General Considerations (ONC-14.1)

ON.GI.0014.1.A

- A biopsy is not always required for the diagnosis of Hepatocellular carcinoma (HCC). A dedicated triple-phase CT or MRI may be obtained. MRI with contrast is the test of choice for the evaluation of liver masses. It offers soft tissue contrast resolution superior to CT as well as the possibility of using two different contrast agents, one of which if more blood flow based and the other which also is blood flow based and demonstrates hepatobiliary function (Eovist). Classical imaging findings include:
 - Arterial phase hyperenhancement
 - Venous phase washout appearance
 - Capsule appearance
 - o Threshold growth
- For individuals who are high-risk for developing HCC (cirrhosis, chronic Hepatitis B or current or prior HCC), if the liver lesion is >1 cm with 2 classic enhancements on triple-phase CT or MRI, the diagnosis is confirmatory and biopsy is not needed.
- For lesions less than 1 cm or with less than 2 classical enhancements or for any liver lesions in individuals who are not high-risk, a biopsy is needed for histological confirmation. PET/CT scan is considered experimental, investigational, or unproven for the diagnosis or staging of HCC

Hepatocellular Carcinoma (HCC) – Suspected/Diagnosis (ONC-14.2)

ON.GI.0014.2.A

- See: <u>Chronic Liver Disease</u>, <u>Cirrhosis and Screening for HCC (AB-26.1)</u> in the Abdomen Imaging Guidelines
- See: Liver Lesion Characterization (AB-29.1) in the Abdomen Imaging Guidelines

Hepatocellular Carcinoma (HCC) – Initial Work-up/Staging (ONC-14.3)

ON.GI.0014.3.A

Indication	Imaging Study
All individuals	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250)
	And ONE of the following:
	CT Abdomen with contrast (CPT® 74160)
	CT Abdomen without and with contrast (CPT® 74170)
	CT Abdomen and Pelvis with contrast (CPT® 74177) or without and with contrast (CPT® 74178)
	MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast

Hepatocellular Carcinoma (HCC) - Restaging/Recurrence (ONC-14.4)

ON.GI.0014.4.C

Indication	Imaging Study
 ONE of the following: After initial therapy For suspected recurrence Individuals receiving systemic therapy (every 2 cycles) 	 CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) ONE of the following: CT Abdomen with contrast (CPT® 74160) CT Abdomen without and with contrast (CPT® 74170) CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Hepatocellular Carcinoma treated with embolization	 CTA Abdomen (CPT® 74175) can be approved immediately prior to embolization ONE of the following, immediately prior to and 1 month post-ablation: CT Abdomen without and with contrast (CPT® 74170) MRI Abdomen without and with contrast (CPT® 74183) See: Liver Metastases (ONC-31.2) for imaging studies indicated prior to and post-embolization
Hepatocellular Carcinoma awaiting liver transplant	See: Liver Transplant, Pre-Transplant (AB-42.1) in the Abdomen Imaging Guidelines

Hepatocellular Carcinoma (HCC) – Surveillance/Follow-up (ONC-14.5)

ON.GI.0014.5.A

Indication	Imaging Study
Hepatocellular Carcinoma: Treated with surgical resection Treated with embolization Being monitored off therapy	 Every 3 months for 2 years, then every 6 months until year 5: CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) And ONE of the following: CT Abdomen with contrast (CPT® 74160) CT Abdomen without and with contrast (CPT® 74170) CT Abdomen and Pelvis with contrast (CPT® 74177) or without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and Pelvis (CPT® 72197) without and with contrast
Hepatocellular Carcinoma treated with liver transplant	See: <u>Liver Transplant, Post-transplant Imaging (AB-42.3)</u> in the Abdomen Imaging Guidelines

Gallbladder and Biliary Tumors – Initial Work-up/Staging (ONC-14.6)

ON.GI.0014.6.A

Indication	Imaging Study
All individuals	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250)
	And ONE of the following:
	CT Abdomen with contrast (CPT® 74160)
	CT Abdomen without and with contrast (CPT® 74170)
	CT Abdomen and Pelvis with contrast (CPT® 74177)
	MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Gallbladder and Biliary Tumors – Restaging/Recurrence (ONC-14.7)

ON.GI.0014.7.A

Indication	Imaging Study
 ANY of the following: After initial therapy For suspected recurrence or new liver lesions Individuals receiving systemic chemotherapy (every 2 cycles) 	 CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) And ONE of the following: CT Abdomen with contrast (CPT® 74160) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Gallbladder and Biliary Tumors – Surveillance/Follow-up (ONC-14.8)

ON.GI.0014.8.A

Indication	Imaging Study
All individuals	 Every 6 months for 2 years, and then annually up to year 5: CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) And ONE of the following: CT Abdomen with contrast (CPT® 74160) CT Abdomen without and with contrast (CPT® 74170) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Biliary carcinoma treated with liver transplant	See: Liver Transplant, Post-transplant Imaging (AB-42.3) in the Abdomen Imaging Guidelines

Gastric Cancer – Initial Work-up/Staging (ONC-14.9)

ON.GI.0014.9.A

Indication	Imaging Study
All individuals	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Gastric cancer ≥T2 or higher with no metastatic disease by conventional imaging	• PET/CT (CPT® 78815)

Gastric Cancers - Restaging/Recurrence (ONC-14.10)

ON.GI.0014.10.C

Indication	Imaging Study
 After initial therapy for presumed surgically resectable disease Post curative chemoradiation being treated without surgery For suspected recurrence 	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Monitoring response to chemotherapy (every 2 cycles, ~every 6-8 weeks) for: • Unresected primary disease • Metastatic disease	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for: New/worsening pulmonary symptoms Abnormal chest x-ray findings Known prior pulmonary involvement
New liver lesion(s) and primary site controlled	CT Abdomen without and with contrast (CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183)
 ONE of the following: After neoadjuvant therapy for presumed surgically resectable disease or Post curative chemoradiation being treated without surgery 	CT Chest with contrast (CPT® 71260)and CT Abdomen and Pelvis with contrast (CPT® 74177)
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Gastric Cancer – Surveillance/Follow-up (ONC-14.11)

ON.GI.0014.11.A

Indication	Imaging Study
Stage I (treated with resection alone)	No routine imaging unless clinical signs/ symptoms of recurrence
 ANY of the following: Stage I treated with systemic therapy Stages II-III Stage IV - Metastatic disease (post definitive treatment of all measurable disease or being observed off therapy) 	 Every 6 months for 2 years, and then annually for 3 more years: CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)

References (ONC-14)

- 1. Ajani JA, D'Amico TA, Bentrem DJ et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 March 10, 2023. Gastric cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/gastric.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Gastric cancer V1.2023 March 10, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Benson AB, D'Angelica MI, Abrams T, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023—March 10, 2023, Hepatocellular Carcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/hcc.pdf, Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Hepatocellular Carcinoma V1.2023 3/10/2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Benson AB, D'Angelica MI, Abrams T, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023—May 10, 2023, Biliary Tract Cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/btc.pdf, Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Biliary Tract Cancers V2.2023 5/10/2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org
- Vallböhmer D, Hölscher AH, Schnieder PM, et al. [18F]-fluorodeoxyglucose-positron emission tomography for the assessment of histopathologic response and prognosis after completion of neoadjuvant chemotherapy in gastric cancer. *J Surg Oncol.* 2010;102(2):135-140. doi:10.1002/jso.21592.
- 5. Zou H, Zhao Y. 18FDG PET-CT for detecting gastric cancer recurrence after surgical resection: a meta-analysis. *Surg Oncol*. 2013;22(3):162-166. doi:10.1016/j.suronc.2013.05.001.
- 6. Bridgewater J, Galle PR, Khan SA, et al. Guidelines for the diagnosis and management of intrahepatic cholangiocarcinoma. *J Hepatol.* 2014;60(6):1268-1289. doi:10.1016/j.jhep.2014.01.021.
- Khan SA, Davidson BR, Goldin RD, et al. Guidelines for the diagnosis and treatment of cholangiocarcinoma: an update. Gut. 2012;61(12):1657-1669. doi:10.1136/gutjnl-2011-301748.
- 8. Benson AB 3rd, D'Angelica MI, Abrams TA, et al. Hepatobiliary cancers, version 2.2014. *J Natl Compr Canc Netw.* 2014;12(8):1152-1182.

Neuroendocrine Cancers and Adrenal Tumors (ONC-15)

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General Considerations (ONC-15.1)

ON.NA.0015.1.A

v1.0.2024

This guideline includes low-grade or well-differentiated carcinoid and endocrine tumors of the lung, thymus, pancreas, gastrointestinal tract or unknown primary site; including insulinoma, glucagonoma, VIPoma, gastrinoma, somatostatinoma and others as well as catecholamine-secreting tumors of the adrenal gland such as pheochromocytoma, paraganglioma, adrenocortical carcinoma, and others.

- For poorly-differentiated or high-grade small cell or large cell neuroendocrine tumors arising outside the lung or from an unknown primary site, see:
 Extrathoracic Small Cell and Large Cell Neuroendocrine Tumors (ONC-31.8).
- For poorly-differentiated or high grade neuroendocrine tumors of the lung, see: **Small Cell Lung Cancer (ONC-7)**.
- Neuroblastoma, ganglioneuroblastoma, and ganglioneuroma occurring in adults should be imaged according to <u>Neuroblastoma (PEDONC-6)</u> in the Pediatric Oncology Imaging Guidelines.
- Many are associated with Multiple Endocrine Neoplasia (MEN) familial syndromes.
 See: <u>Multiple Endocrine Neoplasias (MEN) (PEDONC-2.8)</u> in the Pediatric Oncology Imaging Guidelines for screening recommendations.
- Somatostatin receptor (SSR) based imaging is more sensitive and specific for evaluation of well-differentiated neuroendocrine tumors and may be performed using ¹¹¹In DTPA Octreotide scintigraphy or PET/CT scan with SSR radiotracers (such as ⁶⁸Ga–DOTATATE, ⁶⁸Ga-DOTATOC, or ⁶⁴Cu-DOTATATE). This study is not part of evaluation of poorly-differentiated or high-grade neuroendocrine tumors, which are imaged according to: Extrathoracic Small Cell and Large Cell Neuroendocrine Tumors (ONC-31.8).

Gastrointestinal/Pancreatic Neuroendocrine Cancers – Suspected/ Diagnosis (ONC-15.2)

ON.NA.0015.2.A

Indication	Ima	aging Study
 Systemic symptoms suggestive of function neuroendocrine tume. Suspicious findings of imaging studies. Unexplained elevation following: Chromogranin A 5HIAA Insulin VIP Glucagon Gastrin Substance P Serotonin Somatostatin 	on in ANY of the	Y of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) or without and with contrast (CPT® 74178) OR MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) CT with contrast or MRI without and with contrast of any other symptomatic body areas
Continued suspicion negative/inconclusive	e CT or MRI	 E of the following: Octreotide scan Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the follow SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE

Gastrointestinal/Pancreatic Neuroendocrine Cancers – Initial Work-up/Staging (ONC-15.3)

ON.NA.0015.3.A

V1.0.2024		
Indication	Imaging Study	
GI or pancreatic neuroendocrine (carcinoid) tumors	 If not already done: CT Abdomen and Pelvis with contrast (CPT® 74177) or without and with contrast (CPT® 74178) OR MRI Abdomen (CPT® 74183) and Pelvis (CPT® 72197) without and with contrast is indicated CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) 	
Inconclusive CT or MRI scans	 ONE of the following: Octreotide scan (ANY ONE of the following): Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE 	
 ANY of the following: Markers fail to normalize after complete resection AND CT/MRI and somatostatin-receptor based study are negative Biopsy-proven neuroendocrine tumor of unknown primary site AND CT/MRI and somatostatin-receptor based study are negative 	FDG-PET/CT scan (CPT® 78815)	

Gastrointestinal/Pancreatic Neuroendocrine Cancers – Restaging/ Recurrence (ONC-15.4)

ON.NA.0015.4.A

Indication	Imaging Study
All after surgical resection	See: Gastrointestinal/Pancreatic Neuroendocrine Cancers – Surveillance (ONC-15.5)
Unresectable/metastatic disease on treatment with somatostatin analogues	CT of involved body area no more frequently than every 3 months
Unresectable/metastatic disease on treatment with chemotherapy	CT of involved body area every 2 cycles (6 to 8 weeks)
Progression of symptoms or elevation of tumor markers	CT Chest without contrast (CPT® 71250) or CT Chest with contrast (CPT® 71260)
	 And ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Continued suspicion for recurrence with negative or inconclusive CT or MRI	 ONE of the following: Octreotide scan: Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE

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Indication	Imaging Study
To assess candidacy for peptide receptor radionuclide therapy (PRRT) with Lutetium ¹⁷⁷ Lu-dotatate	 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE

Gastrointestinal/Pancreatic Neuroendocrine Cancers – Surveillance (ONC-15.5)

ON.NA.0015.5.A

Indication	Imaging Study
 ANY of the following: Appendix carcinoid ≤2 cm, completely resected Rectal carcinoid <1 cm, completely resected Gastric carcinoid treated with complete endoscopic resection 	Advanced imaging is not routinely indicated for surveillance
Rectal carcinoid 1-2 cm, completely resected	MRI Pelvis without and with contrast (CPT® 72197) at 6 and 12 months post resection. If clear, no further surveillance imaging indicated
All other GI neuroendocrine tumors (stomach, large and small intestine)	CT Abdomen and Pelvis with contrast (CPT® 74177) once at 3 to 12 months postoperatively and annually for 3 years and then every 2 years up to year 10
Unresected GI neuroendocrine tumors being monitored with observation alone	CT Abdomen with contrast (CPT® 74160) once at 3 to 12 months from initial diagnosis then annually up to year 10
Pancreatic neuroendocrine tumors	CT Abdomen with contrast (CPT® 74160) once at 3 to 12 months postoperatively then annually up to year 10
Unresected pancreatic neuroendocrine tumors being monitored with observation alone	CT Abdomen with contrast (CPT® 74160) once at 3 to 12 months from initial diagnosis then annually up to year 10
Measurable metastatic disease on maintenance treatment or off therapy	CT of involved body area no more frequently than every 3 months

Bronchopulmonary or Thymic Carcinoid – Initial Staging (ONC-15.6)

ON.NA.0015.6.A

	v1.0.2024
Indication	Imaging Study
Initial diagnosis	 If not already done: CT Chest with contrast (CPT® 71260) CT Abdomen with contrast (CPT® 74160) or without and with contrast (CPT® 74170) If CT inconclusive, MRI Abdomen (CPT® 74183) without and with contrast is indicated
Inconclusive CT or MRI scans	 ONE of the following: Octreotide scan (ANY ONE of the following): Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE
 ANY of the following: Markers fail to normalize after complete resection AND CT/MRI and somatostatin-receptor based study are negative Biopsy-proven neuroendocrine tumor of unknown primary site AND CT/MRI and somatostatin-receptor based study are negative 	FDG-PET/CT scan (CPT® 78815)

Bronchopulmonary or Thymic Carcinoid – Restaging/Recurrence (ONC-15.7)

ON.NA.0015.7.A

Indication	Imaging Study
All after surgical resection	See: Bronchopulmonary or Thymic Carcinoid - Surveillance (ONC-15.8)
Unresectable/metastatic disease on treatment with somatostatin analogues	CT of involved body area no more frequently than every 3 months
Unresectable/metastatic disease on treatment with chemotherapy	CT of involved body area every 2 cycles (6 to 8 weeks)
Progression of symptoms or elevation of tumor markers	 CT Chest without (CPT® 71250) or CT Chest with contrast (CPT® 71260) And ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and Pelvis (CPT® 72197) without and with contrast
Continued suspicion for recurrence with negative or inconclusive CT or MRI	 ONE of the following: Octreotide scan Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE

Bronchopulmonary or Thymic Carcinoid – Surveillance (ONC-15.8)

ON.NA.0015.8.A

Indication	Imaging Study
Carcinoid tumors of lung or thymus	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) once at 3 to 12 months post resection and then annually for 3 years and then every 2 years up to year 10
Unresected primary tumors being monitored with observation alone	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) once at 3 to 12 months from initial diagnosis then annually for 3 years and then every 2 years up to year 10
Measurable metastatic disease on maintenance treatment or off therapy	CT of involved body area no more frequently than every 3 months

Adrenal Tumors – Suspected/Diagnosis (ONC-15.9)

ON.NA.0015.9.A

- See: <u>Adrenal Cortical Lesions (AB-16.1)</u> in the Abdomen Imaging Guidelines for evaluation of indeterminate adrenal masses.
- Adrenal tumors that involve the adrenal medulla or neural crest tissue outside the adrenal gland include pheochromocytoma, paraganglioma, and paraganglioneuroma
 - These tumors are imaged according to sections <u>ONC-15.10 through ONC-15.12</u>
 - Malignant adrenal tumors that involve the adrenal cortex are addressed in <u>Adrenocortical Carcinoma (ONC-15.13)</u>
- Adrenocortical carcinoma is imaged according to <u>Adrenocortical Carcinoma</u> (ONC-15.13)
- If concern for genetic predisposition syndrome such as MEN, neurofibromatosis, or Von Hippel-Lindau disease, see screening recommendations in <u>Screening</u> <u>Imaging and Cancer Predisposition Syndromes (PEDONC-2)</u> in the Pediatric Oncology Imaging Guidelines.

Adrenal Tumors – Initial Work-up/Staging (ONC-15.10)

ON.NA.0015.10.A

Indication	Imaging Study
For ANY of the following: Pheochromocytoma Paraganglioma Paraganglioneurom a	 If not already done: CT Chest without (CPT® 71250) or CT Chest with contrast (CPT® 71260) And ONE of the following (if not already done): CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast CT with contrast or MRI without and with contrast of any other symptomatic body areas
Continued suspicion with negative/inconclusive CT or MRI	 ONE of the following: Octreotide or MIBG scan: Any one of the following planar imaging codes - CPT® 78801, 78802, 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE
All above studies done and negative/inconclusive	FDG-PET/CT scan (CPT® 78815)

Adrenal Tumors – Restaging/Recurrence (ONC-15.11)

ON.NA.0015.11.A

Indication	Imaging Study
If surgery is primary therapy	CT Abdomen with contrast (CPT® 74160) one time within first year post resection then go to surveillance recommendations
Recurrence, progression of symptoms, or elevation of tumor markers	 CT Chest without contrast (CPT® 71250) or CT Chest with contrast (CPT® 71260) CT with contrast of involved areas And ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Continued suspicion for recurrence with negative or inconclusive CT or MRI	 ONE of the following: Octreotide scan (ANY ONE of the following): Any one of the following planar imaging codes - CPT® 78801, 78802, or 78804 AND Any one of the following SPECT/SPECT-CT codes - CPT® 78803, 78830, 78831, 78832 PET/CT scan (CPT® 78815) with any ONE of the following SSR radiotracers: ⁶⁸Ga-DOTATATE ⁶⁸Ga-DOTATOC ⁶⁴Cu-DOTATATE
All above studies done and negative/inconclusive	FDG-PET/CT scan (CPT® 78815)

Adrenal Tumors – Surveillance (ONC-15.12)

ON.NA.0015.12.A

Indication	Imaging Study
All individuals	Once within 3-12 months post resection and then annually for 10 years:
	CT Chest with contrast (CPT® 71260) or without contrast (CPT® 71250)
	CT Abdomen and Pelvis with contrast (CPT® 74177) or MRI Abdomen and Pelvis without and with contrast (CPT® 74183 and 72197)
	CT with contrast of other involved body areas
Measurable metastatic disease being observed off therapy or on maintenance treatment	CT of involved body area no more frequently than every 3 months for up to 5 years after completion of definitive therapy and annually thereafter

Adrenocortical Carcinoma (ONC-15.13)

ON.NA.0015.13.A

	v1.0.2024
Indication	Imaging Study
Initial Staging	CT Chest without (CPT® 71250) or CT Chest with contrast (CPT® 71260)
	 And ONE of the following (if not already done): CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with
	 contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Suspected recurrence	CT Chest without (CPT® 71250) or CT Chest with contrast (CPT® 71260)
	And ONE of the following:
	CT Abdomen and Pelvis with contrast (CPT® 74177)
	CT Abdomen and Pelvis without and with contrast (CPT® 74178)
	MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
 Solitary adrenal mass >4 cm on conventional imaging and plans for aggressive surgical resection Inconclusive findings on conventional imaging 	FDG PET/CT scan (CPT® 78815)
Surveillance after complete response to definitive treatment	 Annually for 5 years: CT Chest with contrast (CPT® 71260), CT Abdomen with contrast (CPT® 74160), and CT of other involved body areas with contrast
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 Every 3 months for up to 5 years after completion of definitive therapy: CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and CT with contrast of other involved body areas

References (ONC-15)

- 1. Bergsland E, Goldner WS, Benson III AB, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 August 2, 2023. Neuroendocrine and Adrenal tumors, available at: https://www.nccn.org/professionals/physician_gls/pdf/neuroendocrine.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Neuroendocrine and Adrenal tumors V1.2023 August 2, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Qadan M, Ma Y, Visser BC, et al. Reassessment of the current American Join Committee on Cancer staging system for pancreatic neuroendocrine tumors. *J Am Coll Surg*. 2014;218(2):188-195. doi:10.1016/j.jamcollsurg.2013.11.001.
- 3. Lenders JWM, Duh Q-Y, Eisenhofer G, et al. Pheochromocytoma and paraganglioma: an Endocrine Society clinical practice guideline. *J Clin Endocrinol Metab.* 2014;99(6):1915-1942. doi:10.1210/jc.2014-1498
- Ruys AT, Bennink RJ, van Westreenen HL, et al. FDG-positron emission tomography/computed tomography and standardized uptake value in the primary diagnosis and staging of hilar cholangiocarcinoma. HPB (Oxford). 2011;13(4):256-262. doi:10.1111/j.1477-2574.2010.00280.x.
- 5. Ter-Minassian M, Chan JA, Hooshmand SM, et al. Clinical presentation, recurrence, and survival in patients with neuroendocrine tumors: results from a prospective institutional database. *Endocr Relat Can.* 2013;20(2):187-196. doi:10.1530/ERC-12-0340.
- 6. Murray SE, Lloyd RV, Sippel RS, Chen H, Oltmann SC. Postoperative surveillance of small appendiceal carcinoid tumors. *Am J Surg.* 2014;207(3):342-345. doi:10.1016/j.amjsurg.2013.08.038.
- 7. Thakker RV, Newey PJ, Walls GV, et al. Clinical practice guidelines for multiple endocrine neoplasia type 1 (MEN1). *J Clin Endocrinol Metab*. 2012;97(9):2990-3011. doi:10.1210/jc.2012-1230.
- 8. Singh S, Moody L, Chan DL, et al. Follow-up recommendations for completely resected gastroenteropancreatic neuroendocrine tumors. *JAMA Oncol.* 2018;4(11):1597-1604. doi:10.1001/jamaoncol.2018.2428.

Colorectal and Small Bowel Cancer (ONC-16)

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Colorectal Cancer – General Considerations (ONC-16.0)

ON.CC.0016.0.A

- Neuroendocrine tumors of the bowel are covered in: Neuroendocrine Cancers and Adrenal Tumors. (ONC-15)
- Appendiceal adenocarcinoma (including pseudomyxoma peritonei) follows imaging guidelines for colorectal cancer.
- For squamous cell carcinoma of the rectum, see: Anal Carcinoma (ONC-24)

Colorectal Cancer – Suspected/Diagnosis (ONC-16.1)

ON.CC.0016.1.A

- See: GI Bleeding (AB-22) or CT Colonography (CTC) (AB-25.1) in the Abdomen Imaging Guidelines for evaluation of suspected colorectal malignancies.
- See: <u>Abnormal Findings on Endoscopy/Colonoscopy (AB-13.3)</u> in the Abdomen Imaging Guidelines for evaluation of abnormal findings on endoscopy/colonoscopy.
- If findings on colonoscopy are suspicious for colon cancer, see: <u>Colorectal Cancer</u> <u>Initial Work-up/Staging (ONC-16.2)</u>

Colorectal Cancer – Initial Work-up/Staging (ONC-16.2)

ON.CC.0016.2.A

Indication	Imaging Study
Carcinoma within a polyp that is completely removed	No advanced imaging needed
 Biopsy proven invasive adenocarcinoma Colonoscopy findings suspicious for colon cancer 	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
 Further evaluation of an inconclusive liver lesion seen on CT Potentially resectable liver metastases 	MRI Abdomen without and with contrast (CPT® 74183)
Rectal adenocarcinoma	MRI Pelvis without and with contrast (CPT® 72197) or MRI Pelvis without contrast (CPT® 72195) (can be obtained in addition to CT scans for initial staging)
Rectal adenocarcinoma with ANY one of the following: Rectal MRI is contraindicated Rectal MRI is inconclusive Superficial lesions	Endorectal ultrasound (CPT® 76872)
 ONE of the following: Isolated metastatic lesion(s) on other imaging and individual is a candidate for aggressive surgical resection or other localized treatment to metastasis for curative intent Inconclusive conventional imaging 	• PET/CT (CPT® 78815)

Colorectal Cancer – Restaging/Recurrence (ONC-16.3)

ON.CC.0016.3.C

Indication	Imaging Study
 Complete resection Individuals receiving post- operative adjuvant chemotherapy 	See: Surveillance/Follow-up (ONC-16.4)
Recurrence suspected	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
After completion of planned neoadjuvant therapy	 Prior to surgical resection in individuals with nonmetastatic rectal cancer: CT Chest with contrast (CPT® 71260) and Any ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen with contrast (CPT® 74160) and MRI Pelvis without and with contrast (CPT® 72197)
Unresectable disease or metastatic disease on chemotherapy	 Every 2 cycles of chemotherapy treatment and at the completion of chemotherapy: CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
 Further evaluation of an inconclusive liver lesion seen on CT Potentially resectable liver metastases 	MRI Abdomen without and with contrast (CPT® 74183)

Indication	Imaging Study
 ONE of the following: Postoperative elevated or rising CEA or LFTs with negative recent conventional imaging Isolated metastatic lesion(s) on other imaging and individual is a candidate for aggressive surgical resection or other localized treatment to metastasis for curative intent Differentiate local tumor recurrence from postoperative and/or post-radiation scarring 	• PET/CT (CPT® 78815)
New or worsening pelvic pain and recent CT imaging negative or inconclusive	MRI Pelvis without and with contrast (CPT® 72197)

Colorectal Cancer – Surveillance/Follow-Up (ONC-16.4)

ON.CC.0016.4.C

Indication	Imaging/Lab Study
Colon and rectal adenocarcinoma: • Stage I	No routine advanced imaging indicated
Colon and rectal adenocarcinoma: • Stage II-III	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) after completion of surgery and then every 6 months for 5 years
 Colon and rectal adenocarcinoma: Stage IV or distant metastatic disease (post definitive treatment of all measurable disease or being observed off therapy) 	 CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) every 3 months for 2 years and then every 6 months for 3 years
Measurable metastatic disease on maintenance therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Rectal cancer treated with transanal excision alone	 Endorectal ultrasound (CPT® 76872) every 6 months for 5 years MRI Pelvis without and with contrast (CPT® 72197) for: Abnormal findings on ultrasound Endorectal ultrasound is not feasible New signs/symptoms concerning for local recurrence
Stage II-III rectal cancer treated with chemoradiation alone (no surgical treatment)	In addition to the above stage-specific surveillance: • MRI Pelvis (CPT® 72197) without and with contrast every 6 months for 3 years

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Indication	Imaging/Lab Study
Pseudomyxoma peritonei	 ONE of each of the following, every 3 months for first year, then every 6 months for 4 more years: • CT Chest with (CPT® 71260) or without (CPT® 71250) contrast • CT Abdomen and Pelvis with contrast (CPT® 74177) or MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast

Small Bowel Cancer – Initial Work-up/Staging (ONC-16.5)

ON.CC.0016.5.A

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This section provides imaging guidelines for small bowel adenocarcinoma arising from the duodenum, jejunum, and ileum.

Indication	Imaging/Lab Study
Carcinoma within a polyp that is completely removed	No advanced imaging needed
Invasive adenocarcinoma	 CT Chest with contrast (CPT[®] 71260) and CT Abdomen and Pelvis (CPT[®] 74177) with contrast
	 MRI Abdomen without and with contrast (CPT® 74183) and MRI Pelvis without and with contrast (CPT® 72197) if CT is inconclusive or cannot be performed

Small Bowel Cancer – Restaging/Recurrence (ONC-16.6)

ON.CC.0016.6.A

Indication	Imaging Study
Complete resection	See Surveillance below
Recurrence suspected	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
Unresected primary disease or metastatic disease on chemotherapy	 Every 2 cycles of chemotherapy: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Further evaluation of an inconclusive liver lesion seen on CT	MRI Abdomen without and with contrast (CPT® 74183)
 ONE of the following: Postoperative elevated or rising CEA or LFTs with negative recent conventional imaging Isolated metastatic lesion(s) on other imaging and individual is a candidate for aggressive surgical resection or other localized treatment to metastasis for curative intent 	• PET/CT (CPT® 78815)

Small Bowel Cancer – Surveillance/Follow-up (ONC-16.7)

ON.CC.0016.7.A

Indication	Imaging/Lab Study
Stage I-III	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast after completion of surgery, and then annually for 5 years
Stage IV - Metastatic disease (post definitive treatment of all measurable disease, or being observed off therapy)	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast every 6 months for 2 years and then annually for 3 years
Measurable metastatic disease on maintenance therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)

References (ONC-16)

- 1. Benson III AB, Venook AP, Al-Hawary MM, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 April 25, 2023. Colon cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/colon.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Colon cancer V2.2023 April 25, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Benson III AB, Venook AP, Al-Hawary MM, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 4.2023 July 25, 2023. Rectal cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/rectal.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Rectal cancer V4.2023 July 25, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Benson AB, Venook AP, Pedersen K, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 January 9, 2023. Small Bowel Adenocarcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/small_bowel.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Small Bowel Adenocarcinoma V1.2023 January 9, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 4. ACR Appropriateness Criteria. Pretreatment Staging of Colorectal Cancer. Rev. 2011.
- 5. Bailey CE, Hu C-Y, You YN et al. Variation in positron emission tomography use after colon cancer resection. *J Oncol Pract*. 2015;11(3):e363-e372. doi:10.1200/JOP.2014.001933.
- 6. Lu YY, Chen JH, Ding HJ, Chien CR, Lin WY, Kao CH. A systematic review and meta-analysis of pretherapeutic lymph node staging of colorectal cancer by 18F-FDG PET or PET/CT. *Nucl Med commun*. 2012;33(11):1127-1133. doi:10.1097/MNM0b013e328357b2d9.
- 7. Moulton CA, Gu CS, Law CH, et al. Effect of PET before liver resection on surgical management for colorectal adenocarcinoma metastases: a randomized clinical trial. *JAMA*. 2014;311(18):1863-1869. doi:10.1001/jama.2014.3740.
- Steele SR, Chang GJ, Hendren S, et al. Practice guideline for the surveillance of patients after curative treatment of colon and rectal cancer. *Dis Colon Rectum*. 2015;58(8):713-725. doi:10.1097/DCR.0000000000001010.
- van de Velde CJ, Boelens PG, Borras JM, et al. EURECCA colorectal: multidisciplinary management: European concensus conference colon & rectum. Eur J Cancer. 2014;50(1):e1-e34. doi:10.1016/j.ejca.2013.06.048.
- 10. Akce M, El-Rayes BF. Nonsurgical management of rectal cancer. Journal of Oncology Practice. 2019;15(3):123-131. doi:10.1200/JOP.18.00769.

Renal Cell Cancer (RCC) (ONC-17)

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Renal Cell Cancer (RCC) – General Considerations (ONC-17.0)

ON.RC.0017.0.A

- PET considered experimental, investigational, or unproven for initial diagnosis, staging or restaging of renal cell cancer.
- A minority of adult individuals with renal cell cancer (RCC) will have translocations in TFE3 or TFEB, which have a different natural history than "adult type" RCC. Individuals of any age with TFE3 or TFEB translocated RCC should be imaged according to guidelines in <u>Pediatric Renal Cell Carcinoma (RCC) (PEDONC-7.4)</u> in the Pediatric Oncology Imaging Guidelines.
- Individuals of any age with Wilms tumor should be imaged according to guidelines in section <u>Unilateral Wilms Tumor (UWT) (PEDONC-7.2)</u> or <u>Bilateral Wilms</u> <u>Tumor (BWT) (PEDONC-7.3)</u> in the Pediatric Oncology Imaging Guidelines.
- Oncocytoma in individuals of all ages should be imaged according to these guidelines.

Renal Cell Cancer (RCC) – Suspected/ Diagnosis (ONC-17.1)

ON.RC.0017.1.A

Indication	Imaging Study
Solitary renal mass suspicious for renal cell cancer	 See: Indeterminate Renal Lesion (AB-35.1) in the Abdomen Imaging Guidelines for evaluation of suspected renal malignancies Chest x-ray or CT Chest with contrast with (CPT® 71260) or without contrast (CPT® 71250)

Renal Cell Cancer (RCC) – Initial Work-Up/Staging (ONC-17.2)

ON.RC.0017.2.C

Indication	Imaging Study
All individuals	 If not done previously: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen and Pelvis, contrast as requested
 ANY of the following: Extension of tumor into the vena cava by other imaging Inconclusive findings on CT 	MRI Abdomen without and with contrast (CPT® 74183)
Bone pain	Bone scan
 EITHER of the following: Signs/symptoms suspicious for brain metastases Newly diagnosed stage IV/metastatic RCC 	MRI Brain without and with contrast (CPT® 70553)

Renal Cell Cancer (RCC) – Restaging/Recurrence (ONC-17.3)

ON.RC.0017.3.A

Indication	Imaging Study
Unresectable disease or metastatic disease on systemic therapy	 Every 2 cycles of treatment (commonly every 6 to 8 weeks: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of other involved or symptomatic areas
Recurrence suspected	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
EITHER of the following: • Biopsy-proven recurrent/metast atic disease • Signs or symptoms concerning for brain metastases	MRI Brain without and with contrast (CPT® 70553)

Renal Cell Cancer (RCC) – Surveillance (ONC-17.4)

ON.RC.0017.4.C

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Indication	Imaging Study
RCC on active surveillance of renal mass <1 cm	 ONE of the following, once within 6 months of surveillance initiation and annually thereafter: CT Abdomen without and with contrast (CPT® 74170) MRI Abdomen without and with contrast (CPT® 74183) See: Indeterminate Renal Lesion (AB-35.1) in the Abdomen Imaging Guidelines Chest x-ray (in addition to abdominal imaging) CT Chest with contrast (CPT® 71260) or without contrast (CPT® 71250) may be obtained for one of the following: New chest x-ray abnormalities
	Pulmonary signs/symptoms
RCC on active surveillance of renal mass ≥1 cm	 ONE of the following, every 3 months for year 1, every 6 months for years 2 and 3 and annually thereafter: CT Abdomen without and with contrast (CPT® 74170) MRI Abdomen without and with contrast (CPT® 74183) Chest x-ray (in addition to abdominal imaging) CT Chest with contrast (CPT® 71260) or without contrast (CPT® 71250) may be obtained for one of the following: New chest x-ray abnormalities Pulmonary signs/symptoms
Follow up after post- ablation therapy of RCC	 EITHER of the following, at 1 to 3 months, 6 months, and 12 months post-ablation and then annually thereafter: CT Abdomen without and with contrast (CPT® 74170) or MRI Abdomen without and with contrast (CPT® 74183) AND Annually for 5 years: Chest x-ray (in addition to abdominal imaging) or CT Chest with contrast (CPT® 71260) or without contrast (CPT® 71250)

Indication	Imaging Study
Stage I RCC, treated with partial or radical nephrectomy	 ONE of each of the following, 3 to 12 months post-resection: CT Chest with (CPT® 71260) or CT Chest without (CPT® 71250) contrast CT Abdomen with (CPT® 74160) or CT Abdomen without contrast (CPT® 74150) or MRI Abdomen without and with contrast (CPT® 74183) Annually for 5 years: Chest x-ray or CT Chest with (CPT® 71260) or without (CPT® 71250) contrast Abdominal imaging with any ONE of the following: CT Abdomen with (CPT® 74160) or without (CPT® 74150) contrast MRI Abdomen without and with contrast (CPT® T1200)
Stage II RCC, post-nephrectomy	 ONE of each of the following, 3 to 6 months post-resection: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen with (CPT® 74160) or without (CPT® 74150) contrast or MRI Abdomen without and with contrast (CPT® 74183) ONE of each of the following, every 6 months for 2 years, then annually until year 5: Chest x-ray or CT Chest with (CPT® 71260) or without (CPT® 71250) contrast Abdominal imaging with any ONE of the following: CT Abdomen with (CPT® 74160) or without (CPT® 74150) contrast MRI Abdomen without and with contrast (CPT® 74183)

Indication	Imaging Study
Stage III RCC, post- nephrectomy	 ONE of each of the following, 3 to 6 months post-resection: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen with (CPT® 74160) or without (CPT® 74150) contrast or MRI Abdomen without and with contrast (CPT® 74183) ONE of each of the following, every 3 months for 3 years, then annually to year 5: CT Chest with (CPT® 71260) or without (CPT® 71250) contrast CT Abdomen with (CPT® 74160) or without (CPT® 74150) contrast or MRI Abdomen without and with contrast (CPT® 74183)
Stage IV/metastatic disease on maintenance therapy or being observed off therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast CT with contrast of other involved or symptomatic areas

References (ONC-17)

- 1. Motzer RJ, Jonasch E, Agarwal N, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2024 June 21, 2023. Kidney cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/kidney.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Kidney cancer V1.2024 June 21, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. ACR Appropriateness Criteria. Post-treatment follow up of renal cell carcinoma. Rev. 2013.
- 3. Herts BR, Silverman SG, Hindman NM, et al. *Management of the incidental renal mass on CT: a white paper of the ACR incidental findings committee*. J Am Coll Radiol. 2018;15(2):264-273. doi:10.1016/j.jacr.2017.04.028.
- Finelli A, Ismaila N, Bro B, et al. Management of small renal masses. American Society of Clinical Oncology clinical practice guideline. *Journal of Clinical Oncology*. 2017;35(6):668-680. doi:10.1200/JCO.2016.69.9645.
- Davenport MS, Caoili EM, Cohan RH, et al. MRI and CT characteristics of successfully ablated renal masses: imaging surveillance after radiofrequency ablation. AJR Am J Roentgenol. 2009;192:1571-1578. doi:10.2214/AJR.08.1303.
- Clark TW, Millward SF, Gervais DA, et al. Reporting standards for percutaneous thermal ablation of renal cell carcinoma. J Vasc Interv Radiol. 2009;20(7 Suppl):S409-S416. doi:10.1016/j.jvir.2009.04.013.
- Rais-Bahrami S, Guzzo TJ, Jarrett TW, Kavoussi LR, Allaf ME. Incidentally discovered renal masses: oncological and perioperative outcomes in patients with delayed surgical intervention. *BJU Int*. 2009;103(10):1355-1358. doi:10.1111/j.1464-410X.2008.08242.x.
- 8. Wang HY, Ding HJ, Chen JH, Chao CH, Lu YY, Lin WY, Kao CH. Meta-analysis of the diagnostic performance of [18F]FDG-PET and PET/CT in renal cell carcinoma. *Cancer Imaging*. 2012 October;12:464-474. doi:10.1102/1470-7330.2012.0042.
- 9. Kim EH, Strope SA. Postoperative surveillance imaging for patients undergoing nephrectomy for renal cell carcinoma. *Urol Oncol.* 2015;33(12):499-502. doi:10.1016/j.urolonc.2015.08.008.
- 10. Sankineni S, Brown A, Cieciera M, Choyke PL, Turkbey B. Imaging of renal cell carcinoma. *Urol Oncol.* 2016;34(3):147-155. doi:10.1016/j.urolonc.2015.05.020.
- 11. ACR Appropriateness Criteria. Renal cell carcinoma staging. Rev. 2015.
- 12. Campbell S, Uzzo R, Allaf M, et al. Renal mass and localized renal cancer: AUA guideline. *J Urol.* 2017:198(3):520-529. doi:10.1016/j.juro.2017.04.100.

Transitional Cell Cancer (ONC-18)

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Transitional Cell Cancer – General Considerations (ONC-18.0)

ON.TS.0018.0.C

- Transitional cell cancers can include: tumors of the bladder, ureters, prostate, urethra, or renal pelvis. For primary cancer of the kidney, see: <u>Renal Cell Cancer</u> (<u>RCC</u>) (<u>ONC-17</u>).
- The most common histology of bladder cancer is transitional cell (TCC) or urothelial carcinoma (UCC). Rare histologies include adenocarcinoma, squamous cell (imaged according to: <u>Transitional Cell Cancer (ONC-18)</u>), or small cell (imaged according to: <u>Extrathoracic Small Cell and Large Cell Neuroendocrine Tumors (ONC-31.8)</u>).
- Urachal cancer is a rare type of bladder cancer; the most common histology is adenocarcinoma. These are imaged according to muscle invasive bladder cancer.

Transitional Cell Cancer – Suspected/Diagnosis (ONC-18.1)

ON.TS.0018.1.A

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• See: <u>Hematuria and Hydronephrosis (AB-39)</u> in the Abdomen Imaging Guidelines for evaluation of suspected transitional cell malignancies.

Transitional Cell Cancer – Initial Workup/Staging (ONC-18.2)

ON.TS.0018.2.A

Indication	Imaging Study
All individuals	 ONE of the following: CT Abdomen and Pelvis without and with contrast (CPT® 74178) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast if contraindication to CT contrast
	CT Abdomen and Pelvis without contrast (CPT® 74176) with retrograde pyelogram or renal ultrasound (CPT® 76770 or CPT® 76775) in individuals who cannot receive either CT or MRI contrast
 ANY of the following: Muscle invasive bladder carcinoma Urethral carcinoma Urothelial carcinoma of the prostate 	CT Chest without (CPT® 71250) or with (CPT® 71260) contrast
Individuals without metastatic disease, when requested by operating surgeon for operative planning	CT with contrast or MRI without and with contrast of all operative sites
To evaluate inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Transitional Cell Cancer – Restaging/Recurrence (ONC-18.3)

ON.TS.0018.3.A

Indication	Imaging Study
After definitive surgery	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen and Pelvis without and with contrast (CPT® 74178) for post-operative baseline
Recurrence suspicion	 CT Abdomen and Pelvis with contrast (CPT® 74177) or with and without contrast (CPT® 74178) CT Chest with contrast (CPT® 71260) for ANY of the following: Signs/symptoms of pulmonary disease Abnormal chest x-ray Prior involvement of the chest
After neoadjuvant therapy and before resection	CT Chest with contrast (CPT® 71260) and CT Urogram (CPT® 74178)
Monitoring therapy for metastatic disease	 Every 2 cycles of therapy: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for ANY of the following: Signs/symptoms of pulmonary disease Prior involvement of the chest Abnormal chest x-ray
To evaluate inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Transitional Cell Cancer – Surveillance/Follow-up (ONC-18.4)

ON.TS.0018.4.A

7110		
Indication	Imaging Study	
 ANY of the following: Papillary urothelial neoplasm of low malignant potential Low risk lesions Solitary Ta lesions ≤3cm Intermediate risk lesions Low-grade >3 cm Low-grade multifocal T1 lesions High-grade solitary Ta ≤3cm 	Advanced imaging is not routinely indicated for surveillance	
ANY of the following high-risk non-muscle invasive transitional cell carcinoma of the bladder or upper tracts: • Multifocal high-grade lesions • High-grade lesions >3 cm • Superficial and minimally invasive (Tis and T1) • BCG unresponsive • Lymphovascular invasion • Prostatic urethral invasion	 CT Urogram (CPT® 74178) every 2 years for 10 years MR Urogram (CPT® 74183 and CPT® 72197) may be obtained for renal insufficiency or CT dye allergy 	
Non-muscle-invasive transitional carcinoma of the bladder treated with cystectomy	 CT Urogram (CPT® 74178) at 3 and 12 months post-cystectomy, and then annually for years 2-5 MR Urogram (CPT® 74183 and CPT® 72197) may be obtained for renal insufficiency or CT dye allergy 	

Indication	Imaging Study
Muscle invasive lower and upper genitourinary tumors treated with cystectomy or chemoradiation	 Every 6 months for 2 years, then annually for 3 more years: CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250), and CT Abdomen and Pelvis with contrast (CPT® 74177) or without and with contrast (CPT® 74178)
	 OR CT Chest with contrast (CPT[®] 71260) or CT Chest without contrast (CPT[®] 71250), and MR Urogram (CPT[®] 74183 and CPT[®] 72197)
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Urogram (CPT® 74178)
Urethral cancers (high-risk T1 or greater) and urothelial carcinoma of the prostate	 Every 6 months for 2 years, then annually: CT Abdomen and Pelvis with contrast (CPT[®] 74177) or without and with contrast (CPT[®] 74178)
	OR • MR Urogram (CPT® 74183 and CPT® 72197)
	AND • Chest x-ray
	CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) if abnormal signs/symptoms of pulmonary disease or abnormal chest x-ray

References (ONC-18)

- 1. Flaig TW, Spiess PE, Abern M, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 May 25, 2023. Bladder cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/bladder.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Bladder cancer V3.2023 May 25, 2023. ®2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Verma S, Rajesh A, Prasad SR et al, Urinary bladder cancer: role of MR imaging. *Radiographics*. 2012;32(2):371-387. doi:10.1148/rg.322115125.
- 3. Lu YY, Chen JH, Liang JA. Clinical value of FDG PET or PET/CT in urinary bladder cancer: a systematic review and meta-analysis. *Eur J Radiol*. 2012;81(9):2411-2416. doi:10.1016/j.ejrad.2011.07.018.
- 4. Witjes JA, Comperat E, Cowan NC, et al. EAU guidelines on muscle-invasive and metastatic bladder cancer: summary of the 2013 guidelines. *Eur Urol.* 2014;65(4):778-792. doi:10.1016/j.eururo.2013.11.046.
- 5. Gakis G, Witjes JA, Comperat E, et al. EAU guidelines on primary urethral carcinoma. *Eur Urol.* 2013;64(5):823-830. doi:10.1016/j.eururo.2013.03.044.
- 6. Rouprêt M, Babjuk M, Compérat E, et al. European guidelines on upper tract urothelial carcinomas: 2013 update. *Eur Urol.* 2013;63(6):1059-1071. doi:10.1016/j.eururo.2013.03.032.

Prostate Cancer (ONC-19)

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Prostate Cancer – General Considerations (ONC-19.0)

ON.PR.0019.0.C

- Prostate cancer screening begins at age 45 for individuals at average risk of prostate cancer. However, individuals at high-risk may begin screening at age 40. High-risk features include:
 - African ancestry
 - Germline mutations (BRCA1 or 2, HOXB13, ATM, CHEK2, or mismatch repair genes - MLH1, MSH2, MSH6, PMS2) that increase the risk of prostate cancer
 - Family history of first or second-degree relative with prostate, male breast, colorectal, pancreatic, endometrial or female breast cancer at age <45 years.
- Treatment of benign prostatic hyperplasia with 5-alfa reductase inhibitors (such as finasteride and dutasteride) can falsely reduce the measured PSA levels by 50%.
 Thus, the reported PSA level should be doubled when prostate cancer is suspected in individuals on these medications
- Individuals with high-risk adverse clinical and pathological factors may benefit from a more aggressive diagnostic and therapeutic approach at the time of relapse after initial treatment. These factors include pre-treatment Gleason score of ≥8, pretreatment clinical stage of cT3b or higher, positive surgical margins, post-treatment PSA doubling time of <3 months, and an interval to biochemical failure of <3 years after initial treatment.
- PET/CT scans using ¹⁸F-FDG radiotracer are considered investigational and experimental for evaluation of prostate cancer.
- ¹¹C Choline, ¹⁸F-Fluciclovine (AXUMIN®), and PSMA-specific radiopharmaceuticals have recently gained FDA approval for evaluation of prostate cancer. Optimal detection rates for these radiotracers vary greatly with PSA levels. False positive rate is high and histological confirmation of positive sites is recommended.
- PSMA-specific PET radiopharmaceuticals that are currently FDA-approved and indicated in prostate cancer are: ⁶⁸Ga PSMA-11 (UCSF & UCLA), ¹⁸F Piflufolastat (Pylarify®), ¹⁸F Flotufolastat (Posluma®), and ⁶⁸Ga Gozetotide (Illuccix® and Locametz®).
- While early detection of low-volume recurrence after treatment of prostate cancer using PET/CT scans may influence therapeutic decisions; there is lack of evidence that this approach has any meaningful impact on overall survival.
- MR Spectroscopy (CPT® 76390) is considered experimental, investigational, unproven in the evaluation of prostate cancer at this time.
- As laser prostate ablation is considered investigational and experimental at this time, advanced imaging for treatment planning and/or surveillance of laser prostate ablation is considered not medically necessary.
- Monitoring an elevated prostate-specific antigen level (PSA) with serial MRI is not indicated for suspected prostate cancer.

 Requests for imaging based on PSA must provide a recent (within the last 60 days) PSA.

ISUP Prostate Cancer Grade Groups³⁰

Grade Group	Gleason Score	Gleason Pattern
1	≤6	≤3+3
2	7	3+4
3	7	4+3
4	8	4+4, 3+5, 5+3
5	9 or 10	4+5, 5+4, or 5+5

NCCN Initial Risk Stratification

- Very Low Risk
 - o ALL of the following features are present:
 - Tumor not clinically palpable, but present on one or both lobes on biopsy (cT1a, cT1b, or cT1c)
 - PSA (ng/mL) <10
 - Gleason Grade Group = 1
 - <3 prostate biopsy cores positive, ≤50% cancer in each core</p>
 - PSA Density <0.15 ng/mL/g
- Low Risk
 - ALL of the following features are present but does not qualify for very low risk:
 - Clinical T Stage = cT1-cT2a (palpable tumor limited to ≤1/2 of one side)
 - PSA (ng/mL) <10
 - Gleason Grade Group = 1
- Favorable Intermediate Risk
 - O ALL of the following features are present:
 - Gleason Grade Group = 1 or 2
 - <50% biopsy cores positive (e.g., <6 of 12 cores)</p>
 - And only ONE of the following features is present:
 - Clinical T Stage = cT2b-cT2c (palpable disease confined to one or both lobes of the prostate)
 - PSA (ng/mL) = 10-20
- Unfavorable Intermediate Risk
 - o Any one of the following are present:
 - Gleason grade group = 3
 - ≥50% biopsy cores positive (e.g., ≥6 of 12 cores)
 - Presence of at least two of the following three features:

- PSA (ng/mL) = 10-20
- Gleason Grade Group = 2 or 3
- Clinical T Stage = cT2b-cT2c (palpable disease confined to one or both lobes of the prostate)
- High-Risk
 - Only ONE of the following high-risk features is present:
 - Clinical T Stage = cT3a (unilateral or bilateral extra-prostatic extension that is not fixed and does not invade the seminal vesicles)
 - PSA (ng/mL) >20
 - Gleason Grade Group = 4 or 5
- Very High-Risk
 - At least ONE of the following features is present:
 - Clinical T stage = cT3b-cT4 (extension into the seminal vesicles or invasion into adjacent structures)
 - Primary Gleason Pattern = 5
 - Gleason Grade Group = 4 or 5 in >4 cores
 - Presence of 2 or 3 high-risk features (noted above)

3D Rendering of MRI for MRI/Ultrasound Fusion Biopsy:

- When specific target lesion(s) is (are) detected on mpMRI (multi-parametric MRI) and classified as PIRADS 4 or 5, 3D Rendering (CPT® 76377) to generate prostate segmentation data image set for target identification on MRI/Transrectal ultrasound (TRUS) fusion biopsy is approvable as:
 - Subsequent separate standalone request; or
 - As retrospective request for medical necessity.
- For MRI/TRUS fusion biopsy of a PIRADS 1-3 lesion, approval of 3D rendering at independent workstation (CPT® 76376 or CPT® 76377) can be considered on a case-by-case basis.
- If there is no target lesion identified on MRI then 3D rendering and MRI/TRUS fusion biopsy is not generally indicated.
- 3D Rendering for the TRUS component of a fusion is a part of the UroNav Fusion Equipment Software and an additional CPT® code CPT® 76377 should not be approved.

Suspected Prostate Cancer (ONC-19.1)

ON.PR.0019.1.A

Inc	dication	Imaging Study
	IY of the following: Age 40-75 years with PSA >3 ng/ml or very suspicious DRE and ONE of the following high-risk features: • African ancestry • Germline mutations that increase the risk of prostate cancer • Family history of first or second-degree relative with prostate, male breast, colorectal, pancreatic, endometrial or female breast cancer at age <45 years Age 45-75 years and ONE of the following: • PSA >3 ng/ml • Very suspicious DRE Age >75 years and ONE of the following: • PSA ≥4 ng/ml • Very suspicious DRE At least one negative/nondiagnostic TRUS biopsy and ANY of the following: • Rising PSA • Abnormal DRE • Need for confirmatory MR/US fusion biopsy	ANY of the following: Transrectal ultrasound (CPT® 76872) TRUS-guided biopsy (CPT® 76942) MRI Pelvis without and with contrast (CPT® 72197) or MRI Pelvis without contrast (CPT® 72195) if an MR/US guided fusion biopsy is planned MRI/US fusion biopsy (CPT® 76942)
•	PIRADS 4 or 5 lesion identified on recent diagnostic MRI Pelvis (CPT® 72195 or CPT® 72197) and planning for biopsy to be done by MRI/TRUS fusion technique	3D Rendering (CPT® 76376 or CPT® 76377)

Indication	Imaging Study
 ANY of the following: Multifocal (3 or more lesions) high-grade prostatic intraepithelial neoplasia (PIN) Atypia on biopsy 	Extended pattern rebiopsy within 6 months by TRUS-guided biopsy (CPT® 76942)
Focal PIN (1-2 lesions)	 ONE of the following: MRI Pelvis without contrast (CPT® 72195) MRI Pelvis without and with contrast (CPT® 72197) MRI/US fusion biopsy (CPT® 76942) MRI guided biopsy (CPT® 77021)

Prostate Cancer – Initial Work-up/Staging (ONC-19.2)

ON.PR.0019.2.C

	V1.U.2U24
Indication	Imaging Study
Localized prostate cancer with any of the following risk groups (see: Prostate Cancer – General Considerations (ONC-19.0) for definition of risk groups): Very low risk Low risk Favorable intermediate risk	Advanced imaging is not routinely indicated for initial staging If not already performed prior to biopsy, MRI Pelvis without and with contrast (CPT® 72197) is appropriate for any of the following: • Prior to planned treatment (surgery and/or radiation therapy) • To establish candidacy for active surveillance
Localized prostate cancer with any of the following risk groups (see: Prostate Cancer – General Considerations (ONC-19.0) for definition of risk groups): • Unfavorable intermediate risk • High-risk • Very high-risk	Any ONE of the following combinations, not all (may be obtained in addition to mpMRI prostate): • CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and Bone scan • CT Chest with contrast (CPT® 71260), CT Abdomen with contrast (CPT® 74160), MRI Pelvis without and with contrast (CPT® 72197) if not previously performed, and Bone scan • PSMA PET/CT scan (CPT® 78815 or CPT® 78816) using any one of the following radiotracers: • 68 Ga-PSMA-11 • 18 F Piflufolastat (Pylarify®) • 68 Ga Gozetotide (Illuccix® and Locametz®) • 18 F Flotufolastat (Posluma®)
Known or clinically suspected metastatic prostate cancer (including prior to prostate biopsy)	CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and Bone scan
Inconclusive bone scan	CT with contrast or MRI without and with contrast of involved body site

Indication	Imaging Study
 For ANY of the following: Inconclusive bone findings on both CT/MRI and bone scan Conventional imaging studies (CT and bone scan) suggests oligo- or low volume metastatic disease that need further confirmation 	PET/CT scan (CPT ® 78815 or CPT ® 78816) using any one of the following radiotracers: 18F Fluciclovine 11C Choline 68Ga-PSMA-11 18F Piflufolastat (Pylarify®) 68Ga Gozetotide (Illuccix® and Locametz®) 18F Flotufolastat (Posluma®)

Prostate Cancer – Restaging/Recurrence (ONC-19.3)

ON.PR.0019.3.C

	V1.0.2024
Indication	Imaging Study
 For ANY of the following: Obvious progression by DRE with plans for prostatectomy or radiation therapy Repeat TRUS biopsy for rising PSA shows progression to a higher Gleason's score with plans for prostatectomy or radiation therapy Inconclusive findings on CT scan 	MRI Pelvis without and with contrast (CPT® 72197)
Non-metastatic prostate cancer previously treated with prostatectomy, radiation therapy, ablation, hormonal therapy or chemotherapy and any one of the following: Clinical suspicion of relapse/recurrence PSA fails to become undetectable post prostatectomy Palpable anastomotic recurrence PSA rises above post-treatment baseline to >0.2 ng/mL but <0.5 ng/mL on two consecutive measurements	 Any ONE of the following combinations: CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and Bone scan CT Chest with contrast (CPT® 71260), CT Abdomen with contrast (CPT® 74160), MRI Pelvis without and with contrast (CPT® 72197), and Bone scan

Indication	Imaging Study
 Non-metastatic prostate cancer previously treated with prostatectomy or radiation therapy, and all of the following are met: PSA rises on two consecutive measurements above post-treatment baseline and PSA ≥0.5 ng/mL and Individual is a candidate for salvage local therapy 	 Any ONE of the following combinations, not both: CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and Bone scan CT Chest with contrast (CPT® 71260), CT Abdomen with contrast (CPT® 74160), MRI Pelvis without and with contrast (CPT® 72197), and Bone scan PSMA PET/CT scan (CPT® 78815 or CPT® 78816) using any one of the following radiotracers: 68 Ga-PSMA-11 18 F Piflufolastat (Pylarify®) 68 Ga Gozetotide (Illuccix® and Locametz®) 18 F Flotufolastat (Posluma®)
 Non-metastatic prostate cancer previously treated with prostatectomy or radiation therapy, and all of the following are met: PSA rises on two consecutive measurements above post-treatment baseline and PSA ≥1 ng/mL and Recent CT scan and bone scan are negative for metastatic disease and Individual is a candidate for salvage local therapy Suspected progression of known metastatic disease based on: New or worsening signs/symptoms Rising PSA levels 	 PET/CT scan (CPT ® 78815 or CPT ® 78816) using any ONE of the following radiotracers: 18F-Fluciclovine 68Ga-PSMA-11 18F Piflufolastat (Pylarify®) 68Ga Gozetotide (Illuccix® and Locametz®) 18F Flotufolastat (Posluma®) CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and Bone scan CT with contrast of any involved or symptomatic body part

Indication	Imaging Study
Metastatic prostate cancer receiving treatment with chemotherapy	 CT Abdomen and Pelvis with contrast (CPT® 74177) and CT scan with contrast of any involved body part every 2 cycles (6 to 8 weeks) while on chemotherapy Bone scan may be obtained every 3-6 months
Metastatic prostate cancer receiving anti- androgen therapy	 CT Abdomen and Pelvis with contrast (CPT® 74177) and CT scan of any involved body part every 3 months while on anti-androgen therapy Bone scan may be obtained every 3-6 months
Previously treated metastatic prostate cancer progressed on conventional imaging and being considered for ¹⁷⁷ Lu-PSMA-617 (Pluvicto®) treatment ^{31, 32}	 PSMA PET/CT scan (CPT® 78815 or CPT® 78816) with one of the following agents: 68Ga PSMA-11 18F Piflufolastat (Pylarify®) 68Ga Gozetotide (Illuccix® and Locametz®) 18F Flotufolastat (Posluma®)
Prior to start of Xofigo (Radium-223) therapy	ONE time CT Chest, Abdomen and Pelvis with contrast (CPT® 71260 and CPT® 74177)
Inconclusive bone scan	CT with contrast or MRI without and with contrast of involved body site
 For ANY of the following: Inconclusive bone findings on both CT/MRI and bone scan Conventional imaging studies (CT and bone scan) suggests oligo- or low volume metastatic disease that needs further confirmation 	 PET/CT scan (CPT ® 78815 or CPT ® 78816) using any one of the following radiotracers: 18F Fluciclovine 11C Choline 68Ga-PSMA-11 18F Piflufolastat (Pylarify®) 68Ga Gozetotide (Illuccix® and Locametz®) 18F Flotufolastat (Posluma®)

Prostate Cancer – Follow-up On Active Surveillance (ONC-19.4)

ON.PR.0019.4.A

v1.0.2024

Active surveillance is being increasingly utilized in prostate cancer, and this therapeutic option involves regimented monitoring of an individual with known diagnosis of low risk prostate cancer for disease progression, without specific anticancer treatment. While being treated with active surveillance, an individual is generally considered a potential candidate for curative intent treatment approaches in the event that disease progression occurs.

It is important to distinguish active surveillance from watchful waiting (or observation), which is generally employed in individuals with limited life expectancy. Watchful waiting involves cessation of routine monitoring and treatment is initiated only if symptoms develop.

Current active surveillance guidelines suggest the following protocol:

- PSA every 6 months
- Digital Rectal Exam (DRE) every 12 months
- Repeat prostate biopsy every 12 months
- Repeat mpMRI (CPT® 72195 or CPT® 72197) no more often than every 12 months

Indication	Imaging Study
Routine monitoring on active surveillance protocol	MRI Pelvis without (CPT® 72195) or without and with contrast (CPT® 72197) at initiation of active surveillance, and every 12 months thereafter
 For ANY of the following: Progression is suspected based on DRE changes or rising PSA and a recent TRUS biopsy was negative Repeat TRUS biopsy shows progression to a higher Gleason score 	MRI Pelvis without (CPT® 72195) or MRI Pelvis without and with contrast (CPT® 72197)
Individuals on active surveillance who are noted to have progression and have plans to initiate treatment	 Imaging studies for initial staging as per ONC- 19.2

Surveillance/Follow-up For Treated Prostate Cancer (ONC-19.5)

ON.PR.0019.5.C

Indication	Imaging/Lab Study
 An individual with ALL of the following: Asymptomatic or stable chronic symptoms Stable DRE findings Stable PSA levels 	Advanced imaging not routinely indicated
 An individual with ANY of the following: New or worsening symptoms Change in DRE findings Rising PSA 	 ANY of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Pelvis without contrast (CPT® 72195) or without and with contrast (CPT® 72197)

References (ONC-19)

- 1. Schaeffer E, Srinivas S, Adra N, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 July 17, 2023. Prostate cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/prostate.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Prostate cancer V2.2023 July 17, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Moses KA, Sprenkle PC, Bahler C, et al. National Comprehensive Cancer Network (NCCN) Guidelines V1.2023 January 09, 2023. Prostate Cancer Early Detection available at: https://www.nccn.org/professionals/physician_gls/pdf/prostate detection.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Prostate Cancer Early Detection V1.2023 January 09, 2023 ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Jadvar H, Calais J, Fanti S, et. al. Appropriate use criteria for prostate-specific membrane antigen PET imaging, Society for Nuclear Medicine and Molecular Imaging. https://www.snmmi.org/ClinicalPractice/content.aspx?ItemNumber=38657.
- Hofman MS, Lawrentschuk N, Francis RJ, et. al. Prostate-specific membrane antigen PET-CT in patients with high-risk prostate cancer before curative-intent surgery or radiotherapy (proPSMA): a prospective, randomised, multicentre study. *Lancet*. 2020;395(10231):1208-1216. doi:10.1016/S0140-6736(20)30314-7.
- Pienta KJ, Gorin MA, Rowe SP, et. al. A phase 2/3 prospective multicenter study of the diagnostic accuracy of prostate specific membrane antigen PET/CT with ¹⁸F-DCFPyL in prostate cancer patients (OSPREY). J Urol. 2021;206(1):52-61. doi:10.1097/JU.00000000001698.
- 6. Artibani W, Porcaro AB, De Marco V, et. al. Management of biochemical recurrence after primary curative treatment for prostate cancer: A review. *Urol Int.* 2018;100:251–262. doi:10.1159/000481438.
- 7. Zumsteg ZS, Spratt DE, Romesser PB, et. al. The natural history and predictors of outcome following biochemical relapse in the dose escalation era for prostate cancer patients undergoing definitive external beam radiotherapy. *Eur Urol.* 2015;67(6):1009-1016. doi:10.1016/j.eururo.2014.09.028.
- 8. Trabulsi EJ, Rumble RB, Jadvar H, et al. Optimum imaging strategies for advanced prostate cancer: ASCO guideline. *J Clin Oncol*. 2020 Jan 15. doi:10.1200/JCO/19.02757 (Epub ahead of print).
- 9. Andriole G, Siegel B, LOCATE Study Group. PD60-12 Sites of prostate cancer recurrence delineated with 18F-Flucicloved positron emission tomography in patients with negative or equivocal conventional imaging. *Journal of Urology*. 2019;201(4):e1100-e1101. doi:10.1097/01.JU.0000557289.21741.20.
- ACR Appropriateness Criteria. Prostate cancer pretreatment detection, surveillance, and staging. Rev. 2016.
- Schoots IG, Nieboer D, Giganti F, Moore CM, Bangma CH, Roobol MJ. Is magnetic resonance imaging-targeted biopsy a useful addition to systematic confirmatory biopsy in men on active surveillance for low risk prostate cancer? A systematic review and meta-analysis. *BJU Int*. 2018;122(6):946-958. doi:10.1111/bju.14358.
- 12. Mullins J, Bodenkamp D, Landis P, et al. Multiparametric magnetic resonance imaging findings in men with low-risk prostate cancer followed by active surveillance. *BJU Int.* 2013;111(7):1037-1045. doi:10.1111/i.1464-410X.2012.11641.x.
- 13. Sanda MG, Chen RC, Crispino T, et al. *AUA/ASTRO/SUO guidelines for clinically localized prostate cancer.* Linthicum, MD: American Urological Association; 2017.
- 14. Lu-Yao GL, Albertsen PC, Moore DF, et al. Outcomes of localized prostate cancer following conservative management. *JAMA*. 2009;302(11):1202-1209. doi:10.1001/jama.2009.1348.
- 15. Chen RC, Rumble RB, Loblaw DA, et al. Active surveillance for the management of localized prostate cancer (Cancer Care Ontario guideline): American Society of Clinical Oncology clinical practice guideline endorsement. *J Clin Oncol.* 2016;34(18):2182-2190. doi:10.1200/JCO.2015.65.7759.
- 16. Liu D, Lehmann HP, Frick KD, Carter HB. Active surveillance versus surgery for low risk prostate cancer: a clinical decision analysis. *J Urol*. 2012;187(4):1241-1246. doi:10.1016/j.juro/2011.12.015.
- Klotz L, Zhang L, Lam A, Nam R, Mamedov A, Loblaw A. Clinical results of long-term follow-up of a large, active surveillance cohort with localized prostate cancer. *J Clin Oncol*. 2010;28(1):126-131. doi:10.1200/JCO.2009.24.2180.

- 18. Blomqvist L, Carlsson S, Gjertsson P, et al. Limited evidence for the use of imaging to detect prostate cancer: a systematic review. *Eur J Radiol.* 2014;83(9):1601–1606. doi:10.1016.j.ejrad.2014.06.028.
- 19. Schoots IG, Petrides N, Giganti F, et al. Magnetic resonance imaging in active surveillance of prostate cancer: a systematic review. *Eur Urol.* 2015;67(4):627-636. doi:10.1016/j.eururo.2014.10.050.
- Quentin M, Blondin D, Arsov C, et al. Prospective evaluation of magnetic resonance imaging guided inbore prostate biopsy versus systematic transrectal ultrasound guided prostate biopsy in biopsy naïve men with elevated prostate specific antigen. *J Urol*. 2014;192(5):1374-1379. doi:10.1016/j.juro.2014.05.090.
- 21. Klotz L, Vesprini D, Sethukavalan P, et al. Long-term follow-up of a large active surveillance cohort of patients with prostate cancer. *J Clin Oncol*. 2015;33(3):272-277. doi:10.1200/JCO.2014.55.1192.
- 22. Cooperberg MR. Long-term active surveillance for prostate cancer: answers and questions. *J Clin Oncol*. 2015;33(3):238-240. doi:10.1200/JCO.2014.59.2329.
- 23. Risko R, Merdan S, Womble PR, et al. Clinical predictors and recommendations for staging CT scan among men with prostate cancer. *Urology*. 2014;84(6):1329-1334. doi:10.1016.j.urology.2014.07.051.
- 24. Heck MM, Souvatzoglou M, Retz M, et al. Prospective comparison of computed tomography, diffusion-weighted magnetic resonance imaging and [11C]choline positron emission tomography/computed tomography for preoperative lymph node staging in prostate cancer patients. *Eur J Nucl Med Mol Imaging*. 2014;41(4):694-701. doi:10.1007/s00259-013-2634-1.
- 25. Armstrong JM, Martin CR, Dechet C, et al. ¹⁸F-fluciclovine PET CT detection of biochemical recurrent prostate cancer at specific PSA thresholds after definitive treatment. *J Urol Onc.* 2020;38(7):636.e1-636.e6. doi:10.1016/j.urolonc.2020.03.021.
- Baruch B, Lovrec P, Solanki A, et al. Fluorine 18 labeled fluciclovine PET/CT in clinical practice: factors affecting the rate of detection of recurrent prostate cancer. *AJR*. 2019;213(4):851-858. doi:10.2214/AJ.19.21153.
- 27. Marcus C, Butler P, Bagrodia A, et al. Fluorine-18-labeled fluciclovine PET/CT in primary and biochemical recurrent prostate cancer management. *AJR*. 2020:1-10. doi:10.2214/AJR.19.22404.
- 28. Trabulsi EJ, Rumble BR, Jadvar H, et. al. Optimum imaging strategies for advanced prostate cancer: ASCO quideline. J Clin Oncol. 2020;38:1963-1996. doi:10.1200/JCO.19.02757.
- 29. Lowrance WT, Breau RH, Chou R, et. al. Advanced Prostate Cancer: AUA/ASTRO/SUO Guideline PART I. *J Urol.* 2021;205:14.
- Epstein JI, Egevad L, Amin MB, Delahunt B, Srigley JR, Humphrey PA. The 2014 International Society of Urological Pathology (ISUP) consensus conference on gleason grading of prostatic carcinoma: definition of grading patterns and proposal for a new grading system. *Am J Surg Pathol.* 2016;40(2):244-52. doi:10.1097/PAS.00000000000000530.
- 31. FDA Oncology Center of Excellence. FDA approves Pluvicto for metastatic castration-resistant prostate cancer. 2022. https://www.fda.gov/drugs/resources-information-approved-drugs/fda-approves-pluvicto-metastatic-castration-resistant-prostate-cancer.
- 32. Sartor O, de Bono J, Chi KN, et. al. Lutetium-177-PSMA-617 for metastatic castration-resistant prostate cancer. *N Engl J Med*. 2021;385:1091-1103. doi:10.1056/NEJMoa2107322.

Testicular, Ovarian and Extragonadal Germ Cell Tumors (ONC-20)

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Testicular, Ovarian and Extragonadal Germ Cell Tumors – General Considerations (ONC-20.0)

ON.TO.0020.0.C

- This section applies to primary germ cell tumors occurring outside the central
 nervous system if individual's age >15 years at the time of initial diagnosis.
 Individuals age ≤15 years at diagnosis should be imaged according to pediatric
 guidelines in: Pediatric Germ Cell Tumors (PEDONC-10) in the Pediatric
 Oncology Imaging Guidelines.
- These guidelines are for germ cell tumors of the testicle, ovary, and extragonadal sites as well as malignant sex cord stromal tumors (granulosa cell and Sertoli-Leydig cell tumors).
- Requests for imaging must state the histologic type of the cancer being evaluated.
- Classified as pure seminomas (dysgerminomas, 40%) or Non-seminomatous germ cell tumors (NSGCT, 60%):
 - Pure seminomas are defined as pure seminoma histology with a normal serum concentration of alpha fetoprotein (AFP). Seminomas with elevated AFP are by definition Mixed
 - Required for TNM staging are the tumor marker levels indicated by "S" (TNMS)
 - Mixed tumors are treated as NSGCTs, as they tend to be more aggressive.
 - The NSGCT histologies include:
 - Yolk-Sac tumors
 - Immature (malignant) teratomas
 - Choriocarcinomas (<1%)
 - Embryonal cell carcinomas (15% to 20%)
 - Endodermal Sinus Tumors (ovarian)
 - Combinations of all of the above (mixed)
- PET/CT is considered experimental, investigational, or unproven for the evaluation of non-seminomatous germ cell tumors.
- Active surveillance in testicular cancer refers to treatment with surgery (orchiectomy) alone without any additional post-operative treatment such as chemotherapy or radiotherapy

Testicular, Ovarian and Extragonadal Germ Cell Tumors – Initial Work-Up/Staging (ONC-20.1)

ON.TO.0020.1.A

Indication	Imaging Study
Orchiectomy/oophorectomy is both diagnostic and therapeutic	 All individuals, following orchiectomy or oophorectomy: CT Abdomen and Pelvis with contrast (CPT® 74177)
 For ANY of the following: Non-seminoma histology Ovarian germ cell tumor Abdominal lymphadenopathy noted on CT scan Abnormal Chest X-ray or signs/symptoms suggestive of chest involvement 	CT Chest with contrast (CPT® 71260)
Extragonadal Germ Cell Tumor	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)

Testicular, Ovarian and Extragonadal Germ Cell Tumors – Restaging/Recurrence (ONC-20.2)

ON.TO.0020.2.A

Indication	Imaging Study
Treatment response for stage II-IV individuals with measurable disease on CT	CT with contrast of previously involved body areas every 2 cycles
Seminoma with residual mass >3 cm after completion of chemotherapy	• PET/CT (CPT® 78815)
End of therapy evaluation for NSGCT post chemotherapy or post retroperitoneal lymph node dissection (RPLND)	CT Abdomen and Pelvis with contrast (CPT® 74177)
Recurrence suspected, including increased tumor markers	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast Ultrasound (CPT® 76856 or CPT® 76857) of the remaining gonad if applicable
Unexplained pulmonary symptoms despite a negative chest x-ray, or new findings on chest x-ray	CT Chest with contrast (CPT® 71260)
All others	See: Surveillance (ONC-20.3)

Testicular, Ovarian and Extragonadal Germ Cell Tumors – Surveillance (ONC-20.3)

ON.TO.0020.3.C

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MRI in place of CT scans to reduce risk of secondary malignancy is not supported by the peer-reviewed literature. CT scans are indicated for surveillance and is the preferred modality of imaging to assess for recurrence.

Indication	Imaging Study
Stage I Seminoma treated with orchiectomy alone (no radiotherapy or chemotherapy, also called active surveillance)	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) once at 4-6 months and 12 months post-orchiectomy, then every 6 months for year 2 and 3, and then annually until year 5
Stage I Seminoma treated with radiotherapy and/or chemotherapy	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) annually for 3 years
Stage IIA and non-bulky Stage IIB Seminomas treated with radiotherapy or chemotherapy	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) once at 3 months then once at 9-12 months after completion of therapy, then annually for 2 additional years

Indication	Imaging Study
Bulky Stage IIB, IIC, and III Seminomas treated with chemotherapy	For individuals with ≤3 cm residual mass: • CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) every 4 months for 1 year, every 6 months for 1 year and then annually for 2 additional years
	For individuals with >3 cm residual mass and negative PET scan:
	 CT Abdomen and Pelvis with contrast (CPT[®] 74177) or CT Abdomen with contrast (CPT[®] 74160) at 6 and 12 months after completion of therapy, then annually until year 5
	 For individuals with thoracic disease: CT Chest with contrast (CPT® 71260) every 2 months for 1 year, then every 3 months for 1 year, then annually until year 5 after completion of therapy
Stage IA Non-Seminomatous germ cell tumors treated with orchiectomy alone (without risk factors)	 CT Abdomen and Pelvis with contrast (CPT[®] 74177) or CT Abdomen with contrast (CPT[®] 74160) every 6 months for 2 years and then annually for year 3
Stage IB Non-Seminomatous germ cell tumors treated with orchiectomy alone (with risk factors – lymphovascular invasion or invasion into spermatic cord/scrotum)	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) every 4 months for 1 year, then every 6 months for 2 years, then annually until year 4
Stage IA/IB Non-Seminomatous germ cell tumors treated with chemotherapy and/or primary RPLND	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) annually for 2 years
Stage II-III Non-Seminomatous germ cell tumors with complete response to chemotherapy +/-post-chemotherapy RPLND	CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) once at 6, 12, 24, and 36 months after completion of therapy
	For individuals with thoracic disease:
	 CT Chest with contrast (CPT® 71260) every 6 months for 2 years, then annually until year 4 after completion of therapy

Indication	Imaging Study
Stage IIA or IIB Non- Seminomatous germ cell tumors treated with post-primary RPLND and adjuvant chemotherapy	 CT Abdomen and Pelvis with contrast (CPT[®] 74177) or CT Abdomen with contrast (CPT[®] 74160) once at 4 months after completion of RPLND
Stage IIA or IIB Non- Seminomatous germ cell tumors treated with post-primary RPLND without adjuvant chemotherapy	 CT Abdomen and Pelvis with contrast (CPT® 74177) or CT Abdomen with contrast (CPT® 74160) once at 3 to 4 months after completion of therapy and repeat annually for 1 year
All stages of ovarian dysgerminoma germ cell tumors	CT Abdomen and Pelvis with contrast (CPT® 74177) every 4 months for 1 year, every 6 months for 1 year and then annually for 3 years after completion of therapy
All ovarian non-dysgerminoma germ cell tumors:	 CT Abdomen and Pelvis with contrast (CPT® 74177) every 4 months for 1 year, every 6 months for 1 year and then annually for 3 years after completion of therapy CT Chest with contrast (CPT® 71260) every 4 months for 1 year and every 6 months for 1 year after completion of therapy
Sex cord stromal tumors (male and female)Mature teratoma	No routine advanced imaging indicated unless elevated tumor markers or clinical signs/symptoms of recurrence
Extragonadal germ cell tumors	CT of the involved region every 3 months for one year and every 6 months for one year

References (ONC-20)

- 1. Gilligan T, Lin DW, Aggarwal R, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 January 26, 2023. Testicular cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/testicular.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Testicular cancer V1.2023 January 26, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Armstrong DK, Alvarez RD, Backes FJ, et. al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 June 2, 2023. Ovarian cancer, including fallopian tube cancer and primary peritoneal cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/ovarian.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Ovarian cancer V2.2023 June 2, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines ™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Salani R, Backes FJ, Fung MF, et al. Post treatment surveillance and diagnosis of recurrence in women with gynecologic malignancies: Society of Gynecologic Oncologists recommendations. *Am J Obstet Gynecol*. 2011;204(6):466-478. doi:10.1016/j.ajog.2011.03.008.
- Gershenson DM. Management of ovarian germ cell tumors. J Clin Oncol. 2007;25(20):2938-2943. doi:10.1200/JCO.2007.10.8738.
- 5. Colombo N, Parma G, Zanagnolo V, Insinga A. Management of ovarian stromal cell tumors. *J Clin Oncol*. 2007;25(20):2944-2951. doi:10.1200/JCO.2007.11.1005.
- 6. Cadron I, Leunen K, Van Gorp T, Amant F, Neven P, Vergote I. Management of Borderline Ovarian Neoplasms. *J Clin Oncol*. 2007;25(20):2928-2937. doi:10.1200/JCO/2007.10.8076.
- 7. del Carmen MG, Birrer M, Schorge JO. Carcinosarcoma of the ovary: a review of the literature. *Gynecol Oncol.* 2012;125(1):271-277. Doi:10.1016/j.ygyno.2011.12.418.
- Kollmannsberger C, Tandstad T, Bedard PL, et al. Patterns of relapse in patients with clinical stage I testicular cancer managed with active surveillance. *J Clin Oncol*. 2015;33(1):51-57. doi:10.1200/JCO.2014.56.2116.
- Oechsle K, Hartmann M, Brenner W, et al. [18F]Fluorodeoxyglucose positron emission tomography in nonseminomatous germ cell tumors after chemotherapy: the German multicenter positron emission tomography study group. *J Clin Oncol*. 2008;26(36):5930-5935. doi:10.1200/JCO.2008.17.1157.
- Daugard G, Gundgaard MG, Mortensen MS, et al. Surveillance for stage I non seminoma testicular cancer: outcomes and long term follow-up in a population based cohort. *J. Clin Oncol.* 2014;32(34):3817-3823. doi:10.1200/JCO.2013.53.5831.
- 11. Zuniga A, Kakiashvilli D, Jewett MA. Surveillance in stage I nonseminomatous germ cell tumours of the testis. *BJU Int.* 2009;104:1351-1356. doi:10.1111/j.1464-410X.2009.08858.x.

Ovarian Cancer (ONC-21)

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Ovarian Cancer – General Considerations (ONC-21.0)

ON.OC.0021.0.A

- Ovarian cancers include: epithelial ovarian cancers, ovarian cancers of low malignant potential and mixed Müllerian tumors, primary peritoneal and fallopian tube cancers.
 - o There are five main types of epithelial ovarian cancers:
 - High-grade serous carcinoma (HGSC) (70%)
 - Endometrioid carcinoma (EC) (10%)
 - Clear cell carcinoma (CCC) (10%)
 - Mucinous carcinoma (MC) (3%)
 - Low-grade serous carcinoma (LGSC) (<5%)
- Borderline tumors (formerly referred to as tumors of low malignant potential) usually have some feature of carcinoma when they recur.
- Fallopian tube and primary peritoneal are usually serous carcinoma.
- Germ cell tumors and sex cord stromal tumors (granulosa cell tumors), are imaged according to <u>Testicular</u>, <u>Ovarian and Extragonadal Germ Cell Cancer</u> (<u>ONC-20</u>).

Screening for Ovarian Cancer (ONC-21.1)

ON.OC.0021.1.C

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• The use of advanced imaging in ovarian cancer screening is not recommended by the USPSTF, and is considered experimental and investigational.

Ovarian Cancer – Suspected/Diagnosis (ONC-21.2)

ON.OC.0021.2.C

Indication	Imaging/Lab Study
Pelvic symptoms (pelvic pain, abdominal bloating)Palpable pelvic mass	Transvaginal (TV) ultrasound (CPT® 76830) and/or Pelvic ultrasound (CPT® 76856 or CPT® 76857)
 Ultrasound shows complex and/or solid adnexal mass suspicious for ovarian malignancy or Any suspicious signs/symptoms for ovarian malignancy (ascites, abdominal symptoms such as distension or tenderness, elevated CA-125, elevated LFTs, obstructive uropathy* 	 ANY of the following: MRI Pelvis without and with contrast (CPT® 72197) MRI Pelvis without contrast (CPT® 72195) if contrast is contraindicated CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen and Pelvis without and with contrast (CT Urogram – CPT® 74178) only for symptoms of obstructive uropathy*

Ovarian Cancer – Initial Work-Up/Staging (ONC-21.3)

ON.OC.0021.3.A

Indication	Imaging Study
Clinical stage II disease or higher	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for: Abnormal signs/symptoms of pulmonary disease Abnormal chest x-ray
 ANY of the following: Primary peritoneal disease with biopsy-proven malignancy consistent with ovarian carcinoma Elevated tumor markers with negative or inconclusive CT imaging 	• PET/CT (CPT® 78815)

Ovarian Cancer – Restaging/Recurrence (ONC-21.4)

ON.OC.0021.4.A

Indication	Imaging Study
Completely resected or definitively treated with chemotherapy and normal(ized) tumor markers	No advanced imaging needed
 ANY of the following: Unresected disease Unknown preoperative markers Difficult or abnormal examination Elevated LFTs Elevated tumor markers (CA-125, inhibin) Signs or symptoms of recurrence 	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for ANY of the following: Known prior thoracic disease New or worsening pulmonary symptoms New or worsening chest x-ray findings Rising tumor markers (CA-125, inhibin)
Monitoring response to treatment (every 2 cycles, or ~every 6 to 8 weeks)	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for ANY of the following: Known prior thoracic disease New or worsening pulmonary symptoms New or worsening chest x-ray findings
 CT negative or inconclusive and CA-125 continues to rise or elevated LFTs Conventional imaging failed to demonstrate tumor or if persistent radiographic mass with rising tumor markers 	• PET/CT (CPT® 78815)

Ovarian Cancer – Surveillance (ONC-21.5)

ON.OC.0021.5.A

Indication	Imaging Study
Stages I-III	Advanced imaging is not routinely indicated for surveillance
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 Every 3 months for up to 5 years after completion of active treatment: CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved body areas

References (ONC-21)

- 1. Armstrong DK, Alvarex RD, Backes FJ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 June 2, 2023. Ovarian Cancer Including Fallopian Tube Cancer and Primary Peritoneal Cancer available at: https://www.nccn.org/professionals/physician_gls/pdf/ovarian.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Ovarian Cancer Including Fallopian Tube Cancer and Primary Peritoneal Cancer V2.2023 June 2, 2023. [®]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Daly MB, Pal T, AlHilli Z, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version V3.2023 February 13, 2023. Genetic/Familial High-Risk Assessment: Breast, Ovarian, and Pancreatic available at: https://www.nccn.org/professionals/physician_gls/pdf/genetics_bop.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Ovarian cancer V3.2023 February 13, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Moyer VA, U.S. Preventive Services Task Force. Screening for ovarian cancer: U.S. Preventive Services
 Task Force reaffirmation recommendation statement. *Ann Intern Med.* 2012;157(12):900-904.
 doi:10.7326/0003-4819-157-11-201212040-00539.
- Cadron I, Leunen K, Van Gorp T, Amant F, Neven P, Vergote I. Management of borderline ovarian neoplasms. J Clin Oncol. 2007;25(20):2928-2937. doi:10.1200/JCO.2007.10.8076.
- 5. ACR Appropriateness Criteria. Ovarian cancer screening. Rev. 2017.
- Rosenthal AN, Fraser LSM, Phipott S. Evidence of stage shift in women diagnosed with ovarian cancer during phase II of the United Kingdom familial ovarian cancer screening study. *J Clin Oncol*. 2017;35(13):13:1411-1420. doi:10.1200/JCO.2016.69.9330.
- Shinagare AB, O'Neill AC, Cheng S, et al. Advanced high-grade serous ovarian cancer: frequency and timing of thoracic metastases and the implications for chest imaging follow-up. *Radiology*. 2015;277(3):733-740. doi:10.1148/radiol.2015142467.
- Musto A, Grassetto G, Marzola MC, et al. Management of epithelial ovarian cancer from diagnosis to restaging: an overview of the role of imaging techniques with particular regard to the contribution of 18F-FDG PET/CT. *Nucl Med Commun.* 2014;35(6):588-597. doi:10.1097/MNM.00000000000000091.
- 9. Fischerova D, Burgetova A. Imaging techniques for the evaluation of ovarian cancer. *Best Pract Res Clin Obstet Gynaecol.* 2014;28(5):697-720. doi:10.1016/j.bpobgyn.2014.04.006.

Uterine Cancer (ONC-22)

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Uterine Cancer – General Considerations (ONC-22.0)

ON.UC.0022.0.C

- Gestational trophoblastic neoplasia (GTN) see: Molar Pregnancy and
 <u>Gestational Trophoblastic Neoplasia (GTN) (PV-16.1)</u> in the Pelvic Imaging
 Guidelines.
- Imaging not routinely indicated pre-operatively for laparoscopic/minimally invasive surgery unless initial staging criteria are met. Pelvic and para-aortic lymphadenectomy can still be performed.

Uterine Cancer – Suspected/Diagnosis (ONC-22.1)

ON.UC.0022.1.A

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• See: <u>Abnormal Uterine Bleeding (PV-2.1)</u> in the Pelvic Imaging Guidelines for evaluation of suspected uterine malignancies.

Uterine Cancer – Initial Work-Up/Staging (ONC-22.2)

ON.UC.0022.2.A

	V1.0.2024
Indication	Imaging Study
 ANY of the following: Extrauterine disease suspected To assess local extent of tumor prior to fertility-sparing surgery for well-differentiated Stage IA (grade 1) uterine cancer Poor surgical candidate (due to medical comorbidities) considering medical therapy 	 MRI Pelvis without and with contrast (CPT® 72197) Transvaginal ultrasound (CPT® 76830) if MRI is contraindicated Chest x-ray CT Chest with contrast (CPT® 71260) if chest x-ray is abnormal
 ANY of the following: Abdominal symptoms or abnormal examination findings Elevated LFTs Other imaging studies suggest liver involvement 	 ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen with contrast (CPT® 74160)
ANY of the following high-risk histologies: Papillary serous Clear cell High-grade/poorly differentiated endometrioid carcinoma Uterine sarcomas: Carcinosarcoma Soft tissue sarcoma of the uterus Leiomyosarcoma Rhabdomyosarcoma Undifferentiated sarcoma Endometrial stromal sarcoma	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast

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Indication	Imaging Study
Tumors detected incidentally or incompletely staged surgically and ANY of the following high-risk features: • Myoinvasion >50% • Cervical stromal involvement • Lymphovascular invasion • Tumor >2 cm	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
Inconclusive findings on conventional imaging	PET/CT scan (CPT® 78815)

Uterine Cancer – Restaging/Recurrence (ONC-22.3)

ON.UC.0022.3.C

Indication	Imaging Study
 Unresected disease Medically inoperable disease Incomplete surgical staging Difficult or abnormal examination Elevated LFTs or rising tumor markers Signs or symptoms of recurrence 	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Monitoring response to chemotherapy (every 2 cycles, ~every 6-8 weeks) for: • Unresected primary disease • Metastatic disease	 CT Abdomen and Pelvis with contrast (CPT® 74177) CT Chest with contrast (CPT® 71260) for: New/worsening pulmonary symptoms Known prior pulmonary involvement
ANY of the following: • After fertility sparing treatment • Inconclusive CT scan findings	MRI Pelvis without and with contrast (CPT® 72197)
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Uterine Cancer – Surveillance (ONC-22.4)

ON.UC.0022.4.A

Indication	Imaging Study
Stage I-III of uterine carcinoma	Advanced imaging is not routinely indicated for surveillance
Measurable metastatic disease on maintenance therapy or being monitored off therapy	 Every 3 months for up to 5 years after completion of definitive treatment: CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved body areas
 All stages of uterine sarcoma: Soft tissue sarcoma of the uterus Leiomyosarcoma Adenosarcoma Carcinosarcoma Rhabdomyosarcoma Undifferentiated sarcoma Endometrial stromal sarcoma 	CT Chest (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) every 3 months for 2 years, every 6 months for 3 years, and then every 1-2 years until year 10

Gestational Trophoblastic Neoplasia (GTN) (ONC-22.5)

ON.UC.0022.5.A

- The most common form of gestational trophoblastic disease (GTD) is hydatidiform mole (HM), a benign form, also known as molar pregnancy.
 - See: Molar Pregnancy and GTN (PV-16.1)
- Gestational trophoblastic neoplastic disorders including a malignant form of GTD, and can present as invasive mole, choriocarcinoma, placental site trophoblastic tumor (PSTT), or epithelioid trophoblastic tumor (ETT). GTN cells are malignant and can metastasize to other organs such as lungs, brain, bone and vagina. These tumors have a high likelihood of cure and treatment with methotrexate usually allows for fertility preservation.
- Surveillance is generally with serial monitoring of HCG levels, and advanced imaging is reserved for high-risk histologies where HCG levels may not be a reliable marker.

marker.		
Indication	Imaging Study	
Initial staging	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 	
 EITHER of the following: Pulmonary metastases noted on CT scan Signs/symptoms of CNS involvement 	MRI Brain without and with contrast (CPT® 70553)	
 EITHER of the following: Monitoring response to systemic therapy (every 2 cycles, i.e., 6-8 weeks Suspected progression 	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 	
Surveillance for any of the following high risk histologies: Placental site trophoblastic tumor (PSTT) Epithelioid trophoblastic tumor (ETT)	 Annually for 2 years: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 	

References (ONC-22)

- 1. Abu-Rustum NR, Yashar CM, Arend R, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 April 28, 2023. Uterine Neoplasms, available at: https://www.nccn.org/professionals/physician_gls/pdf/uterine.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Uterine Neoplasms 2.2023 April 28, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Fader AN, Boruta D, Olawaiye AB, Gehrig PA. Updates on uterine papillary serous carcinoma. *Expert Rev Obstet Gynecol*. 2009;4(6):647-657. doi:10.1586/eog.09.49.
- Boruta DM 2nd, Gehrig PA, Fader AN, Olawaiye AB. Management of women with uterine papillary serous cancer: A Society of Gynecologic Oncology (SGO) review. *Gynecol Oncol*. 2009;115(1):142-153. doi:10.1016/j.ygyno.2009.06.011.
- Olawaiye AB, Boruta DM 2nd. Management of women with clear cell endometrial cancer: a Society of Gynecologic Oncology (SGO) review. *Gynecol Oncol*. 2009;113(2):277-283. doi:10.1016/j.ygyno.2009.02.003.
- 5. Salani R, Backes FJ, Fung MF et al. Post treatment surveillance and diagnosis of recurrence in women with gynecologic malignancies: Society of Gynecologic Oncologists recommendations. *Am J Obstet Gynecol.* 2011;204(6):466-478. doi:10.1016/j.ajog.2011.03.008.
- Reinhold C, Ueno Y, Akin EA, et. al. ACR Appropriateness Criteria[®] Evaluation and follow-up of endometrical cancer. Available at https://acsearch.acr.org/docs/69459/Narrative/. American College of Radiology. Accessed 7/29/2020.
- 7. Abu-Rustum NR, Yashar CM, Bradley, K, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2021 March 31, 2021. Gestational Trophoblastic Neoplasia, available at: https://www.nccn.org/professionals/physician_gls/pdf/cns.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™ for Gestational Trophoblastic Neoplasia, March 31, 2021 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.

Cervical Cancer (ONC-23)

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Cervical Cancer – General Considerations (ONC-23.0)

ON.CV.0023.0.A

- Primary histology for cervical cancer is squamous cell. Other, less common histologies are adenosquamous and adenocarcinoma. If biopsy is consistent with one of these less common histologies, it is necessary to clarify that tumor is not of primary uterine origin.
- If the primary histology is uterine in origin, follow imaging recommendations for uterine cancer, see: <u>Uterine Cancer (ONC-22)</u>.

Cervical Cancer – Suspected/Diagnosis (ONC-23.1)

ON.CV.0023.1.A

Indication	Imaging Study
All	Biopsy should be performed prior to imaging

Cervical Cancer – Initial Work-Up/Staging (ONC-23.2)

ON.CV.0023.2.A

Indication	Imaging Study
Stage IB1 or higher stages	 ANY of the following combinations, not both: PET/CT (CPT® 78815) or CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Any size cervical cancer incidentally found in a hysterectomy specimen	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
 ANY of the following: To assess local extent of disease prior to fertility sparing treatment To assess residual pelvic disease post-operatively Inconclusive CT findings 	MRI Pelvis without and with contrast (CPT® 72197)
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Cervical Cancer – Restaging/Recurrence (ONC-23.3)

ON.CV.0023.3.A

Indication	Imaging Study
If primary therapy was surgery	See: Cervical Cancer – Surveillance (ONC-23.4)
If primary therapy radiation therapy ± chemotherapy	 ANY of the following, not both: PET/CT (CPT® 78815) at least 12 weeks after completion of treatment
(no surgery)	OR CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
Unresectable disease or metastatic disease on systemic treatment	 Every 2 cycles of treatment (commonly every 6 to 8 weeks): CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of other involved or symptomatic areas
Suspected or biopsy proven recurrence	ANY of the following, not both: • PET/CT (CPT® 78815) OR • CT Chest (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Inconclusive findings on CT scan	MRI Pelvis without and with contrast (CPT® 72197)

Cervical Cancer – Surveillance (ONC-23.4)

ON.CV.0023.4.A

Indication	Imaging Study
Stage I disease treated with fertility sparing approach	 MRI Pelvis without and with contrast (CPT® 72197) at 6 months after surgery and then annually for 2 years
All individuals	No routine advanced imaging needed in asymptomatic individuals.

References (ONC-23)

- 1. Abu-Rustum NR, Yashar CM, Arend R, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 April 28, 2023. Cervical Cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/cervical.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Cervical Cancer V1.2023 April 28, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Salani R, Backes FJ, Fung MF et al. Post treatment surveillance and diagnosis of recurrence in women with gynecologic malignancies: Society of Gynecologic Oncologists recommendations. Am J Obstet Gynecol. 2011;204(6):466-478. doi:10.1016/j.ajog.2011.03.008.
- 3. Zanagnolo V, Ming L, Gadducci A, et al. Surveillance procedures for patients with cervical carcinoma: a review of the literature. *Int J Gynecol Cancer.* 2009;19(3):194-201. doi:10.1111/IGC.0b013e3181a130f3.
- 4. Elit L, Fyles AW, Devries MC, et al. Follow-up for women after treatment for cervical cancer: A systematic review. *Gynecol Oncol.* 2009;114(3):528-535. doi:10.1016/j.ygyno.2009.06.001.
- 5. Schwarz JK, Siegel BA, Dehdashti F, Grigsby PW. Association of posttherapy positron emission tomography with tumor response and survival in cervical carcinoma. *JAMA*. 2007;298(19):2289-2295. doi:10.1001/jama.298.19.2289.
- Meads C, Davenport C, Malysiak S, et al. Evaluating PET-CT in the detection and management of recurrent cervical cancer: systematic reviews of diagnostic accuracy and subjective elicitation. *BJOG*. 2014;121(4):398-407. doi:10.1111/1471-0528.12488.

Anal Cancer & Cancers of the External Genitalia (ONC-24)

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Anal Carcinoma – General Considerations (ONC-24.0)

ON.AN.0024.0.A

- Most are squamous cell carcinomas, although some transitional and cloacogenic carcinomas are seen.
- Adenocarcinoma of the anal canal is managed as rectal cancer according to <u>Colorectal and Small Bowel Cancer (ONC-16)</u>
- Squamous cell carcinoma of the perianal region (up to 5 cm radius from the anal verge) are imaged according to anal carcinoma guidelines.
- Bowen's disease and Paget's disease of the perianal and perigenital skin are considered non-invasive/in-situ conditions and do not routinely require advanced imaging. See: <u>Non-Melanoma Skin Cancers – Initial Work-up/Staging (ONC-5.6)</u>

Anal Carcinoma – Suspected/Diagnosis (ONC-24.1)

ON.AN.0024.1.A

Indication	Imaging Study
All	Advanced imaging prior to biopsy is not needed

Anal Carcinoma – Initial Work-up/Staging (ONC-24.2)

ON.AN.0024.2.A

	V 1.U.ZUZ4
Indication	Imaging Study
All individuals	CT Chest with contrast (CPT® 71260) and
	 Any ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) CT Abdomen with contrast (CPT® 74160) and MRI Pelvis without and with contrast (CPT® 72197)
 Stage II-III Squamous Cell Carcinoma of the Anal Canal and no evidence of metastatic disease by conventional imaging Inconclusive findings on conventional imaging 	• PET/CT (CPT® 78815)

Anal Carcinoma – Restaging/Recurrence (ONC-24.3)

ON.AN.0024.3.A

Indication	Imaging Study
Stage I treated with complete surgical resection	See: Anal Carcinoma – Surveillance (ONC-24.4) for surveillance guidelines
Stages I, II and III – post chemoradiation evaluation	 Any ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Metastatic (stage IV) disease	 CT Abdomen and Pelvis with contrast (CPT® 74177) every 2 cycles (generally 6 to 8 weeks) on treatment CT Chest with contrast (CPT® 71260) if Chest X-ray is abnormal or if symptoms of chest involvement
 Difficult or abnormal examination Elevated LFTs Signs or symptoms of recurrence Biopsy proven recurrence 	 CT Chest (CPT® 71260) with contrast and Any ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Anal Carcinoma – Surveillance (ONC-24.4)

ON.AN.0024.4.A

Indication	Imaging Study
Stage I	Advanced imaging is not routinely indicated for surveillance
 Stage II Stage III Local recurrence treated definitively 	 CT Chest (CPT® 71260) with contrast or CT Chest without contrast (CPT® 71250) annually for 3 years And ANY one of the following annually for three years: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen without and with contrast (CPT® 74183) and MRI Pelvis without and with contrast (CPT® 72197)
Stage IV – measurable metastatic disease on maintenance treatment or being observed off treatment	 Every 3 months for up to 5 years after completion of all treatment: CT Chest (CPT® 71260) with contrast CT Abdomen and Pelvis with contrast (CPT® 74177)

Cancers of External Genitalia – General Considerations (ONC-24.5)

ON.AN.0024.5.A

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• These imaging guidelines are applicable for squamous cell carcinomas arising from the vulva, vagina, penis, urethra, and scrotum

Cancers of External Genitalia – Initial Work-Up/Staging (ONC-24.6)

ON.AN.0024.6.A

Indication	Imaging Study
Clinical node negative vulvar cancer with ANY of the following: • Lesion >2 cm • Any size with stromal invasion >1 mm	 For planned sentinel lymph node (SLN) biopsy: Lymph system imaging (lymphoscintigraphy, CPT® 78195) SPECT/CT (CPT® 78830) is indicated as an add on code if requested
For stage II or higher	 ONE of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) OR CT Abdomen with contrast (CPT® 74160) and MRI Pelvis without and with contrast (CPT® 72197) CT Chest with contrast (CPT® 71260) is indicated only for: Signs/symptoms suggestive of chest involvement Abnormal findings on chest x-ray
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Cancers of External Genitalia – Restaging/Recurrence (ONC-24.7)

ON.AN.0024.7.A

Indication	Imaging Study
 Difficult or abnormal examination Elevated LFTs Signs or symptoms of recurrence Biopsy proven recurrence 	 CT Chest with contrast (CPT® 71260) And ANY one of the following: CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast
Individuals receiving systemic treatment	 CT Abdomen and Pelvis with contrast (CPT® 74177) every 2 cycles (generally 6 to 8 weeks) during treatment and at the end of planned chemotherapy treatment CT Chest with contrast (CPT® 71260) if chest x-ray is abnormal or if symptoms of chest involvement
Inconclusive findings on conventional imaging	• PET/CT (CPT® 78815)

Cancers of External Genitalia – Surveillance (ONC-24.8)

ON.AN.0024.8.A

Indication	Imaging Study
All stages of vulvar and vaginal cancers	Routine advanced imaging is not indicated for asymptomatic surveillance
Penile Cancer: stage I- IIIA	Routine advanced imaging is not indicated for asymptomatic surveillance
Penile cancer: stages IIIB and higher	CT Abdomen and Pelvis with contrast (CPT® 74177) every 3 months for year 1, and then every 6 months for year 2, then no further routine advanced imaging indicated

References (ONC-24)

- 1. Benson III AB, Venook AP, Al-Hawary MM, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 April 28, 2023. Anal Carcinoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/anal.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for anal carcinoma V2.2023– April 28, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Flaig TW, Spiess PE, Agarwal N. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 December 1, 2022. Penile Cancer, available at: https://www.nccn.org/professionals/physician_gls/pdf/penile.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Penile Cancer V1.2023 December 1, 2022. [©]2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Abu-Rustum NR, Yashar CM, Arend R. et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 December 22, 2022. Vulvar Cancer (Squamous Cell Carcinoma), available at: htt ps://www.nccn.org/professionals/physician_gls/pdf/vulvar.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Vulvar Cancer (Squamous Cell Carcinoma) V1.2023 December 22, 2022. ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 4. Bhuva NJ, Glynne-Jones R, Sonoda L, Wong WL, Harrison MK. To PET or not to PET? That is the question. Staging in anal cancer. *Ann Oncol.* 2012;23(8):2078-2082. doi:10.1093/annonc/mdr599.
- Mistrangelo M, Pelosi E, Bellò M, et al. Role of positron emission tomography-computed tomography in the management of anal cancer. *Int J Radiat Oncol Biol Phys.* 2012;84(1):66-72. doi:10.1016/j.ijrobp.2011.10.048.
- Jones M, Hruby G, Solomon M, Rutherford N, Martin J. The role of FDG-PET in the initial staging and response assessment of anal cancer: a systematic review and meta-analysis. *Ann Surg Oncol*. 2015;22(11):3574-3581. doi:10.1245/s10434-015-4391-9.
- 7. Moncrieff M, Pywell S, Snelling A, et. al. Effectiveness of SPECT/CT imaging for sentinel node biopsy staging of primary cutaneous melanoma and patient outcomes. *Ann Surg Oncol*. 2022;29(2):767-775. doi:10.1245/s10434-021-10911-4.
- 8. Quartuccio N, Garau LM, Arnone A, et. al. Comparsion of 99mTc-labeled colloid SPECT/CT and planar lymphoscintigraphy in sentinel lymph node detection in patients with melanoma: a meta-analysis. *J Clin Med*. 2020;9(6):1680. doi:10.3390/jcm9061680.
- 9. Bennie G, Vorster M, Buscombe J, Sathekge M. The added value of a single-photon emission computed tomography-computed in sentinel lymph node mapping in patients with breast cancer and malignant melanoma. *World J Nucl Med*. 2015;14(01):41-46. doi:10.4103/1450-1147.150543

Multiple Myeloma and Plasmacytomas (ONC-25)

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Multiple Myeloma and Plasmacytomas – General Considerations (ONC-25.0)

ON.MM.0025.0.A

- Multiple myeloma (MM) is a neoplastic disorder characterized by the proliferation of a single clone of plasma cells derived from B cells which grows in the bone marrow and adjacent bone, producing skeletal destruction.
- Multiple myeloma group of disorders can be classified as below, which influence imaging modality of choice.

Condition	Monoclonal protein	Bone marrow plasma cells	CRAB criteria**
Solitary Plasmacytoma (biopsy proven tumor containing plasma cells)	<3 gm/dL	Absent	Absent
Monoclonal Gammopathy of Unknown Significance (MGUS)	<3 gm/dL	<10%	Absent
Smoldering Myeloma (SMM) (stage I MM or asymptomatic MM)	≥3 gm/dL	10% - 60%	Absent
Multiple Myeloma (MM)	≥3 gm/dL	≥10%	Present

- **CRAB criteria = hypercalcemia, renal insufficiency, anemia, lytic bony lesions
- Diagnosis and monitoring of response to therapy is primarily with laboratory studies that include urine and serum monoclonal protein levels, serum free light chain levels, LDH and beta-2 microglobulin. Routine advanced imaging to monitor response to treatment is not indicated.
- Rarely, (<5%), an individual may have Nonsecretory Myeloma, which does not produce measurable M-protein. These individuals require imaging as primary method to monitor disease.
- Other conditions that may present with Monoclonal Gammopathy include:
 - <u>POEMS syndrome</u>: Polyneuropathy, Organomegaly, Endocrinopathy, Monoclonal protein and Skin Changes – may also have sclerotic bone lesions and Castleman's disease. See: <u>Multiple Myeloma and Plasmacytomas –</u> <u>Initial Work-up/Staging (ONC-25.2)</u> for imaging recommendations.
 - <u>Waldenström's Macroglobulinemia</u>: IgM monoclonal protein along with bone marrow infiltration of small lymphocytes. See: <u>Waldenström</u> <u>Macroglobulinemia or Lymphoplasmacytic Lymphoma (ONC-27.10)</u> for imaging recommendations.
 - Systemic Light chain Amyloidosis: light chain monoclonal protein in serum or urine with clonal plasma cells in bone marrow, systemic involvement of the kidneys, liver, heart, gastrointestinal tract or peripheral nerves due to amyloid

deposition. See: <u>Multiple Myeloma and Plasmacytomas – Initial</u> <u>Work-up/Staging (ONC-25.2)</u> and <u>Cardiac Amyloidosis (CD-3.8)</u> for imaging recommendations for systemic light chain amyloidosis.

Multiple Myeloma and Plasmacytomas –Suspected/Diagnosis (ONC-25.1)

ON.MM.0025.1.A

Indication	Imaging Study
All	X-ray skeletal series

Multiple Myeloma and Plasmacytomas – Initial Work-Up/Staging (ONC-25.2)

ON.MM.0025.2.A

	v1.0.2024
Indication	Imaging Study
ANY of the following:Abnormal skeletal surveyAbnormal myeloma labsSigns/symptoms of multiple myeloma	Whole-body low-dose skeletal CT (CPT® 76497)
 ANY of the following: If skeletal CT is negative, inconclusive, or not feasible Suspected solitary bone/osseous plasmacytoma 	 ONE of the following: MRI Bone Marrow Blood Supply (CPT® 77084) MRI Cervical (CPT® 72141), Thoracic (CPT® 72146), Lumbar spine (CPT® 72148), and Pelvis (CPT® 72195) without contrast MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), Lumbar spine (CPT® 72158), and Pelvis (CPT® 72197) without and with contrast CT contrast as requested of a specific area to determine radiotherapy or surgical candidacy, or for suspected extra-osseous plasmacytoma
 ANY of the following (after above tests completed): Determine if plasmacytoma is truly solitary Suspected extra-osseous plasmacytomas Suspected progression of MGUS or SMM to a more malignant form and CT/MRI imaging are negative Whole-body skeletal CT and MRI Bone Marrow are negative, inconclusive, or not feasible 	• PET/CT (CPT® 78815 or CPT® 78816)
ANY of the following: Systemic light chain amyloidosis POEMS syndrome	 CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)

Multiple Myeloma and Plasmacytomas – Restaging/Recurrence (ONC-25.3)

ON.MM.0025.3.A

Indication	Imaging Study
Extra-osseous plasmacytoma response to initial therapy	 ONE of the following: CT of any previously involved area, contrast as requested MRI of any previously involved area, contrast as requested
Known spine involvement with new neurological signs/symptoms or worsening pain	MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), Lumbar spine (CPT® 72158) without and with contrast
 Treatment response assessment After completion of primary therapy Non-secretory multiple myeloma To determine therapy response with inconclusive labs 	 ONE of the following: Whole-body low-dose skeletal CT scan (CPT® 76497) MRI Bone Marrow Blood Supply (CPT® 77084) MRI Cervical (CPT® 72141), Thoracic (CPT® 72146), Lumbar spine (CPT® 72148), and Pelvis (CPT® 72195) without contrast MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), Lumbar spine (CPT® 72158), and Pelvis (CPT® 72197) without and with contrast MRI without contrast, or MRI without and with contrast for any previously involved bony area or symptomatic area
CAR-T cell therapy	Once before treatment and once 30-60 days after completion of treatment: • PET/CT (CPT® 78815 or CPT® 78816)

Indication	Imaging Study
 ANY of the following: Suspected relapse/recurrence Suspected progression of MGUS or SMM to a more malignant form 	 ONE of the following: Whole-body low-dose skeletal CT (CPT® 76497) MRI Bone Marrow Blood Supply (CPT® 77084) MRI Cervical (CPT® 72141), Thoracic (CPT® 72146), Lumbar spine (CPT® 72148), and Pelvis (CPT® 72195) without contrast MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), Lumbar spine (CPT® 72158), and Pelvis (CPT® 72197) without and with contrast MRI without contrast, or MRI without and with contrast for any previously involved bony area or symptomatic area
 ANY of the following: Negative PET will allow change in management from active treatment to maintenance or surveillance. Inconclusive findings on conventional imaging 	PET/CT (CPT® 78815 or CPT® 78816)
Stem cell transplant recipients	 ONE of the following, once before transplant and once within 30-100 days after transplant: Whole-body low-dose skeletal CT scan (CPT® 76497) MRI Bone Marrow Blood Supply (CPT® 77084) MRI Cervical (CPT® 72141), Thoracic (CPT® 72146), Lumbar spine (CPT® 72148), and Pelvis (CPT® 72195) without contrast MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), Lumbar spine (CPT® 72158), and Pelvis (CPT® 72197) without and with contrast

Multiple Myeloma and Plasmacytomas – Surveillance (ONC-25.4)

ON.MM.0025.4.A

Indication	Study
PlasmacytomasSmoldering myelomaMultiple myeloma	 ANY ONE of the following annually for 5 years: Whole-body low-dose skeletal CT (CPT® 76497) MRI Bone Marrow Blood Supply (CPT® 77084)

References (ONC-25)

- 1. Kumar SK, Callander NS, Adekola K, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 December 8, 2022. Myeloma, available at: https://www.nccn.org/professionals/physician_gls/pdf/myeloma.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Myeloma V3.2023 December 8, 2022. ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Hillengass J, Usmani S, Rajkumar SV, Durie BGM, Mateos M, Lonial S. International myeloma working group consensus recommendations on imaging in monoclonal plasma cell disorders. The Lancet. 2019:20(6):PE302-E312. doi:10.1016/S1470-2045(19)30309-2.
- 3. Kyle RA, Remstein ED, Therneau TM, et al. Clinical course and prognosis of smoldering (asymptomatic) multiple myeloma. *N Engl J Med*. 2007;356:2582-2590. doi:10.1056/NEJMoa070389.
- 4. Dimopoulos M, Terpos E, Comenzo RL, et al. International myeloma working group consensus statement and guidelines regarding the current role of imaging techniques in the diagnosis and monitoring of multiple myeloma. *Leukemia*. 2009;23(9):1545-1556. doi:10.1038/leu.2009.89.
- 5. ACR Committee on Drugs and Contrast Media. ACR Manual on Contrast Media, version 10.3. Reston, VA: American College of Radiology; 2018.
- Mulligan ME, Badros AZ. PET/CR and MR imaging in myeloma. Skeletal Radiol. 2007;36(1):5-16. doi:10.1007/s00256-006-0184-3.
- 7. Dimopoulos MA, Hillengrass J, Usmani S, et al. Role of magnetic resonance imaging in the management of patients with multiple myeloma: a consensus statement. *J Clin Oncol.* 2015;33(6):657-664. doi:10.1200/JCO.2014.57.9961.
- 8. Dimopoulos M, Terpos E, Comenzo RL, et al. International myeloma working group consensus statement and guidelines regarding the current role of imaging techniques in the diagnosis and monitoring of multiple myeloma. *Leukemia*. 2009;23(9):1545-1556. doi:10.1038.leu.2008.89.
- Dammacco F, Rubini G, Ferrari C, Vacca A, Racanelli V. 18F-FDG PET/CT: a review of diagnostic and prognostic features in multiple myeloma and related disorders. *Clin Exp Med*. 2015;15(1):1-18. doi:10.1007/s10238-014-0308-3.
- Ferraro R, Agarwal A, Martin-Macintosh EL, Peller PJ, Subramaniam RM. MR imaging and PET/CT in diagnosis and management of multiple myeloma. *Radiographics*. 2015;35(2):438-454. doi:10.1148/rg.352140112.
- 11. Rajkumar SV, Kumar S. Multiple myeloma: diagnosis and treatment. *Mayo Clin Proc.* 2016;91(1):101-119. doi:10.1016/j.mayocp.2015.11.007.
- 12. Westerland O, Amlani A, Kelly-Morland C, et. al. Comparison of the diagnostic performance and impact on management of 18F-FDG PET/CT and whole-body MRI in multiple myeloma. *Eur J Nucl ed Mol Imaging*. 2021. doi:10.1007/s00259-020-05182-2

Leukemias, Myelodysplasia and Myeloproliferative Neoplasms (ONC-26)

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Leukemias, Myelodysplasia and Myeloproliferative Neoplasms – General Considerations (ONC-26.1)

ON.LM.0026.1.A

- PET imaging is considered investigational, experimental, or unproven for all indications in acute lymphoblastic leukemia, acute myeloid leukemia, and chronic myeloid leukemia.
- Routine advanced imaging is not indicated for the evaluation and management of Hairy cell leukemia in the absence of specific localizing clinical symptoms.

Acute Leukemias (ONC-26.2)

ON.LM.0026.2.A

- Imaging indications for acute lymphoblastic leukemia in adult individuals are identical to those for pediatric individuals. See: <u>Acute Lymphoblastic Leukemia</u> (<u>ALL</u>) (<u>PEDONC-3.2</u>) in the Pediatric Oncology Imaging Guidelines.
- Imaging indications for acute myeloid leukemia in adult individuals are identical to those for pediatric individuals. See: <u>Acute Myeloid Leukemia (AML) (PEDONC-3.3)</u> in the Pediatric Oncology Imaging Guidelines.

Chronic Myeloid Leukemias, Myelodysplastic Syndrome and Myeloproliferative Disorders (ONC-26.3)

ON.LM.0026.3.A

- Routine advanced imaging is not indicated in the evaluation and management of chronic myeloid leukemias, myelodysplastic syndromes or myeloproliferative disorders in the absence of specific localizing clinical symptoms or clearance for hematopoietic stem cell transplantation.
- See: <u>Hematopoietic Stem Cell Transplantation (ONC-29)</u> for imaging guidelines related to transplant.
- For work-up of elevated blood counts, see: **Paraneoplastic Syndromes General Considerations (ONC-30.3)**.

Chronic Lymphocytic Leukemia (CLL)/ Small Lymphocytic Lymphoma (SLL) (ONC-26.4)

ON.LM.0026.4.A

- PET imaging is not indicated in the evaluation of CLL/SLL with the exception of suspected Richter's transformation (See Suspected transformation, below).
- CLL/SLL is monitored with serial laboratory studies. Routine advanced imaging is not indicated for monitoring treatment response or surveillance, except when initial studies reveal bulky disease involvement.
- Bulky disease is defined as lymph node mass >10 cm or spleen >6 cm below costal margin

margin		
Indication	Imaging Study	
Initial Staging/Diagnosis	Advanced imaging is not routinely indicated for initial evaluation of asymptomatic individuals	
 For ANY of the following: Bulky lymph node mass (>10 cm) Splenomegaly >6 cm below costal margin Presence of B symptoms Progressive anemia and thrombocytopenia Prior to planned systemic therapy 	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 	
Treatment Response	 For individuals with bulky nodal disease at diagnosis, CT with contrast of previously involved area(s) every 2 cycles of therapy Routine imaging is not indicated for individuals without bulky nodal disease at diagnosis 	
End of Therapy Evaluation	For individuals with bulky nodal disease at diagnosis, CT with contrast of previously involved area(s)	
Suspected Progression	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) 	

Indication	Imaging Study
Suspected transformation (Richter's) from a low-grade lymphoma to a more aggressive type based on one or more of the following: New B symptoms Rapidly growing lymph nodes Extranodal disease develops Significant recent rise in LDH above normal range	• PET/CT (CPT® 78815 or CPT® 78816)
Surveillance	 For individuals with bulky nodal disease at diagnosis, every 6 months for two years, then annually: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) Routine imaging is not indicated for individuals without bulky nodal disease at diagnosis

References (ONC-26)

- 1. Wierda WG, Brown J, Abramson JS, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 June 12, 2023. Chronic Lymphocytic Leukemia/Small Lymphocytic Lymphoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/cll.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Chronic Lymphocytic Leukemia/Small Lymphocytic Lymphoma V3.2023 June 12, 2023. [©]2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Conte MJ, Bowen DA, Wiseman GA, et al. Use of positron emission tomography-computed tomography in the management of patients with chronic lymphocytic leukemia/small lymphocytic lymphoma. *Leuk Lymphoma*. 2014;55(9):2079-2084. doi:10.3109/10428194.2013.869801.
- Mauro FR, Chauvie S, Paoloni F, et al. Diagnostic and prognostic role of PET/CT in patients with chronic lymphocytic leukemia and progressive disease. *Leukemia*. 2015;29(6):1360-1365. doi:10.1038/leu.2015.21.
- Nabhan C, Rosen ST. Chronic lymphocytic leukemia: a clinical review. JAMA. 2014;312(21):2265-2276. doi:10.1001/jama.2014.14553.
- 5. Patnaik MM, Tefferi A. Chronic myelomonocytic leukemia: focus on clinical practice. *Mayo Clin Proc.* 2016;91(2):259-272. doi:10.1016/j.mayocp.2015.11.011.
- American Society of Hematology. Choosing Wisely: Don't perform baseline or routine surveillance computed tomography (CT) scans in patients with asymptomatic, early-stage chronic lymphocytic leukemia (CLL). 2014. https://www.choosingwisely.org/clinician-lists/american-society-hematology-baseline-or-routine-surveillance-ct-scans-for-asymptomatic-early-stage-chronic-lymphocytic-leukemia/.

Non-Hodgkin Lymphomas (ONC-27)

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Non-Hodgkin Lymphomas – General Considerations (ONC-27.1)

ON.NH.0027.1.A

- Lymphoma is often suspected when individuals have any of the following:
 - Bulky lymphadenopathy (lymph node mass >10 cm in size), hepatomegaly or splenomegaly
 - The presence of systemic symptoms (fever, drenching night sweats or unintended weight loss of >10%, called "B symptoms")
- Individuals with AIDS-related lymphoma should be imaged according to the primary lymphoma histology
- See: <u>Castleman's Disease (unicentric and multicentric) (ONC-31.11)</u> for guidelines covering Castleman's disease.
- See: <u>Chronic Lymphocytic Leukemia (CLL)/Small Lymphocytic Lymphoma (SLL) (ONC-26.4)</u> for guidelines covering Chronic Lymphocytic Leukemia (CLL)/Small Lymphocytic Lymphoma (SLL).

(CLL)/Smail Lymphocytic Lymphoma (SLL).		
Indication	Imaging Study	
 Biopsy proven lymphoma or suspected lymphoma with one of the following: Bulky lymphadenopathy (LN mass >10 cm) Hepatomegaly Splenomegaly B symptom: Unexplained fever, drenching night sweats, unintended weight loss >10% total body weight 	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast MRI without and with contrast for individuals who cannot tolerate CT contrast due to allergy or impaired renal function 	
Signs or symptoms of disease involving the neck	CT Neck with contrast (CPT® 70491)	
Signs or symptoms suggesting CNS involvement with lymphoma.	 MRI Brain without and with contrast (CPT® 70553) See: CNS Lymphoma (also known as Microglioma) (ONC-2.7) 	
Known or suspected bone involvement with lymphoma	 MRI without and with contrast of symptomatic or previously involved bony areas Bone scan is inferior to MRI for evaluation of known or suspected bone involvement with lymphoma 	

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Indication	Imaging Study
Determine a more favorable site for biopsy when a relatively inaccessible site is contemplated	 PET/CT (CPT® 78815 or CPT® 78816) PET/CT is not indicated for all other indications prior to histological confirmation of lymphoma
CAR-T cell therapy	Once before treatment and once 30-60 days after completion of treatment: • PET/CT (CPT® 78815 and CPT® 78816)

Diffuse Large B Cell Lymphoma (DLBCL) (ONC-27.2)

ON.NH.0027.2.C

- Grey zone lymphomas, primary mediastinal B cell lymphomas, and Grade 3 (high) follicular lymphoma and double-hit or triple-hit lymphomas should also be imaged according to these guidelines.
- PET/CT scan is not generally supported for interim restaging (monitoring response
 to treatment) due to increased false-positive results. Treatment intensification based
 on positive interim PET/CT scan does not improve outcomes. Any positive findings
 noted on an interim PET/CT scan should be biopsied before changing treatment.

Indication	Imaging Study
Initial Staging/	ONE of the following may be approved:
Diagnosis	• PET/CT (CPT® 78815 or CPT® 78816)
	OR
	CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment	ANY one of the following may be approved, not both:
response for all stages	CT with contrast of previously involved area(s) may be approved every 2 cycles (6-8 weeks) of therapy
	OR
	PET/CT (CPT® 78815 or CPT® 78816) after 3-4 cycles of chemotherapy
End of	ANY or ALL of the following may be approved:
Chemotherapy	PET/CT (CPT® 78815 or CPT® 78816) may be approved at the
and/or Radiation Therapy	end of chemotherapy and again at the end of radiation
Evaluation	CT with contrast of previously involved area(s)
Suspected	ANY or ALL of the following may be approved:
Recurrence	CT Chest with contrast (CPT® 71260)
	CT Abdomen and Pelvis with contrast (CPT® 74177)
	CT with contrast of previously involved area(s)
	PET/CT can be considered in rare circumstances (e.g. bone involvement).
Biopsy-proven recurrence	PET/CT (CPT® 78815 or CPT® 78816)

Indication	Imaging Study
CAR-T cell therapy	Once before treatment and once 30-60 days after completion of treatment: • PET/CT (CPT® 78815 or CPT® 78816)
Surveillance for ANY of the following: • All stages of DLBCL • Relapsed lymphoma • Primary mediastinal large B cell lymphoma • Primary cutaneous diffuse large B cell lymphoma	 Every 6 months for 2 years after completion of treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)

Follicular Lymphoma (ONC-27.3)

ON.NH.0027.3.C

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 This section applies to follicular lymphomas with WHO grade of 1 (low) or 2 (intermediate). Grade 3 (high) follicular lymphomas should be imaged according to guidelines found in: <u>Diffuse Large B Cell Lymphoma (DLBCL)(ONC-27.2)</u>.

Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
 For ANY of the following: If radiation therapy is being considered for stage I or II disease If systemic therapy is planned Pediatric-type follicular lymphoma in adults 	• PET/CT (CPT® 78815 or CPT® 78816)
Treatment Response	CT with contrast of previously involved area(s) every 2 cycles of therapy
End of Chemotherapy Evaluation	 ONE of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) OR CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)

Indication	Imaging Study
Suspected transformation (Richter's) from a low grade lymphoma to a more aggressive type based on one or more of the following: New B symptoms Rapidly growing lymph nodes Extranodal disease develops Significant recent rise in LDH above normal range	• PET/CT (CPT® 78815 or CPT® 78816)
Surveillance for ANY of the following: • After completion of active treatment • On maintenance treatment • Observation without any treatment	For all stages, every 6 months for two years, then annually: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Surveillance of pediatric-type follicular lymphoma in adults	Advanced imaging is not indicated routinely after complete response

Marginal Zone Lymphomas (ONC-27.4)

ON.NH.0027.4.A

- MALT lymphomas in any location should also be imaged according to these guidelines
- Splenic Marginal Zone Lymphoma is diagnosed with splenomegaly, peripheral blood flow cytometry and bone marrow biopsy. Splenectomy is diagnostic and therapeutic. PET scan is not routinely indicated prior to splenectomy.

Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
EITHER of the following: If radiation therapy is being considered for stage I or II disease If systemic therapy is planned	• PET/CT (CPT® 78815 or CPT® 78816)
Treatment Response	CT with contrast of previously involved area(s) every 2 cycles of therapy
End of Therapy Evaluation	 ONE of the following may be approved: CT with contrast of previously involved area(s) PET/CT (CPT® 78815 or CPT® 78816)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) PET/CT can be considered in rare circumstances (e.g. bone involvement).

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Indication	Imaging Study
Surveillance of all stages of nodal marginal zone lymphoma for any of the following: • After completion of active treatment • On maintenance treatment • Observation without any treatment	 Every 6 months for two years, then annually: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Surveillance of all stages of extranodal marginal zone lymphoma	Advanced imaging is not routinely indicated for surveillance of asymptomatic individuals

Mantle Cell Lymphoma (ONC-27.5)

ON.NH.0027.5.A

Indication	Imaging Study
Initial Staging/Diagnosis	 ONE of the following may be approved: CT Chest with contrast (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) OR PET/CT (CPT® 78815 or CPT® 78816)
Treatment Response	 CT with contrast of previously involved area(s) every 2 cycles of therapy PET/CT is not indicated for monitoring treatment response but can be considered in rare circumstances when CT did not show disease (e.g. bone).
End of Therapy Evaluation	 ONE of the following may be approved: CT with contrast of previously involved area(s) PET/CT (CPT® 78815 or CPT® 78816)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) PET/CT can be considered in rare circumstances (e.g. bone involvement).
Surveillance for all stages	 Every 6 months for 2 years, and then annually: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)

Burkitt's Lymphomas (ONC-27.6)

ON.NH.0027.6.A

Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment Response	 CT with contrast of previously involved area(s) every 2 cycles of therapy PET/CT is not indicated for monitoring treatment response but can be considered in rare circumstances when CT did not show disease (e.g. bone).
End of Therapy Evaluation	 ANY or ALL of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) may be approved at the end of chemotherapy and again at the end of radiation CT with contrast of previously involved area(s)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) PET/CT can be considered in rare circumstances (e.g. bone involvement).
Surveillance	 Every 6 months for 2 years after completion of treatment: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)

Lymphoblastic Lymphomas (ONC-27.7)

ON.NH.0027.7.A

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 Individuals with lymphoblastic lymphoma (even those with bulky nodal disease) are treated using the leukemia treatment plan appropriate to the cell type (B or T cell). Imaging indications in adult individuals are identical to those for pediatric individuals. See: <u>Acute Lymphoblastic Leukemia (ALL) (PEDONC-3.2)</u> in the Pediatric Oncology Imaging Guidelines.

Cutaneous Lymphoma and T Cell Lymphomas (ONC-27.8)

ON.NH.0027.8.A

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 Includes Primary Cutaneous B Cell Lymphomas, Peripheral T-Cell Lymphomas, Mycosis Fungoides/Sézary Syndrome, Anaplastic Large Cell Lymphoma, Angioimmunoblastic lymphoma, and Primary Cutaneous CD30+T Cell Lymphoproliferative Disorders

Indication	Imaging Study
Initial Staging/Diagnosis	ANY or ALL of the following may be approved: • PET/CT (CPT® 78815 or CPT® 78816) • CT Chest with contrast (CPT® 71260) • CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment Response	Any ONE of the following may be approved after 3-4 cycles: • PET/CT (CPT® 78815 or 78816) or • CT Chest with contrast (CPT® 71260), and • CT Abdomen and Pelvis with contrast (CPT® 74177) and • CT with contrast of previously involved area(s)
End of Therapy Evaluation	Any ONE of the following may be approved at the end of chemotherapy and again at the end of radiation therapy: • PET/CT (CPT® 78815 or CPT® 78816) or • CT Chest with contrast (CPT® 71260), and • CT Abdomen and Pelvis with contrast (CPT® 74177), and • CT with contrast of previously involved area(s)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) PET/CT can be considered in rare circumstances (e.g., bone involvement).
Surveillance, all stages	 Every 6 months for 2 years, then annually for 5 years: CT Chest with contrast (CPT® 71260), CT Abdomen and Pelvis with contrast (CPT® 74177), and CT of previously involved areas

Post-Transplant Lymphoproliferative Disorders (ONC-27.9)

ON.NH.0027.9.A

- Post-transplant lymphoproliferative disorder (PTLD) or viral-associated lymphoproliferative disorder can rarely occur following solid organ or hematopoietic stem cell transplantation, or in primary immunodeficiency. When reduction of immunosuppression is unsuccessful, these are often treated with chemoimmunotherapy similar to high-grade NHL.
- This section applies to Monomorphic (B-cell type) PTLD and Polymorphic PTLD.
- For Hodgkin-lymphoma subtype of PTLD, see: <u>Hodgkin Lymphomas (ONC-28)</u> for imaging recommendations.

inaging recommendations.			
Indication	Imaging Study		
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 		
Treatment Response	ANY or ALL of the following may be approved after 4 weeks of reducing immunosuppression or every 2 cycles (6-8 weeks) of chemo/immunotherapy: CT Chest with contrast (CPT® 71260), and CT Abdomen and Pelvis with contrast (CPT® 74177), and CT with contrast of previously involved area(s)		
End of Therapy Evaluation	ANY one of the following may be approved at the end of treatment: • PET/CT (CPT® 78815 or CPT® 78816) or • CT Chest with contrast (CPT® 71260), and • CT Abdomen and Pelvis with contrast (CPT® 74177), and • CT with contrast of previously involved area(s)		
Suspected recurrence	 CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) 		
Surveillance	Advanced imaging is not routinely indicated for surveillance		

Waldenström Macroglobulinemia or Lymphoplasmacytic Lymphoma (ONC-27.10)

ON.NH.0027.10.A

Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment Response	CT with contrast of previously involved area(s) every 2 cycles of therapy
End of Therapy Evaluation	CT with contrast of previously involved area(s)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Surveillance	Advanced imaging is not routinely indicated for surveillance

References (ONC-27)

- 1. Zelenetz AD, Gordon LI, Abramson JS, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 5.2023 July 7, 2023. B-cell lymphomas, available at: https://www.nccn.org/professionals/physician_gls/pdf/B-CELL.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for B-cell lymphomas V5.2023 July 7, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Horwitz SM, Ansell S, Ai WZ, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 January 5, 2023. T-cell lymphomas, available at: https://www.nccn.org/professionals/physician_gls/pdf/T-CELL.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for T-cell lymphomas V1.2023 January 5, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Cheson BD, Fisher RI, Barrington SF, et al. Recommendations for initial evaluation, staging, and response assessment for Hodgkin and Non-Hodgkin lymphoma: the Lugano classification. *J Clin Oncol*. 2014;32(27):3059-3067. doi:10.1200/JCO.2013.54.8800.
- 4. Barrington SF, Mikhaeel NG, Kostakoglu L, et al. Role of imaging in the staging and response assessment of lymphoma: consensus of the International Conference on Malignant Lymphomas Imaging Working Group. *J Clin Oncol.* 2014;32(27):3048-3058. doi:10.1200/JCO.2013.53.5229.
- Thompson CA, Ghesquieres H, Maurer MJ, et al. Utility of routine post-therapy surveillance imaging in diffuse Large B-Cell Lymphoma. J Clin Oncol. 2014;32(31):3506-3512. doi:10.1200/JCO.2014.55.7561.
- El-Galaly TC, Jakobsen LH, Hutchings M, et al. Routine imaging for diffuse Large B-Cell Lymphoma in first complete remission does not improve post-treatment survival: a Danish-Swedish population-based study. J Clin Oncol. 2015;33(34):3993-3998. doi:10.1200/JCO.2015.62.0229.
- Huntington SF, Svoboda J, Doshi JA. Cost-effectiveness analysis of routine surveillance imaging of patients with diffuse Large B-Cell Lymphoma in first remission. *J Clin Oncol*. 2015;33(13):1467-1474. doi:10.1200/JCO.2014.58.5729.
- 8. Mamot C, Klingbiel D, Hitz F, et al. Final results of a prospective evaluation of the predictive value of interim positron emission tomography in patients with diffuse large B-cell lymphoma treated with R-CHOP-14 (SAKK 38/07). *J Clin Oncol*. 2015;33(23):2523-2529. doi:10.1200/JCO.2014.58.9846.
- Mylam KJ, Nielsen AL, Pedersen LM, Hutchings M. Fluorine-18-fluorodeoxyglucose positron emission tomography in diffuse large B-cell lymphoma. PET Clin. 2014;9(4):443-455. doi:10.1016/i.cpet.2014.06.001.
- 10. Avivi I, Zilberlicht A, Dann EJ, et al. Strikingly high false positivity of surveillance FDG-PET/CT scanning among patients with diffuse large cell lymphoma in the rituximab era. *Am J Hematol*. 2013;88(5):400-405. doi:10.1002/ajh.23423.
- 11. Ulrich Dührsen, Stefan Müller, Bernd Hertenstein, et al. Positron emission tomography-guided therapy of aggressive non-Hodgkin lymphomas (PETAL): a multicenter, randomized phase III trial. *J Clin Oncol*. 2018;36(20):2024-2034. doi:10.1200/JCO.2017.76.8093.

Hodgkin Lymphoma (ONC-28)

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Hodgkin Lymphoma – General Considerations (ONC-28.1)

ON.HL.0028.1.A

- Lymphoma is often suspected when individuals have any of the following:
 - Bulky lymphadenopathy (lymph node mass >10 cm in size), hepatomegaly or splenomegaly
 - The presence of systemic symptoms (fever, drenching night sweats or unintended weight loss of >10%, called "B symptoms")
- Individuals with AIDS-related lymphoma should be imaged according to the primary lymphoma histology
- The Deauville Criteria are internationally accepted criteria, which utilize a five-point scoring system for the FDG avidity of a Hodgkin's lymphoma or Non-Hodgkin's lymphoma tumor mass as seen on FDG PET.
 - Score 1: No uptake above the background
 - Score 2: Uptake ≤mediastinum
 - Score 3: Uptake >mediastinum but ≤liver
 - Score 4: Uptake moderately increased compared to the liver at any site
 - Score 5: Uptake markedly increased compared to the liver at any site
 - Score X: New areas of uptake unlikely to be related to lymphoma

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Indication	Imaging Study		
Biopsy proven lymphoma or suspected lymphoma with one of the following: • Bulky lymphadenopathy (LN mass >10 cm) • Hepatomegaly • Splenomegaly • B symptom: Unexplained fever, drenching night sweats, unintended weight loss >10% total body weight	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast MRI without and with contrast for individuals who cannot tolerate CT contrast due to allergy or impaired renal function 		
Signs or symptoms of disease involving the neck	CT Neck with contrast (CPT® 70491)		
Signs or symptoms suggesting CNS involvement with lymphoma	 MRI Brain without and with contrast (CPT[®] 70553) See: CNS Lymphoma (also known as Microglioma) (ONC-2.7) 		

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Indication	Imaging Study	
Known or suspected bone involvement with lymphoma	 MRI without and with contrast of symptomatic or previously involved bony areas Bone scan is inferior to MRI for evaluation of known or suspected bone involvement with lymphoma 	
Determine a more favorable site for biopsy when a relatively inaccessible site is contemplated	PET/CT (CPT® 78815 or CPT® 78816) PET/CT is medically unnecessary for all other indications prior to histological confirmation of lymphoma	
CAR-T cell therapy	Once before treatment and once 30-60 days after completion of treatment: • PET/CT (CPT® 78815 or CPT® 78816)	

Classical Hodgkin Lymphoma (ONC-28.2)

ON.HL.0028.2.A

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• This section applies to nodular sclerosis, mixed cellularity, lymphocyte-depleted and lymphocyte-rich subtypes of Hodgkin lymphoma.

Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment Response	PET/CT (CPT® 78815 or CPT® 78816) as frequently as every 2 cycles
End of Chemotherapy and/or Radiation Therapy Evaluation	PET/CT (CPT® 78815 or CPT® 78816) may be approved at the end of chemotherapy and again at the end of radiation (at least 12 weeks after completion of radiation therapy)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Biopsy proven recurrence	• PET/CT (CPT® 78815 or CPT® 78816)
Surveillance	 ANY or ALL of the following may be approved every 6 months for 2 years after completion of therapy: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) In addition to the above studies: A single follow-up PET/CT may be approved at three months if end of therapy PET/CT shows Deauville 4 or 5 FDG avidity

Nodular Lymphocyte – Predominant Hodgkin Lymphoma (ONC-28.3)

ON.HL.0028.3.A

	V1.U.2U24
Indication	Imaging Study
Initial Staging/Diagnosis	 ANY or ALL of the following may be approved: PET/CT (CPT® 78815 or CPT® 78816) CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177)
Treatment Response	CT with contrast of previously involved areas as frequently as every 2 cycles
End of Chemotherapy and/or Radiation Therapy Evaluation	PET/CT (CPT® 78815 or CPT® 78816) may be approved at the end of chemotherapy and again at the end of radiation (at least 12 weeks after completion of radiation therapy)
Suspected Recurrence	 ANY or ALL of the following may be approved: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s)
Biopsy proven recurrence	• PET/CT (CPT® 78815 or CPT® 78816)
Suspected transformation (Richter's) from a low-grade lymphoma to a more aggressive type based on one or more of the following: New B symptoms Rapidly growing lymph nodes Extranodal disease develops Significant recent rise in LDH above normal range	• PET/CT (CPT® 78815 or CPT® 78816)

Indication	Imaging Study
Surveillance	 ANY or ALL of the following may be approved every 6 months for 2 years after completion of therapy: CT Neck with contrast (CPT® 70491) CT Chest with contrast (CPT® 71260) CT Abdomen and Pelvis with contrast (CPT® 74177) CT with contrast of previously involved area(s) In addition to the above studies: A single follow-up PET/CT may be approved at three months if end of therapy PET/CT shows Deauville 4 or 5 FDG avidity

References (ONC-28)

- 1. Hoppe RT, Advani RH, Ambinder RF, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 2.2023 November 8, 2022. Hodgkin lymphoma, available at: https://www.nccn.org/professionals/physician_gls/pdf/hodgkins.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Hodgkins Lymphoma V2.2023 November 8, 2022. [©]2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Cheson BD, Fisher RI, Barrington SF, et al. Recommendations for initial evaluation, staging, and response assessment for Hodgkin and Non-Hodgkin lymphoma: the Lugano classification. J Clin Oncol. 2014;32(27):3059-3067. doi:10.1200/JCO.2013.54.8800.
- 3. Barrington SF, Mikhaeel NG, Kostakoglu L, et al. Role of imaging in the staging and response assessment of lymphoma: consensus of the International Conference on Malignant Lymphomas Imaging Working Group. J Clin Oncol. 2014;32(27):3048-3058. doi:10.1200/JCO.2013.53.5229.
- 4. Pingali SR, Jewell SW, Havlat L, et al. Limited utility of routine surveillance imaging for classical Hodgkin lymphoma patients in first complete remission. *Cancer*. 2014;120:2122-2129.
- Ha CS, Hodgson DC, Advani R, et al. Follow-up of Hodgkin lymphoma. ACR Appropriateness Criteria[®] 2014:1-16.
- 6. Picardi M, Pugliese N, Cirillo, M et al. Advanced-stage Hodgkin lymphoma: US/Chest radiography for detection of relapse in patients in first complete remission—a randomized trial of routine surveillance imaging procedures. *Radiology*. 2014;272:262-274.
- 7. Gallamini A, and Kostakoglu L. Interim FDG-PET in Hodgkin lymphoma: a compass of a safe navigation in clinical trials? *Blood*. 2012;120(25):4913-4920.
- 8. Biggi A, Gallamini A, Chauvie S, et al. International validation study for interim PET in ABVD-treated, advanced-stage Hodgkin lymphoma: interpretation criteria and concordance rate among reviewers. *J Nucl Med*. 2013; 54(5):683-690.
- 9. Gallamini A, Barrington SF, Biggi, et al. The predictive role of interim positron emission tomography for Hodgkin lymphoma treatment outcome is confirmed using the interpretation criteria of the Deauville five-point scale. *Haematologica*. 2014; 99(6):1107-1113.
- 10. El-Galaly TC, Mylam KJ, Brown P, et al. Positron emission tomography/computed tomography surveillance in patients with Hodgkin lymphoma in first remission has a low positive predictive value and high costs. *Haematologica*. 2012;97(6):931-936.

Hematopoietic Stem Cell Transplantation (ONC-29)

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General Considerations for Stem Cell Transplant (ONC-29.1)

ON.HT.0029.1.A

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Transplant types:

Allogeneic ("allo"): The donor and recipient are different people, and there are multiple types depending on the source of the stem cells and degree of match between donor and recipient. This is most commonly used in diseases originating in the hematopoietic system, such as leukemias and lymphomas, and bone marrow failure syndromes or metabolic disorders. Common types are:

- Matched sibling donor (MSD or MRD): Donor and recipient are full siblings and HLA-matched
- Matched unrelated donor (MUD): Donor and recipient are HLA matched but not related to each other
- Cord blood: Donor stem cells come from frozen umbilical cord blood not related to the recipient, sometimes from multiple different donors at once
- Haploidentical transplant (haplo): Donor is a half-HLA match to the recipient, usually a parent

Autologous ("auto"): The donor and recipient are the same person. The process involves delivery of high dose chemotherapy that is ablative to the bone marrow, followed by an infusion of one's own harvested stem cells.

Allogeneic HSCT results in a much greater degree of immunosuppression than autologous HSCT because of the need to allow the new immune system to chimerize with the recipient's body. Immune reconstitution commonly takes more than a year for individuals who receive allogeneic HSCT, and individuals remain at high- risk for invasive infections until that has occurred.

Pre-Transplant Imaging in HSCT:

 Pre-transplant imaging in HSCT generally takes place within 30 days prior to transplant and involves a reassessment of the individual's disease status as well as infectious disease clearance.

Indication	Imaging
Immediate pre-transplant period	 Chest x-ray CT Chest with contrast (CPT® 71260) or CT Chest without contrast (CPT® 71250) for new findings on chest x-ray, or new/worsening signs/symptoms CT Sinus (CPT® 70486) for any clinical signs or symptoms

Indication	Imaging
Assess cardiac function	Echocardiogram (CPT® 93306, CPT® 93307 or CPT® 93308)
	 MUGA scan (CPT® 78472) may be indicated in specific circumstances, see: <u>Oncologic</u> <u>Indications for Cancer Therapeutics-Related</u> <u>Cardiac Dysfunction (CTRCD) (CD-12.1)</u> in the Cardiac Imaging Guidelines
Assess pulmonary function	Pulmonary function tests
Assess primary disease status	See disease-specific guidelines for end of therapy response assessment

Post-Transplant Imaging in HSCT:

- There are many common complications from HSCT, including infection, acute and chronic graft versus host disease (GVHD), hepatic sinusoidal obstruction syndrome, restrictive lung disease, among others.
- Disease response generally takes place at ~Day +30 (autos and some allos) or ~Day +100 (allos) post-transplant.

Indication	Imaging
Assess known or suspected HSCT complications	Site-specific imaging should generally be approved
Suspected hepatic GVHD (elevated liver enzymes)	Abdominal US (CPT® 76700 or CPT® 76705)
Suspected Bronchiolitis Obliterans Syndrome (BOS)	CT Chest without contrast (CPT® 71250)
Assess primary disease status post-transplant	See disease-specific guidelines for end of therapy evaluation and surveillance
Individuals receiving tandem auto transplants (2-4 autos back-to-back, spaced 6 to 8 weeks apart)	Guideline recommended imaging can be repeated after each transplant

Reference (ONC-29)

v1.0.2024

1. Loren AW, Mielcarek M, Bolaños-Meade J, et. al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 – March 31, 2023. Hematopoietic Cell Transplantation, available at: https://www.nccn.org/professionals/physician_gls/pdf/hct.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines®) for Hematopoietic Cell Transplantation V1.2023 – March 31, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines® and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines®, go online to NCCN.org.

Medical Conditions with Cancer in the Differential Diagnosis (ONC-30)

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Fever of Unknown Origin (FUO) (ONC-30.1)

ON.MC.0030.1.A

- FUO is defined as a persistent fever ≥101°F and ≥3 weeks with unidentified cause.
- While fever is a classic "B" symptom of advanced lymphoma, a cancer-related fever presenting in isolation without any other signs or symptoms of neoplastic disease is rare.

Indication	Imaging Study
If physical examination, Chest X-ray, and laboratory studies are non-diagnostic	 Echocardiogram (CPT® 93306) Abdominal ultrasound (CPT® 76700) MRI Brain without and with contrast (CPT® 70553)
Above studies (including PE/ENT exam, pelvic exam, and DRE with laboratory studies) have failed to demonstrate site of infection	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast Radiopharmaceutical localization of tumor or distribution of radiopharmaceutical agent(s): CPT® 78800, CPT® 78801, or CPT® 78802, CPT® 78804, CPT® 78803 or CPT® 78831 (SPECT), or CPT® 78830, or CPT® 78832 (SPECT/CT)
"B" symptoms	See: Non-Hodgkin Lymphomas (ONC- 27)
Any CNS sign/symptom accompanied by fever	MRI Brain without and with contrast (CPT® 70553)
All individuals	PET is not indicated in the work-up of individuals with FUO

Unexplained Weight Loss (ONC-30.2)

ON.MC.0030.2.A

- Unintentional weight loss is defined as loss of ≥10 lbs. or ≥5% of body weight over
 6 months or less, without an identifiable reason.
- Initial workup for all individuals may include appropriate detailed history, physical exam, baseline laboratory studies (e.g., CBC, CMP, HgbA1c, ESR/CRP, infectious workup, stool hemoccult, endocrine evaluation to rule out thyroid, pituitary, or gonadal dysfunction, etc.), chest x-ray, age-appropriate cancer screening, and neurological evaluation to rule out depression/dementia.
- Additional workup is directed to evaluate specific signs, symptoms, red flags, or abnormalities detected on initial workup. See condition-specific imaging guidelines for additional details.
- PET is not appropriate in the work-up of individuals with unexplained weight loss.

Indication	Imaging Study
CNS symptoms or abnormal pituitary hormones	MRI Brain or Sella Turcica without and with contrast (CPT® 70553)
Abnormal thyroid function	Thyroid ultrasound (CPT® 76536)
Abnormal liver function	Abdominal ultrasound (CPT® 76700)
Abnormal kidney function	 Ultrasound kidney and bladder (CPT[®] 76770 or CPT[®] 76775)
Suspected cardiac dysfunction	Echocardiogram (CPT® 93306)
Non-smokers	 Chest x-ray CT Chest with contrast (CPT® 71260) to evaluate abnormalities on chest x-ray
Current or former smokers	CT Chest with contrast (CPT® 71260)
Dysphagia or early satiety	See: <u>Dysphagia and Esophageal</u> <u>Disorders (NECK-3)</u>
GI bleeding	See: Gl Bleeding (AB-22)
Abdominal pain without red flag signs	See: Abdominal Pain (AB-2)

Indication	Imaging Study
Suspected pancreatic cancer in individuals aged ≥60 years with weight loss and at least one of the following ¹³ : • Diarrhea • Back pain • Abdominal pain • Nausea/vomiting • Constipation • New onset diabetes • Abnormal labs (CA 19-9, LFTs) • Non-diagnostic or negative abdominal ultrasound	 Any ONE of the following may be obtained: CT Abdomen with contrast (CPT® 74160) CT Abdomen and Pelvis with contrast (CPT® 74177) MRI Abdomen without and with contrast (CPT® 74183) See also: Epigastric Pain and Dyspepsia (AB-2.5)
If all of the above do not identify cause of weight loss	 Any of the following, if not previously performed: CT Chest (CPT® 71260) CT Abdomen and Pelvis (CPT® 74177) with contrast

Paraneoplastic Syndromes (ONC-30.3)

ON.MC.0030.3.A

- Paraneoplastic syndromes are metabolic and neuromuscular disturbances. These
 syndromes are not directly related to a tumor or to metastatic disease. There may
 be a lead time between initial finding of a possible paraneoplastic syndrome and
 appearance of the cancer with imaging. Limited studies suggest annual imaging for
 2 years after diagnosis of possible paraneoplastic syndrome may detect cancer,
 however benefit after 2 years is not well documented.
- The following are the most common symptoms of paraneoplastic syndromes known to arise from various malignancies:
 - Hypertrophic Pulmonary Osteoarthropathy: Often presents as a constellation of rheumatoid-like polyarthritis, periostitis of long bones, and clubbing of fingers and toes
 - o Amyloidosis
 - o Hypercalcemia
 - Hypophosphatemia
 - Cushing's Syndrome
 - Somatostatinoma syndrome (vomiting, abdominal pain, diarrhea, cholelithiasis)
 - Syndrome of inappropriate antidiuretic hormone secretion (SIADH)
 - o Polymyositis/dermatomyositis
 - o Opsoclonus
 - Paraneoplastic sensory neuropathy
 - Subacute cerebellar degeneration
 - Eaton-Lambert syndrome (a myasthenia-like syndrome)
 - Second event of unprovoked thrombosis
 - Disseminated Intravascular Coagulation
 - Migratory thrombophlebitis
 - Polycythemia
 - Chronic leukocytosis and/or thrombocytosis
 - Elevated tumor markers
 - o Cryptogenic stroke (see also: HD-21.3)
- See: <u>Muscle Disorders (PN-6)</u> in the Peripheral Nerve Disorders Imaging Guidelines.
- See: <u>Multiple Myeloma and Plasmacytomas (ONC-25)</u> for evaluation of possible multiple myeloma.

Indication	Imaging Study
Initial evaluation	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
 ANY of the following: Abnormality on conventional imaging difficult to biopsy Inconclusive conventional imaging Documented paraneoplastic antibody and conventional imaging fails to demonstrate primary site 	• PET/CT (CPT® 78815 or CPT® 78816)
Subsequent evaluation for known paraneoplastic syndrome	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast may be repeated every 6 months for 2 years after initial imaging for Lambert-Eaton Myasthenia syndrome CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast may be repeated every 6 months for 4 years for all other paraneoplastic syndromes
Systemic mastocytosis	 ANY ONE of the following: CT Abdomen and Pelvis (CPT® 74177) with contrast MRI Abdomen (CPT® 74183) and MRI Pelvis (CPT® 72197) without and with contrast is indicated PET/CT scan is not indicated for evaluation of mastocytosis
First episode of unprovoked DVT/VTE	Imaging to evaluate for malignancy is not indicated
Second unprovoked DVT/PE	Imaging may be considered in the setting of a negative work-up for inherited thrombophilia and antiphospholipid syndrome
Thyroid US is recomme	nded for elevated CEA, and upper/lower endoscopy is

recommended for elevated CEA or CA 19-9.

References (ONC-30)

- 1. Carrier M, Lazo-Langner A, Shivakumar S, et al. Screening for occult cancer in unprovoked venous thromboembolism. *N Engl J Med*. 2015 June;373:697-704. doi:10.1056/NEJMoa1506623.
- 2. Sioka C, Fotopoulos A, Kyritsis AP. Paraneoplastic neurological syndromes and the role of PET imaging. *Oncology.* 2010;78(2):150–156. doi:10.1159/000312657.
- 3. Schramm N, Rominger A, Schmidt C, et al. Detection of underlying malignancy in patients with paraneoplastic neurological syndromes: comparison of 18F-FDG PET/CT and contrast-enhanced CT. *Eur J Nucl Med Mol Imaging*. 2013;40(7):1014-1024. doi:10.1007/s00259-013-2372-4.
- Qiu L, Chen Y. The role of 18F-FDG PET or PET/CT in the detection of fever of unknown origin. Eur J Radiol. 2012;81(11):3524-3529. doi:10.1016/j.ejrad.2012.05.025.
- 5. Pelosof LC, Gerber DE. Paraneoplastic syndromes: an approach to diagnosis and treatment. *Mayo Clin Proc.* 2010;85(9):838-854. doi:10.4065/mcp.2010.0099.
- 6. Wong CJ. Involuntary weight loss. *Med Clin North Am.* 2014;98(3):625-43. doi:10.1016/j.mcna.2014.01.012.
- 7. Titulaer MJ, Soffieti R, Dalmau J, et al. Screening of tumours in paraneoplastic syndromes: report of an EFNS task force. *Eur J Neurol.* 2011;18(1):19–e3. doi:10.1111/j.1468-1331.2010.03220.x.
- 8. Lancaster E. Paraneoplastic disorders. *Continuum (Minneap Minn)*. 2017;23(6, Neuro-oncology):1653-1679. doi:10.1212/CON.00000000000542.
- 9. Gerds AT, Gotlib J, Ali H, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2022 – April 14, 2022. Systemic Mastocytosis, available at: https://www.nccn.org/professionals/physician_gls/pdf/mastocytosis.pdf. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Systemic Mastocytosis V1.2022 – April 14, 2022. ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 10. Saver JL. Cryptogenic stroke. N Engl J Med. 2016;374:2065-2074. doi:10.1056/NEJMcp1503946.
- Schwarzbach CJ, Schaefer A, Ebert A, et. al. Stroke and cancer: the importance of cancer-assocaited hypercoagulation as a possible stroke etiology. *Stroke*. 2012;43(11):3029-3034. doi:10.1161/STROKEAHA.112.658625.
- 12. Kamel H, Merkler AE, Iadecola C, Gupta A, Navi B. Tailoring the approach to embolic stroke of undetermined source: a review. *JAMA Neurol*. 2019;76(7):855-861. doi:10.1001/jamaneurol.2019.0591.
- National Institute for Health and Care Excellence (NICE). Upper gastroinestinal tract cancers. In: Suspected cancer: recognition and referral. 2015. https://www.nice.org.uk/guidance/ng12/chapter/Recommendations-organised-by-site-of-cancer#upper-gastrointestinal-tract-cancers

Metastatic Cancer, Carcinoma of Unknown Primary Site, and Other Types of Cancer (ONC-31)

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General Guidelines (ONC-31.0)

ON.UP.0031.0.A

- Guideline sections <u>Lung Metastases (ONC-31.1)</u> through <u>Bone (including Vertebral) Metastases (ONC-31.5)</u> should only be used for individuals with metastatic cancer in the following circumstances:
 - The primary diagnosis section does not address a particular metastatic site that is addressed in these sections
 - The cancer type is rare and does not have its own diagnosis-specific imaging guidelines

Lung Metastases (ONC-31.1)

ON.UP.0031.1.A

Indication	Imaging Study
New or worsening signs or symptoms suggestive of metastatic lung involvement or new or worsening chest x-ray abnormality	CT Chest with contrast (CPT® 71260) CT Chest without contrast (CPT® 71250) can be approved if there is a contraindication to CT contrast or only parenchymal lesions are being evaluated
Chest wall or brachial plexus involvement	MRI Chest without and with contrast (CPT® 71552)
 ONE of the following and no diagnosis-specific guideline regarding PET imaging: Lung nodule(s) ≥8 mm Confirm solitary metastasis amenable to resection on conventional imaging 	 PET/CT (CPT® 78815) When primary cancer known, PET request should be reviewed by primary cancer guideline
Previous or current malignancy and pulmonary nodule(s) that would reasonably metastasize to the lungs	CT Chest with contrast (CPT® 71260) at 3, 6, 12, and 24 months from the first study

Liver Metastases (ONC-31.2)

ON.UP.0031.2.A

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 Ablation of liver metastases or primary HCC may be performed utilizing chemical, chemotherapeutic, radiofrequency, or radioactive isotope. Regardless of the modality of ablation, PET is not indicated for assessing response to this mode of therapy.

Indication	Imaging Study
New or worsening signs or symptoms suggestive of metastatic liver involvement or new elevation in LFTs	CT Abdomen with (CPT® 74160) or without and with (CPT® 74170) contrast
ANY of the following:Considering limited resectionInconclusive CT findings	MRI Abdomen without and with contrast (CPT® 74183)
 ONE of the following and no diagnosis-specific guideline regarding PET imaging: Confirm solitary metastasis amenable to resection on conventional imaging LFT's and/or tumor markers continue to rise and CT and MRI are negative 	 PET/CT (CPT® 78815) When primary cancer known, PET request should be reviewed by primary cancer guideline
Monitoring of liver metastases that have been surgically resected	Review according to primary cancer guideline
Evaluation of hepatic artery chemotherapy infusion or TACE (transarterial chemoembolization)	 ONE of the following studies, if not previously done: CT Abdomen without and with contrast (CPT® 74170) MRI Abdomen without and with contrast (CPT® 74183)

Indication	Imaging Study
Evaluation for hepatic artery radioembolization with Y-90 radioactive spheres (TheraSphere or SIR Spheres) for liver metastases or primary liver tumors	To assess hepatic vascular anatomy before the procedure, any ONE of the following: • 3D Rendering (CPT® 76377) if conventional hepatic angiogram is being performed • CTA Abdomen (CPT® 74175)
or primary invertainers	ONE of the following studies may be approved PRE-treatment based upon provider preference:
	 Liver Imaging Planar (CPT® 78201) Radiopharmaceutical Localization Limited Area (CPT® 78800 or CPT® 78801)
	• SPECT or SPECT/CT (CPT® 78803, 78831, 78830, or 78832)
	ONE of the following studies may be approved POST-treatment based upon provider preference:
	Liver Planar Imaging (CPT® 78201)
	Radiopharmaceutical Localization Limited Area (CPT® 78800 or CPT® 78801)
	• SPECT or SPECT/CT (CPT® 78803, 78831, 78830, or 78832)
	Please note: liver-lung shunt calculation is included in the pre-treatment Liver Scan and does not require additional Lung Perfusion Scan
Monitoring of ablated liver metastases or primary tumors	ONE of the following, immediately prior to ablation, 1 month post-ablation, then every 3 months for 2 years, and then every 6 months until year 5:
	 CT Abdomen without and with contrast (CPT® 74170)
	MRI Abdomen without and with contrast (CPT® 74183)

Brain Metastases (ONC-31.3)

ON.UP.0031.3.A

Indication	Imaging Study
Individual with cancer and signs or symptoms of CNS disease or known brain metastasis with new signs or symptoms.	MRI Brain without and with contrast (CPT® 70553)
To determine candidacy for SRS, and a diagnostic thin-slice MRI Brain has not been performed in the preceding 30 days	MRI Brain without and with contrast (CPT® 70553)
Stereotactic radiosurgery planning	Unlisted MRI for treatment planning purposes (CPT® 76498)
Monitoring of brain metastases treated with surgery or radiation therapy	Post-treatment, then every 3 months for 1 year and every 6 months thereafter: • MRI Brain without and with contrast (CPT® 70553) ***Individuals treated with stereotactic radiosurgery alone may have MRI Brain without and with contrast (CPT® 70553) immediately after stereotactic radiosurgery, then every 2 months for the first year, and then every 6 months thereafter
Brain metastases treated with radiation therapy, with recent MRI Brain indeterminate in distinguishing radiation necrosis vs. tumor progression	MRI Perfusion imaging (CPT® 70553)
Brain metastases treated with radiation therapy, with recent MRI Brain and MR Perfusion studies both unable to distinguish radiation necrosis vs. tumor progression	PET Metabolic Brain (CPT® 78608)

Indication	Imaging Study
 Any of the following: Solitary brain metastasis suspected in individual with prior diagnosis of cancer and no diagnosis-specific guideline regarding PET imaging Brain metastases and no known primary tumor 	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast Mammography for female individuals PET/CT (CPT® 78815 or CPT® 78816) is indicated for ANY of the following: Inconclusive conventional imaging Confirm either stable systemic disease or absence of other metastatic disease When primary cancer known, PET request should be reviewed by primary cancer guideline
Primary brain tumors	See: Primary Central Nervous System Tumors (ONC-2)
MR Spectroscopy (CPT® 76390) is considered investigational and experimental for evaluation of metastatic brain cancer	

Adrenal Gland Metastases (ONC-31.4)

ON.UP.0031.4.A

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Indication	Imaging Study
Differentiate benign adrenal adenoma from metastatic disease	See: Adrenal Cortical Lesions (AB-16.1) in the Abdomen Imaging Guidelines
Known cancer and no known systemic metastases: New adrenal mass Enlarging adrenal mass Inconclusive findings on recent CT	 If not done previously, ANY of the following may be obtained: CT Abdomen without contrast (CPT® 74150) CT Abdomen without and with contrast (CPT® 74170, adrenal protocol) MRI Abdomen without contrast (CPT® 74181) MRI Abdomen without and with contrast (CPT® 74183) CT-directed needle biopsy (CPT® 77012)
One of the following and no diagnosis-specific guideline regarding PET imaging: Biopsy is not feasible or is non-diagnostic Isolated metastasis on conventional imaging and individual is a candidate for aggressive surgical management	PET/CT (CPT® 78815) When primary cancer known, PET request should be reviewed by primary cancer guideline
Known extra-adrenal malignancy and undiagnosed adrenal mass being monitored off treatment	See: Phases of Oncology Imaging and General Phase-Related Considerations (ONC-1.2)

Bone (including Vertebral) Metastases (ONC-31.5)

ON.UP.0031.5.C

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• Individuals with stage IV cancer with new onset back pain can forgo a bone scan (and plain films) in lieu of an MRI without and with contrast of the spine.

Indication	Imaging Study
ANY of the following in an individual with a current or prior malignancy: Bone pain Rising tumor markers Elevated alkaline phosphatase	Bone scan supplemented by plain x-rays is the initial diagnostic imaging study of choice
Indeterminate findings on bone scan	MRI without and with contrast or CT without and with contrast of the involved body site
 ANY of the following: Any individual with stage IV cancer with new onset back pain Bone scan is not feasible or readily available Continued suspicion despite inconclusive or negative bone scan or other imaging modalities Neurological compromise Soft tissue component suggested on other imaging modalities or physical exam Differentiate neoplastic disease from Paget's disease of Bone Suspected leptomeningeal involvement 	 ANY of the following may be approved: MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), and Lumbar Spine (CPT® 72158) without and with contrast CT Cervical (CPT® 72127), Thoracic (CPT® 72130), and Lumbar Spine (CPT® 72133) without and with contrast can be approved if MRI is contraindicated or not readily available CT without contrast can be approved if there is a contraindication to CT contrast
Monitoring untreated spinal metastases	MRI without and with contrast or CT without and with contrast of the involved spinal level every 3 months for 1 year.
	**Imaging beyond 1 year is based on any new clinical signs/symptoms

Indication	Imaging Study
Monitoring metastases within the spine treated with surgery and/or radiation therapy	 MRI without and with contrast or CT without and with contrast of the involved spinal level once within 3 months post treatment and then every 3 months for 1 year. **Imaging beyond 1 year is based on any
	new clinical signs/symptoms
Leptomeningeal involvement with cancer	 On active treatment: MRI Brain without and with contrast (CPT® 70553) MRI Cervical (CPT® 72156), Thoracic (CPT® 72157), and Lumbar spine (CPT® 72158) without and with contrast every 2 cycles Once treatment completed:
	Routine advanced imaging not indicated for surveillance in asymptomatic individuals
Bone pain when both bone scan and either CT or MRI are inconclusive	¹⁸ F-FDG-PET/CT (CPT® 78815 or CPT® 78816) on a case-by-case basis
Suspected metastatic bone disease and negative work-up for myeloma	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
No prior cancer history with suspected pathologic fracture on plain x-ray	See: Carcinoma of Unknown Primary Site (ONC-31.7)
Signs/symptoms concerning for spinal cord compression	See: Spinal Cord Compression (ONC- 31.6)

Spinal Cord Compression (ONC-31.6)

ON.UP.0031.6.A

	V1.0.2024
Indication	Imaging Study
 ANY of the following in a current or former cancer individual: Any individual with stage IV cancer with new onset back pain New back pain persisting over two weeks Back pain that is rapidly progressive or refractory to aggressive pain management Signs or symptoms of neurological compromise at the spinal cord level Unexpected, sudden loss of bowel or bladder control Sudden loss of ability to ambulate Complete loss of pinprick sensation corresponding to a specific vertebral level Loss of pain at a site that had previously been refractory to pain management 	 ANY or ALL of the following may be approved: MRI Cervical (CPT® 72156), MRI Thoracic (CPT® 72157), and MRI Lumbar Spine (CPT® 72158) without and with contrast Post myelogram CT Cervical (CPT® 72126), CT Thoracic (CPT® 72129), and CT Lumbar (CPT® 72132) Spine
Any current or former cancer individual with radicular symptoms suggestive of nerve root involvement but not consistent with cord compression and one of the following: Unilateral weakness Unilateral change of reflexes Pain unrelieved by change in position Age >70 years Unintentional weight loss Night pain Severe and worsening spinal pain despite a reasonable (generally after 1 week) trial of provider-directed treatment with reevaluation	 ONE of the following: MRI without and with contrast of involved spinal level MRI without contrast of the involved spinal level CT without contrast of the involved spinal level if MRI contraindicated

Carcinoma of Unknown Primary Site (ONC-31.7)

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General Considerations

- Defined as carcinoma found in a lymph node or in an organ known not to be the primary for that cell type (e.g., adenocarcinoma arising in the brain or in a neck lymph node).
- This guideline also applies to a pathologic fracture that is clearly due to metastatic neoplastic disease in an individual without a previous cancer history.
- Detailed history and physical examination including pelvic and rectal exams and laboratory tests to be performed before advanced imaging.
- Individuals presenting with a thoracic squamous cell carcinoma described as metastatic appearing on chest imaging, or in lymph nodes above the clavicle, should undergo a detailed head and neck examination by a clinician skilled in laryngeal and pharyngeal examinations, especially in smokers.
- Individuals with suspected unknown primary based on only suspicious lytic bone lesions should be considered for serum protein electrophoresis (SPEP); urine protein electrophoresis (UPEP) and serum free light chains prior to consideration of extensive imaging.

Indication	Imaging Study
Carcinoma found in a lymph node or in an organ known not to be primary	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast CT Neck with contrast (CPT® 70491) if cervical or supraclavicular involvement CT with contrast or MRI without and with contrast of any other symptomatic site For female individuals: Diagnostic (not screening) mammogram and full pelvic exam MRI Breast Bilateral (CPT® 77049) if pathology consistent with breast primary and mammogram is inconclusive

Indication	Imaging Study
Sebaceous carcinoma of the skin (can be associated with underlying primary malignancy)	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast CT Neck with contrast (CPT® 70491) if cervical or supraclavicular involvement CT with contrast or MRI without and with contrast of any other symptomatic site
Axillary adenocarcinoma	 Diagnostic (not screening) mammogram and full pelvic exam MRI Breast Bilateral (CPT® 77049) if pathology consistent with breast primary and mammogram is inconclusive If the above are non-diagnostic for primary site: CT Neck (CPT® 70491), CT Chest (CPT® 71260), and CT Abdomen (CPT® 74160) with contrast CT with contrast or MRI without and with contrast of any other symptomatic site
Carcinoma found within a bone lesion	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis with contrast (CPT® 74177) Bone Scan CT with contrast or MRI without and with contrast of any symptomatic site
 ANY of the following: Above studies have failed to demonstrate site of primary CT scans reveal isolated metastatic disease for which definitive curative therapy is planned 	PET/CT (CPT® 78815 or CPT® 78816)
Post-treatment surveillance	Advanced imaging is not indicated for routine surveillance of asymptomatic individuals after treatment completion

Extrathoracic Small Cell and Large Cell Neuroendocrine Tumors (ONC-31.8)

ON.UP.0031.8.C

v1.0.2024

All poorly-differentiated or high-grade, small cell and large cell neuroendocrine tumors arising outside the lungs or of unknown primary origin are imaged according to these guidelines.

Indication	Imaging Study
Initial staging	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast
Inconclusive findings on conventional imaging studies	• PET/CT (CPT® 78815)
ANY of the following: Poorly differentiated neuroendocrin e cancers of the head or neck Signs or symptoms of CNS involvement	MRI Brain without and with contrast (CPT® 70553)
Restaging during treatment	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) and any known sites of disease with contrast every 2 cycles
Suspected Recurrence	 ANY or ALL of the following are indicated: CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast MRI Brain without and with contrast (CPT® 70553) Bone scan PET imaging is generally not indicated but can be considered for rare circumstances.

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Indication	Imaging Study
Surveillance	CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast every 3 months for 1 year, then every 6 months for 4 additional years, then annually

Primary Peritoneal Mesothelioma (ONC-31.9)

ON.UP.0031.9.A

Indication	Imaging Study
Initial staging	 CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast PET/CT (CPT® 78815) if there is no evidence of metastatic disease or conventional imaging is inconclusive
Recurrence/ Restaging	 If there is known prior disease, CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast PET for inconclusive finding on conventional imaging
Surveillance	CT Abdomen and Pelvis with contrast (CPT® 74177) every 3 months for 2 years, then every year of life

Kaposi's Sarcoma (ONC-31.10)

ON.UP.0031.10.A

Indication	Imaging Study
Kaposi's Sarcoma	 Advanced imaging is not generally indicated since disease is generally localized to skin. CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast can be approved at initial diagnosis. If initial scans are negative then future imaging would be based on signs or symptoms.

Castleman's Disease (Unicentric and Multicentric) (ONC-31.11)

ON.UP.0031.11.A

Indication	Imaging Study
Initial staging	 Either CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast or PET/CT (CPT® 78815) CT Neck with contrast (CPT® 70491) if cervical or supraclavicular involvement If CT scans were utilized initially and suggested unicentric disease, and surgical resection is being considered, PET/CT (CPT® 78815) can be approved to confirm unicentric disease If unicentric disease is surgically removed, proceed to Surveillance section
Restaging: • Multicentric disease or surgically unresected unicentric disease on chemotherapy	 ONE of the following every 2 cycles: CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast PET/CT (CPT® 78815)
ANY of the following: • Suspected recurrence • Recurrent B symptoms • Rising LDH/IL-6/VEGF levels	ONE of the following: • CT Chest (CPT® 71260) and CT Abdomen and Pelvis (CPT® 74177) with contrast • PET/CT (CPT® 78815)
Surveillance	CT with contrast of involved areas no more than every 6 months up to 5 years

References (ONC-31)

- 1. Ettinger DS, Stevenson MM, Ahn D, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 3.2023 December 21, 2022. Occult primary, available at: https://www.nccn.org/professionals/physician_gls/pdf/occult.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Occult Primary V3.2023 December 21, 2022 ©2022 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 2. Nabors BL, Portnow J, Baehring J, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 March 24, 2023. Central Nervous System Cancers, available at: https://www.nccn.org/professionals/physician_gls/pdf/cns.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for CNS Cancer V1.2023 March 24, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 3. Zelenetz AD, Gordon LI, Abramson JS, et al, National Comprehensive Cancer Network (NCCN) Guidelines Version 5.2023 July 7, 2023. B-cell lymphomas, available at: https://www.nccn.org/professionals/physician_gls/pdf/B-CELL.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for B-cell lymphomas V5.2023 July 7, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- Mayo-Smith WM, Song JH, Boland GL, et al. Management of incidental adrenal masses: a white paper of the ACR Incidental Findings Committee. *J Am Coll Radiol*. 2017;14(8):1038-1044. doi:10.1016/j.jacr.2017.05.001.
- 5. ACR Appropriateness Criteria®. Incidentally discovered adrenal mass. Rev. 2012.
- 6. Braat AJ, Smits ML, Braat MN, et al. 90Y hepatic radioembolization: an update on current practice and recent developments. *J Nucl Med*. 2015;56(7):1079–1087. doi:10.2967/jnumed.115.157446.
- Pawaskar AS, Basu S. Role of 2-fluoro-2-deoxyglucose PET/computed tomography in carcinoma of unknown primary. PET Clin. 2015;10(3):297-310. doi:10.1016/j.cpet.2015.03.004.
- 8. Avram AM. Radioiodine scintigraphy with SPECT/CT: an important diagnostic tool for thyroid cancer staging and risk stratification. *J Nucl Med*. 2012;53(5): 754-764. doi:10.2967/jnumed.111.104133.
- Mayo-Smith WW, Song JH, Boland GL, et al. Management of incidental adrenal masses: a white paper of the ACR Incidental Findings Committee. J Am Coll Radiol. 2017;14(8):1038-1044. doi:10.1016/j.jacr.2017.05.001.
- 10. Vaidya A, Hamrahian A, Bancos I, Fleseriu M, Ghayee HK. The evaluation of incidentally discovered adrenal masses. *Endocrine Practice*. 2019:25(2);178-192. doi: 10.4158/DSCR-2018-0565.
- 11. Bergsland E, Goldner WS, Benson III AB, et al. National Comprehensive Cancer Network (NCCN) Guidelines Version 1.2023 August 2, 2023. Neuroendocrine and Adrenal Tumors, available at: https://www.nccn.org/professionals/physician_gls/pdf/neuroendocrine.pdf Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines™) for Neuroendocrine and Adrenal Tumors V1.2023 August 2, 2023. ©2023 National Comprehensive Cancer Network, Inc. All rights reserved. The NCCN Guidelines™ and illustrations herein may not be reproduced in any form for any purpose without the express written permission of the NCCN. To view the most recent and complete version of the NCCN Guidelines™, go online to NCCN.org.
- 12. Furuse M, Nonoguchi N, Yamada K, et. al. Radiological diagnosis of brain radiation necrosis after cranial irradiation for brain tumor: a systematic review. *Radiat Oncol.* 2019;14(28). doi:10.1186/s13014-019-1228-x.
- 13. American College of Radiology. ACR practice parameter for the performance of stereotactic radiosurgery. 2016; https://www.acr.org/-/media/ACR/Files/Practice-Parameters/stereobrain.pdf
- 14. Soffietti R, Abacioglu U, Baumert B, et. al. Diagnosis and treatment of brain metastases from solid tumors: guidelines from the European Assocation of Neuro-Oncology (EANO). *Neuro-Oncology*. 2017;19(2):162-174. doi:10.1093/neuonc/now241.
- 15. Mehrabian H, Detsky J, Soliman H, Sahgal A, Stanisz GJ. Advanced magnetic resonance imaging techniques in management of brain metastases. *Front Oncol.* 2019;9(440). doi:10.3389/fonc.2019.00440.

- 16. Murthy R, Nunez R, Szklaruk J, et. al. Yttrium-90 microsphere therapy for hepatic malignancy: devices, indications, technical considerations and potential complications. *RadioGraphics*. 2005;25:S41–S55. doi:10.1148/rg.25si055515.
- 17. Lencioni R, Petruzzi P, Crocetti L. Chemoembolization of hepatocellular carcinoma. *Semin Intervent Radiol.* 2013;30(1):3-11. doi:10.1055/s-0033-1333648.