



## FEP Medical Policy Manual

### FEP 6.01.51 Interim Positron Emission Tomography Scanning in Oncology to Detect Early Response During Treatment

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**Related Policies:**

- 6.01.06 - Miscellaneous (Noncardiac, Nononcologic) Applications of Fluorine 18 Fluorodeoxyglucose Positron Emission Tomography
- 6.01.20 - Cardiac Applications of Positron Emission Tomography Scanning
- 6.01.26 - Oncologic Applications of Positron Emission Tomography Scanning (Genitourinary)
- 6.01.61 - Oncologic Applications of Positron Emission Tomography Scanning (Gastrointestinal and Pancreatic)
- 6.01.62 - Oncologic Applications of Positron Emission Tomography Scanning (Breast and Gynecologic)
- 6.01.63 - Oncologic Applications of Positron Emission Tomography Scanning (Bone Sarcoma and Soft Tissue Sarcoma)
- 6.01.64 - Oncologic Applications of Positron Emission Tomography Scanning (Hematologic)
- 6.01.65 - Oncologic Applications of Positron Emission Tomography Scanning (Lung)
- 6.01.66 - Oncologic Applications of Positron Emission Tomography Scanning (Thyroid, Neuroendocrine, Head and Neck)
- 6.01.67 - Oncologic Applications of Positron Emission Tomography Scanning (Brain, Melanoma, Unknown Primary)

## Interim Positron Emission Tomography Scanning in Oncology to Detect Early Response During Treatment

### Description

#### Description

Positron emission tomography (PET) scanning has many established roles in oncology. One potential use of PET scanning is to assess treatment response early in the course of therapy, with the intent of potentially altering the regimen based on PET scan results. While several types of PET scanning are used for interim detection of cancer, this review refers to fluorine 18 fluorodeoxyglucose positron emission tomography (FDG-PET) unless otherwise noted.

## OBJECTIVE

The objective of this evidence review is to evaluate the clinical validity and clinical utility of interim positron emission tomography in assessing early response to treatment in individuals with various types of cancer.

## POLICY STATEMENT

The use of interim fluorine 18 fluorodeoxyglucose positron emission tomography scans to determine response to tyrosine kinase inhibitor treatment in individuals with gastrointestinal stromal tumors is considered **medically necessary**.

The use of interim fluorine 18 fluorodeoxyglucose positron emission tomography scans to assess response during treatment for advanced (stages IIB to 4) Hodgkin lymphoma is considered **medically necessary** (see Policy Guidelines).

The use of positron emission tomography scans to determine early response to treatment (positron emission tomography scans done during a planned course of chemotherapy and/or radiotherapy) in individuals with gastrointestinal stromal tumors on palliative or adjuvant therapy, as well as all other cancers, is considered **investigational**.

## POLICY GUIDELINES

None

## BENEFIT APPLICATION

Experimental or investigational procedures, treatments, drugs, or devices are not covered (See General Exclusion Section of brochure).

## FDA REGULATORY STATUS

A number of PET scan platforms have been cleared by the U.S. Food and Drug Administration (FDA) through the 510(k) process since the Penn-PET scanner was approved in 1989. These systems are intended to aid in detecting, localizing, diagnosing, staging, and restaging of lesions, tumors, disease, and organ function for the evaluation of diseases and disorders such as, but not limited to, cardiovascular disease, neurologic disorders, and cancer. The images produced by the system can aid in radiotherapy treatment planning and interventional radiology procedures.

PET radiopharmaceuticals have been evaluated and approved as drugs by the FDA for use as diagnostic imaging agents. These radiopharmaceuticals are approved for specific conditions. In December 2009, the FDA issued guidance for Current Good Manufacturing Practice for PET drug manufacturers<sup>2</sup>, and, in August 2011, issued similar Current Good Manufacturing Practice Guidance for small businesses compounding radiopharmaceuticals.<sup>3</sup> An additional final guidance document issued in December 2012 required all PET drug manufacturers and compounders to operate under an approved new drug application, abbreviated new drug application, or investigational new drug application, by December 12, 2015.<sup>4</sup>

Table 1 lists some of the radiopharmaceuticals granted FDA approval for use with PET for oncologic-related indications.

**Table 1. Radiopharmaceuticals Approved for Use With PET for Carcinoma-Related Indications**

Agent	Brand Name	Manufacturer	Date Approved	NDA No.	Carcinoma-Related Indication With PET
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Carbon 11 choline	NA	Various	2012	203155	Suspected prostate cancer recurrence based on elevated blood PSA after therapy and noninformative bone scintigraphy, CT, or MRI
Copper 64 dotatate	Detectnet™	Curium	2020	213227	Localization of somatostatin receptor-positive NETs in adult patients
Fluorine 18 fluorodeoxyglucose	NA	Various	2000	20306	Suspected or existing diagnosis of cancer, all types
Fluorine 18 fluciclovine	Axumin™	Blue Earth Diagnostics	2016	208054	Suspected prostate cancer recurrence based on elevated blood PSA levels after treatment
Fluorine 18 fluoroestradiol	CERIANNA™	Zionexa	2020	212155	Detection of ER-positive lesions as an adjunct to biopsy in patients with recurrent or metastatic breast cancer
Gallium 68 dotatate	NETSPOT™	Advanced Accelerator Applications	2016	208547	Localization of somatostatin receptor-positive NETs in adult and pediatric patients
Gallium 68 dotatoc	NA	University of Iowa	2019	210828	Localization of somatostatin receptor-positive NETs in adult and pediatric patients
Gallium 68 PSMA-11	NA	University of California, Los Angeles and the University of California, San Francisco	2020	212642	PSMA positive lesions in men with prostate cancer with suspected metastasis who are candidates for initial definitive therapy or with suspected recurrence based on elevated serum PSA level
Piflufolostat fluorine-18	Pylarify	Progenics Pharmaceuticals, Inc	2021	214793	PSMA positive lesions in men with prostate cancer with suspected metastasis who are candidates for initial definitive therapy or with suspected recurrence based on elevated serum PSA level

CT: computed tomography; ER: estrogen receptor; MRI: magnetic resonance imaging; NA: not applicable; NDA: new drug application; NETs: neuroendocrine tumors; PET: positron emission tomography; PSA: prostate-specific antigen; PSMA: prostate-specific membrane antigen.

## RATIONALE

### Summary of Evidence

#### Breast Cancer

For individuals with breast cancer who receive interim fluorine 18 fluorodeoxyglucose positron emission tomography (FDG-PET) as an adjunct to interim computed tomography (CT), the evidence consists of several systematic reviews, randomized controlled trials (RCTs), and many observational studies. Relevant outcomes are overall survival (OS), disease-specific survival, change in disease status, quality of life (QOL), morbid events, and treatment-related morbidity. Results from systematic reviews have shown wide ranges in sensitivities, specificities, and negative (NPV) and positive predictive values (PPV). The wide ranges might be due to small sample sizes, the use of various definitions of the outcome measure (pathologic complete response), and differences in breast cancer subtype populations. Two RCTs were identified in which therapy decisions were guided by FDG-PET results. In the first RCT, nonresponders, determined by positron emission tomography (PET) measures, were given more intensive chemotherapy. Although the results showed initially higher response rates in the more intensive treatment group, this did not translate to long-term improvements in disease-free survival. The second RCT found that patients receiving less intensive initial treatment who were determined to be responders by PET measures had significantly higher response rates to treatment. By year 3, 94.8% of those PET-responders treated with less-intensive treatment had experienced disease-free survival, without any serious adverse events. This PET-based, pathologic complete response-adapted strategy identified about a third of patients in the clinical trial with HER2-positive early breast cancer who could safely omit chemotherapy and was associated with an excellent 3-year disease free survival. However, further clinical investigation is warranted to determine the appropriateness of interim FDG-PET in dictating treatment. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

#### Esophageal Cancer

For individuals with esophageal cancer who receive interim FDG-PET as an adjunct to interim CT, the evidence includes meta-analyses, nonrandomized studies, and retrospective studies. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. Results on clinical validity were inconsistent across the studies. The meta-analysis reported low pooled sensitivities and specificities, while a subgroup analysis including only patients with squamous cell carcinoma and 2 studies published after the meta-analysis reported an adequate potential in predicting responders to neoadjuvant therapy. No evidence was identified that examined the clinical utility of PET for patients with esophageal cancer. Evidence for clinical utility of FDG-PET for patients with esophageal cancer consists of 1 meta-analysis and 1 RCT. The meta-analysis found that patients considered to be responders early in therapy based on FDG-PET assessment were found to have improvements in progression-free survival (PFS) and OS compared to nonresponders. A single RCT found that PET-guided therapy led to improvements in PFS, but not OS, in patients considered nonresponders to initial therapy. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

#### Gastrointestinal Stromal Tumors

For individuals with gastrointestinal stromal tumors receiving palliative or adjuvant therapy who receive interim FDG-PET as an adjunct to interim CT, the evidence includes a systematic review. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. The systematic review included 19 studies, 2 of which reviewed FDG-PET scans more than 6 months after the start of treatment. CT is currently recommended for standard long-term follow-up and surveillance of gastrointestinal stromal tumors. FDG-PET is equivalent to CT in the detection of treatment response when follow-up is long-term. No studies were identified that tested outcomes following PET-guided treatment. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

For individuals with gastrointestinal stromal tumors treated with tyrosine kinase inhibitors (TKIs) for 6 months or less who receive interim FDG-PET as an adjunct to interim CT, the evidence includes a systematic review. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. The systematic review included 19 studies, 17 of which showed that FDG-PET detected an early response to TKI therapy, which was a strong predictor of clinical outcomes. FDG-PET detected treatment response as early as 1 week after initiation of treatment. While CT detects anatomic changes in the tumor, PET detects changes in the metabolic activity of the tumor. Because metabolic changes precede anatomic changes by several weeks or sometimes months, PET can detect treatment response earlier than CT. PET is therefore preferred if a rapid read-out of response to targeted therapy is needed to guide treatment decisions (eg, change in targeted therapy or surgery). While no studies were identified that tested outcomes following PET-guided treatment, it is possible to construct a chain of evidence demonstrating improved patient outcomes. The evidence is sufficient to determine that the technology results in an improvement in the net health outcome.

## Head and Neck Cancer

For individuals with head and neck cancer who receive interim FDG-PET as an adjunct to CT, the evidence includes several systematic reviews. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. There was an overlap of studies among the systematic reviews. Most studies included in the reviews showed that FDG-PET used during radiotherapy, with or without chemotherapy, can adequately predict disease-free and OS. Meta-analyses to determine response could not be performed in any of the systematic reviews due to the heterogeneity in the methods across the studies. Most studies used maximum standardized uptake value (SUVmax), however, threshold values to determine response varied across studies. No studies were identified that provided evidence for the clinical utility of PET. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

## Hodgkin Lymphoma

For individuals with Hodgkin lymphoma (HL) who receive interim FDG-PET as an adjunct to interim CT, the evidence includes systematic reviews with meta-analyses and RCTs. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. The systematic review evaluating the validity of interim FDG-PET in lymphomas did not identify any studies in HL. Evidence for the clinical utility of interim PET for guiding treatment in patients with HL consists of 2 Cochrane reviews and several RCTs. A 2025 Cochrane review identified 10 RCTs evaluating interim PET in patients with HL. The evidence in early- and intermediate-stage was limited; however, the authors concluded that interim-PET has the potential to increase OS without negatively affecting progression-free survival (PFS) and long-term adverse events in patients with advanced-stage HL. Another Cochrane review found moderate-certainty evidence that interim PET scan results predict OS, and very low-certainty evidence that interim PET scan results predict PFS in treated individuals with HL. The evidence is sufficient to determine that the technology results in an improvement in the net health outcome for patients with advanced (stages IIB to 4) HL; however, evidence is insufficient to determine that the technology results in an improvement in the net health outcome for patients with early or intermediate stages of HL.

## Non-Hodgkin Lymphoma

For individuals with non-Hodgkin lymphoma (NHL) who receive interim FDG-PET as an adjunct to interim CT, the evidence includes systematic reviews with meta-analyses and RCTs. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. The systematic review evaluating the validity of interim FDG-PET showed high false-positive rates for NHL. Evidence for the clinical utility of interim PET for guiding treatment in patients with NHL consists of a systematic review and 2 RCTs. The systematic review included only observational studies. The phase 3 RCT evaluated treatment intensification in patients who were PET2-positive and found intensification did not improve survival and increased toxicity. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

## Non-Small-Cell Lung Cancer

For individuals with non-small-cell lung cancer (NSCLC) who receive interim FDG-PET as an adjunct to interim CT, the evidence includes numerous small observational studies. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. While most studies showed correlations between FDG-PET measurements and progression-free and OS, the generalizability of the results is limited. The studies were small, with most population sizes fewer than 50 patients. The studies were also heterogeneous, including patients at different stages of the disease, undergoing different treatment regimens, and receiving PET at different times during treatment cycles. No studies were identified that evaluated outcomes after PET-guided therapy. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

## Ovarian Cancer

For individuals with ovarian cancer who receive interim FDG-PET as an adjunct to interim CT, the evidence includes a systematic review. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and treatment-related morbidity. The systematic review identified 9 studies that calculated hazard ratios for various FDG-PET parameters (eg, SUVmax, metabolic tumor volume, tumor lesion glycolysis). The only parameter consistently showing prognostic value was tumor lesion glycolysis. Additionally, no studies were identified that evaluated outcomes after PET-guided therapy. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

## Other Cancers

For individuals with other malignant solid tumors (eg, bladder, colorectal, prostate, thyroid) who receive FDG-PET as an adjunct to interim CT, the evidence includes a systematic review, National Comprehensive Cancer Network (NCCN) task force report, and single-arm observational studies published after the task force report. Relevant outcomes are OS, disease-specific survival, change in disease status, QOL, morbid events, and

treatment-related morbidity. Results have been inconsistent on the use of interim FDG-PET among the various cancers. While some have reported associations between interim FDG-PET and recurrence or survival, there is a lack of comparative trials evaluating outcomes in patients whose treatments were altered based on PET measurements. The evidence is insufficient to determine that the technology results in an improvement in the net health outcome.

## SUPPLEMENTAL INFORMATION

### Practice Guidelines and Position Statements

Guidelines or position statements will be considered for inclusion in 'Supplemental Information' if they were issued by, or jointly by, a US professional society, an international society with US representation, or National Institute for Health and Care Excellence (NICE). Priority will be given to guidelines that are informed by a systematic review, include strength of evidence ratings, and include a description of management of conflict of interest.

#### American College of Radiology et al

The American College of Radiology (ACR) and the Society for Pediatric Radiology (2016; amended 2023 ) updated their joint practice parameter for performing fluorine 18 fluorodeoxyglucose positron emission tomography (FDG-PET) coupled with computed tomography (CT) in oncology.<sup>113</sup> The practice parameter states that examples of indications for FDG-PET/CT include, but are not limited to, the following:

- "Staging on presentation for guiding initial treatment strategy in patients with a known malignancy;
- Monitoring response to therapy to include determining whether residual abnormalities identified with another imaging modality represent persistent viable tumor or posttreatment changes (inflammation, fibrosis, or necrosis);
- Restaging in the setting of relapse;
- Attempting to localize the site of primary tumor when metastatic disease is the initial manifestation of malignancy;
- Verifying and localizing "occult" disease, especially in the presence of clinical indicators such as elevated tumor markers;
- Evaluating an abnormality considered "indeterminate" by another imaging modality to determine whether glucose metabolism in that abnormality favors a benign or malignant process;
- Guiding treatment goals, such as curative versus palliative therapy;
- Guiding biopsy and radiation therapy planning."

#### European Association of Nuclear Medicine

The European Association of Nuclear Medicine (EANM; 2021) published guidelines on FDG-PET/CT in the management of ovarian cancer, which are endorsed by the American College of Nuclear Medicine, the Society of Nuclear Medicine and Molecular Imaging (SNMMI), and the International Atomic Energy Agency.<sup>114</sup> The guidelines acknowledge the lack of clinical trials evaluating the role of FDG-PET scanning when used for assessment of response to therapy in patients with ovarian cancer (Level of evidence, II; grade B recommendation). Further recommendations are not provided.

The EANM published 2024 guidelines for FDG-PET/CT in diagnosis, staging, prognostication, therapy assessment, and restaging of plasma cell disorders.<sup>115</sup> The guidelines state the following for therapy assessment in multiple myeloma, "[18F]FDG PET/CT is able to distinguish between metabolically active MM lesions and inactive fibrous residual osteolytic lesions, with an earlier and higher rate of scan normalization than MRI after therapy initiation" per the International Myeloma Working Group.

In 2024, the EANM and SNMMI published joint guidelines on FDG-PET/CT in the management of breast cancer, which are endorsed by the ACR.<sup>116</sup> Relevant recommendations are summarized in Table 2.

#### Table 2. EANM-SNMMI Recommendations for FDG-PET/CT in Breast Cancer

Recommendation	LOE	Grade of recommendation
<i>Indications for FDG-PET/CT</i>		
FDG PET/CT is not recommended in stage I	II	B
FDG PET/CT may be useful in patients with clinical stage IIA (T1N1 or T2N0), but there is not enough strong data to recommend routine use in this subgroup	III	C
FDG PET/CT can be recommended for baseline staging of stage IIB (preferably before surgery) and stage III (including inflammatory breast cancer)	II	B
FDG PET/CT can be done instead of, and not in combination with, conventional imaging modalities for staging (combination of bone scan, chest X-ray or CT-chest, and ultrasound of the liver or CT-abdomen)	II	B
FDG PET/CT is recommended in baseline treatment planning and may improve radiation therapy planning	III	C
FDG PET/CT can be useful for determining the extent of metastatic disease (outside the brain) and improving treatment planning	III	C
FDG PET/CT can be done instead of, and not in addition to separate conventional imaging modalities (combination of bone scan, chest X-ray or CT-chest, and ultrasound of the liver or CT-abdomen)	II	B
<i>Assessment of treatment response</i>		
FDG PET/CT may be used to assess early metabolic response in non-metastatic breast cancer, particularly in TNBC and HER2 +	II	B
FDG PET/CT may play a role in monitoring treatment response in metastatic breast cancer	III	C
FDG PET/CT may be particularly useful to assess bone metastases and enable early response to treatment evaluation	III	C
<i>Assessment of recurrence</i>		
FDG PET/CT is useful to detect the site and extent of recurrence when conventional imaging methods are equivocal	I	A
FDG PET/CT can be recommended:		
In patients with signs or symptoms suggestive of metastatic disease	I	A
In patients with rising serum tumour markers	II	B
To guide site of biopsy	IV	D
To improve radiation therapy planning	III	C
FDG PET/CT can substitute for CT and/or bone scan in the detection of bone metastases	II	B

CT: computed tomography; EANM: European Association of Nuclear Medicine; FDG: fluorine 18 fluorodeoxyglucose; HER2: human epidermal growth factor receptor 2; LOE: level of evidence; PET: positron emission tomography; SNMMI: Society of Nuclear Medicine and Molecular Imaging; TNBC: triple-negative breast cancer.

## National Comprehensive Cancer Network

Current National Comprehensive Cancer Network recommendations for interim PET scanning during treatment to assess early response in a variety of cancers are summarized in Table 3.

The policies contained in the FEP Medical Policy Manual are developed to assist in administering contractual benefits and do not constitute medical advice. They are not intended to replace or substitute for the independent medical judgment of a practitioner or other health care professional in the treatment of an individual member. The Blue Cross and Blue Shield Association does not intend by the FEP Medical Policy Manual, or by any particular medical policy, to recommend, advocate, encourage or discourage any particular medical technologies. Medical decisions relative to medical technologies are to be made strictly by members/patients in consultation with their health care providers. The conclusion that a particular service or supply is medically necessary does not constitute a representation or warranty that the Blue Cross and Blue Shield Service Benefit Plan covers (or pays for) this service or supply for a particular member.

**Table 3. Recommendations for Interim PET Scanning**

Guideline	Version	Recommendation
Bladder cancer <sup>117</sup> ,	1.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Breast cancer <sup>118</sup> ,	4.2025 4	"Studies of functional imaging, such as radionuclide bone scans and PET imaging, are particularly challenging when used to assess response... PET imaging is challenging because of the absence of a reproducible, validated, and widely accepted set of standards for disease activity assessment."
CNS cancers <sup>119</sup> ,	2.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Cervical cancer <sup>120</sup> ,	4.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Colon cancer <sup>121</sup> ,	4.2025	"The panel strongly discourages the routine use of PET/CT scanning for staging, baseline imaging, or routine follow-up." "PET/CT scans should not be used to assess response to chemotherapy because a PET/CT scan can become transiently negative after chemotherapy. False-positive PET/CT scan results can occur in the presence of tissue inflammation after surgery or infection."
Esophageal and EGJ cancers <sup>122</sup> ,	4.2025	"Regardless of the cut-off values used,...studies...concluded that FDG-PET is predictive of pathologic response and survival in patients with esophageal cancer who undergo preoperative treatment." "Increased FDG uptake due to radiation-induced inflammation limits the use of FDG-PET for early response assessment of esophageal carcinomas. To reduce the incidence of false-positive results due to inflammation, the guidelines recommend that FDG-PET/CT (preferred) or FDG-PET should be performed at least 5 to 8 weeks after the completion of preoperative therapy. However, the guidelines caution that post-treatment FDG-PET results should not be used to select patients for surgery since FDG-PET cannot distinguish microscopic residual disease."
Soft tissue sarcoma <sup>123</sup> ,	1.2025	Interim PET for assessing response to ongoing treatment is not addressed. "FDG-PET/CT scan may be useful in staging, prognostication, grading, and determining response to neoadjuvant therapy."
Head and neck cancers <sup>124</sup> ,	5.2025	Short-term (<6 months) locoregionally advanced disease: "FDG PET/CT should be performed within 3 to 6 months of definitive radiation of systemic therapy/RT for assessment of treatment response and to identify any residual tumor." "Early FDG-PET/CT scans before 12 weeks are associated with significant false-positive rates and should be avoided in the absence of signs of recurrence or progression." "The optimal timing of PET scans after radiation treatment appears to be at the 3- to 6-month window. A negative PET at this time point predicts improved overall survival at 2 years."
Hepatocellular Carcinoma <sup>125</sup> ,	1.2025	Interim PET for assessing response to ongoing treatment is not addressed. "PET/CT has limited sensitivity but high specificity, and may be considered when there is an equivocal finding. When HCC is detected by CT or MRI and has increased metabolic activity on PET/CT, higher intralésional standardized uptake value is a marker of biologic aggressiveness and might predict less optimal response to locoregional therapies."
Biliary Tract Cancers <sup>126</sup> ,	2.2025	Interim PET for assessing response to ongoing treatment is not addressed. "PET/CT has limited sensitivity but high specificity and may be considered when there is an equivocal finding or on a case-by-case basis. The routine use of PET/CT in the preoperative setting has not been established in prospective trials"
Hodgkin lymphoma <sup>127</sup> ,	2.2025	"Interim FDG-PET scans can be prognostic and are increasingly being used to assess treatment response during therapy as they can inform treatment adaptation, including treatment escalation and de-escalation. Early interim FDG-PET imaging after chemotherapy has been shown to be a sensitive prognostic indicator of treatment outcome in patients with advanced-stage disease. Interim FDG-PET scans may also be useful to identify a subgroup of patients with early- and advanced-stage disease that can be treated with chemotherapy alone. The NCCN Guidelines emphasize that the value of interim FDG-PET scans remains unclear for some clinical scenarios, and all measures of response should be considered in the context of management decisions."

		It is important that the Deauville score be incorporated into the nuclear medicine FDG-PET scan report, since subsequent management is often dependent upon that score."
Cutaneous melanoma <sup>128,</sup>	2.2025	Interim PET for assessing response to ongoing treatment is not addressed. "Recent studies in patients with stage III or IV melanoma... indicated that additional information provided by PET/CT may impact treatment decisions in up to 30% of patients, with the greatest impact seen in surgical management."
Malignant pleural mesothelioma <sup>129,</sup>	2.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Multiple myeloma <sup>128,</sup>	2.2026	Interim PET for assessing response to ongoing treatment is not addressed.
Non-Hodgkin lymphoma: B-cell <sup>130,</sup>	3.2025	"Further prospective studies are warranted to determine whether interim PET scans have a role in guiding post-induction therapeutic interventions." "A negative FDG-PET scan after 2 to 4 cycles of induction therapy has been associated with significantly higher EFS and OS rates in several studies. However, interim FDG-PET scans can produce false-positive results and chemoimmunotherapy is associated with a favorable long-term outcome despite a positive interim FDG-PET scan."
Non-Hodgkin lymphoma: T-cell <sup>131,</sup>	2.2025	"The guidelines recommend interim restaging with PET/CT (preferred) or CT after 3 to 4 cycles of chemotherapy."
Primary Cutaneous Lymphomas <sup>132,</sup>	3.2025	Interim PET for assessing response to ongoing treatment is not addressed. "FDG-PET/CT or chest/abdomen/pelvis CT with contrast at the end of treatment may be needed to assess response or if there is clinical suspicion of progressive disease."
NSCLC <sup>133,</sup>	8.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Ovarian cancer <sup>134,</sup>	3.2025	Interim PET for assessing response to ongoing treatment is not addressed. Primary chemotherapy regimens include monitoring with chest/abdominal/pelvic CT or MRI with contrast, PET/CT (skull base to mid-thigh), or PET as indicated <sup>a</sup>
Pancreatic adenocarcinoma <sup>135,</sup>	2.2025	Interim PET for assessing response to ongoing treatment is not addressed. "PET/CT or PET/MRI scan may be considered after formal pancreatic CT protocol in patients with high-risk to detect extrapancreatic metastases. It is not a substitute for high-quality, contrast-enhanced CT."
Prostate cancer <sup>136,</sup>	2.2026	"F-18 FDG-PET should not be used routinely, because data are limited in patients with prostate cancer and suggest that its sensitivity is significantly lower than that seen with the above described tracers."
Rectal cancer <sup>137,</sup>	3.2025	"Chest/abdomen/pelvis CT with contrast or chest CT and abdomen/pelvis MRI with contrast to monitor progress of therapy. PET/CT should not be used. "
SCLC <sup>138,</sup>	2.2026	"The panel maintains that FDG-PET/CT is not recommended for routine follow-up unless contrast CT chest/abdomen/pelvis or MRI is contraindicated."
Thyroid carcinoma <sup>139,</sup>	1.2025	Interim PET for assessing response to ongoing treatment is not addressed.
Uterine neoplasms <sup>140,</sup>	3.2025	Interim PET for assessing response to ongoing treatment is not addressed

CNS: central nervous system; CT: computed tomography; EFS: event-free survival; EGJ: esophagogastric junction; FDG: fluorine 18 fluorodeoxyglucose; HCC: hepatocellular carcinoma; MRI: magnetic resonance imaging; NCCN: National Comprehensive Cancer Network; NSCLC: non-small-cell lung cancer; OS: overall survival; PCBC: primary cutaneous B-cell lymphoma; PET: positron emission tomography; SCLC: small-cell lung cancer; SUV: standardized uptake value.

<sup>a</sup> This statement is a footnote to epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer treatment recommendations

## U.S. Preventive Services Task Force Recommendations

Not applicable.

## Medicare National Coverage

The national coverage determination on FDG-PET for oncologic conditions (220.6.17) makes the following coverage decisions:<sup>141</sup>.

"Three FDG PET scans are nationally covered when used to guide subsequent management of anti-tumor treatment strategy after completion of initial anti-cancer therapy. Coverage of more than three FDG PET scans to guide subsequent management of anti-tumor treatment strategy after completion of initial anti-cancer therapy shall be determined by the local Medicare Administrative Contractors."

## REFERENCES

- Hillner BE, Siegel BA, Shields AF, et al. The impact of positron emission tomography (PET) on expected management during cancer treatment: findings of the National Oncologic PET Registry. *Cancer*. Jan 15 2009; 115(2): 410-8. PMID 19016303
- Food and Drug Administration (FDA). PET Drugs - Current Good Manufacturing Practice (CGMP). 2009; <https://www.fda.gov/downloads/Drugs/GuidanceComplianceRegulatoryInformation/Guidances/UCM070306.pdf>. Accessed September 14, 2025.
- Food and Drug Administration (FDA). PET Drugs - Current Good Manufacturing Practice (CGMP) Small Entity Compliance Guide. 2011; <https://www.fda.gov/downloads/Drugs/GuidanceComplianceRegulatoryInformation/Guidances/UCM266640.pdf>. Accessed September 15, 2025.
- Food and Drug Administration (FDA). Guidance: Investigational New Drug Applications for Positron Emission Tomography (PET) Drugs. 2012; <https://www.fda.gov/downloads/Drugs/GuidanceComplianceRegulatoryInformation/Guidances/UCM291573.pdf>. Accessed September 16, 2025.
- Li H, Yao L, Jin P, et al. MRI and PET/CT for evaluation of the pathological response to neoadjuvant chemotherapy in breast cancer: A systematic review and meta-analysis. *Breast*. Aug 2018; 40: 106-115. PMID 29758503
- Lindenberg MA, Miquel-Cases A, Retl VP, et al. Imaging performance in guiding response to neoadjuvant therapy according to breast cancer subtypes: A systematic literature review. *Crit Rev Oncol Hematol*. Apr 2017; 112: 198-207. PMID 28325260
- Chen L, Yang Q, Bao J, et al. Direct comparison of PET/CT and MRI to predict the pathological response to neoadjuvant chemotherapy in breast cancer: a meta-analysis. *Sci Rep*. Aug 16 2017; 7(1): 8479. PMID 28814795
- Boers-Sonderen MJ, de Geus-Oei LF, Desar IM, et al. Temsirolimus and pegylated liposomal doxorubicin (PLD) combination therapy in breast, endometrial, and ovarian cancer: phase Ib results and prediction of clinical outcome with FDG-PET/CT. *Target Oncol*. Dec 2014; 9(4): 339-47. PMID 24577626
- Groheux D, Hindi E, Giacchetti S, et al. Early assessment with 18F-fluorodeoxyglucose positron emission tomography/computed tomography can help predict the outcome of neoadjuvant chemotherapy in triple negative breast cancer. *Eur J Cancer*. Jul 2014; 50(11): 1864-71. PMID 24841218
- Humbert O, Cochet A, Riedinger JM, et al. HER2-positive breast cancer: <sup>18</sup>F-FDG PET for early prediction of response to trastuzumab plus taxane-based neoadjuvant chemotherapy. *Eur J Nucl Med Mol Imaging*. Aug 2014; 41(8): 1525-33. PMID 24647576
- Andrade WP, Lima EN, Osrio CA, et al. Can FDG-PET/CT predict early response to neoadjuvant chemotherapy in breast cancer?. *Eur J Surg Oncol*. Dec 2013; 39(12): 1358-63. PMID 24120422
- Mghanga FP, Lan X, Bakari KH, et al. Fluorine-18 fluorodeoxyglucose positron emission tomography-computed tomography in monitoring the response of breast cancer to neoadjuvant chemotherapy: a meta-analysis. *Clin Breast Cancer*. Aug 2013; 13(4): 271-9. PMID 23714689
- Humbert O, Riedinger JM, Charon-Barra C, et al. Identification of Biomarkers Including 18FDG-PET/CT for Early Prediction of Response to Neoadjuvant Chemotherapy in Triple-Negative Breast Cancer. *Clin Cancer Res*. Dec 15 2015; 21(24): 5460-8. PMID 26130460
- Humbert O, Riedinger JM, Vrigneaud JM, et al. 18F-FDG PET-Derived Tumor Blood Flow Changes After 1 Cycle of Neoadjuvant Chemotherapy Predicts Outcome in Triple-Negative Breast Cancer. *J Nucl Med*. Nov 2016; 57(11): 1707-1712. PMID 27103025
- Lee HW, Lee HM, Choi SE, et al. The Prognostic Impact of Early Change in 18F-FDG PET SUV After Neoadjuvant Chemotherapy in Patients with Locally Advanced Breast Cancer. *J Nucl Med*. Aug 2016; 57(8): 1183-8. PMID 27033896
- Luo J, Zhou Z, Yang Z, et al. The Value of 18F-FDG PET/CT Imaging Combined With Pretherapeutic Ki67 for Early Prediction of Pathologic Response After Neoadjuvant Chemotherapy in Locally Advanced Breast Cancer. *Medicine (Baltimore)*. Feb 2016; 95(8): e2914. PMID 26937935
- Pahk K, Kim S, Choe JG. Early prediction of pathological complete response in luminal B type neoadjuvant chemotherapy-treated breast cancer patients: comparison between interim 18F-FDG PET/CT and MRI. *Nucl Med Commun*. Sep 2015; 36(9): 887-91. PMID 25932536
- Lin NU, Guo H, Yap JT, et al. Phase II Study of Lapatinib in Combination With Trastuzumab in Patients With Human Epidermal Growth Factor Receptor 2-Positive Metastatic Breast Cancer: Clinical Outcomes and Predictive Value of Early [<sup>18</sup>F]Fluorodeoxyglucose Positron Emission Tomography Imaging (TBCRC 003). *J Clin Oncol*. Aug 20 2015; 33(24): 2623-31. PMID 26169615
- Kitajima K, Miyoshi Y, Yamano T, et al. Assessment of tumor response to neoadjuvant chemotherapy in patients with breast cancer using MRI and FDG-PET/CT-RECIST 1.1 vs. PERCIST 1.0. *Nagoya J Med Sci*. May 2018; 80(2): 183-197. PMID 29915436
- Kitajima K, Nakatani K, Yamaguchi K, et al. Response to neoadjuvant chemotherapy for breast cancer judged by PERCIST - multicenter study in Japan. *Eur J Nucl Med Mol Imaging*. Sep 2018; 45(10): 1661-1671. PMID 29754160

21. Yoon HJ, Kim Y, Chung J, et al. Predicting neo-adjuvant chemotherapy response and progression-free survival of locally advanced breast cancer using textural features of intratumoral heterogeneity on F-18 FDG PET/CT and diffusion-weighted MR imaging. *Breast J.* May 2019; 25(3): 373-380. PMID 29602210
22. Groheux D, Biard L, Giacchetti S, et al. <sup>18</sup>F-FDG PET/CT for the Early Evaluation of Response to Neoadjuvant Treatment in Triple-Negative Breast Cancer: Influence of the Chemotherapy Regimen. *J Nucl Med.* Apr 2016; 57(4): 536-43. PMID 26697967
23. Groheux D, Majdoub M, Sanna A, et al. Early Metabolic Response to Neoadjuvant Treatment: FDG PET/CT Criteria according to Breast Cancer Subtype. *Radiology.* Nov 2015; 277(2): 358-71. PMID 25915099
24. van Ramshorst MS, Teixeira SC, Koolen BB, et al. Additional value of 18 F-FDG PET/CT response evaluation in axillary nodes during neoadjuvant therapy for triple-negative and HER2-positive breast cancer. *Cancer Imaging.* May 25 2017; 17(1): 15. PMID 28545563
25. Schmitz AMT, Teixeira SC, Pengel KE, et al. Monitoring tumor response to neoadjuvant chemotherapy using MRI and 18F-FDG PET/CT in breast cancer subtypes. *PLoS One.* 2017; 12(5): e0176782. PMID 28531188
26. Riedl CC, Pinker K, Ulaner GA, et al. Comparison of FDG-PET/CT and contrast-enhanced CT for monitoring therapy response in patients with metastatic breast cancer. *Eur J Nucl Med Mol Imaging.* Aug 2017; 44(9): 1428-1437. PMID 28462446
27. Coudert B, Pierga JY, Mouret-Reynier MA, et al. Use of [(18)F]-FDG PET to predict response to neoadjuvant trastuzumab and docetaxel in patients with HER2-positive breast cancer, and addition of bevacizumab to neoadjuvant trastuzumab and docetaxel in [(18)F]-FDG PET-predicted non-responders (AVATAXHER): an open-label, randomised phase 2 trial. *Lancet Oncol.* Dec 2014; 15(13): 1493-1502. PMID 25456368
28. Coudert B, Pierga JY, Mouret-Reynier MA, et al. Long-term outcomes in patients with PET-predicted poor-responsive HER2-positive breast cancer treated with neoadjuvant bevacizumab added to trastuzumab and docetaxel: 5-year follow-up of the randomised Avataxher study. *EClinicalMedicine.* Nov 2020; 28: 100566. PMID 33205032
29. Prez-Garca JM, Gebhart G, Ruiz Borrego M, et al. Chemotherapy de-escalation using an 18 F-FDG-PET-based pathological response-adapted strategy in patients with HER2-positive early breast cancer (PHERGain): a multicentre, randomised, open-label, non-comparative, phase 2 trial. *Lancet Oncol.* Jun 2021; 22(6): 858-871. PMID 34019819
30. Prez-Garca JM, Corts J, Ruiz-Borrego M, et al. 3-year invasive disease-free survival with chemotherapy de-escalation using an 18 F-FDG-PET-based, pathological complete response-adapted strategy in HER2-positive early breast cancer (PHERGain): a randomised, open-label, phase 2 trial. *Lancet.* Apr 27 2024; 403(10437): 1649-1659. PMID 38582092
31. Han S, Kim YI, Woo S, et al. Prognostic and predictive values of interim 18 F-FDG PET during neoadjuvant chemoradiotherapy for esophageal cancer: a systematic review and meta-analysis. *Ann Nucl Med.* Apr 2021; 35(4): 447-457. PMID 33471289
32. Cong L, Wang S, Gao T, et al. The predictive value of 18F-FDG PET for pathological response of primary tumor in patients with esophageal cancer during or after neoadjuvant chemoradiotherapy: a meta-analysis. *Jpn J Clin Oncol.* Dec 2016; 46(12): 1118-1126. PMID 27702836
33. van Rossum PSN, Fried DV, Zhang L, et al. The value of 18 F-FDG PET before and after induction chemotherapy for the early prediction of a poor pathologic response to subsequent preoperative chemoradiotherapy in oesophageal adenocarcinoma. *Eur J Nucl Med Mol Imaging.* Jan 2017; 44(1): 71-80. PMID 27511188
34. Hagen PV, Heijl MV, van Berge Henegouwen MI, et al. Prediction of disease-free survival using relative change in FDG-uptake early during neoadjuvant chemoradiotherapy for potentially curable esophageal cancer: A prospective cohort study. *Dis Esophagus.* Feb 01 2017; 30(2): 1-7. PMID 27001344
35. Odawara S, Kitajima K, Katsuura T, et al. Tumor response to neoadjuvant chemotherapy in patients with esophageal cancer assessed with CT and FDG-PET/CT - RECIST 1.1 vs. PERCIST 1.0. *Eur J Radiol.* Apr 2018; 101: 65-71. PMID 29571803
36. Manoharan V, Lee S, Chong S, et al. Serial imaging using [18F]Fluorodeoxyglucose positron emission tomography and histopathologic assessment in predicting survival in a population of surgically resectable distal oesophageal and gastric adenocarcinoma following neoadjuvant therapy. *Ann Nucl Med.* May 2017; 31(4): 315-323. PMID 28299585
37. Goodman KA, Ou FS, Hall NC, et al. Randomized Phase II Study of PET Response-Adapted Combined Modality Therapy for Esophageal Cancer: Mature Results of the CALGB 80803 (Alliance) Trial. *J Clin Oncol.* Sep 01 2021; 39(25): 2803-2815. PMID 34077237
38. Treglia G, Mirk P, Stefanelli A, et al. 18F-Fluorodeoxyglucose positron emission tomography in evaluating treatment response to imatinib or other drugs in gastrointestinal stromal tumors: a systematic review. *Clin Imaging.* 2012; 36(3): 167-75. PMID 22542374
39. Podoloff DA, Advani RH, Allred C, et al. NCCN task force report: positron emission tomography (PET)/computed tomography (CT) scanning in cancer. *J Natl Compr Canc Netw.* May 2007; 5 Suppl 1: S1-22; quiz S23-2. PMID 17509259
40. Helsen N, Van den Wyngaert T, Carp L, et al. FDG-PET/CT for treatment response assessment in head and neck squamous cell carcinoma: a systematic review and meta-analysis of diagnostic performance. *Eur J Nucl Med Mol Imaging.* Jun 2018; 45(6): 1063-1071. PMID 29478080
41. Min M, Lin P, Liney G, et al. A review of the predictive role of functional imaging in patients with mucosal primary head and neck cancer treated with radiation therapy. *J Med Imaging Radiat Oncol.* Feb 2017; 61(1): 99-123. PMID 27469298
42. Castelli J, De Bari B, Depeursinge A, et al. Overview of the predictive value of quantitative 18 FDG PET in head and neck cancer treated with chemoradiotherapy. *Crit Rev Oncol Hematol.* Dec 2016; 108: 40-51. PMID 27931839
43. Dos Anjos RF, Dos Anjos DA, Vieira DL, et al. Effectiveness of FDG-PET/CT for evaluating early response to induction chemotherapy in head and neck squamous cell carcinoma: A systematic review. *Medicine (Baltimore).* Aug 2016; 95(32): e4450. PMID 27512861
44. Adams HJA, Kwee TC. Proportion of false-positive lesions at interim and end-of-treatment FDG-PET in lymphoma as determined by histology: Systematic review and meta-analysis. *Eur J Radiol.* Nov 2016; 85(11): 1963-1970. PMID 27776647
45. Sickinger MT, von Tresckow B, Kobe C, et al. Positron emission tomography-adapted therapy for first-line treatment in individuals with Hodgkin lymphoma. *Cochrane Database Syst Rev.* Jan 09 2015; 1(1): CD010533. PMID 25572491
46. Radford J, Illidge T, Counsell N, et al. Results of a trial of PET-directed therapy for early-stage Hodgkin's lymphoma. *N Engl J Med.* Apr 23 2015; 372(17): 1598-607. PMID 25901426

47. Picardi M, De Renzo A, Pane F, et al. Randomized comparison of consolidation radiation versus observation in bulky Hodgkin's lymphoma with post-chemotherapy negative positron emission tomography scans. *Leuk Lymphoma*. Sep 2007; 48(9): 1721-7. PMID 17786707
48. Raemaekers JM, Andr MP, Federico M, et al. Omitting radiotherapy in early positron emission tomography-negative stage I/II Hodgkin lymphoma is associated with an increased risk of early relapse: Clinical results of the preplanned interim analysis of the randomized EORTC/LYSA/FIL H10 trial. *J Clin Oncol*. Apr 20 2014; 32(12): 1188-94. PMID 24637998
49. Kreuzberger N, Goldkuhle M, von Tresckow B, et al. Positron emission tomography-adapted therapy for first-line treatment in adults with Hodgkin lymphoma. *Cochrane Database Syst Rev*. Mar 26 2025; 3(3): CD010533. PMID 40135712
50. Aldin A, Umlauff L, Estcourt LJ, et al. Interim PET-results for prognosis in adults with Hodgkin lymphoma: a systematic review and meta-analysis of prognostic factor studies. *Cochrane Database Syst Rev*. Jan 13 2020; 1(1): CD012643. PMID 31930780
51. Deniz K, O'Mahony S, Ross G, et al. Breast cancer in women after treatment for Hodgkin's disease. *Lancet Oncol*. Apr 2003; 4(4): 207-14. PMID 12681264
52. Travis LB, Gospodarowicz M, Curtis RE, et al. Lung cancer following chemotherapy and radiotherapy for Hodgkin's disease. *J Natl Cancer Inst*. Feb 06 2002; 94(3): 182-92. PMID 11830608
53. Galper SL, Yu JB, Mauch PM, et al. Clinically significant cardiac disease in patients with Hodgkin lymphoma treated with mediastinal irradiation. *Blood*. Jan 13 2011; 117(2): 412-8. PMID 20858859
54. Swerdlow AJ, Higgins CD, Smith P, et al. Myocardial infarction mortality risk after treatment for Hodgkin disease: a collaborative British cohort study. *J Natl Cancer Inst*. Feb 07 2007; 99(3): 206-14. PMID 17284715
55. Fuchs M, Goergen H, Kobe C, et al. Positron Emission Tomography-Guided Treatment in Early-Stage Favorable Hodgkin Lymphoma: Final Results of the International, Randomized Phase III HD16 Trial by the German Hodgkin Study Group. *J Clin Oncol*. Nov 01 2019; 37(31): 2835-2845. PMID 31498753
56. Borchmann P, Pfltschow A, Kobe C, et al. PET-guided omission of radiotherapy in early-stage unfavourable Hodgkin lymphoma (GHSG HD17): a multicentre, open-label, randomised, phase 3 trial. *Lancet Oncol*. Feb 2021; 22(2): 223-234. PMID 33539742
57. Borchmann P, Goergen H, Kobe C, et al. PET-guided treatment in patients with advanced-stage Hodgkin's lymphoma (HD18): final results of an open-label, international, randomised phase 3 trial by the German Hodgkin Study Group. *Lancet*. Dec 23 2017; 390(10114): 2790-2802. PMID 29061295
58. Gallamini A, Rossi A, Patti C, et al. Consolidation Radiotherapy Could Be Safely Omitted in Advanced Hodgkin Lymphoma With Large Nodal Mass in Complete Metabolic Response After ABVD: Final Analysis of the Randomized GITIL/FIL HD0607 Trial. *J Clin Oncol*. Nov 20 2020; 38(33): 3905-3913. PMID 32946355
59. Ricardi U, Levis M, Evangelista A, et al. Role of radiotherapy to bulky sites of advanced Hodgkin lymphoma treated with ABVD: final results of FIL HD0801 trial. *Blood Adv*. Nov 09 2021; 5(21): 4504-4514. PMID 34597375
60. Casasnovas RO, Bouabdallah R, Brice P, et al. PET-adapted treatment for newly diagnosed advanced Hodgkin lymphoma (AHL2011): a randomised, multicentre, non-inferiority, phase 3 study. *Lancet Oncol*. Feb 2019; 20(2): 202-215. PMID 30658935
61. Johnson P, Federico M, Kirkwood A, et al. Adapted Treatment Guided by Interim PET-CT Scan in Advanced Hodgkin's Lymphoma. *N Engl J Med*. Jun 23 2016; 374(25): 2419-29. PMID 27332902
62. Luminari S, Fossa A, Trotman J, et al. Long-Term Follow-Up of the Response-Adjusted Therapy for Advanced Hodgkin Lymphoma Trial. *J Clin Oncol*. Jan 01 2024; 42(1): 13-18. PMID 37883739
63. Kreissl S, Goergen H, Buehnen I, et al. PET-guided eBEACOPP treatment of advanced-stage Hodgkin lymphoma (HD18): follow-up analysis of an international, open-label, randomised, phase 3 trial. *Lancet Haematol*. Jun 2021; 8(6): e398-e409. PMID 34048679
64. Gallamini A, Tarella C, Viviani S, et al. Early Chemotherapy Intensification With Escalated BEACOPP in Patients With Advanced-Stage Hodgkin Lymphoma With a Positive Interim Positron Emission Tomography/Computed Tomography Scan After Two ABVD Cycles: Long-Term Results of the GITIL/FIL HD 0607 Trial. *J Clin Oncol*. Feb 10 2018; 36(5): 454-462. PMID 29360414
65. Raemaekers JM. Early FDG-PET adapted treatment improved the outcome of early FDG-PET positive patients with stages I/II Hodgkin lymphoma (HL): final results of the randomized Intergroup EORTC/LYSA/FIL H10 trial. Paper presented at: 13th International Conference on Malignant Lymphoma; 2015; Lugano, Switzerland.
66. Andr MPE, Girinsky T, Federico M, et al. Early Positron Emission Tomography Response-Adapted Treatment in Stage I and II Hodgkin Lymphoma: Final Results of the Randomized EORTC/LYSA/FIL H10 Trial. *J Clin Oncol*. Jun 01 2017; 35(16): 1786-1794. PMID 28291393
67. Fonseca JM, Zloic SK, Ayogu CI, et al. Deauville Score-Based Evaluation of Interim PET/CT in Follicular Lymphoma: A Prognostic Factor Systematic Review and Meta-Analysis. *Cureus*. Dec 2024; 16(12): e75169. PMID 39759650
68. Casasnovas RO, Ysebaert L, Thieblemont C, et al. FDG-PET-driven consolidation strategy in diffuse large B-cell lymphoma: final results of a randomized phase 2 study. *Blood*. Sep 14 2017; 130(11): 1315-1326. PMID 28701367
69. Dharsen U, Miller S, Hertenstein B, et al. Positron Emission Tomography-Guided Therapy of Aggressive Non-Hodgkin Lymphomas (PETAL): A Multicenter, Randomized Phase III Trial. *J Clin Oncol*. Jul 10 2018; 36(20): 2024-2034. PMID 29750632
70. Dharsen U, Bockisch A, Hertenstein B, et al. Response-guided first-line therapy and treatment of relapse in aggressive lymphoma: 10-year follow-up of the PETAL trial. *Blood Neoplasia*. Sep 2024; 1(3): 100018. PMID 40453056
71. Kanazu M, Maruyama K, Ando M, et al. Early pharmacodynamic assessment using <sup>18</sup>F-fluorodeoxyglucose positron-emission tomography on molecular targeted therapy and cytotoxic chemotherapy for clinical outcome prediction. *Clin Lung Cancer*. May 2014; 15(3): 182-7. PMID 24518101
72. Stefano A, Russo G, Ippolito M, et al. Evaluation of erlotinib treatment response in non-small cell lung cancer using metabolic and anatomic criteria. *Q J Nucl Med Mol Imaging*. May 09 2014. PMID 24809275
73. Tiseo M, Ippolito M, Scarlattei M, et al. Predictive and prognostic value of early response assessment using <sup>18</sup>FDG-PET in advanced non-small cell lung cancer patients treated with erlotinib. *Cancer Chemother Pharmacol*. Feb 2014; 73(2): 299-307. PMID 24258456

74. Tsuchida T, Morikawa M, Demura Y, et al. Imaging the early response to chemotherapy in advanced lung cancer with diffusion-weighted magnetic resonance imaging compared to fluorine-18 fluorodeoxyglucose positron emission tomography and computed tomography. *J Magn Reson Imaging*. Jul 2013; 38(1): 80-8. PMID 23239463
75. Usmanij EA, de Geus-Oei LF, Troost EG, et al. 18F-FDG PET early response evaluation of locally advanced non-small cell lung cancer treated with concomitant chemoradiotherapy. *J Nucl Med*. Sep 2013; 54(9): 1528-34. PMID 23864719
76. Podoloff DA, Ball DW, Ben-Josef E, et al. NCCN task force: clinical utility of PET in a variety of tumor types. *J Natl Compr Canc Netw*. Jun 2009; 7 Suppl 2: S1-26. PMID 19555588
77. Grootjans W, Usmanij EA, Oyen WJ, et al. Performance of automatic image segmentation algorithms for calculating total lesion glycolysis for early response monitoring in non-small cell lung cancer patients during concomitant chemoradiotherapy. *Radiother Oncol*. Jun 2016; 119(3): 473-9. PMID 27178141
78. Han EJ, Yang YJ, Park JC, et al. Prognostic value of early response assessment using 18F-FDG PET/CT in chemotherapy-treated patients with non-small-cell lung cancer. *Nucl Med Commun*. Dec 2015; 36(12): 1187-94. PMID 26375438
79. Nygrd L, Vogelius IR, Fischer BM, et al. Early lesion-specific (18)F-FDG PET response to chemotherapy predicts time to lesion progression in locally advanced non-small cell lung cancer. *Radiother Oncol*. Mar 2016; 118(3): 460-4. PMID 26806265
80. Mattoli MV, Massacesi M, Castelluccia A, et al. The predictive value of 18 F-FDG PET-CT for assessing the clinical outcomes in locally advanced NSCLC patients after a new induction treatment: low-dose fractionated radiotherapy with concurrent chemotherapy. *Radiat Oncol*. Jan 05 2017; 12(1): 4. PMID 28057034
81. Crandall JP, Tahari AK, Juergens RA, et al. A comparison of FLT to FDG PET/CT in the early assessment of chemotherapy response in stages IB-IIIa resectable NSCLC. *EJNMMI Res*. Dec 2017; 7(1): 8. PMID 28102506
82. Romine PE, Martins RG, Eaton KD, et al. Long term follow-up of neoadjuvant chemotherapy for non-small cell lung cancer (NSCLC) investigating early positron emission tomography (PET) scan as a predictor of outcome. *BMC Cancer*. Jan 14 2019; 19(1): 70. PMID 30642285
83. Suppiah S, Chang WL, Hassan HA, et al. Systematic Review on the Accuracy of Positron Emission Tomography/Computed Tomography and Positron Emission Tomography/Magnetic Resonance Imaging in the Management of Ovarian Cancer: Is Functional Information Really Needed?. *World J Nucl Med*. 2017; 16(3): 176-185. PMID 28670174
84. Ko WS, Kim SJ. Predictive Value of 18 F-FDG PET/CT for Assessment of Tumor Response to Neoadjuvant Chemotherapy in Bladder Cancer. *Clin Nucl Med*. Jul 01 2023; 48(7): 574-580. PMID 36976654
85. Singh S, Poon R, Wong R, et al. 68Ga PET Imaging in Patients With Neuroendocrine Tumors: A Systematic Review and Meta-analysis. *Clin Nucl Med*. Nov 2018; 43(11): 802-810. PMID 30247209
86. Beckers RCJ, Lambregts DMJ, Lahaye MJ, et al. Advanced imaging to predict response to chemotherapy in colorectal liver metastases - a systematic review. *HPB (Oxford)*. Feb 2018; 20(2): 120-127. PMID 29196021
87. Facey K, Bradbury I, Laking G, et al. Overview of the clinical effectiveness of positron emission tomography imaging in selected cancers. *Health Technol Assess*. Oct 2007; 11(44): iii-iv, xi-267. PMID 17999839
88. Engelmann BE, Loft A, Kjr A, et al. Positron emission tomography/computed tomography and biomarkers for early treatment response evaluation in metastatic colon cancer. *Oncologist*. Feb 2014; 19(2): 164-72. PMID 24451199
89. Hong YS, Kim HO, Kim KP, et al. 3'-Deoxy-3'-18F-fluorothymidine PET for the early prediction of response to leucovorin, 5-fluorouracil, and oxaliplatin therapy in patients with metastatic colorectal cancer. *J Nucl Med*. Aug 2013; 54(8): 1209-16. PMID 23804324
90. Li C, Lan X, Yuan H, et al. 18F-FDG PET predicts pathological response to preoperative chemoradiotherapy in patients with primary rectal cancer: a meta-analysis. *Ann Nucl Med*. Jun 2014; 28(5): 436-46. PMID 24623152
91. Memon S, Lynch AC, Akhurst T, et al. Systematic review of FDG-PET prediction of complete pathological response and survival in rectal cancer. *Ann Surg Oncol*. Oct 2014; 21(11): 3598-607. PMID 24802909
92. Formiga MN, Fanelli MF, Dettino AL, et al. Is early response by (18)F-2-fluoro-2-deoxy-D-glucose positron emission tomography-computed tomography a predictor of long-term outcome in patients with metastatic colorectal cancer?. *J Gastrointest Oncol*. Jun 2016; 7(3): 365-72. PMID 27284468
93. Hendlisz A, Deleporte A, Delaunoy T, et al. The Prognostic Significance of Metabolic Response Heterogeneity in Metastatic Colorectal Cancer. *PLoS One*. 2015; 10(9): e0138341. PMID 26421426
94. Kim SJ, Chang S. Volumetric parameters changes of sequential 18F-FDG PET/CT for early prediction of recurrence and death in patients with locally advanced rectal cancer treated with preoperative chemoradiotherapy. *Clin Nucl Med*. Dec 2015; 40(12): 930-5. PMID 26204222
95. Koo PJ, Kim SJ, Chang S, et al. Interim Fluorine-18 Fluorodeoxyglucose Positron Emission Tomography/Computed Tomography to Predict Pathologic Response to Preoperative Chemoradiotherapy and Prognosis in Patients With Locally Advanced Rectal Cancer. *Clin Colorectal Cancer*. Dec 2016; 15(4): e213-e219. PMID 27316919
96. Garca Vicente AM, Soriano Castrejn A, Len Martn A, et al. Early and delayed prediction of axillary lymph node neoadjuvant response by (18)F-FDG PET/CT in patients with locally advanced breast cancer. *Eur J Nucl Med Mol Imaging*. Jul 2014; 41(7): 1309-18. PMID 24744045
97. Koolen BB, Valds Olmos RA, Wesseling J, et al. Early assessment of axillary response with <sup>18</sup>F-FDG PET/CT during neoadjuvant chemotherapy in stage II-III breast cancer: implications for surgical management of the axilla. *Ann Surg Oncol*. Jul 2013; 20(7): 2227-35. PMID 23456316
98. Giannatempo P, Alessi A, Miceli R, et al. Interim fluorine-18 fluorodeoxyglucose positron emission tomography for early metabolic assessment of therapeutic response to chemotherapy for metastatic transitional cell carcinoma. *Clin Genitourin Cancer*. Dec 2014; 12(6): 433-9. PMID 24787972
99. Truong MT, Viswanathan C, Godoy MB, et al. Malignant pleural mesothelioma: role of CT, MRI, and PET/CT in staging evaluation and treatment considerations. *Semin Roentgenol*. Oct 2013; 48(4): 323-34. PMID 24034264
100. Francis RJ, Byrne MJ, van der Schaaf AA, et al. Early prediction of response to chemotherapy and survival in malignant pleural mesothelioma using a novel semiautomated 3-dimensional volume-based analysis of serial 18F-FDG PET scans. *J Nucl Med*. Sep 2007; 48(9): 1449-58. PMID 17704250

101. Bhatnagar P, Subesinghe M, Patel C, et al. Functional imaging for radiation treatment planning, response assessment, and adaptive therapy in head and neck cancer. *Radiographics*. 2013; 33(7): 1909-29. PMID 24224586
102. Hoang JK, Das SK, Choudhury KR, et al. Using FDG-PET to measure early treatment response in head and neck squamous cell carcinoma: quantifying intrinsic variability in order to understand treatment-induced change. *AJNR Am J Neuroradiol*. Jul 2013; 34(7): 1428-33. PMID 23391836
103. Lalami Y, Garcia C, Flamen P, et al. Phase II trial evaluating the efficacy of sorafenib (BAY 43-9006) and correlating early fluorodeoxyglucose positron emission tomography-CT response to outcome in patients with recurrent and/or metastatic head and neck cancer. *Head Neck*. Mar 2016; 38(3): 347-54. PMID 25332069
104. Wong KH, Panek R, Welsh L, et al. The Predictive Value of Early Assessment After 1 Cycle of Induction Chemotherapy with 18F-FDG PET/CT and Diffusion-Weighted MRI for Response to Radical Chemoradiotherapy in Head and Neck Squamous Cell Carcinoma. *J Nucl Med*. Dec 2016; 57(12): 1843-1850. PMID 27417648
105. Wilson JM, Mukherjee S, Brunner TB, et al. Correlation of 18 F-Fluorodeoxyglucose Positron Emission Tomography Parameters with Patterns of Disease Progression in Locally Advanced Pancreatic Cancer after Definitive Chemoradiotherapy. *Clin Oncol (R Coll Radiol)*. Jun 2017; 29(6): 370-377. PMID 28190636
106. Evangelista L, Zucchetta P, Moletta L, et al. The role of FDG PET/CT or PET/MRI in assessing response to neoadjuvant therapy for patients with borderline or resectable pancreatic cancer: a systematic literature review. *Ann Nucl Med*. Jul 2021; 35(7): 767-776. PMID 34047926
107. Eary JF, Conrad EU, O'Sullivan J, et al. Sarcoma mid-therapy [F-18]fluorodeoxyglucose positron emission tomography (FDG PET) and patient outcome. *J Bone Joint Surg Am*. Jan 15 2014; 96(2): 152-8. PMID 24430415
108. Hyun O J, Lubner BS, Leal JP, et al. Response to Early Treatment Evaluated with 18F-FDG PET and PERCIST 1.0 Predicts Survival in Patients with Ewing Sarcoma Family of Tumors Treated with a Monoclonal Antibody to the Insulinlike Growth Factor 1 Receptor. *J Nucl Med*. May 2016; 57(5): 735-40. PMID 26795289
109. Farnebo J, Grybck P, Harmenberg U, et al. Volumetric FDG-PET predicts overall and progression-free survival after 14 days of targeted therapy in metastatic renal cell carcinoma. *BMC Cancer*. Jun 06 2014; 14: 408. PMID 24906441
110. Chen JL, Appelbaum DE, Kocherginsky M, et al. FDG-PET as a predictive biomarker for therapy with everolimus in metastatic renal cell cancer. *Cancer Med*. Aug 2013; 2(4): 545-52. PMID 24156027
111. Gilles R, de Geus-Oei LF, Mulders PF, et al. Immunotherapy response evaluation with (18)F-FDG-PET in patients with advanced stage renal cell carcinoma. *World J Urol*. Aug 2013; 31(4): 841-6. PMID 21739122
112. Horn KP, Yap JT, Agarwal N, et al. FDG and FLT-PET for Early measurement of response to 37.5 mg daily sunitinib therapy in metastatic renal cell carcinoma. *Cancer Imaging*. Sep 03 2015; 15(1): 15. PMID 26335224
113. American College of Radiology (ACR)-ACNM-SNMMI-SPR. ACR-ACNM-SNMMI-SPR practice parameter for performing FDG-PET/CT in oncology, amended 2023. <https://gravitas.acr.org/PPTS/GetDocumentView?docId=173>. Accessed September 16, 2025.
114. Delgado Bolton RC, Aide N, Colletti PM, et al. EANM guideline on the role of 2-[ 18 F]FDG PET/CT in diagnosis, staging, prognostic value, therapy assessment and restaging of ovarian cancer, endorsed by the American College of Nuclear Medicine (ACNM), the Society of Nuclear Medicine and Molecular Imaging (SNMMI) and the International Atomic Energy Agency (IAEA). *Eur J Nucl Med Mol Imaging*. Sep 2021; 48(10): 3286-3302. PMID 34215923
115. Nanni C, Deroose CM, Balogova S, et al. EANM guidelines on the use of [ 18 F]FDG PET/CT in diagnosis, staging, prognostication, therapy assessment, and restaging of plasma cell disorders. *Eur J Nucl Med Mol Imaging*. Dec 2024; 52(1): 171-192. PMID 39207486
116. Vaz SC, Woll JPP, Cardoso F, et al. Joint EANM-SNMMI guideline on the role of 2-[ 18 F]FDG PET/CT in no special type breast cancer : (endorsed by the ACR, ESSO, ESTRO, EUSOBI/ESR, and EUSOMA). *Eur J Nucl Med Mol Imaging*. Jul 2024; 51(9): 2706-2732. PMID 38740576
117. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Bladder Cancer. Version 1.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/bladder.pdf](https://www.nccn.org/professionals/physician_gls/pdf/bladder.pdf). Accessed September 13, 2025.
118. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Breast Cancer. Version 4.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/breast.pdf](https://www.nccn.org/professionals/physician_gls/pdf/breast.pdf). Accessed September 16, 2025.
119. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Central Nervous System Cancers. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/cns.pdf](https://www.nccn.org/professionals/physician_gls/pdf/cns.pdf). Accessed September 12, 2025.
120. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Cervical Cancer. Version 4.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/cervical.pdf](https://www.nccn.org/professionals/physician_gls/pdf/cervical.pdf). Accessed September 11, 2025.
121. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Colon Cancer. Version 4.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/colon.pdf](https://www.nccn.org/professionals/physician_gls/pdf/colon.pdf). Accessed September 10, 2025.
122. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Esophageal and Esophagogastric Junction Cancers. Version 4.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/esophageal.pdf](https://www.nccn.org/professionals/physician_gls/pdf/esophageal.pdf). Accessed September 8, 2025.
123. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Soft Tissue Sarcoma. Version 1.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/sarcoma.pdf](https://www.nccn.org/professionals/physician_gls/pdf/sarcoma.pdf). Accessed August 27, 2025.
124. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Head and Neck Cancers. Version 5.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/head-and-neck.pdf](https://www.nccn.org/professionals/physician_gls/pdf/head-and-neck.pdf). Accessed September 7, 2025.
125. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Hepatocellular Carcinoma. Version 1.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/hcc.pdf](https://www.nccn.org/professionals/physician_gls/pdf/hcc.pdf). Accessed September 6, 2025.
126. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Biliary Tract Cancers. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/btc.pdf](https://www.nccn.org/professionals/physician_gls/pdf/btc.pdf). Accessed September 14, 2025.
127. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Hodgkin Lymphoma. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/hodgkins.pdf](https://www.nccn.org/professionals/physician_gls/pdf/hodgkins.pdf). Accessed September 5, 2025.

128. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Cutaneous Melanoma. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/cutaneous\\_melanoma.pdf](https://www.nccn.org/professionals/physician_gls/pdf/cutaneous_melanoma.pdf). Accessed September 9, 2025.
129. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Malignant Pleural Mesothelioma. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/meso\\_pleural.pdf](https://www.nccn.org/professionals/physician_gls/pdf/meso_pleural.pdf). Accessed September 17, 2025.
130. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: B-Cell Lymphomas. Version 3.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/b-cell.pdf](https://www.nccn.org/professionals/physician_gls/pdf/b-cell.pdf). Accessed September 15, 2025.
131. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: T-Cell Lymphomas. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/t-cell.pdf](https://www.nccn.org/professionals/physician_gls/pdf/t-cell.pdf). Accessed August 26, 2025.
132. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Primary Cutaneous Lymphomas. Version 3.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/primary\\_cutaneous.pdf](https://www.nccn.org/professionals/physician_gls/pdf/primary_cutaneous.pdf). Accessed August 31, 2025.
133. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Non-Small Cell Lung Cancer. Version 8.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/nscl.pdf](https://www.nccn.org/professionals/physician_gls/pdf/nscl.pdf). Accessed September 3, 2025.
134. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Ovarian Cancer. Version 3.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/ovarian.pdf](https://www.nccn.org/professionals/physician_gls/pdf/ovarian.pdf). Accessed September 2, 2025.
135. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Pancreatic Adenocarcinoma. Version 2.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/pancreatic.pdf](https://www.nccn.org/professionals/physician_gls/pdf/pancreatic.pdf). Accessed September 1, 2025.
136. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Prostate Cancer. Version 2.2026. [https://www.nccn.org/professionals/physician\\_gls/pdf/prostate.pdf](https://www.nccn.org/professionals/physician_gls/pdf/prostate.pdf). Accessed August 30, 2025.
137. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Rectal Cancer. Version 3.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/rectal.pdf](https://www.nccn.org/professionals/physician_gls/pdf/rectal.pdf). Accessed August 29, 2025.
138. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Small Cell Lung Cancer. Version 2.2026. [https://www.nccn.org/professionals/physician\\_gls/pdf/scl.pdf](https://www.nccn.org/professionals/physician_gls/pdf/scl.pdf). Accessed August 28, 2025.
139. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Thyroid Carcinoma. Version 1.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/thyroid.pdf](https://www.nccn.org/professionals/physician_gls/pdf/thyroid.pdf). Accessed August 25, 2025.
140. National Comprehensive Cancer Network (NCCN). NCCN Clinical Practice Guidelines in Oncology: Uterine Neoplasms. Version 3.2025. [https://www.nccn.org/professionals/physician\\_gls/pdf/uterine.pdf](https://www.nccn.org/professionals/physician_gls/pdf/uterine.pdf). Accessed August 24, 2025.
141. Centers for Medicare & Medicaid Services. National Coverage Determination (NCD) for Positron Emission Tomography (FDG) for Oncologic Conditions (220.6.17). 2014; <https://www.cms.gov/medicare-coverage-database/details/ncd-details.aspx?ncdid=331>. Accessed September 16, 2025.

## POLICY HISTORY - THIS POLICY WAS APPROVED BY THE FEP® PHARMACY AND MEDICAL POLICY COMMITTEE ACCORDING TO THE HISTORY BELOW:

Date	Action	Description
June 2012	New policy	
December 2013	Replace policy	Policy updated with literature review. References 1-3, 5-11, 18, 20, 21, and 23 added. Others removed or renumbered. No change in policy statement.
December 2014	Replace policy	Policy updated with literature search; adding references 3, 9-12, 17-25, and 27-67; updating references 13, 16, and 68; references 1-11 (trial registrations) deleted. No change to policy statements. Title revised, added "Interim,,"
December 2015	Replace policy	Policy updated with literature review through July 8, 2015; references 13- 16, 18-9, 25, and 27 (NCCN) deleted; references 3-5, 24, and 36 added; reference 58 updated. Policy statement unchanged.
December 2016	Replace policy	Policy updated with literature review; references 1-2, 5-10, 16-19, 23, 28- 31, 42-44, 55-60, 68-69, 71, 74, and 78 were added. Policy statement unchanged.
December 2017	Replace policy	Policy updated with literature review through July 21, 2017; references 5, 19, 20, 24-27, 30-34, 44-56, 66-68, 71, 84, and 94-95 were added. The following policy statement was added: The use of interim positron emission tomography scans to determine response to tyrosine kinase inhibitor treatment in patients with gastrointestinal stromal tumors is considered medically necessary.
December 2018	Replace policy	Policy updated with literature review through July 26, 2018; references 5, 7, 19-21, 26, 31, 34, 39, 80, 134 added; references 38 and 111-133 updated. Policy statement unchanged.

The policies contained in the FEP Medical Policy Manual are developed to assist in administering contractual benefits and do not constitute medical advice. They are not intended to replace or substitute for the independent medical judgment of a practitioner or other health care professional in the treatment of an individual member. The Blue Cross and Blue Shield Association does not intend by the FEP Medical Policy Manual, or by any particular medical policy, to recommend, advocate, encourage or discourage any particular medical technologies. Medical decisions relative to medical technologies are to be made strictly by members/patients in consultation with their health care providers. The conclusion that a particular service or supply is medically necessary does not constitute a representation or warranty that the Blue Cross and Blue Shield Service Benefit Plan covers (or pays for) this service or supply for a particular member.

Date	Action	Description
December 2019	Replace policy	Policy updated with literature review through July 8, 2019; references added, references on NCCN updated. Policy statements unchanged.
December 2020	Replace policy	Policy updated with literature review through July 30, 2020; references added. Policy statements unchanged.
December 2021	Replace policy	Policy updated with literature review through August 6, 2021; references added. Policy statements unchanged.
December 2022	Replace policy	Policy updated with literature review through August 1, 2022; reference added. Minor editorial refinements to policy statements; intent unchanged.
December 2023	Replace policy	Policy updated with literature review through August 2, 2023; references added. Policy statements unchanged.
December 2024	Replace policy	Policy updated with literature review through July 23, 2024; references added. Policy statements unchanged.
March 2026	Replace policy	Policy updated with literature review through September 17, 2025; references added. Split lymphoma PICO to 2 PICOs - Hodgkin and non-Hodgkin. Medically necessary policy statement added for advanced Hodgkin lymphoma.

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