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# Percutaneous Tumor Ablation Tools: Microwave, Radiofrequency, or Cryoablation—What Should You Use and Why?<sup>1</sup>

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**Abbreviations:** HCC = hepatocellular carcinoma, RCC = renal cell carcinoma, RF = radiofrequency

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#### SA-CME LEARNING OBJECTIVES

After completing this journal-based SA-CME activity, participants will be able to:

• Describe the physical properties of the most commonly used thermal ablation modalities.

Discuss the potential advantages and disadvantages of each thermal ablation modality.

■ Apply decision-making algorithms to select an appropriate thermal ablation modality for common clinical indications.

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**TEACHING POINTS** See last page Image-guided thermal ablation is an evolving and growing treatment option for patients with malignant disease of multiple organ systems. Treatment indications have been expanding to include benign tumors as well. Specifically, the most prevalent indications to date have been in the liver (primary and metastatic disease, as well as benign tumors such as hemangiomas and adenomas), kidney (primarily renal cell carcinoma, but also benign tumors such as angiomyolipomas and oncocytomas), lung (primary and metastatic disease), and soft tissue and/or bone (primarily metastatic disease and osteoid osteomas). Each organ system has different underlying tissue characteristics, which can have profound effects on the resulting thermal changes and ablation zone. Understanding these issues is important for optimizing clinical results. In addition, thermal ablation technology has evolved rapidly during the past several decades, with substantial technical and procedural improvements that can help improve clinical outcomes and safety profiles. Staying up to date on these developments is challenging but critical because the physical properties underlying the different ablation modalities and the appropriate use of adjuncts will have a tremendous effect on treatment results. Ultimately, combining an understanding of the physical properties of the ablation modalities with an understanding of the thermal kinetics in tissue and using the most appropriate ablation modality for each patient are key to optimizing clinical outcomes. Suggested algorithms are described that will help physicians choose among the various ablation modalities for individual patients.

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## Introduction

Image-guided thermal ablation is becoming increasingly accepted for the treatment of certain benign and malignant tumors of the lungs, liver, kidneys, bone, and soft tissues (1–6). Numerous thermal and nonthermal ablation modalities are available, including radiofrequency (RF) ablation, microwave ablation, cryoablation, high-intensity focused ultrasonography (US), laser ablation, irreversible electroporation, chemical ablation (with ethanol and acetic acid), and brachytherapy. This article focuses on the thermal ablation devices used most frequently in the United States: RF ablation, microwave ablation, and cryoablation devices. The number of available ablation modalities, the rapid changes in the associated technology (especially recently), the relative lack of randomized controlled or comparative trials, the wide discrepancies in published results (with associated difficulties in interpreting the results), and the variability of practice in different centers have led to confusion about best practice in patients. In this article, some of the issues are elucidated, and the present and future of image-guided tumor ablation are clarified.

The mechanisms of action of various ablation modalities are at the center of many of the relative advantages, disadvantages, and limitations encountered in clinical practice. A basic understanding of the underlying physical processes is critical to determining the most advantageous modality. Therefore, we start by discussing the physical properties of the various modalities and then address the reasons why particular technologies may succeed or fail when applied clinically.

## Physical Properties of Thermal Ablation

#### **Heat-based Ablation**

Heat has long been recognized as a mechanism for inducing cellular damage. The relationship between cellular temperatures and cellular death is a complex one affected by temperature, time of exposure, and tissue-level effects related to such factors as perfusion, vascular coagulation, adjunctive treatments, and cell fragility (7). For tumor ablation, however, the typical goal is to reach temperatures higher than 55°C in the target tissue to create coagulation necrosis. Temperatures would ideally decrease rapidly outside the target volume to avoid unwanted collateral damage.

**RF Ablation.**—The heating mechanism of RF ablation has been described in detail elsewhere (8). Briefly, an alternating electric current is applied to the target tissue through use of an interstitial electrode; in monopolar RF ablation, the current returns through grounding pads on the skin surface. The electric current rapidly oscillates tissue ions, creating frictional heating in areas of high current density adjacent to the electrode. Additional growth of the ablation zone can be attributed primarily to thermal diffusion (9).

Two primary issues associated with this heating mechanism have limited the ability of RF ablation to effectively treat the growing number and diversity of tumors being considered for minimally invasive therapies. First, heating tissue with electric current is a self-limiting process. Water vapor, desiccation, and charring created during RF ablation progressively increase tissue impedance. An increase in impedance limits the application of additional current and, hence, limits temperatures

that can be achieved in the ablation zone. Second, conductive heat transfer is relatively slow in most tissues and in the in vivo setting has limited ability to overcome relatively minor competing processes, such as perfusion and ventilation. As a result, RF ablation zones can vary widely according to the local tissue environment (10,11). For example, aerated lung is characterized by high impedance to electric current flow and poor heat transfer compared with most solid organs. The high impedance of normal lung tissue surrounding a tumor can reduce power delivery, whereas poor thermal diffusion limits growth of the ablation zone (12,13). Some of these technical limitations can be overcome by increases in power and time, but, overall, RF ablation is a suboptimal choice for ablation of lung tumor (14).

Numerous attempts have been made to overcome the physical limitations of RF ablation, with varying degrees of success. Developments that have improved the size of the ablation zone include the following: (a) internal electrode cooling to limit charring in the tissues immediately adjacent to the electrode (15); (b) expandable, multi-tined, or clustered electrodes to increase the electrode surface area and facilitate greater power delivery (16); and (c) rapidly switching multiple-electrode approaches that allow the simultaneous production of separate or overlapping ablation zones (17–19). Other innovations include the use of saline infusion and high-power generators (20). Unfortunately, although several of these techniques have improved the overall performance of RF ablation, many are associated with an increased risk for complications related to an increase in applicator size or number, increased current with the potential for skin burns, and escalation in procedural complexity and the probability of collateral damage.

Microwave Ablation.—Cell death in microwave ablation is nearly identical to that observed in RF ablation (21). However, the mechanism of microwave heating may have advantages in certain clinical applications. Polar molecules (primarily water) continuously realign with the oscillating microwave field, effectively increasing kinetic energy and tissue temperature. In contrast to electric currents, microwaves radiate through all biological tissues, including those with high impedance to electric current, such as bone, lung, and charred or desiccated tissues (22). This allows microwaves to continuously generate heat in a much larger volume of tissue surrounding the applicator (23). As a result, microwave energy can produce faster, hotter, and larger ablation zones in multiple tissue types compared with RF current (Fig 1). The clinical result is that, on

Figure 1. Graph shows the comparison of temperatures obtained with a third-generation microwave ablation system versus a similar RF ablation system. Microwave energy produces faster and hotter ablation zones than does RF current.

Figure 2. Diagram compares the electromagnetic fields induced by three different microwave ablation antenna designs. The differences in electromagnetic fields directly affect the size and shape of the resulting ablation zone. (Scale is in watts per kilogram.)





average, fewer applicators are needed, and ablative margins are easier to obtain.

Microwave systems can be divided into three types: (a) First-generation systems lack active antenna cooling and are therefore limited to low power and short durations; (b) second-generation systems have antenna cooling but limited generator power; and (c) third-generation systems incorporate antenna cooling and highpower generators. Each system is characterized by a different combination of antenna diameter, number of antennae, frequency and phase control, generated power, and power loss between the generator and applicator tip. System performance can vary widely, so it is critical that physicians understand the ablation-zone shapes and sizes created by different time and power combinations with a particular system.

The volume of microwave ablation zones depends on the power applied, antenna design, number and orientation of antennae, and microwave frequency.

Microwave antenna design is a critical factor that affects the size and shape of the ablation zone, and dozens of antennae have been described to control energy delivery (Fig 2)

(24,25). Many antennae produce an elongated ablation zone (as long as 6 cm) that may increase the risk for burns to the body wall and other structures, so newer designs aim for more rounded and forward-weighted heating that is amenable to treating smaller tumors (Fig 3) (26,27). Some systems also include antenna designs to transfer energy more optimally to different organ types, such as the lungs (28). As with other ablation modalities, multiple antennae can also be used in concert to create larger, more conformal and confluent ablation zones (29,30). In addition, the driving phase of each antenna can be controlled to create zones of "constructive" and "destructive" wave interaction that can have a profound effect on the ablation result (31 - 33).

Frequency is another issue that differentiates systems. Although investigators in previous studies have suggested that 915 MHz may create larger ablation zones than can 2.45 GHz (because of increased energy penetration), several acknowledged limitations make those conclusions suspect (34). Most notably, these investigators did not control for power delivered to the tissue, and the assumption that penetration is greater at

RadioGraphics



**Figure 3.** Computed tomographic (CT) images in a 77-year-old man with cirrhosis and a 2.6-cm hepatocellular carcinoma (HCC) in the right hepatic lobe. The HCC was targeted for microwave ablation with a single, specially designed "precision" 17-gauge gas-cooled antenna at 65 W for 7.5 minutes. The resulting ablation zone (arrows) shown on axial (a) and coronal (b) contrast material–enhanced postablation CT images is round and measures approximately 3.4 cm in all dimensions. Microwave ablation allowed destruction of the HCC with minimal associated damage to the hepatic parenchyma. Note the artificial ascites used to protect the diaphragm (arrowhead in **b**).

lower frequencies is not valid in the near-field of microwave antennae (35). Adequately powered 915-MHz and 2.45-GHz systems can create relatively large ablation zones, but the longer wavelength at 915 MHz typically results in a longer ablation zone (28,34,36–38). In general, the greatest heating from any system occurs within 1 cm of the antenna.

Overall, the potential advantages of microwave over RF ablation include faster ablations (typically 2-8 minutes with high-powered systems), higher temperatures without the limitations related to electric impedance, less sensitivity to tissue type with more consistent results (which may be beneficial in treating some tumors that are resistant to treatment with RF, such as sarcomas and hemangiopericytomas), a relative insensitivity to "heat sinks" compared with RF, and the ability to create much larger ablation zones if needed (ablation zones as long as 8 cm have been achieved without applicator repositioning) (22,25,33,39,40). The continued challenges of microwave ablation are as follows: (a) Because microwave ablation is relatively new, clinical data and experience are minimal; (b) a learning curve is associated with using microwave ablation safely because of the potentially larger ablation zones compared with RF ablation; and (c) the clinical systems are heterogeneous in terms of antenna design, wavelength, frequency, power, and cooling, which leads to different performance characteristics and confusion regarding the interpretation of clinical results and the predictability of results between systems from different manufacturers.

#### Cryoablation

The mechanism of cell death with cold temperatures is different from that with heat. The freezing process results in both intracellular and extracellular ice formation, both of which can result in cellular death, but by different mechanisms. The location of ice formation and, therefore, the mechanism of cell death vary with the freezing rate and final tissue temperature. Faster freezing to lower temperatures promotes the formation of intracellular ice crystals, which results in cell death because of direct damage to the cell membrane and organelles (41). Slower rates of freezing favor the formation of extracellular ice crystals, which leads to a change of osmolality within the extracellular space and, in turn, to cell dehydration and death (42). In addition, apoptosis induced by freezing contributes to this process, a role that is poorly understood (43). The mechanism of cell death with freezing is associated with cell membrane disruption and an associated release of intracellular contents. Because the ablation zone is reperfused after the ice ball melts, the result is a rapid release of cellular debris into the systemic circulation. This probably explains the systemic complications that are seen with cryoablation (ie, cryoshock) but are rare with heat-based ablation (44,45). The temperature necessary to cause reliable cellular necrosis depends on the cell type and thermal history of the tissue, with

estimates of  $-35^{\circ}$ C to  $-20^{\circ}$ C (46,47). The lower estimates suggest that cell death may only occur approximately 8 mm deep to the edge of the visualized ice ball; although there is a phase change from liquid to solid at the 0°C isotherm along the edge of the ice ball, this phase change is not necessarily associated with cell death (48). One of the biggest benefits of cryoablation is that the expanding ice ball (predictive of the ablation zone) is well visualized at US, CT, and magnetic resonance imaging, thus allowing relatively more precise monitoring of the ablation zone than is possible with many heat-based systems (48,49).

Cryoablation systems use the Joule-Thomson theory of expanding gases within a needlelike cryoprobe. As the cryogen (typically argon) moves from an internal feed line into an internal expansion chamber, it produces a heat sink near the antenna tip that cools the probe to temperatures of -160°C or colder. Heat transfer from the tissue into the cryoprobe is governed by passive thermal diffusion; unlike RF and microwave ablation devices, cryoablation devices provide no zone of direct or active cooling. As a result, the surface area of the cryoprobe limits cooling efficiency; smaller cryoprobe diameters are associated with lower cooling capacity and, consequently, smaller ablation zones. Therefore, several cryoprobes are required to treat most tumors in clinical practice, and ablation times are typically on the order of 25-30 minutes, much longer than the average times seen with microwave ablation (approximately 5 minutes) or RF ablation (12-30 minutes) (50).

#### **Clinical Decision Making**

Arguably the most critical component for a successful ablation practice is the operator. The importance of excellent training and a thorough understanding of the indications, imaging options, techniques, complications, ablation modalities, and specific system being used cannot be stressed enough. In addition, the ability to accurately place needles by using US and CT guidance is paramount, as is a thorough knowledge of the expected appearance of successful and unsuccessful ablations and complications at immediate postprocedural imaging.

In deciding which ablation device to use for a specific patient, many factors should be considered, not the least of which is availability. The unfortunate reality is that many institutions have only one ablation system available. Nonetheless, it is optimal to be familiar with and have available both a heat- and a cold-based system. Herein, we discuss a variety of algorithms for determining which modality may be optimal for a given patient, but some rules of thumb to keep in mind

include the following: (a) The preponderance of the data suggests that current microwave systems have inherent advantages over RF ablation and that these advantages can give similar or superior results in the clinical setting in less time (note that the increased power does come with the potential for increased complications related to collateral damage as well, but thus far, this has not proven to be the case in clinical practice [51,52]); (b) the organ involved, tumor type, and underlying comorbid conditions can affect the safety and effectiveness of different modalities; (c) the precision that is possible with cryoablation can be important for treatment of tumors adjacent to vulnerable structures (49,53); and (d) vascular mediated cooling can dramatically affect the size and shape of the ablation zone and should be considered in determining both the modality and the positioning and number of applicators (54–56).

In determining the optimal ablation modality for a particular case, the decision usually comes down to "heat versus cold." If heat is selected, then our group generally favors microwave ablation because of the associated benefits of decreased procedure time, larger ablation zones (leading to use of fewer applicators), higher temperatures, less influence from vascular heat sinks, and increased predictability (23,57). If microwave ablation is not available, or if the available system is suboptimal, then RF ablation can be used effectively; however, its relative limitations should be understood. The decision of whether to use a heat- or a cold-based ablation modality can be a complex one, and in our practice, we base the decision on organ-specific considerations, tumor type, tumor location (especially proximity to vulnerable structures), and patient factors (eg, comorbid conditions). We describe these considerations according to organ system.

### **Organ-specific Considerations**

**Liver.**—In the liver, the primary indications for image-guided tumor ablation are HCC and metastatic disease. In addition, ablation modalities are increasingly used to treat benign hepatic masses, including giant hemangiomas, hepatic adenomas, and cysts (58–62). The indications for treatment of benign tumors include controlling mass-related symptoms and, in the case of hepatic adenomas, preventing tumoral hemorrhage and potential malignant degeneration (63,64). The option of treating benign hepatic tumors with percutaneous ablation rather than with open or laparoscopic surgical resection (procedures with increased morbidity compared with ablation) should be presented to patients when applicable.



Teaching

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5b.

**Figures 4, 5.** (4) CT images in a 64-year-old male patient with cirrhosis and HCC. (a) Axial arterial phase biphasic CT image shows a 1.8-cm HCC in the right hepatic lobe (arrows). This was targeted for microwave ablation with a single 17-gauge gas-cooled antenna. A 3-minute ablation was performed at 90 W. (b) Axial nonenhanced CT image obtained during ablation shows that the mass has essentially been replaced by gas (arrow). (c) Axial CT image obtained immediately after ablation shows a  $3.0 \times 2.8$ -cm ablation zone (arrows), with complete ablation of the tumor and only the expected benign periablational enhancement (arrowhead). The high attenuation seen centrally in the ablation zone is desiccated dense tissue, which is more prominent with microwave ablation than with RF ablation. (5) CT images in a 55-year-old male patient with cirrhosis and HCC. (a) Axial nonenhanced CT image obtained during transarterial chemoembolization shows a  $5.4 \times 5.2$ -cm HCC in the right hepatic lobe, with intense uptake of ethiodized oil (arrows) and less intense nontarget uptake in the remainder of the liver. The tumor was targeted for microwave ablation by using US guidance. Three 17-gauge gas-cooled microwave antennae were positioned in a triangular configuration within the tumor, and 65 W was applied to all antennae simultaneously, for a total ablation time of 10 minutes. (b) Sagittal reconstructed contrast-enhanced CT image obtained immediately after ablation shows that the ablation was technically successful, with an adequate circumferential ablative margin (arrows). Note the necrosis in the nontarget liver related to transarterial chemoembolization (arrowhead).

Heat is the preferred method for treating HCC in patients with cirrhosis because of the increased bleeding risk and disseminated intravascular coagulation-like reaction, called *cryoshock*, associated with cryoablation (44,65,66). Heat-based ablation provides improved hemostasis compared with cryoablation and essentially no risk for cryoshock (67). Therefore, although many studies have shown that smallvolume cryoablation is feasible in patients with cirrhosis and HCC, it is difficult to justify the additional risk of cryoablation in these patients when viable heat-based alternatives are available (44). Small HCCs (up to 2–3 cm in diameter) are effectively treated with RF or microwave ablation alone, depending on tumor location. Treatment of larger tumors (>3.0 cm) with RF or microwave monotherapy is associated with high rates of local recurrence (68). Combination treatment of large HCCs (>3.0 cm in diameter) with transarterial chemoembolization followed by RF or microwave ablation results in fewer



Figure 6. CT images in a symptomatic 37-year-old woman. (a) Axial contrast-enhanced portal venous phase CT image shows an  $8.8 \times 8.7$ -cm giant hemangioma in the right hepatic lobe (arrows). (b) Sagittal unenhanced CT image shows the hemangioma targeted for microwave ablation, with three 17-gauge gas-cooled microwave antennae distributed evenly in the hemangioma (arrows). (c) Axial contrast-enhanced CT image obtained after 15 minutes of ablation at 65 W shows complete ablation of the tumor, with a small margin of normal hepatic tissue (arrow). Note the hydrodissection fluid (arrowhead), which was used to protect the abdominal wall and diaphragm.

local failures and is well tolerated, with few serious adverse effects (Figs 4, 5) (69,70).

For patients with normal background liver, heat-based ablation or cryoablation can be considered. The results in terms of tumor control and survival are expected to be similar, but cryoablation is associated with an increased risk for complications (67). In our experience, cryoablation is rarely needed in the liver and is limited to central tumors for which precision is critical because of proximity to important bile ducts that cannot be sacrificed. Some data indicate that cryoablation is less damaging to ducts than are heat-based therapies (71,72). In general, larger tumors that may have previously been treated only with cryoablation (because of the ability to place numerous cryoprobes simultaneously) can now reasonably be treated with microwave ablation because third-generation systems can produce large confluent ablation zones (Fig 6).

Of note, the absence of a specific Current Procedural Terminology code for microwave ablation somewhat limited early adoption of this modality. However, both the American College of Radiology and the Society of Interventional Radiology have subsequently addressed this issue and specifically do not recommend the use





c.

of unlisted procedure codes. Rather, because microwaves are part of the RF spectrum, they recommend the use of the appropriate RF ablation code based on anatomic location.

A proposed decision-making algorithm for liver ablation is shown in Figure 7.

Kidney.—Most tumors treated with ablation are renal cell carcinomas (RCCs), although a subset of patients with benign disease, such as oncocytoma and angiomyolipoma, may benefit from ablation for symptom control and prevention of complications (eg, mass effect and tumor-related hemorrhage) (73,74).

There are many ways to successfully ablate a renal tumor, and both RF ablation and cryoablation have proven successful for renal ablation (75). However, the most appropriate modality to use in the kidney depends on tumor size, tumor location, and proximity of vulnerable structures. As seen with the liver, microwave ablation has several theoretical advantages that make it our modality of choice for many renal tumors (57). With small exophytic masses, microwave ablation has the advantage of speed and the need for fewer applicators (Fig 8). However, central tumors and tumors adjacent to vulnerable struc-



Figure 7. Flowchart shows the recommended modality choices for liver ablation. Cryo = cryoablation, MW =microwave.





c.

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Figure 8. CT and US images in a 63-year-old male patient with RCC. (a) Axial contrast-enhanced nephrographic phase CT image shows a 2.9-cm posterior exophytic RCC (arrow) that was confirmed at biopsy. The lesion was targeted for microwave ablation by using a single 17-gauge gas-cooled microwave antenna with US guidance. (b) Transverse US image during a 3-minute ablation with 90 W shows complete replacement of the tumor by hyperechoic gas bubbles (arrow). (c) Axial contrast-enhanced nephrographic phase postablation CT image shows complete ablation of the lesion, with an adequate ablative margin (arrows). Note the nonenhancing "tumor ghost" in the ablation zone (arrowhead).

tures may benefit from the precision and relative sparing of the collecting system and ureter that are associated with cryoablation (71,72,76). However, recall that the zone of cellular necrosis does not extend to the margin of the ice ball

and, depending on multiple factors, may extend to 8 mm inside of the edge of the ice ball (50). Therefore, defining margins and performing a complete treatment may be more challenging than is generally appreciated. Tumors larger than 3-4 cm can be difficult to completely treat with ablation. These tumors may benefit from



c.

Figure 9. CT images in a 69-year-old woman with biopsy-proven RCC. (a, b) Axial CT images obtained with the patient in the supine (a) and decubitis (b) positions show a  $4.0 \times 3.5$ -cm RCC in the left anterior kidney (arrows). The pancreas (arrowhead) is closely adjacent to the RCC in **a**, which increases the risk associated with heat-based ablation, and the relationship does not change in **b** with the patient in the decubitus position. The RCC was targeted for cryoablation with US guidance. (c) Axial contrast-enhanced CT image shows that hydrodissection (high-attenuation fluid) was used to displace the pancreas (arrowhead), and the RCC was ablated successfully with three 15-gauge cryoprobes and a 10-, 5-, 10-minute freeze-thaw-freeze protocol. Note that the visibility of the ice ball (arrows) makes the ablation precise. (d) On an axial 6-month follow-up CT image, the RCC (arrows) shows no substantial enhancement and has decreased in size, as expected. The patient had no evidence of pancreatitis or other complications.

the larger ablation zones that are possible with both microwave ablation and cryoablation; therefore, the choice of modality may hinge on the ability to perform the ablation safely based on the proximity of adjacent structures or the ability to hydrodissect effectively (Fig 9). Note that hydrodissection plays a much larger role in renal ablations than in hepatic ablations because of the relatively close proximity of the bowel, pancreas, and ureter. In fact, more than 50% of our renal ablations require hydrodissection to allow safe ablation (77). The treatment of tumors adjacent to the ureter may benefit from the placement of a ureteral stent and pyeloperfusion in addition to hydrodissection.

The primary indication for the treatment of renal angiomyolipomas is to decrease the associated risk for future hemorrhage (73,74). In this setting, we prefer heat-based ablation because of the associated cautery effect, even with relatively

large vessels. In our experience, RF is limited by the high impedance associated with these tumors, presumably related to the insulative effects of fat. When ablation of angiomyolipomas is performed, the solid soft-tissue nodules and larger feeding vessels can be specifically targeted and ablated, markedly reducing the risk for future hemorrhage (Fig 10).

In summary, renal tumors smaller than 3 cm in diameter can be ablated successfully with both RF ablation and cryoablation. Emerging data on microwave ablation for RCC is promising and is the preferred method in our practice for treating peripheral renal tumors because of speed, high temperatures, and cautery. For central renal tumors, we favor cryoablation because of the theoretical decreased risk for injury to the collecting system and ureter.

A proposed decision-making algorithm for kidney ablation is shown in Figure 11.



Figure 10. CT images in a 55-year-old woman with a history of right renal angiomyolipoma with associated massive retroperitoneal hemorrhage that required immediate embolization. (a) Coronal reconstructed image from follow-up contrast-enhanced CT shows continued avid enhancement of the lesion (arrows). Note the closely adjacent colon (\*). After successful hydrodissection for displacement of the colon, the lesion was targeted for microwave ablation with three 17-gauge gas-cooled antennae at 65 W for 5 minutes. (b) Coronal reconstructed image from contrast-enhanced CT 4 months after ablation shows minimal, if any, residual enhancement of the lesion (arrows). The patient has remained asymptomatic.



Figure 11. Flowchart shows the recommended modality choices for kidney ablation. AML = angiomyolipoma, Cryo = cryoablation, MW = microwave.

Lung.-Most patients are referred for imageguided ablation of nonoperable non-small-cell lung carcinoma or metastases to the lung, wherein tumor eradication would increase survival or control symptoms. The choice between stereotactic body radiation therapy and tumor ablation for T1a non-small-cell lung carcinoma remains controversial, and few investigators have compared these two modalities (78). Patients with non-small-cell lung carcinoma often have associated cardiopulmonary disease due to a long smoking history, and percutaneous proce-

dures in patients with emphysema are associated with a higher complication rate than that seen in patients with normal background lung tissue. However, most studies of percutaneous RF ablation, microwave ablation, and cryoablation in patients with non-small-cell lung carcinoma demonstrate low rates of serious complications (79, 80).

Most studies of tumor ablation in the lung have concerned RF ablation, although several investigators have assessed microwave ablation and cryoablation (4,79,81). The results with

Figure 12. CT images in a 54-year-old woman with a history of metastatic colorectal cancer. (a) Axial CT image shows a 6-mm biopsy-proven metastatic focus in the left lower lobe of the lung (arrow) that was targeted for RF ablation. A single 17-gauge water-cooled electrode was positioned in the medial aspect of the nodule, and a 12-minute ablation was performed. (b) Axial CT image obtained 6 months after ablation shows rapid local tumor progression, with the metastatic focus now measuring 1.6 cm. This area was targeted for repeat RF ablation by using a water-cooled cluster electrode. The repeat ablation resulted in long-term local control but was complicated by a large hemothorax that ultimately required surgical intervention. The hemothorax was at least partially related to the increased invasiveness of the cluster electrode.

RF ablation in the lung have been mixed, perhaps because of the inherent limitations to the conduction of electric current in the presence of aerated lung tissue. Air is an insulator, resulting in high electric impedance that limits current flow and tissue heating. These physical limitations can produce poor clinical results, even with correctly placed electrodes in small tumors (Fig 12) (82). Cryoablation can effectively ablate tumors in the lung and has the advantage of being relatively resistant to the cold-sink effects of ventilation, compared with RF ablation (83,84). The effect has not been studied in microwave ablation as of yet, but according to animal studies, microwave ablation seems to be less susceptible to these effects and creates a larger and more confluent zone of ablation compared with RF ablation and cryoablation (28). Another advantage of cryoablation is seen with tumors that are relatively central or closely adjacent to or invasive into the mediastinum or chest wall (Fig 13). Although patients who have undergone previous radiation therapy and experience recurrent or new disease in the radiation field are part



b.

of an interesting potential patient population for this procedure, the ablation zone may be larger than expected in these patients (probably because of abnormal perfusion), and the complication rate may be significantly higher (because of disordered vasculature and poor tissue healing). Therefore, although these patients may benefit from ablation, it is important to be aware of the increased risk. Cryoablation is associated with minimal, if any, postprocedural pain, even when the ablation extends into the chest wall. The relative rate of complications and the oncologic effectiveness of the various modalities have not been established definitively.

Microwave ablation has the advantage of creating predictable and large ablation zones, even in the setting of aerated lung (28). The advantage of a faster procedure is also true in the lungs—most ablations require approximately 5 minutes after antenna placement. Physicians should keep in mind several important considerations when performing RF and microwave ablation of the lungs. First, unlike standard lung biopsy technique, in which the shortest possible



b.

Teaching

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Figure 13. Imaging findings in a 71-year-old woman with a history of non-small-cell lung carcinoma who had undergone radiation therapy 2 years previously. (a) Axial positron emission tomography (PET)/CT image shows recurrent disease in the radiation field (arrows). The patient was referred for ablation. Cryoablation was chosen because of the close proximity of the mass to the mediastinum and the associated risk for injury to the phrenic nerve. (b) Axial CT image shows the four cryoprobes (arrow) used to perform the ablation, which involved a 3-, 5-, 7-, 5-, 5-minute freeze-thaw protocol. This resulted in excellent coverage with the ice ball (arrowhead). Phrenic nerve palsy developed, with associated diaphragmatic paralysis. Combined with poor baseline lung function, this complication resulted in a prolonged postablation hospitalization. Within 5 weeks, however, both diaphragmatic function and respiratory status returned to baseline levels. (c) Axial nonenhanced CT image obtained 18 months after ablation shows calcification of the treated mass (arrows), with no evidence of local or distant recurrence.

path to the tumor is often desirable, a longer approach that traverses normal lung is better for two reasons: (a) The relatively long ablation zone associated with certain microwave ablation devices (as long as 6 cm, depending on the antenna design) means that a direct puncture can be associated with ablation zones that extend into the chest wall, resulting in pain and potential skin burns, and (b) the ablated tumor or lung becomes dehydrated and noncompliant, which, in combination with volume loss, can result in retraction of the antenna or electrode insertion site from the pleural surface, potentially resulting in persistent air leaks and bronchopleural fistulas (85). Therefore, an indirect approach that leaves an unablated tract of at least 2 cm of aerated lung is preferred (Fig 14). Note that tract cautery should not be performed



A proposed decision-making algorithm for lung ablation is shown in Figure 16.

Musculoskeletal System.—Pain is the most common indication for percutaneous ablation of musculoskeletal tumors. The use of ablation is widely accepted for both painful disease metastatic to bone and osteoid osteomas (5,86,87). Soft-tissue ablation is most frequently performed





#### b.

Figure 14. CT images in a 65-year-old female patient with a history of non-small-cell lung cancer. (a) Axial nonenhanced CT image shows an 8-mm focus of recurrent disease in the right upper lobe of the lung. Although it would be reasonable to perform a biopsy of the lesion with an approach perpendicular to the pleural surface (black arrow), thus minimizing the lung parenchyma traversed, or with a more tangential approach, it is important with microwave ablation to use an approach that allows for an adequate tract within the lung parenchyma (white arrow). This minimizes the likelihood of a persistent air leak or bronchopleural fistula. (b) Axial nonenhanced CT image shows the area of recurrent disease that was targeted for microwave ablation with CT guidance. A single 17-gauge gas-cooled microwave ablation antenna was positioned in the nodule (arrow), and a 5-minute ablation at 40-55 W was performed. Note the area of scarring from a microwave ablation performed 11 months previously (arrowhead). (c) Axial nonenhanced CT image shows that the procedure resulted in technically successful ablation, with complete coverage of the nodule by ground-glass opacity (arrow). A pneumothorax that developed at ablation was treated successfully during the procedure with a pleural blood patch. A chest tube was not required.

for metastatic disease and for palliative or curative intent, depending on the scenario (86).

Metastatic disease to the bone with associated intractable pain is regrettably common. Although radiation treatment is often considered the first-line therapy, it is expensive, inconvenient (requiring numerous treatments), and associated with a delay in onset of effect. In addition, repeat treatment with radiation is often limited by dose, whereas all ablation modalities can be repeated as needed. This and other factors make ablation an attractive alternative, and its effectiveness has been evaluated in many trials. The results of these trials suggest that although both heat-based abla-

tion and cryoablation can be effective, cryoablation is associated with more rapid and durable pain relief. Therefore, cryoablation should be preferred in this setting and can be associated with exceptional results (Fig 17) (86,87).

Percutaneous ablation of osteoid osteomas is now considered standard therapy because of a high success rate with minimal complications (5). Treatment of osteoid osteomas is almost always performed with RF ablation because of the ability to create a small and short ablation zone (approximately 1 cm), which is adequate for optimal outcomes and results in almost no associated collateral damage. Percutaneous ablation for osteoid osteoma

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Figure 15. CT images in a 73-year-old man with a history of non-small-cell lung carcinoma who had undergone several resections. (a) Axial CT image shows a  $2.0 \times 1.2$ -cm PET-positive, biopsy-proven focus of recurrent disease (arrow). The area was targeted for microwave ablation with a single 17-gauge gas-cooled antenna at 140 W for 10 minutes. The longer time and high power were used because of previous difficulties with achieving complete ablation, even with small tumors, by using RF ablation. (b) Axial CT image shows the resulting nearcomplete vaporization of the small tumor and a  $6.4 \times 3.5$ -cm ablation zone (arrowheads). (c) Axial CT image obtained 5 months after ablation shows no evidence of local tumor progression, but the aggressive ablation led to the formation of a large area of parenchymal cavitation (arrows). This was associated with hemoptysis that lasted more than 6 months. As a result of this experience and subsequent animal studies, we have decreased the power and time used for microwave ablation of the lung.





in certain anatomic situations. Heat-based ablation does not allow as definitive a delineation of the ablation zone and can therefore be associated with damage to adjacent vulnerable structures (Fig 19). The advantages of microwave ablation for the treatment of soft-tissue tumors include speed, the ability to create a large ablation zone, and the ability to heat and ablate nearly every tumor type. This characteristic of microwave ablation may be particularly important in the treatment of fibrous tumors, such as sarcomas, which may be resistant to cryoablation, and tumors with closely adjacent bone or vascular heat sinks (Fig 20).

#### Conclusion

Image-guided tumor ablation is becoming increasingly accepted for treatment of both benign and malignant tumors in a variety of organs. However, the large number of percutaneous ablation tools that are now clinically available has led to confusion about which device should be used to treat particular tumors. In general, we favor the use of high-powered microwave ablation for treatment of tumors in the liver, periphery of the kidney, and lung. The use of any heat-based ablation modality in the lung can be associated with prolonged air leaks or bronchopleural fistulas, and, thus, the operator must avoid both tract

can even be performed in difficult-to-access locations with minimal morbidity (Fig 18). Microwave ablation may have a role in the future as antennae with shorter ablation zones become available.

For soft-tissue ablation, the decision about which modality to use primarily depends on the location of the tumor and adjacent structures. Encompassing a nerve in a cryoablation zone is associated with regeneration of nerve function over time (88). As is seen with ablation in other tissues, the visibility of the ice ball can be an advantage



Figure 16. Flowchart shows the recommended modality choices for lung ablation. Cryo = cryoablation, MW = microwave, ptx = pneuomothorax.



Figure 17. CT images in a 48-year-old man with a history of non-small-cell carcinoma of the lung who presented with a painful area of metastasis to the right iliac wing. (a) Axial nonenhanced CT image obtained before ablation shows a lytic expansile mass in the right iliac wing (arrows). Note the metastasis in the left posterior iliac wing (arrowhead), which was also targeted for cryoablation. Four 15-gauge cryoprobes were placed in the lesion with CT fluoroscopic guidance, and a 10-, 5-, 10-minute double freeze-thaw cycle was performed. (b) Axial nonenhanced CT image obtained during ablation shows the tumor completely enveloped in the ice ball (arrows). The patient's pain rapidly decreased from a score of 5 of 10 on a visual analogue scale to a score of 2 of 10, and the patient survived another 18 months relatively free of pain from this site of disease. (Case courtesy of Kirkland Davis, MD, Department of Radiology, University of Wisconsin School of Medicine and Public Health, Madison, Wis.)

cautery and direct puncture of peripheral tumors. RF ablation has substantial physical limitations compared with microwave ablation for tissue heating and is thus reserved for treating osteoid osteomas and for other situations that require a short ablation zone. We use cryoablation to treat central renal tumors because of the theoretical risk for damage to the central collecting system and ureter, although we recognize that this is not the experience in all practices. Cryoablation can

also be useful for treating some lung tumors close to the chest wall and mediastinum and musculoskeletal tumors, for which pain palliation is the primary objective.

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b.

**Figure 18.** CT images in a 16-year-old male patient with lateral ankle pain. (a) Coronal nonenhanced CT image shows an osteoid osteoma of the lateral talar dome (arrow). (b) Because a surgical approach would be associated with substantial potential morbidity, the osteoid osteoma was targeted for treatment with RF ablation. As shown on an axial CT image, a single 17-gauge electrode with a 1-cm active tip (arrow) was placed in the lesion with CT fluoroscopic guidance, and a 6-minute ablation was performed. The symptoms completely resolved, and the patient remains asymptomatic. (Case courtesy of Kirkland Davis, MD.)

c.



**Figure 19.** CT images in a 62-year-old woman with a history of multiple episodes of locally recurrent RCC and two previous surgical resections. (a) Axial nonenhanced CT image obtained before image-guided tumor ablation shows a  $4.5 \times 2.6$ -cm site of recurrent disease (arrows) positioned immediately between two loops of colon filled with positive oral contrast material (\*) near the hepatic flexure. (b) Axial CT image shows that aggressive hydrodissection with two

separate needles (arrows) increased the distance between the mass (arrowhead) and the adjacent loops of colon (\*). The tumor was targeted for microwave ablation with two 17-gauge gas-cooled antennae at 65 W for 3 minutes. Other retroperitoneal sites of disease were also treated during the same session. Fever, flank pain, and erythema developed in the days after the procedure. (c) Planar fluoroscopic spot image obtained during an enema examination with Gastrografin (Bracco Diagnostic, Princeton, NJ) helped confirm a contained colon leak (arrows). This was treated successfully with percutaneous drainage.

b.



b.

**Figure 20.** CT images in a 71-year-old woman with a history of RCC of the left kidney that was resected 13 years previously. (a) Axial contrast-enhanced CT image shows recurrent retroperitoneal disease (arrows). The tumor is partially encasing and wrapping anterior to the aorta (arrowhead). This would be problematic with RF ablation because of heat-sink effects and associated tumor sparing. The tumor was targeted for ablation with three microwave antennae for 5 minutes at 50 W. (b) Axial contrast-enhanced CT image obtained at 1-month follow-up shows complete ablation of the tumor, with an adequate margin (arrows). In addition, the tumor that was encasing and anterior to the aorta has been stripped off the aorta and completely ablated (arrowhead).

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## Percutaneous Tumor Ablation Tools: Microwave, Radiofrequency, or Cryoablation—What Should You Use and Why?

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System performance can vary widely, so it is critical that physicians understand the ablation-zone shapes and sizes created by different time and power combinations with a particular system.

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(a) The preponderance of the data suggests that current microwave systems have inherent advantages over RF ablation and that these advantages can give similar or superior results in the clinical setting in less time (note that the increased power does come with the potential for increased complications related to collateral damage as well, but thus far, this has not proven to be the case in clinical practice); (b) the organ involved, tumor type, and underlying comorbid conditions can affect the safety and effectiveness of different modalities; (c) the precision that is possible with cryoablation can be important for treatment of tumors adjacent to vulnerable structures; and (d) vascular mediated cooling can dramatically affect the size and shape of the ablation zone and should be considered in determining both the modality and the positioning and number of applicators.

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The option of treating benign hepatic tumors with percutaneous ablation rather than with open or laparoscopic surgical resection (procedures with increased morbidity compared with ablation) should be presented to patients when applicable.

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Therefore, although many studies have shown that small-volume cryoablation is feasible in patients with cirrhosis and HCC, it is difficult to justify the additional risk of cryoablation in these patients when viable heat-based alternatives are available.

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Therefore, an indirect approach that leaves an unablated tract of at least 2 cm of aerated lung is preferred.