American College of Radiology ACR Appropriateness Criteria[®] Post-Treatment Follow-up and Active Surveillance of Clinically Localized Renal Cell Carcinoma

Va	riant	1.

Follow-up for clinically localized renal cell carcinoma; post radical or partial nephrectomy.

Procedure	Appropriateness Category	Relative Radiation Level
CT abdomen with IV contrast	Usually Appropriate	€€
MRI abdomen without and with IV contrast	Usually Appropriate	0
CT abdomen without and with IV contrast	Usually Appropriate	���
CT abdomen and pelvis with IV contrast	May Be Appropriate	���
CT abdomen and pelvis without and with IV contrast	May Be Appropriate	€€€
CT chest with IV contrast	May Be Appropriate	${\mathfrak G}{\mathfrak G}{\mathfrak G}$
MRI abdomen without IV contrast	May Be Appropriate	0
Radiography chest	May Be Appropriate	۲
US abdomen with IV contrast	May Be Appropriate	0
CT abdomen without IV contrast	May Be Appropriate	${\bf \mathfrak{D}}{\bf \mathfrak{D}}{\bf \mathfrak{D}}$
CT chest without IV contrast	May Be Appropriate	���
MRI abdomen and pelvis without and with IV contrast	May Be Appropriate (Disagreement)	0
MRI abdomen and pelvis without IV contrast	May Be Appropriate	0
CT abdomen and pelvis without IV contrast	May Be Appropriate	${\mathfrak G}{\mathfrak G}{\mathfrak G}$
FDG-PET/CT skull base to mid-thigh	May Be Appropriate	$\textcircled{\begin{tabular}{lllllllllllllllllllllllllllllllllll$
US kidney retroperitoneal	May Be Appropriate	0
CT chest without and with IV contrast	Usually Not Appropriate	$\textcircled{\black}{\textcircled{\black}{\black}}$
Bone scan whole body	Usually Not Appropriate	ଡ଼ଡ଼ଡ଼
CT head with IV contrast	Usually Not Appropriate	���
CT head without and with IV contrast	Usually Not Appropriate	***
CT head without IV contrast	Usually Not Appropriate	���
CTU without and with IV contrast	Usually Not Appropriate	€€€€
Radiography intravenous urography	Usually Not Appropriate	���
MRI head without and with IV contrast	Usually Not Appropriate	0
MRI head without IV contrast	Usually Not Appropriate	0
MRU without and with IV contrast	Usually Not Appropriate	0
MRU without IV contrast	Usually Not Appropriate	0
Radiography abdomen	Usually Not Appropriate	�€
Radiography skeletal survey	Usually Not Appropriate	€€€

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Variant 2:

Follow-up for clinically localized renal cell carcinoma; post ablation.

Procedure	Appropriateness Category	Relative Radiation Level	
CT abdomen without and with IV contrast	Usually Appropriate	€€	
MRI abdomen without and with IV contrast	Usually Appropriate	0	
CT abdomen with IV contrast	Usually Appropriate	€€	
CT abdomen and pelvis without and with IV contrast	May Be Appropriate	€€€	
CT chest with IV contrast	May Be Appropriate	€€€	
MRI abdomen and pelvis without and with IV contrast	May Be Appropriate	0	
MRI abdomen without IV contrast	May Be Appropriate	0	
Radiography chest	May Be Appropriate	•	
US abdomen with IV contrast	May Be Appropriate	0	
CT abdomen and pelvis with IV contrast	May Be Appropriate	♥♥♥	
CT chest without IV contrast	May Be Appropriate	♥♥♥	
MRI abdomen and pelvis without IV contrast	May Be Appropriate	0	
CT abdomen and pelvis without IV contrast	May Be Appropriate	♥♥♥	
CT abdomen without IV contrast	May Be Appropriate	♥♥♥	
US kidney retroperitoneal	May Be Appropriate	0	
CT chest without and with IV contrast	Usually Not Appropriate	€€	
FDG-PET/CT skull base to mid-thigh	Usually Not Appropriate	€€€€	
Bone scan whole body	Usually Not Appropriate	♥♥♥	
CT head with IV contrast	Usually Not Appropriate	���	
CT head without and with IV contrast	Usually Not Appropriate	���	
CT head without IV contrast	Usually Not Appropriate	���	
CTU without and with IV contrast	Usually Not Appropriate	€€€€	
Radiography intravenous urography	Usually Not Appropriate	€€	
MRI head without and with IV contrast	Usually Not Appropriate	0	
MRI head without IV contrast	Usually Not Appropriate	0	
MRU without and with IV contrast	Usually Not Appropriate	0	
MRU without IV contrast	Usually Not Appropriate	0	
Radiography abdomen	Usually Not Appropriate	*	
Radiography skeletal survey	Usually Not Appropriate	€€	

Variant 3:

Follow-up for clinically localized renal cell carcinoma; active surveillance.

Procedure	Appropriateness Category	Relative Radiation Level	
MRI abdomen without and with IV contrast	Usually Appropriate	0	
CT abdomen with IV contrast	Usually Appropriate	♥♥♥	
CT abdomen without and with IV contrast	Usually Appropriate	₸₽₽₽	
US abdomen with IV contrast	Usually Appropriate	0	
CT abdomen and pelvis with IV contrast	May Be Appropriate	♥♥♥	
CT abdomen and pelvis without and with IV contrast	May Be Appropriate	€€€	
MRI abdomen and pelvis without and with IV contrast	May Be Appropriate	0	
MRI abdomen without IV contrast	May Be Appropriate	0	
US kidney retroperitoneal	May Be Appropriate	0	
CT abdomen without IV contrast	May Be Appropriate	***	
CT chest with IV contrast	May Be Appropriate	$\textcircled{\black}{\includegraphics{\black}{\textcircled{\black}{\includegraphics{\black}{\textcircled{\black}{\includegraphics{\black}{\textcircled{\black}{\includegraphics{\black}{\b$	
CT chest without IV contrast	May Be Appropriate	♥♥♥	
MRI abdomen and pelvis without IV contrast	May Be Appropriate	0	
Radiography chest	May Be Appropriate	۲	
CT abdomen and pelvis without IV contrast	May Be Appropriate	♥♥♥	
CT chest without and with IV contrast	Usually Not Appropriate	♥♥♥	
FDG-PET/CT skull base to mid-thigh	Usually Not Appropriate	$\mathbf{\hat{\mathbf{x}}}\mathbf{\hat{\mathbf{x}}}\mathbf{\hat{\mathbf{x}}}$	
Bone scan whole body	Usually Not Appropriate	♥♥♥	
CT head with IV contrast	Usually Not Appropriate	♥♥♥	
CT head without and with IV contrast	Usually Not Appropriate	♥♥♥	
CT head without IV contrast	Usually Not Appropriate	♥♥♥	
CTU without and with IV contrast	Usually Not Appropriate	♦€€€	
Radiography intravenous urography	Usually Not Appropriate	₸₽₽₽	
MRI head without and with IV contrast	Usually Not Appropriate	0	
MRI head without IV contrast	Usually Not Appropriate	0	
MRU without and with IV contrast	Usually Not Appropriate	0	
MRU without IV contrast	Usually Not Appropriate	0	
Radiography abdomen	Usually Not Appropriate	•••	
Radiography skeletal survey	Usually Not Appropriate	♦♥♥	

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POST-TREATMENT FOLLOW-UP AND ACTIVE SURVEILLANCE OF CLINICALLY LOCALIZED RENAL CELL CARCINOMA

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Summary of Literature Review

Introduction/Background

According to the American Cancer Society, approximately 73,750 new cases of kidney and renal pelvis cancer will be diagnosed in the United States in 2020, and approximately 14,830 people will die of this disease [1]. Renal cell carcinoma (RCC) accounts for 85% of all malignant renal tumors and represents approximately 2% to 3% of all malignancies in adults [2]. RCC is also considered the most lethal of all urologic cancers.

Surgical resection with curative intent, including radical nephrectomy (RN) or partial nephrectomy (PN), continues to be the standard of care for clinically localized RCC [2]. Ablative therapies such as radiofrequency ablation, microwave ablation, and cryoablation have been shown to be effective and safe alternatives for the treatment of small localized RCCs [3-7]. In some patients with small localized RCCs, treatment may also be deferred, with management instead consisting of active surveillance protocols [8].

For follow-up of patients with treated or untreated RCC and those with neoplasms suspected to represent RCC, radiologic imaging is the most useful component of surveillance, because most relapses and cases of disease progression are identified when patients are asymptomatic [9,10]. There is currently no consensus regarding surveillance protocols; however, various guidelines and strategies have been developed by international oncologic and urologic societies, such as the National Comprehensive Cancer Network, the American Urological Association, and the European Association of Urology, using both patient- and tumor-specific characteristics [2,9,11,12]. Although imaging is the centerpiece in all of these guidelines, the recommendations vary regarding the timing, frequency, and duration of follow-up, as well as the selection of imaging modalities for follow-up [12,13]. Understanding the strengths and limitations of the various imaging modalities for the detection of disease recurrence or progression is important when planning follow-up regimens.

In this document, we provide an update on the appropriate use of imaging examinations for asymptomatic patients who have been treated for RCC with RN, PN, or ablative therapies. We also address the appropriate imaging examinations for asymptomatic patients with localized biopsy-proven or suspected RCC who are undergoing active surveillance. As in the previous version, this document does not address the imaging of complications from treatment and does not discuss the follow-up of patients with known residual or recurrent cancer.

Special Imaging Considerations

CT urography (CTU) is an imaging study that is tailored to improve visualization of both the upper and lower urinary tracts. There is variability in the specific parameters, but it usually involves unenhanced images followed by intravenous (IV) contrast-enhanced images, including nephrographic and excretory phases acquired at least 5 minutes after contrast injection. Alternatively, a split-bolus technique uses an initial loading dose of IV contrast and then obtains a combined nephrographic-excretory phase after a second IV contrast dose; some sites include arterial phase. CTU should use thin-slice acquisition. Reconstruction methods commonly include maximum intensity projection or 3-D volume rendering. For the purposes of this document, we make a distinction between CTU and CT abdomen and pelvis without and with IV contrast. CT abdomen and pelvis without and with IV contrast.

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defined as any protocol not specifically tailored for evaluation of the upper and lower urinary tracts and without both the precontrast and excretory phases.

MR urography (MRU) is also tailored to improve imaging of the urinary system. Unenhanced MRU relies upon heavily T2-weighted imaging of the intrinsic high signal intensity from urine for evaluation of the urinary tract. IV contrast is administered to provide additional information regarding obstruction, urothelial thickening, focal lesions, and stones. A contrast-enhanced T1-weighted series should include a corticomedullary, nephrographic, and excretory phase. Thin-slice acquisition and multiplanar imaging should be obtained. For the purposes of this document, we make a distinction between MRU and MRI abdomen and pelvis without and with IV contrast. MRI abdomen and pelvis without and with IV contrast is defined as any protocol not specifically tailored for evaluation of the upper and lower urinary tracts, without both the precontrast and excretory phases, and without heavily T2-weighted images of the urinary tract.

Discussion of Procedures by Variant

Variant 1: Follow-up for clinically localized renal cell carcinoma; post radical or partial nephrectomy.

Many tumor- and patient-specific characteristics have been shown to be predictive of local recurrence or distant metastasis of RCC after treatment [2,14-19]. In addition to these characteristics, the timing and location of tumor recurrence and the type of treatment (ie, RN versus PN) are considered in the development of imaging surveillance strategies that aim to identify asymptomatic solitary or oligometastatic disease that may benefit from early potentially curative or life-prolonging salvage treatment [10,16,20].

Among the tumor characteristics predictive of tumor recurrence, the tumor, node, and metastases staging system has been the most extensively researched; associations between pathologic T stage and both the risk and patterns of tumor recurrence have been demonstrated in many studies [14,15,17,21]. Patient symptoms, tumor size, tumor necrosis, and microvascular invasion are some of the other factors that have been evaluated and integrated into risk stratification models that separate patients into low-, intermediate-, or high-risk groups according to the probability of local recurrence or distant metastases [11,14,18,19]. Most recurrences occur within 3 years after treatment, with a median time to relapse of 1 to 2 years; thus, most surveillance guidelines address follow-up for up to 5 years after treatment [2,9,11,22]. In patients with high-risk tumors (ie, pT2 and pT3 tumors), especially those patients without a significant competing risk for non-RCC death, follow-up beyond 5 years may also be considered because of a nonnegligible incidence of late recurrence [14,18].

Patients who have undergone PN have a similar or slightly higher incidence of local recurrence compared with those who have undergone RN [11,23]. In some guidelines, a more rigorous follow-up protocol is recommended to assess for local recurrence in those who have undergone PN [2,9]. However, more commonly, recurrence manifests as distant metastases [10,20,24,25]. The lungs are the most common site affected by metastases, followed by the lymph nodes, bones, liver, adrenal glands, and brain. Other less common sites include the spleen, pancreas, diaphragm, heart, skin, and connective tissues. Apart from bone and brain metastases, most metastases and local recurrences are identified in asymptomatic patients [10,15,18,26].

In addition to a detailed clinical history, a thorough physical examination and laboratory workup are needed, and guidelines from major international urological and oncological societies recommend that imaging surveillance of the chest and abdomen be performed after primary treatment for RCC [2,9,11]. Because of the lack of high-level evidence assessing the various surveillance protocols, these guidelines vary in the recommended imaging modalities and timelines. For the chest, both radiographs and CT are used. For the abdomen, CT and MRI are more frequently used than ultrasound (US). In general, more frequent follow-up is performed for the surveillance of intermediate-and high-risk tumors than for tumors with a low risk [12,13]. These posttreatment follow-up strategies can be summarized as follows:

- For low-risk/T1 tumors:
 - Chest imaging: Every 12 to 24 months for 3 to 5 years
 - Abdominal imaging: Some recommend performing a baseline study between 3 and 12 months, especially after PN, then yearly for 3 to 5 years
- For intermediate-risk/T2 primary tumors:
 - Chest and abdominal imaging: Some recommend performing a baseline study at 3 months, then at 6 and 12 months, followed by every 6 to 24 months for 5 years

- For high-risk/T3 tumors:
 - Chest and abdominal imaging: Some recommend performing a baseline study at 3 months, then at 6 and 12 months, followed by every 6 to 12 months for 5 years

Radiography Chest

A chest radiograph is a low-yield diagnostic tool for detecting pulmonary metastasis in patients after surgical excision of RCC, particularly in those with low-risk tumors, irrespective of the treatment modality (RN, PN, or ablation) [27,28]. In a retrospective analysis of 221 patients with pT1-3N0M0 RCC, only 0.85% of the follow-up chest radiographs detected pulmonary metastases in asymptomatic patients [28]. The yield of a chest radiograph increased to 1.9% when used in patients with intermediate-risk (T2) or high-risk (T3) tumors. In more than half of the patients, pulmonary metastases were detected when patients became symptomatic outside of the routine followup. In a second retrospective analysis of 258 patients who had undergone resection or ablation for low-risk (T1a) RCC, pulmonary metastases developed in 3 patients (1.2%), but in only 1 patient (0.4%) was this metastasis diagnosed with surveillance chest radiographs [27]. In a more recent study, only 2 of 384 patients (0.005%) with T1a RCC were found to have pulmonary metastases after surgical therapy, and in both cases, the pulmonary metastases were not detected by surveillance chest radiographs [24]. In the same study, 10 of 184 patients (5.4%) with T1b RCC had suspicious pulmonary lesions found on surveillance radiography of the chest; only 2 of these patients had biopsy-confirmed pulmonary metastasis. However, according to guidelines from urologic and oncologic societies, a chest radiograph is still the recommended technique for the surveillance of patients with T1a tumors, and this technique is also recommended as an alternative to chest CT for the surveillance of patients with T2 and T3 tumors after an initial negative follow-up chest CT examination [9,11]. This is in part because of concerns about potential false-positive findings with chest CT (ie, intrapulmonary lymph nodes and granulomas) that can lead to further unnecessary and potentially invasive investigations [9,12].

Radiography Abdomen

There is no relevant literature regarding the use of abdominal radiographs in the follow-up of patients after surgical excision of RCC, and this method is not recommended by the guidelines [2,9,11].

Radiography Skeletal Survey

There is no relevant literature regarding the use of a radiographic survey of the whole body in the follow-up of patients after surgical excision of RCC, and this method is not included in the guidelines [2,9,11].

Radiography Intravenous Urography

There is no relevant literature regarding the use of IV urography in the follow-up of patients after surgical excision of RCC, and this method is not recommended by the guidelines [2,9,11].

CT Abdomen

CT of the abdomen is the most commonly used method for surveillance after surgical excision of RCC [29]. CT is a sensitive method for the detection of recurrences in the resection bed and in other more common sites of metastases in the abdomen, such as the contralateral kidney, adrenal glands, liver, and lymph nodes, and in the visualized bones [16,17,20,22,30]. Although several studies have advised against routine imaging of the abdomen after resection of low-risk (T1) tumors because of the low frequency of abdominal recurrences [15,17,21,30], CT of the abdomen is commonly performed in this group, particularly after PN, to serve as a baseline for future comparisons and to evaluate postoperative complications [9]. Although CT of the abdomen performed without and with IV contrast may be considered beneficial in cases in which postoperative changes need to be distinguished from residual or recurrent tumors, in general, surveillance protocols in oncology often use a single-phase examination in the portal-venous phase. Because RCC metastases tend to be hypervascular, some authors have also suggested that arterial phase imaging can be used to complement portal-venous imaging for the detection of RCC metastases to the liver, pancreas, or contralateral kidney detected only in the arterial phase, and these findings led to a change in management for 2 patients [31]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen without IV contrast may be considered appropriate.

CT Abdomen and Pelvis

CT of the abdomen is the most commonly used method for surveillance after surgical excision of RCC [29]. CT is a sensitive method for the detection of recurrences in the resection bed and in other more common sites of metastases in the abdomen, such as the contralateral kidney, adrenal glands, liver, and lymph nodes, and in the visualized bones [16,17,20,22,30]. Although several studies have advised against routine imaging of the abdomen after resection of

low-risk (T1) tumors because of the low frequency of abdominal recurrences [15,17,21,30], CT of the abdomen is commonly performed in this group, particularly after PN, to serve as a baseline for future comparisons and to evaluate postoperative complications [9]. Although CT of the abdomen performed without and with IV contrast may be considered beneficial in cases in which postoperative changes need to be distinguished from residual or recurrent tumors, in general, surveillance protocols in oncology often use a single-phase examination in the portal-venous phase. Because RCC metastases tend to be hypervascular, some authors have also suggested that arterial phase imaging can be used to complement portal-venous imaging for the detection of RCC metastases to the liver, pancreas, and contralateral kidney. In a retrospective study including 100 patients, 9 patients had metastases in the liver, pancreas, or contralateral kidney detected only in the arterial phase, and these findings led to a change in management for 2 patients [31]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen true to considered appropriate.

Imaging of the pelvis during surveillance after RCC treatment is considered optional in the guidelines [2,9,11]. Although CT of the pelvis is commonly performed in conjunction with CT of the abdomen, data from 2 retrospective studies suggest that CT of the pelvis has minimal value in this setting. In a study of 493 patients with stages T1 to T3a RCC who underwent RN or PN, 82 patients (16.6%) experienced recurrence, and 78 of these cases (95%) were detected by CT of the chest and abdomen [32]. Limiting the study field to the chest and upper abdomen (to the level of the L3–L4 disc) decreased the sensitivity of the study by only 1% because only 1 case of iliac bone metastasis (which was symptomatic) would have been missed with this protocol [32]. In a second study that included 603 patients with T2 to T4 RCC treated with RN or PN, recurrent or metastatic disease occurred in 163 patients (27%), but pelvic imaging was negative in 97% of the patients [25]. Only 4 patients (0.7%) had positive findings in the pelvis and negative findings in the chest and abdomen, and of these patients, only 2 (0.3%) were asymptomatic [25]. These findings are in line with the results of previous studies, which also demonstrated that CT of the pelvis had limited benefit for the detection of metastases in the initial staging of RCC [33,34].

CTU

There is no relevant literature suggesting that CTU offers any additional benefit over conventional CT of the abdomen in the surveillance of patients after treatment of localized RCC, and this method is not included in the guidelines [2,9,11]. In a retrospective analysis of 23 tumors that progressed locally after ablation, CT or MRIs obtained in the corticomedullary phase were found to be sufficient for diagnosis of recurrence in 100% of the cases; noncontrast, nephrographic, and excretory-phase images, which are typically obtained in a CTU or MRU protocol, were able to detect recurrence in only 11%, 81%, and 44% of cases, respectively [35].

CT Chest

Limited data suggest that CT is more sensitive than radiography for the detection of pulmonary metastases from RCC during staging [27]. Although no direct comparison between the 2 methods has been reported in the posttreatment surveillance setting, one study demonstrated that the overwhelming majority of chest recurrences in asymptomatic cases are detected by chest CT examinations (92.3%) rather than by radiography (7.7%) [36]. In addition to a high sensitivity for the detection of pulmonary metastases, chest CT has a high sensitivity for the detection of pulmonary metastases, chest CT has a high sensitivity for the detection of intrathoracic nodal metastases from RCC; this finding has prognostic implications and may affect surgical planning for metastases resection [37]. The use of IV contrast is optional for chest CT, but it may be beneficial for the detection and characterization of hilar lymph nodes. In patients undergoing surveillance with CT of the abdomen with IV contrast, chest CT should also be performed after IV contrast administration.

Unlike CT of the abdomen, in which images obtained without and with IV contrast may be appropriate in some circumstances, CT of the chest without and with IV contrast does not provide additional information in these patients and is considered inappropriate. Although some consider CT to be the standard chest imaging technique for surveillance after RCC resection [11], there are concerns regarding the risk of false-positive findings (ie, intrapulmonary lymph nodes and granulomas), particularly in patients with T1a RCC, which can lead to further unnecessary and potentially invasive investigations [9]. It is worth noting that in a recent pilot study, the authors suggested that CT of the chest may not be necessary to identify most cases of pulmonary recurrence when a CT examination of the abdomen with coverage of the lung bases to the T7 thoracic level is performed [26].

CT Head

Most patients with metastases to the central nervous system are symptomatic. Thus, surveillance protocols after surgical excision of RCC have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

MRI Abdomen

MRI of the abdomen without and with IV contrast is considered in all major guidelines as an adequate method for surveillance of the abdomen after surgical excision of RCC [2,9,11]. MRI has a high soft-tissue contrast resolution and is an accurate method for detecting metastases in the common sites of RCC recurrences (ie, liver, adrenal glands, lymph nodes, contralateral kidney, and bones) [38]. MRI can also assist in the distinction between residual/recurrent disease and postoperative changes after PN [39]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), MRI of the abdomen without IV contrast may be considered appropriate.

MRI Abdomen and Pelvis

MRI of the abdomen without and with IV contrast is considered in all major guidelines as an adequate method for surveillance of the abdomen after surgical excision of RCC [2,9,11]. MRI has a high soft-tissue contrast resolution and is an accurate method for detecting metastases in the common sites of RCC recurrences (ie, liver, adrenal glands, lymph nodes, contralateral kidney, and bones) [38]. MRI can also assist in the distinction between residual/recurrent disease and postoperative changes after PN [39]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), MRI of the abdomen without IV contrast may be considered appropriate.

Although MRI of the abdomen with IV contrast is considered in all major guidelines as an adequate method for surveillance of the abdomen after surgical excision of RCC, imaging the pelvis during surveillance after RCC treatment is considered optional in the guidelines [2,9,11]. There is no relevant literature regarding the use of MRI of the pelvis in the follow-up of patients after surgical excision of RCC, although data from 2 retrospective studies suggested that imaging of the pelvis with CT had minimal benefit for the detection of metastases in patients after RN or PN for RCC [25,32-34]. Therefore, MRI of the abdomen alone may be preferred over MRI of the abdomen and pelvis in this setting.

MRU

There is no relevant literature suggesting that MRU offers any additional benefit over conventional MRI of the abdomen in the surveillance of patients after treatment of localized RCC, and this method is not included in the guidelines [2,9,11]. In a retrospective analysis of 23 tumors that progressed locally after ablation, CT or MRIs obtained in the corticomedullary phase were found to be sufficient for diagnosis of recurrence in 100% of the cases; noncontrast, nephrographic, and excretory-phase images, which are typically obtained in a CTU or MRU protocol, were able to detect recurrence in only 11%, 81%, and 44% of cases, respectively [35].

MRI Head

Most patients with metastases to the central nervous system are symptomatic. Thus, surveillance protocols for RCC have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

US Kidney Retroperitoneal

The major guidelines include US as another option for imaging surveillance of the abdomen after surgical resection of localized RCC [2,9,11]. Although US may be considered an appropriate alternative for patients with contraindications to CT or MRI, one important consideration is that US is likely to be less sensitive than CT or MRI for the detection of small recurrences or distant visceral and nodal metastases in the abdomen. In one study, among 14 patients who were found to have recurrence after RN or PN for T1-3 RCC, US correctly identified only 1 case of recurrence, whereas CT detected all cases of recurrence [40]. US failed to detect 4 out of 5 recurrences in the kidney after PN [40]. In another study investigating outcomes after PN for T1-2 RCC, CT/MRI detected 96.6% of recurrences in the abdomen, whereas US detected only 3.4% of abdominal recurrences [36].

US Abdomen with IV Contrast

There is no relevant literature regarding the use of contrast-enhanced US (CEUS) in the follow-up of patients after surgical excision of RCC, and this method is not included in the guidelines [2,9,11]. Studies evaluating the performance of CEUS after ablative treatment of renal masses have shown that CEUS has an excellent concordance with CT or MRI with regard to the presence or absence of residual or recurrent tumor after ablation, both immediately after treatment and through long-term follow-up [41-49]. One important consideration is that CEUS would still be less sensitive than CT or MRI for the detection of distant visceral and nodal metastases because the contrast-enhanced portion of the study would be limited to the surgical bed. Nevertheless, in patients at low risk for recurrence, CEUS may be considered an appropriate alternative to CT and MRI.

Bone Scan Whole Body

The prevalence of osseous metastases after treatment for localized RCC has been shown to be low in patients without symptoms (ie, bone pain) or without laboratory abnormalities suggestive of osseous metastases (ie, elevated serum alkaline phosphatase level) [50,51]. Furthermore, the sites commonly involved by osseous metastases, such as the thoracolumbar spine and ribs, are located in areas covered by chest and abdominal imaging. Thus, even though bone scanning can be helpful to confirm clinically or radiographically suspected metastatic disease, current guidelines do not support its routine use in surveillance after treatment for localized RCC [2,9,11].

FDG-PET/CT Skull Base to Mid-Thigh

PET using the tracer fluorine-18-2-fluoro-2-deoxy-D-glucose (FDG-PET)/CT has a low sensitivity and specificity for the initial diagnosis of RCC [52]. This is mainly related to the variable levels of FDG avidity in RCCs; additionally, there is interference from background activity in the renal parenchyma because the kidneys are the major route of excretion of FDG. At present, the guidelines do not recommend FDG-PET/CT for the surveillance of patients after surgical excision of RCC [2,9,11]. However, emerging data suggest that FDG-PET can be useful for detecting metastatic or recurrent RCC [52,53]. In a systematic review and meta-analysis that included 15 studies with a total of 1,168 patients, FDG-PET or PET/CT had a pooled sensitivity of 86% (95% confidence interval [CI]: 0.88–0.93) and a specificity of 88% (95% CI: 0.84–0.91) for the restaging of RCC; in several of the studies, FDG-PET/CT examinations often altered the subsequent management strategy [52]. Because these results are mainly based on retrospective studies with relatively small cohorts of patients and with inconsistent reference standards, more data are needed to support the use of these agents in surveillance after surgical resection of localized RCC.

Preliminary results for other PET tracers are also becoming available. For instance, in a prospective study of 28 patients with RCC undergoing initial staging or restaging, ¹¹C-choline PET/CT was significantly more accurate than FDG-PET/CT (85.7% versus 57.1%). Among 120 lesions detected, ¹¹C-choline PET/CT detected 75 lesions (62.5%), whereas FDG-PET/CT detected 47 lesions (39.2%) [54]. In another prospective study of 10 patients with metastatic RCC, 18F-sodium fluoride (NaF) PET/CT was found to be significantly more sensitive for the detection of RCC skeletal metastases than Tc-99m bone scintigraphy or CT, with sensitivities of 100%, 29%, and 46%, respectively. CT and Tc-99m bone scintigraphy in this study identified only 65% of the metastases detected by NaF-PET/CT [55]. A small series has also shown that ⁶⁸Ga-labeled prostate-specific membrane antigen PET/CT can help to detect metastatic lesions in patients with the clear cell subtype of RCC [56].

Variant 2: Follow-up for clinically localized renal cell carcinoma; post ablation.

Among several techniques available for the ablation of localized RCC, thermal ablation techniques using radiofrequency ablation, microwave ablation, or cryoablation are the most commonly used; these procedures can be performed percutaneously or laparoscopically [57-59]. Ablation therapy is currently considered a less invasive alternative to RN or PN for renal masses measuring <4 cm (ie, T1a tumors) [2,9,11]. There is growing evidence suggesting that ablation of small renal masses produces oncologic outcomes that approach those of surgical excision but with a significantly lower overall complication rate and a significantly lower decline in renal function [5-7,57,60-67]. Because of the higher rate of local recurrence seen with ablation than with surgical excision, ablation requires more frequent use of imaging to monitor tumor involution over time [3,29,65]. Early detection of treatment failure or recurrence is important to maximize retreatment potential [65,68]. Because the risk of local recurrence is greater than the risk of distant metastases in this patient population, surveillance strategies should prioritize evaluation of the treatment bed. Guidelines recommend performing CT or MRI of the abdomen at 3 and 6 months after ablation and yearly thereafter for 5 years [2,9,11]. Guidelines also recommend the use of imaging surveillance with chest radiography or CT annually for up to 5 years after ablation of RCC [2,9,11].

Imaging-guided biopsy of renal masses is encouraged in patients considering ablative therapies [2,9,11,60]. Pretreatment biopsy can help confirm the malignant nature and aggressiveness of the tumors, which in turn can influence the frequency and duration of follow-up. After treatment, biopsy is also indicated for masses that fail to regress or that display findings suggestive of recurrence.

Radiography Chest

Chest radiography is a low-yield diagnostic tool for detecting pulmonary metastasis in patients treated for RCC, particularly in those with low-risk tumors, irrespective of the treatment modality (RN, PN, or ablation) [27,28]. In a retrospective analysis of 258 patients who had undergone resection or ablation of low-risk (T1a) RCC, pulmonary metastases developed in 3 patients (1.2%), but in only 1 patient (0.4%) was this metastasis diagnosed by surveillance chest radiographs [27]. However, according to guidelines from urologic and oncologic societies, chest radiography

is the recommended technique for surveillance of patients after ablation of T1a tumors [2,9,11]. This is in part because of concerns about potential false-positive findings with chest CT (ie, intrapulmonary lymph nodes and granulomas) that can lead to further unnecessary and potentially invasive investigations [9,12].

Radiography Abdomen

There is no relevant literature regarding the use of abdominal radiographs in the follow-up of patients after localized RCC ablation, and this method is not recommended by the guidelines [2,9,11].

Radiography Skeletal Survey

There is no relevant literature regarding the use of a radiographic survey of the whole body in the follow-up of patients after localized RCC ablation, and this method is not recommended by the guidelines [2,9,11].

Radiography Intravenous Urography

There is no relevant literature regarding the use of IV urography in the follow-up of patients after localized RCC ablation, and this method is not recommended by the guidelines [2,9,11].

CT Abdomen

CT of the abdomen is the most commonly used method for imaging surveillance after localized RCC ablation [29]. CT is a sensitive method for the detection of recurrences in the treatment bed and in other more common sites of metastases in the abdomen, such as the contralateral kidney, adrenal glands, liver, and lymph nodes, and in the visualized bones [16,17,20,22,30]. After RCC ablation, CT of the abdomen should be performed without and with IV contrast. A lack of contrast enhancement (ie, increase in attenuation <10-20 Hounsfield units on the postcontrast images) is considered the hallmark of successful treatment, which occurs via disruption of tumor vascularity. However, many completely ablated lesions show enhancement in the immediate posttreatment period, and in some cases, this enhancement may persist for several weeks to months [69,70]. The lack of spontaneous decline in enhancement and involution of the mass over time or the development of new areas of enhancement in the treatment zone or new satellite or port site soft-tissue nodules irrespective of contrast enhancement should raise concern for residual or recurrent disease. In these circumstances, a biopsy could be considered to identify the presence of viable neoplasm [9]. Initial experience with dual-energy CT after ablation of renal masses has suggested that material decomposition techniques can generate adequate virtual noncontrast images that can obviate the need for true noncontrast images. These techniques can also generate iodine-only image data sets that can assist in the evaluation of contrast enhancement of the treated lesions [71]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen without IV contrast may be considered appropriate.

CT Abdomen and Pelvis

CT of the abdomen is the most commonly used method for imaging surveillance after localized RCC ablation [29]. CT is a sensitive method for the detection of recurrences in the treatment bed and in other more common sites of metastases in the abdomen, such as the contralateral kidney, adrenal glands, liver, and lymph nodes, and in the visualized bones [16,17,20,22,30]. After RCC ablation, CT of the abdomen should be performed without and with IV contrast. A lack of contrast enhancement (ie, increase in attenuation <10-20 Hounsfield units on the postcontrast images) is considered the hallmark of successful treatment, which occurs via disruption of tumor vascularity. However, many completely ablated lesions show enhancement in the immediate posttreatment period, and in some cases, this enhancement may persist for several weeks to months [69,70]. The lack of spontaneous decline in enhancement and involution of the mass over time or the development of new areas of enhancement in the treatment zone or new satellite or port site soft-tissue nodules irrespective of contrast enhancement should raise concern for residual or recurrent disease. In these circumstances, a biopsy could be considered to identify the presence of viable neoplasm [9]. Initial experience with dual-energy CT after ablation of renal masses has suggested that material decomposition techniques can generate adequate virtual noncontrast images that can obviate the need for true noncontrast images. These techniques can also generate iodine-only image data sets that can assist in the evaluation of contrast enhancement of the treated lesions [71]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen without IV contrast may be considered appropriate.

Imaging of the pelvis with CT has been found to have limited benefit for the detection of metastases in initial staging and after RN or PN for RCC [25,32-34] and is considered optional in the surveillance guidelines [2,9,11]. Because the risk of distant metastases is significantly lower in patients with localized RCC after ablation, CT of the abdomen is preferred over CT of the abdomen and pelvis.

CT Chest

Limited data suggest that CT is more sensitive than radiography for the detection of pulmonary metastases from RCC during staging [27]. Although no direct comparison between the 2 methods has been reported in the posttreatment surveillance setting, one study demonstrated that the overwhelming majority of chest recurrences in asymptomatic cases are detected by chest CT examinations (92.3%) rather than by radiography (7.7%) [36]. In addition to a high sensitivity for the detection of pulmonary metastases, chest CT has a high sensitivity for the detection of pulmonary metastases, chest CT has a high sensitivity for the detection of intrathoracic nodal metastases from RCC; this finding has prognostic implications and may affect surgical planning for metastases resection [37]. The use of IV contrast is optional for chest CT, but it may be beneficial for the detection and characterization of hilar lymph nodes. In patients undergoing surveillance with CT of the abdomen with IV contrast, chest CT should also be performed after IV contrast administration.

Unlike CT of the abdomen, in which images without and with IV contrast are appropriate, CT of the chest without and with IV contrast does not provide additional information in these patients and is considered inappropriate. Although some consider CT the standard chest imaging technique for surveillance after RCC resection [11], there are concerns about the risk of false-positive findings (ie, intrapulmonary lymph nodes and granulomas), particularly in patients with T1a RCC, which can lead to further unnecessary and potentially invasive investigations [9]. Additionally, some authors suggest that CT of the chest may not be necessary to identify most patients with pulmonary recurrence when CT of the abdomen with coverage of the lung bases at the T7 thoracic level is performed [26].

CT Head

Most patients with metastases to the central nervous system are symptomatic. Thus, surveillance protocols after localized RCC ablation have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

CTU

There is no relevant literature suggesting that CTU offers any additional benefit over conventional CT of the abdomen in the surveillance of patients after treatment of localized RCC, and this method is not included in the guidelines [2,9,11]. In a retrospective analysis of 23 tumors that progressed locally after ablation, CT or MR images in the corticomedullary phase were found to be sufficient for diagnosis of recurrence in 100% of the cases; noncontrast, nephrographic, and excretory-phase images, which are typically obtained in a CTU or MRU protocol, were able to detect recurrence in only 11%, 81%, and 44% of cases, respectively [35].

MRI Abdomen

MRI of the abdomen is commonly used for follow-up after ablation of localized RCC [29]. MRI should be performed without and with IV contrast to assess tumor enhancement. Image data sets generated from subtraction of the precontrast from the postcontrast images can assist with evaluation of residual or recurrent tumor enhancement, especially during the first year of follow-up, because of the high signal intensity background of the ablated tumor on T1-weighted images [72]. However, as with CT, persistent tumor enhancement is common after successful ablation, particularly in patients with clear-cell RCC [73], and this enhancement can last for days to months after treatment [72-74]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), MRI of the abdomen without IV contrast may be considered appropriate.

MRI Abdomen and Pelvis

MRI of the abdomen is commonly used for follow-up after ablation of localized RCC [29]. MRI should be performed without and with IV contrast to assess tumor enhancement. Image data sets generated from subtraction of the precontrast from the postcontrast images can assist with evaluation of residual or recurrent tumor enhancement, especially during the first year of follow-up, because of the high signal intensity background of the ablated tumor on T1-weighted images [72]. However, as with CT, persistent tumor enhancement is common after successful ablation, particularly in patients with clear-cell RCC [73], and this enhancement can last for days to months after treatment [72-74]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reactions), MRI of the abdomen without IV contrast may be considered appropriate.

There is no relevant literature regarding the use of MRI of the pelvis in the follow-up of patients after RCC ablation. Imaging of the pelvis with CT has been found to provide minimal benefit for the detection of metastases in the initial staging and after RN or PN for RCC [25,32-34] and is considered optional in the surveillance guidelines [2,9,11]. Because the risk of distant metastases is significantly lower in patients with localized RCC after ablation, MRI of the abdomen is preferred over MRI of the abdomen and pelvis.

MRU

There is no relevant literature suggesting that MRU offers any additional benefit over conventional MRI of the abdomen in the surveillance of patients after treatment of localized RCC, and this method is not included in the guidelines [2,9,11]. In a retrospective analysis of 23 tumors that progressed locally after ablation, CT or MR images obtained in the corticomedullary phase were found to be sufficient for diagnosis of recurrence in 100% of the cases; noncontrast, nephrographic, and excretory-phase images, which are typically obtained in a CTU or MRU protocol, were able to detect recurrence in only 11%, 81%, and 44% of cases, respectively [35].

MRI Head

Most patients with metastases to the central nervous system are symptomatic. Thus, surveillance protocols for RCC have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

US Kidney Retroperitoneal

There is no relevant literature regarding the use of conventional US of the kidney in follow-up of patients after localized RCC ablation, and the guidelines offer different recommendations. The National Comprehensive Cancer Network considers US an alternative for annual surveillance after negative evaluation with CT or MRI in the first 6 months [2]; the European Association of Urology recommends US only for surveillance after the treatment of RCC with a low-risk profile [11]; and the American Urological Association does not include US in their recommendations regarding follow-up after ablation [9].

US Abdomen with IV Contrast

The use of CEUS after radiofrequency ablation, microwave ablation, and cryoablation of renal masses has been the subject of many studies [41-49]. In these studies, CEUS has been found to have excellent concordance with CT or MRI with regard to the presence or absence of enhancement in renal masses after ablation, both immediately after treatment and through long-term follow-up. In a prospective study including 64 tumors, CEUS and CT were in concordance regarding the presence of residual enhancement in 2 tumors and the presence of complete necrosis in the other 62 tumors at 1 month after radiofrequency ablation. On subsequent follow-up of 61 tumors, CEUS and CT were in concordance for 59 tumors, with 2 false-positive CEUS cases [47]. In another study, enhancement on CEUS and CT/MRI after cryoablation was concordant for 23 of 32 tumors (72%) at 3 months and for 19 of 21 tumors (91%) at 12 months [42]. Researchers in another study reported good interobserver agreement for CEUS among 3 radiologists with ≥ 15 years of experience with US (weighted κ : 0.84 [CI: 0.71–0.93]), although better interobserver agreement was achieved with CT/MRI for 3 radiologists with ≥15 years of experience with CT/MRI (weighted k: 0.94 [CI: 0.88–0.99]) [46]. In a more recent study, CEUS was found to have a high negative predictive value (100%) for local recurrence after thermal ablation of RCC among 20 patients who had either a contraindication to CT or MRI or inconclusive findings with these methods on surveillance imaging [41]. These results suggest that CEUS could be used as an alternative to CT and MRI for the evaluation of treatment response and local recurrence. The performance of CEUS may be limited in a small number of cases in which the ablation cavity is not well visualized on precontrast US images [48]. Additionally, CEUS has limited ability to detect distant RCC metastasis [48].

Bone Scan Whole Body

The prevalence of osseous metastases has been shown to be low in patients without symptoms (ie, bone pain) or without laboratory abnormalities suggestive of osseous metastases (ie, elevated serum alkaline phosphatase level) [50,51]. Furthermore, the sites commonly involved by osseous metastases, such as the thoracolumbar spine and ribs, are located in areas covered by chest and abdominal imaging. Thus, although Tc-99m bone scanning can be helpful in confirming clinically or radiographically suspected metastatic disease, current guidelines do not support its routine use in surveillance after treatment for localized RCC [2,9,11].

FDG-PET/CT Skull Base to Mid-Thigh

There is no relevant literature regarding the use of FDG-PET or PET/CT for the follow-up of patients after localized RCC ablation. At present, the guidelines do not recommend FDG-PET/CT for the surveillance of patients after RCC ablation [2,9,11].

Variant 3: Follow-up for clinically localized renal cell carcinoma; active surveillance.

Active surveillance has been increasingly used for the management of small localized renal masses in a selected group of patients with comorbidities or reduced life expectancy in whom the risks associated with surgical excision or ablative therapies surpass the risk of significant disease progression and cancer-specific mortality [2,9,11,75-80].

Patients on active surveillance undergo rigorous imaging and clinical follow-up, with subsequent surgical or minimally invasive treatment reserved for those with tumors that progress. Available data on active surveillance, which are predominantly based on T1a tumors (ie, tumors ≤ 4 cm in the greatest dimension and confined to the kidney), suggest that this management alternative does not compromise oncologic outcomes, with a risk of metastatic disease progression of 0% to 2% [8,75-78,81-84].

Current guidelines recommend biopsy of the renal masses to define the surveillance strategy [2,9,11]. Researchers have found that biopsy is being increasingly used for T1a tumors and that patients who undergo biopsy are significantly more likely to be treated with nonsurgical management (36.8%) than those who do not undergo biopsy (11.4%) [85]. Of note, small renal mass growth kinetics can vary greatly, especially during the initial 6 to 12 months of active surveillance [82,84]. In a systematic review of the literature, researchers found no significant difference between the growth rates of benign masses (0.3 cm/y) and those of malignant masses (0.35 cm/y) [82]. Furthermore, studies have shown that even masses without growth may be malignant [8,76,77,81]. In spite of this, growth rates are generally accepted as surrogates for aggressive behavior and metastatic potential in small renal masses [76,81]. Therefore, the guidelines recommend defining the growth rate of renal masses with serial imaging of the abdomen with CT or MRI within 6 months of the initiation of active surveillance for masses that are shown to be RCCs or oncocytic neoplasms and for those with indeterminate histology features [2,9]. Imaging should be performed at least annually thereafter with CT, MRI, or US. Imaging surveillance of the chest on a yearly basis (or more frequently depending on clinical behavior) is recommended only in those patients with RCC or tumors with oncocytic features [2,9].

Radiography Chest

Metastatic progression occurs infrequently in patients with T1a renal masses on active surveillance [76,77,81]. Nevertheless, it has been reported that 20% to 30% of T1a tumors have potentially aggressive histologic features, thus requiring surveillance of the chest [9]. No studies have compared chest radiography and chest CT in the setting of active surveillance; however, chest radiography is the most commonly used method for surveillance [2,9]. This is in part because of concerns about potential false-positive findings with chest CT (ie, intrapulmonary lymph nodes and granulomas) that can lead to further unnecessary and potentially invasive investigations [9,12].

Radiography Abdomen

There is no relevant literature regarding the use of abdominal radiographs in the surveillance of small localized renal masses, and this method is not recommended by the guidelines [2,9,11].

Radiography Skeletal Survey

There is no relevant literature regarding the use of a radiographic survey of the whole body in the surveillance of small localized renal masses, and this method is not recommended by the guidelines [2,9,11].

Radiography Intravenous Urography

There is no relevant literature regarding the use of IV urography in the surveillance of small localized renal masses, and this method is not recommended by the guidelines [2,9,11].

CT Abdomen

CT of the abdomen is the most common method by which small renal masses are detected and is also the most commonly used method for surveillance of small localized renal masses. CT of the abdomen performed without and with IV contrast is considered appropriate if there is a need for initial characterization of the enhancement pattern of the renal mass. Subsequent follow-up to monitor tumor growth could be achieved with CT of the abdomen with IV contrast. The maximum diameter of the mass is frequently used to assess tumor growth, although interobserver and intraobserver variabilities on the order of ± 3.1 and ± 2.3 mm, respectively, have been reported [82]. In one study, researchers found that 2-D and 3-D measurements had greater accuracy for the detection of tumor growth than the measurement of the single largest diameter or gestalt visual assessment [86]. After the initial follow-up, once the growth rate of the mass has been established, alternating the follow-up with MRI or US has been suggested [2,9,78]. It is important to note that in addition to interobserver and intraobserver variability, the use of different modalities can result in inconsistent measurements that can ultimately have an effect on patient care [82]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen without IV contrast may be considered appropriate.

CT Abdomen and Pelvis

CT of the abdomen is the most common method by which small renal masses are detected and is also the most commonly used method for surveillance of small localized renal masses. CT of the abdomen performed without

and with IV contrast is considered appropriate if there is a need for initial characterization of the enhancement pattern of the renal mass. Subsequent follow-up to monitor tumor growth could be achieved with CT of the abdomen with IV contrast. The maximum diameter of the mass is frequently used to assess tumor growth, although interobserver and intraobserver variabilities on the order of ± 3.1 and ± 2.3 mm, respectively, have been reported [82]. In one study, researchers found that 2-D and 3-D measurements had greater accuracy for the detection of tumor growth than the measurement of the single largest diameter or gestalt visual assessment [86]. After the initial follow-up, once the growth rate of the mass has been established, alternating the follow-up with MRI or US has been suggested [2,9,78]. It is important to note that in addition to interobserver and intraobserver variability, the use of different modalities can result in inconsistent measurements that can ultimately have an effect on patient care [82]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), CT of the abdomen without IV contrast may be considered appropriate.

Although CT of the abdomen is the most commonly used method for surveillance of small localized renal masses, the benefit of imaging the pelvis during surveillance has not yet been defined and is considered optional in the guidelines [2,9,11]. Data from 2 retrospective studies evaluating RCC staging with CT suggested that imaging of the pelvis had limited benefit for the detection of metastases [33,34]. Because metastatic progression occurs infrequently in patients on active surveillance with T1a renal masses [8,75-78,81-84], CT of the abdomen is preferred over CT of the abdomen and pelvis.

CTU

There is no relevant literature regarding the use of CTU in the surveillance of small localized renal masses, and this method is not recommended by the guidelines [2,9,11].

CT Chest

Chest CT is listed as an alternative to radiography for surveillance of small localized renal masses by the National Comprehensive Cancer Network guidelines [2]. Limited data have demonstrated that CT is more sensitive than radiography for the detection of pulmonary metastases from RCC during staging [27]. However, no comparison between radiography and CT has been reported in the active surveillance setting. Despite the higher sensitivity of CT, there are some concerns about the risk of false-positive findings (ie, intrapulmonary lymph nodes and granulomas), particularly in patients with T1a RCC, which can lead to further unnecessary and potentially invasive investigations [9]. Additionally, some authors suggest that CT of the chest may not be necessary to identify most cases of pulmonary recurrence after nephrectomy for RCC when CT of the abdomen with coverage of the lung bases at the T7 thoracic level is performed [26]. The use of IV contrast is optional for CT of the chest but it may be beneficial for detection and characterization of the hilar lymph nodes. In patients undergoing active surveillance with CT of the abdomen who are receiving IV contrast, chest CT can also be performed after IV contrast administration. Unlike CT of the chest without and with IV contrast may be appropriate in some circumstances, CT of the chest without and with IV contrast does not provide additional information in these patients and is considered inappropriate.

CT Head

Most patients with metastases to the central nervous system are symptomatic. Thus, active surveillance protocols for small localized renal masses have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

MRI Abdomen

MRI of the abdomen without and with IV contrast is an accurate method for the detection and characterization of small localized renal masses. Different sequences, including T2-weighted, chemical shift T1-weighted, contrastenhanced T1-weighted, and diffusion-weighted images, can help distinguish RCC from other benign and malignant lesions and distinguish the clear-cell subtype from other subtypes of RCC. Some MRI features of renal masses beyond size and growth rates can also be used to determine tumor aggressiveness and risk of metastatic potential [87]. This may be particularly useful for the characterization of small renal masses that have indeterminate findings on CT and US or when biopsy of these masses is not feasible or is inconclusive. Active surveillance guidelines include MRI and CT as appropriate imaging modalities for the initial evaluation of growth patterns and for subsequent follow-up [2,9,11]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), MRI of the abdomen without IV contrast may be considered appropriate.

MRI Abdomen and Pelvis

MRI of the abdomen without and with IV contrast is an accurate method for the detection and characterization of small localized renal masses. Different sequences, including T2-weighted, chemical shift T1-weighted, contrastenhanced T1-weighted, and diffusion-weighted images, can help distinguish RCC from other benign and malignant lesions and distinguish the clear-cell subtype from other subtypes of RCC. Some MRI features of renal masses beyond size and growth rates can also be used to determine tumor aggressiveness and risk of metastatic potential [87]. This may be particularly useful for the characterization of small renal masses that have indeterminate findings on CT and US or when biopsy of these masses is not feasible or is inconclusive. Active surveillance guidelines include MRI and CT as appropriate imaging modalities for the initial evaluation of growth patterns and for subsequent follow-up [2,9,11]. For patients in whom contrast is contraindicated (eg, previous anaphylactic reaction), MRI of the abdomen without IV contrast may be considered appropriate.

Although MRI of the abdomen can be useful for characterization and follow-up of small localized renal masses undergoing active surveillance, the benefit of imaging the pelvis during surveillance has not yet been defined and is considered optional in the guidelines [2,9,11]. There is no relevant literature regarding the use of MRI of the pelvis in the follow-up of patients on active surveillance, although data from 2 retrospective studies evaluating RCC staging with CT suggested that imaging of the pelvis had limited benefit for the detection of metastases [33,34]. Furthermore, metastatic progression occurs infrequently in patients on active surveillance with T1a renal masses [8,75-78,81,82]; therefore, MRI of the abdomen is preferred over MRI of the abdomen and pelvis.

MRI Head

Most patients with metastases to the central nervous system are symptomatic. Thus, active surveillance protocols for small localized renal masses have not supported routine imaging of the brain to search for metastases in asymptomatic patients. Brain imaging should be performed only in cases with suggestive signs or symptoms [2,9,11].

MRU

There is no relevant literature regarding the use of MRU in the surveillance of small localized renal masses, and this method is not recommended by the guidelines [2,9,11].

US Kidney Retroperitoneal

US of the kidney is an acceptable imaging modality for follow-up of small localized renal masses on active surveillance, especially once the growth rate of the renal mass has been established with CT or MRI [2,9,11,78]. US is an excellent method for characterizing cystic lesions and often provides supplementary information to the other imaging modalities. However, unenhanced US has an overall diagnostic accuracy for characterizing renal masses of only 30% [88]. Some concerns also exist regarding the reproducibility of measurements obtained with US and their correlation with measurements obtained with CT and MRI; any discrepancies could suggest a falsely positive or negative growth rate [82]. When US is used, some authors have recommended that any discrepancy in tumor size or growth rate or qualitative changes in tumor appearance should prompt imaging with CT or MRI [78].

US Abdomen with IV Contrast

CEUS is an accurate method for the detection and characterization of small renal masses, which in theory may be beneficial for patients on active surveillance. In a large series of CEUS for the evaluation of 1,018 indeterminate renal masses in 721 patients followed for as long as 10 years, the sensitivity of CEUS was 100% (95% CI: 97.1%–100%) with a specificity of 95% (95% CI: 89.9%–98.0%), a positive predictive value of 91.5%, and a negative predictive value of 100% [89]. Multiple additional studies, including a recent meta-analysis of 17 studies with 1,142 lesions, have found that CEUS is more sensitive but slightly less specific than CT and MRI in detecting and characterizing renal masses [90]. The performance of CEUS in active renal mass surveillance may be limited in a small number of cases in which the renal mass is not well visualized on precontrast US [73].

Bone Scan Whole Body

The incidence of metastatic progression in patients with small localized renal masses on active surveillance is low (0%-2%). Furthermore, the prevalence of osseous metastases has been shown to be low in patients without symptoms (ie, bone pain) or without laboratory abnormalities suggestive of osseous metastases (ie, elevated serum alkaline phosphatase level) [50,51]. Therefore, Tc-99m bone scanning is not routinely recommended in active surveillance [2,9,11].

FDG-PET/CT Skull Base to Mid-Thigh

FDG-PET/CT has low sensitivity and specificity for the detection and local staging of RCC [52]. This is mainly related to the variable levels of FDG avidity in RCCs; additionally, there is interference from background activity in the renal parenchyma as the kidneys are the major route of excretion of FDG. At present, given the lack of literature to support the use of FDG-PET/CT, the guidelines do not recommend this technique for active surveillance in patients with renal masses [2,9,11].

Summary of Recommendations

- Variant 1: CT abdomen with IV contrast, CT abdomen without and with IV contrast, or MRI abdomen without and with IV contrast is usually appropriate in the follow-up of patients after surgical excision of RCC. These procedures are equivalent alternatives (ie, only one procedure will be ordered to provide the clinical information to effectively manage the patient's care). The panel did not agree on recommending MRI abdomen and pelvis without and with IV contrast. There is insufficient medical literature to conclude whether the scan is of benefit in this clinical scenario, and its use may be appropriate but controversial.
- Variant 2: CT abdomen with IV contrast, CT abdomen without and with IV contrast, or MRI abdomen without and with IV contrast is usually appropriate in the follow-up of patients after localized RCC ablation. These procedures are equivalent alternatives (ie, only one procedure will be ordered to provide the clinical information to effectively manage the patient's care).
- Variant 3: CT abdomen with IV contrast, CT abdomen without and with IV contrast, MRI abdomen without and with IV contrast, or US abdomen with IV contrast is usually appropriate in the active surveillance of localized RCC. These procedures are equivalent alternatives (ie, only one procedure will be ordered to provide the clinical information to effectively manage the patient's care).

Supporting Documents

The evidence table, literature search, and appendix for this topic are available at <u>https://acsearch.acr.org/list</u>. The appendix includes the strength of evidence assessment and the final rating round tabulations for each recommendation.

For additional information on the Appropriateness Criteria methodology and other supporting documents, go to www.acr.org/ac.

Appropriateness Category Name	Appropriateness Rating	Appropriateness Category Definition
Usually Appropriate	7, 8, or 9	The imaging procedure or treatment is indicated in the specified clinical scenarios at a favorable risk-benefit ratio for patients.
May Be Appropriate	4, 5, or 6	The imaging procedure or treatment may be indicated in the specified clinical scenarios as an alternative to imaging procedures or treatments with a more favorable risk-benefit ratio, or the risk-benefit ratio for patients is equivocal.
May Be Appropriate (Disagreement)	5	The individual ratings are too dispersed from the panel median. The different label provides transparency regarding the panel's recommendation. "May be appropriate" is the rating category and a rating of 5 is assigned.
Usually Not Appropriate	1, 2, or 3	The imaging procedure or treatment is unlikely to be indicated in the specified clinical scenarios, or the risk-benefit ratio for patients is likely to be unfavorable.

Appropriateness Category Names and Definitions

Relative Radiation Level Information

Potential adverse health effects associated with radiation exposure are an important factor to consider when selecting the appropriate imaging procedure. Because there is a wide range of radiation exposures associated with different diagnostic procedures, a relative radiation level (RRL) indication has been included for each imaging examination. The RRLs are based on effective dose, which is a radiation dose quantity that is used to estimate population total radiation risk associated with an imaging procedure. Patients in the pediatric age group are at inherently higher risk from exposure, because of both organ sensitivity and longer life expectancy (relevant to the long latency that appears to accompany radiation exposure). For these reasons, the RRL dose estimate ranges for pediatric examinations are lower as compared with those specified for adults (see Table below). Additional information regarding radiation dose assessment for imaging examinations can be found in the ACR Appropriateness Criteria[®] Radiation Dose Assessment Introduction document [91].

Relative Radiation Level Designations		
Relative Radiation Level*	Adult Effective Dose Estimate Range	Pediatric Effective Dose Estimate Range
0	0 mSv	0 mSv
۲	<0.1 mSv	<0.03 mSv
*	0.1-1 mSv	0.03-0.3 mSv
���	1-10 mSv	0.3-3 mSv
\$\$\$\$	10-30 mSv	3-10 mSv
***	30-100 mSv	10-30 mSv

*RRL assignments for some of the examinations cannot be made, because the actual patient doses in these procedures vary as a function of a number of factors (eg, region of the body exposed to ionizing radiation, the imaging guidance that is used). The RRLs for these examinations are designated as "Varies".

References

- 1. Siegel RL, Miller KD, Jemal A. Cancer statistics, 2020. CA Cancer J Clin 2020;70:7-30.
- 2. Motzer RJ, Jonasch E, Agarwal N, et al. Kidney Cancer, Version 2.2017, NCCN Clinical Practice Guidelines in Oncology. J Natl Compr Canc Netw 2017;15:804-34.
- 3. Andrews JR, Atwell T, Schmit G, et al. Oncologic Outcomes Following Partial Nephrectomy and Percutaneous Ablation for cT1 Renal Masses. Eur Urol 2019;76:244-51.
- Choi SH, Kim JW, Kim JH, Kim KW. Efficacy and Safety of Microwave Ablation for Malignant Renal Tumors: An Updated Systematic Review and Meta-Analysis of the Literature Since 2012. Korean J Radiol 2018;19:938-49.
- 5. Iannuccilli JD, Dupuy DE, Beland MD, Machan JT, Golijanin DJ, Mayo-Smith WW. Effectiveness and safety of computed tomography-guided radiofrequency ablation of renal cancer: a 14-year single institution experience in 203 patients. Eur Radiol 2016;26:1656-64.
- 6. Katsanos K, Mailli L, Krokidis M, McGrath A, Sabharwal T, Adam A. Systematic review and meta-analysis of thermal ablation versus surgical nephrectomy for small renal tumours. Cardiovasc Intervent Radiol 2014;37:427-37.
- Yu J, Zhang X, Liu H, et al. Percutaneous Microwave Ablation versus Laparoscopic Partial Nephrectomy for cT1a Renal Cell Carcinoma: A Propensity-matched Cohort Study of 1955 Patients. Radiology 2020;294:698-706.
- 8. Smaldone MC, Kutikov A, Egleston BL, et al. Small renal masses progressing to metastases under active surveillance: a systematic review and pooled analysis. Cancer 2012;118:997-1006.
- 9. Donat SM, Diaz M, Bishoff JT, et al. Follow-up for Clinically Localized Renal Neoplasms: AUA Guideline. J Urol 2013;190:407-16.
- 10. Merrill SB, Sohl BS, Hamirani A, et al. Capturing Renal Cell Carcinoma Recurrences When Asymptomatic Improves Patient Survival. Clin Genitourin Cancer 2019;17:132-38.
- 11. Ljungberg B, Albiges L, Abu-Ghanem Y, et al. European Association of Urology Guidelines on Renal Cell Carcinoma: The 2019 Update. Eur Urol 2019;75:799-810.

- 12. Williamson TJ, Pearson JR, Ischia J, Bolton DM, Lawrentschuk N. Guideline of guidelines: follow-up after nephrectomy for renal cell carcinoma. BJU Int 2016;117:555-62.
- 13. Dabestani S, Beisland C, Stewart GD, et al. Intensive Imaging-based Follow-up of Surgically Treated Localised Renal Cell Carcinoma Does Not Improve Post-recurrence Survival: Results from a European Multicentre Database (RECUR). Eur Urol 2019;75:261-64.
- 14. Antonelli A, Furlan M, Sodano M, et al. Features, risk factors and clinical outcome of "very late" recurrences after surgery for localized renal carcinoma: A retrospective evaluation of a cohort with a minimum of 10 years of follow up. Int J Urol 2016;23:36-40.
- 15. Levy DA, Slaton JW, Swanson DA, Dinney CP. Stage specific guidelines for surveillance after radical nephrectomy for local renal cell carcinoma. J Urol 1998;159:1163-7.
- 16. Romeo A, Garcia Marchinena P, Jurado AM, Gueglio G. Renal fossa recurrence after radical nephrectomy: Current management, and oncological outcomes. Urol Oncol 2020;38:42 e7-42 e12.
- 17. Stephenson AJ, Chetner MP, Rourke K, et al. Guidelines for the surveillance of localized renal cell carcinoma based on the patterns of relapse after nephrectomy. J Urol 2004;172:58-62.
- 18. Stewart-Merrill SB, Thompson RH, Boorjian SA, et al. Oncologic Surveillance After Surgical Resection for Renal Cell Carcinoma: A Novel Risk-Based Approach. J Clin Oncol 2015;33:4151-7.
- 19. Zisman A, Pantuck AJ, Wieder J, et al. Risk group assessment and clinical outcome algorithm to predict the natural history of patients with surgically resected renal cell carcinoma. J Clin Oncol 2002;20:4559-66.
- 20. Itano NB, Blute ML, Spotts B, Zincke H. Outcome of isolated renal cell carcinoma fossa recurrence after nephrectomy. J Urol 2000;164:322-5.
- 21. Sandock DS, Seftel AD, Resnick MI. A new protocol for the followup of renal cell carcinoma based on pathological stage. J Urol 1995;154:28-31.
- 22. Saidi JA, Newhouse JH, Sawczuk IS. Radiologic follow-up of patients with T1-3a,b,c or T4N+M0 renal cell carcinoma after radical nephrectomy. Urology 1998;52:1000-3.
- 23. Lau WK, Blute ML, Weaver AL, Torres VE, Zincke H. Matched comparison of radical nephrectomy vs nephron-sparing surgery in patients with unilateral renal cell carcinoma and a normal contralateral kidney. Mayo Clin Proc 2000;75:1236-42.
- 24. Chow AK, Kahan AN, Hwang T, Coogan CL, Latchamsetty KC. Should We Separate the Pulmonary Surveillance Protocol for Postsurgical T1a and T1b Renal Cell Carcinoma? A Multicenter Database Analysis. Urology 2018;122:127-32.
- 25. Kaiser A, Davenport MS, Hafez KS, Alva A, Bailey JJ, Francis IR. Utility of Pelvic CT for Surveillance of T2-T4 Renal Cell Carcinoma After Nephrectomy With Curative Intent. AJR Am J Roentgenol 2018;210:1088-91.
- 26. Lee HS, Kang WJ, Cho NH, Park SY. Is Chest Computed Tomography Always Necessary Following Nephrectomy for Renal Cell Carcinoma? A Pilot Study in Single Tertiary Institution. J Comput Assist Tomogr 2019;43:333-37.
- 27. Canvasser NE, Stouder K, Lay AH, et al. The Usefulness of Chest X-Rays for T1a Renal Cell Carcinoma Surveillance. J Urol 2016;196:321-6.
- 28. Doornweerd BH, de Jong IJ, Bergman LM, Ananias HJ. Chest X-ray in the follow-up of renal cell carcinoma. World J Urol 2014;32:1015-9.
- 29. Kowalczyk KJ, Harbin AC, Choueiri TK, et al. Use of surveillance imaging following treatment of small renal masses. J Urol 2013;190:1680-5.
- 30. Hafez KS, Novick AC, Campbell SC. Patterns of tumor recurrence and guidelines for followup after nephron sparing surgery for sporadic renal cell carcinoma. J Urol 1997;157:2067-70.
- 31. Jain Y, Liew S, Taylor MB, Bonington SC. Is dual-phase abdominal CT necessary for the optimal detection of metastases from renal cell carcinoma? Clin Radiol 2011;66:1055-9.
- 32. Gofrit ON, Rabinovich I, Yutkin V, et al. Abbreviated CT protocol for postoperative surveillance of renal cancer. Urol Oncol 2018;36:498 e9-98 e13.
- 33. Fielding JR, Aliabadi N, Renshaw AA, Silverman SG. Staging of 119 patients with renal cell carcinoma: the yield and cost-effectiveness of pelvic CT. AJR Am J Roentgenol 1999;172:23-5.
- 34. Khaitan A, Gupta NP, Hemal AK, Dogra PN, Seth A, Aron M. Is there a need for pelvic CT scan in cases of renal cell carcinoma? Int Urol Nephrol 2002;33:13-5.
- 35. Eiken PW, Atwell TD, Kurup AN, Boorjian SA, Thompson RH, Schmit GD. Imaging following renal ablation: what can we learn from recurrent tumors? Abdom Radiol (NY) 2018;43:2750-55.
- 36. Mouracade P, Chavali JS, Kara O, et al. Imaging strategy and outcome following partial nephrectomy. Urol Oncol 2017;35:660 e1-60 e8.

- 37. Winter H, Meimarakis G, Angele MK, et al. Tumor infiltrated hilar and mediastinal lymph nodes are an independent prognostic factor for decreased survival after pulmonary metastasectomy in patients with renal cell carcinoma. J Urol 2010;184:1888-94.
- 38. Platzek I, Zastrow S, Deppe PE, et al. Whole-body MRI in follow-up of patients with renal cell carcinoma. Acta Radiol 2010;51:581-9.
- 39. Kisa E, Sahin H, Cakmak O, et al. Magnetic resonance imaging characteristics and changes in hemostatic agents after partial nephrectomy. Int Urol Nephrol 2019;51:917-25.
- 40. Quinlan M, Wei G, Davis N, et al. Renal Cell Carcinoma Follow-Up Is it Time to Abandon Ultrasound? Curr Urol 2019;13:19-24.
- 41. Atri M, Alrashed A, Hassan A, Khalili K, Kim TK, Jang HJ. Negative Predictive Value of Contrast-Enhanced Ultrasound of Liver and Kidney Thermal Ablation Sites for Local Tumour Progression During Long-term Follow-up: A Retrospective Consecutive Study. Can Assoc Radiol J 2019;70:434-40.
- 42. Barwari K, Wijkstra H, van Delden OM, de la Rosette JJ, Laguna MP. Contrast-enhanced ultrasound for the evaluation of the cryolesion after laparoscopic renal cryoablation: an initial report. J Endourol 2013;27:402-7.
- 43. Calio BP, Lyshchik A, Li J, et al. Long Term Surveillance of Renal Cell Carcinoma Recurrence Following Ablation using 2D and 3D Contrast-Enhanced Ultrasound. Urology 2018;121:189-96.
- 44. Garbajs M, Popovic P. Contrast-enhanced ultrasound for assessment of therapeutic response after percutaneous radiofrequency ablation of small renal tumors. J BUON 2016;21:685-90.
- 45. Guo F, Hu B, Chen L, Li J. Clinical application of contrast-enhanced ultrasound after percutaneous renal tumor ablation. Br J Radiol 2019;92:20190183.
- 46. Hoeffel C, Pousset M, Timsit MO, et al. Radiofrequency ablation of renal tumours: diagnostic accuracy of contrast-enhanced ultrasound for early detection of residual tumour. Eur Radiol 2010;20:1812-21.
- 47. Kong WT, Zhang WW, Guo HQ, et al. Application of contrast-enhanced ultrasonography after radiofrequency ablation for renal cell carcinoma: is it sufficient for assessment of therapeutic response? Abdom Imaging 2011;36:342-7.
- 48. Sanz E, Hevia V, Arias F, et al. Contrast-enhanced ultrasound (CEUS): an excellent tool in the follow-up of small renal masses treated with cryoablation. Curr Urol Rep 2015;16:469.
- 49. Zeccolini G, Del Biondo D, Cicero C, Casarin A, Guarise A, Celia A. Comparison of Contrast-Enhanced Ultrasound Scan (CEUS) and MRI in the follow-up of cryoablation for small renal tumors. Experience on 25 cases. Urologia 2014;81 Suppl 23:S1-8.
- 50. Blacher E, Johnson DE, Haynie TP. Value of routine radionuclide bone scans in renal cell carcinoma. Urology 1985;26:432-4.
- 51. Koga S, Tsuda S, Nishikido M, et al. The diagnostic value of bone scan in patients with renal cell carcinoma. J Urol 2001;166:2126-8.
- 52. Ma H, Shen G, Liu B, Yang Y, Ren P, Kuang A. Diagnostic performance of 18F-FDG PET or PET/CT in restaging renal cell carcinoma: a systematic review and meta-analysis. Nucl Med Commun 2017;38:156-63.
- 53. Elahmadawy MA, Elazab MSS, Ahmed S, Salama M. Diagnostic value of F-18 FDG PET/CT for local and distant disease relapse surveillance in surgically treated RCC patients: Can it aid in establishing consensus follow up strategy? Nucl Med Rev Cent East Eur 2018;21:85-91.
- 54. Nakanishi Y, Kitajima K, Yamada Y, et al. Diagnostic performance of (11)C-choline PET/CT and FDG PET/CT for staging and restaging of renal cell cancer. Ann Nucl Med 2018;32:658-68.
- 55. Gerety EL, Lawrence EM, Wason J, et al. Prospective study evaluating the relative sensitivity of 18F-NaF PET/CT for detecting skeletal metastases from renal cell carcinoma in comparison to multidetector CT and 99mTc-MDP bone scintigraphy, using an adaptive trial design. Ann Oncol 2015;26:2113-8.
- 56. Sawicki LM, Buchbender C, Boos J, et al. Diagnostic potential of PET/CT using a 68Ga-labelled prostatespecific membrane antigen ligand in whole-body staging of renal cell carcinoma: initial experience. Eur J Nucl Med Mol Imaging 2017;44:102-07.
- 57. Mues AC, Okhunov Z, Haramis G, D'Agostino H, Shingleton BW, Landman J. Comparison of percutaneous and laparoscopic renal cryoablation for small (<3.0 cm) renal masses. J Endourol 2010;24:1097-100.
- 58. Pirasteh A, Snyder L, Boncher N, Passalacqua M, Rosenblum D, Prologo JD. Cryoablation vs. radiofrequency ablation for small renal masses. Acad Radiol 2011;18:97-100.
- 59. Young EE, Castle SM, Gorbatiy V, Leveillee RJ. Comparison of safety, renal function outcomes and efficacy of laparoscopic and percutaneous radio frequency ablation of renal masses. J Urol 2012;187:1177-82.
- 60. Aron M, Kamoi K, Remer E, Berger A, Desai M, Gill I. Laparoscopic renal cryoablation: 8-year, single surgeon outcomes. J Urol 2010;183:889-95.

- 61. Balageas P, Cornelis F, Le Bras Y, et al. Ten-year experience of percutaneous image-guided radiofrequency ablation of malignant renal tumours in high-risk patients. Eur Radiol 2013;23:1925-32.
- 62. Breen DJ, Bryant TJ, Abbas A, et al. Percutaneous cryoablation of renal tumours: outcomes from 171 tumours in 147 patients. BJU Int 2013;112:758-65.
- 63. Georgiades CS, Rodriguez R. Efficacy and safety of percutaneous cryoablation for stage 1A/B renal cell carcinoma: results of a prospective, single-arm, 5-year study. Cardiovasc Intervent Radiol 2014;37:1494-9.
- 64. McDougal WS, Gervais DA, McGovern FJ, Mueller PR. Long-term followup of patients with renal cell carcinoma treated with radio frequency ablation with curative intent. J Urol 2005;174:61-3.
- 65. Shapiro DD, Wells SA, Best SL, et al. Comparing Outcomes for Patients with Clinical T1b Renal Cell Carcinoma Treated With Either Percutaneous Microwave Ablation or Surgery. Urology 2020;135:88-94.
- 66. Wah TM, Irving HC, Gregory W, Cartledge J, Joyce AD, Selby PJ. Radiofrequency ablation (RFA) of renal cell carcinoma (RCC): experience in 200 tumours. BJU Int 2014;113:416-28.
- 67. Zagoria RJ, Pettus JA, Rogers M, Werle DM, Childs D, Leyendecker JR. Long-term outcomes after percutaneous radiofrequency ablation for renal cell carcinoma. Urology 2011;77:1393-7.
- 68. Best SL, Park SK, Yaacoub RF, et al. Long-term outcomes of renal tumor radio frequency ablation stratified by tumor diameter: size matters. J Urol 2012;187:1183-9.
- 69. Javadi S, Ahrar JU, Ninan E, Gupta S, Matin SF, Ahrar K. Characterization of contrast enhancement in the ablation zone immediately after radiofrequency ablation of renal tumors. J Vasc Interv Radiol 2010;21:690-5.
- Nielsen TK, Ostraat O, Andersen G, Hoyer S, Graumann O, Borre M. Computed Tomography Contrast Enhancement Following Renal Cryoablation--Does it Represent Treatment Failure? J Endourol 2015;29:1353-60.
- 71. Park SY, Kim CK, Park BK. Dual-energy CT in assessing therapeutic response to radiofrequency ablation of renal cell carcinomas. Eur J Radiol 2014;83:e73-9.
- 72. Lee HJ, Chung HJ, Wang HK, et al. Evolutionary magnetic resonance appearance of renal cell carcinoma after percutaneous cryoablation. Br J Radiol 2016;89:20160151.
- 73. Takaki H, Nakatsuka A, Cornelis F, et al. False-Positive Tumor Enhancement After Cryoablation of Renal Cell Carcinoma: A Prospective Study. AJR Am J Roentgenol 2016;206:332-9.
- 74. Porter CAt, Woodrum DA, Callstrom MR, et al. MRI after technically successful renal cryoablation: early contrast enhancement as a common finding. AJR Am J Roentgenol 2010;194:790-3.
- 75. Borghesi M, Brunocilla E, Volpe A, et al. Active surveillance for clinically localized renal tumors: An updated review of current indications and clinical outcomes. Int J Urol 2015;22:432-8.
- 76. Mason RJ, Abdolell M, Trottier G, et al. Growth kinetics of renal masses: analysis of a prospective cohort of patients undergoing active surveillance. Eur Urol 2011;59:863-7.
- 77. Patel N, Cranston D, Akhtar MZ, et al. Active surveillance of small renal masses offers short-term oncological efficacy equivalent to radical and partial nephrectomy. BJU Int 2012;110:1270-5.
- 78. Pierorazio PM, Johnson MH, Ball MW, et al. Five-year analysis of a multi-institutional prospective clinical trial of delayed intervention and surveillance for small renal masses: the DISSRM registry. Eur Urol 2015;68:408-15.
- 79. Van Poppel H, Becker F, Cadeddu JA, et al. Treatment of localised renal cell carcinoma. Eur Urol 2011;60:662-72.
- 80. Yang G, Villalta JD, Meng MV, Whitson JM. Evolving practice patterns for the management of small renal masses in the USA. BJU Int 2012;110:1156-61.
- 81. Haramis G, Mues AC, Rosales JC, et al. Natural history of renal cortical neoplasms during active surveillance with follow-up longer than 5 years. Urology 2011;77:787-91.
- 82. Nayyar M, Cheng P, Desai B, et al. Active Surveillance of Small Renal Masses: A Review on the Role of Imaging With a Focus on Growth Rate. J Comput Assist Tomogr 2016;40:517-23.
- 83. Tan WS, Trinh QD, Hayn MH, et al. Delayed nephrectomy has comparable long-term overall survival to immediate nephrectomy for cT1a renal cell carcinoma: A population-based analysis. Urol Oncol 2020;38:74 e13-74 e20.
- 84. Uzosike AC, Patel HD, Alam R, et al. Growth Kinetics of Small Renal Masses on Active Surveillance: Variability and Results from the DISSRM Registry. J Urol 2018;199:641-48.
- 85. Patel HD, Nichols PE, Su ZT, et al. Renal Mass Biopsy is Associated with Reduction in Surgery for Early-Stage Kidney Cancer. Urology 2020;135:76-81.
- 86. Doshi AM, Huang WC, Donin NM, Chandarana H. MRI features of renal cell carcinoma that predict favorable clinicopathologic outcomes. AJR Am J Roentgenol 2015;204:798-803.

- 87. Rosenkrantz AB, Mussi TC, Somberg MB, Taneja SS, Babb JS. Comparison of CT-based methodologies for detection of growth of solid renal masses on active surveillance. AJR Am J Roentgenol 2012;199:373-8.
- 88. Quaia E, Bertolotto M, Cioffi V, et al. Comparison of contrast-enhanced sonography with unenhanced sonography and contrast-enhanced CT in the diagnosis of malignancy in complex cystic renal masses. AJR Am J Roentgenol 2008;191:1239-49.
- 89. Barr RG, Peterson C, Hindi A. Evaluation of indeterminate renal masses with contrast-enhanced US: a diagnostic performance study. Radiology 2014;271:133-42.
- 90. Zarzour JG, Lockhart ME, West J, et al. Contrast-Enhanced Ultrasound Classification of Previously Indeterminate Renal Lesions. J Ultrasound Med 2017;36:1819-27.
- 91. American College of Radiology. ACR Appropriateness Criteria[®] Radiation Dose Assessment Introduction. Available at: <u>https://www.acr.org/-/media/ACR/Files/Appropriateness-Criteria/RadiationDoseAssessmentIntro.pdf</u>. Accessed September 30, 2021.

The ACR Committee on Appropriateness Criteria and its expert panels have developed criteria for determining appropriate imaging examinations for diagnosis and treatment of specified medical condition(s). These criteria are intended to guide radiologists, radiation oncologists and referring physicians in making decisions regarding radiologic imaging and treatment. Generally, the complexity and severity of a patient's clinical condition should dictate the selection of appropriate imaging procedures or treatments. Only those examinations generally used for evaluation of the patient's condition are ranked. Other imaging studies necessary to evaluate other co-existent diseases or other medical consequences of this condition are not considered in this document. The availability of equipment or personnel may influence the selection of appropriate imaging procedures or treatments. Imaging techniques classified as investigational by the FDA have not been considered in developing these criteria; however, study of new equipment and applications should be encouraged. The ultimate decision regarding the appropriateness of any specific radiologic examination or treatment must be made by the referring physician and radiologist in light of all the circumstances presented in an individual examination.