Revisiting Rupture of Benign Thyroid Nodules after Radiofrequency Ablation: Various Types and Imaging Features

Sae Rom Chung¹, Jung Hwan Baek¹, Jin Yong Sung², Ji Hwa Ryu³, So Lyung Jung⁴

¹Department of Radiology and the Research Institute of Radiology, Asan Medical Center, University of Ulsan College of Medicine; ²Department of Radiology and Thyroid Center, Daerim St. Mary’s Hospital, Seoul; ³Department of Radiology, Inje University Haeundae Paik Hospital, Inje University College of Medicine, Busan; ⁴Department of Radiology, Seoul St. Mary’s Hospital, College of Medicine, The Catholic University of Korea, Seoul, Korea

Background: To evaluate the imaging features, clinical manifestations, and prognosis of patients with thyroid nodule rupture after radiofrequency ablation (RFA).

Methods: The records of 12 patients who experienced thyroid nodule rupture after RFA at four Korean thyroid centers between March 2010 and July 2017 were retrospectively reviewed. Clinical data evaluated included baseline patient characteristics, treatment methods, initial presenting symptoms, imaging features, treatment, and prognosis.

Results: The most common symptoms of post-RFA nodule rupture were sudden neck bulging and pain. Based on imaging features, the localization of nodule rupture was classified into three types: anterior, posterolateral, and medial types. The anterior type is the most often, followed by posterolateral and medial type. Eight patients recovered completely after conservative treatment. Four patients who did not improve with conservative management required invasive procedures, including incision and drainage or aspiration.

Conclusion: Thyroid nodule rupture after RFA can be classified into three types based on its localization: anterior, posterolateral, and medial types. Because majority of thyroid nodule ruptures after RFA can be managed conservatively, familiarity with these imaging features is essential in avoiding unnecessary imaging workup or invasive procedures.

Keywords: Thyroid nodule; Ultrasonography; Radiofrequency ablation; Complication; Safety

INTRODUCTION

Radiofrequency ablation (RFA) is a minimally invasive technique that has been used to treat various tumors and has been shown effective in patients with benign thyroid nodules [1-4] and recurrent thyroid cancers [5-7]. Although RFA is relatively safe for treating thyroid nodules, as shown in several systematic reviews and meta-analyses [8-10], it is technically challenging and requires extensive experience [10,11]. Understanding all possible complications of RFA is required for proper diagnosis and treatment.

The most common major complication after RFA is voice...
change [8,10], caused by injury to the recurrent laryngeal or vagus nerve, but this complication may be reduced by the use of a trans-isthmic approach and moving shot technique during RFA [8-10]. The second most common major complication is nodule rupture after RFA [8,10]. To date, however, little is known about the imaging features, clinical course, and risk factors for this adverse event [8,12-14]. This study therefore evaluated the imaging features, clinical manifestations, and prognosis of patients with thyroid nodule rupture after RFA.

METHODS

The Institutional Review Boards of the four participating thyroid centers approved the protocol of this retrospective study (S2017-1436-0001). All patients provided written informed consent prior to RFA.

The records of 12 patients who experienced thyroid nodule rupture after RFA at four Korean thyroid centers between March 2010 and July 2017 were retrospectively reviewed. Clinical data included baseline patient characteristics, treatment method, initial presenting symptoms, imaging features, treatment, and prognosis were evaluated.

Before RFA, all nodules were found to be benign by fine needle aspiration or core needle biopsy [15-17]. Ultrasound (US)-guided RFA was performed by the same radiologist who performed the US assessment. All ablations were performed using a radiofrequency (RF) generators (Cool-Tip RF system, Covielen, Boulder, CO, USA; SSP-2000, Taewoong Medical, Gimpo, Korea; and M-1004, RF Medical, Seoul, Korea) and an 18-gauge internally cooled electrode (Well-Point RF Electrodes, Taewoong Medical, Goyang, Korea; VIVA, STARmed, Goyang, Korea; Big-Tip, RF Medical, Seoul, Korea; Cool-tip RF system, Radiomics, Valleylab, Boulder, CO, USA) with 0.7-, 1-, 1.5-, and 2-cm active tips, respectively, depending on the size of the nodule and the preference of the radiologists. Local anesthesia consisted of injection of 2% lidocaine into the puncture site and perithyroidal area [18]. RFA was performed using the trans-isthmic approach and moving shot technique under US guidance [2,4,11]. If nodules contained fluid, the fluid was aspirated prior to RFA [19,20]. Adjacent structures, including nerves and vessels, were carefully monitored during the procedure to avoid complications [21]. The electrode tip was initially positioned in the deepest and farthest area of the nodule and then moved after a transient hyperechoic zone appeared within 5 to 10 seconds of turning on the RF power [2,4,11]. If a hyperechoic microbubble did not form at the electrode tip within 5 to 10 seconds, the RF power was increased in 10 W increments. If the patient complained of pain during the procedure, the RF power was reduced or turned off for several seconds. The procedure was terminated when the entire visualized area of the nodule had become a transient hyperechoic zone. Repeat ablations were performed when unsatisfactory volume reductions (50%) of treated nodules after 3 months and the presence of an initially untreated portion [12]. All patients suspected of nodule rupture after RFA were examined by US and/or computed tomography (CT).

RESULTS

A review of medical records identified 12 patients, three men and nine women, of mean age 41 years (range, 16 to 75), who experienced post-RFA nodule rupture. Their clinical details are summarized in Table 1. Of these 12 patients, 10 had thyroid nodules with solid and cystic portions and two had solid nodule. The mean volume of the index nodules was 17.0 mL (range, 0.19 to 74.6). One patient required three RFA procedures, two required two procedures, and nine required one ablation. The mean ablation time and mean maximal ablation power per RFA session were 13.4 minutes (range, 4.7 to 40) and 57.5 W (range, 30 to 110), respectively.

Table 2 summarizes the clinical and imaging manifestations of these 12 patients with thyroid nodule rupture. The most common symptoms of post-RFA nodule rupture during follow-up were sudden neck bulging and pain. Only one patient reported fever, cough, and neck discomfort. The mean time from RFA to nodule rupture was 54.6 days (range, 11 to 156).

Following the development of symptoms, 10 patients were evaluated by US, one by CT, and one by US and CT. Based on their localization on imaging features, nodule rupture could be classified into three types, the most common feature being anterior type, observed in nine patients (75%) and characterized by extension of the nodule contents into the anterior extrathyroidal area and sometimes into the strap muscle, with discontinuity of the anterior thyroid capsule (Fig. 1). Two patients (16.7%) had the posterolateral type of nodule rupture, which was characterized by heterogeneous fluid collection around the thyroid capsule (Fig. 2). One patient (8.3%) had the medial type of rupture, which was characterized on CT by soft tissue bulging at the medial aspect of the ablated nodule and protrusion into the lumen of the trachea (Fig. 3). This patient experienced fever, cough, and neck discomfort, unlike the other patients.

All 12 patients were initially treated conservatively, with close observation and/or oral antibiotics. Eight patients recov-
Rupture of Benign Thyroid Nodule after RFA

Table 1. Baseline Characteristics of the 12 Patients with Thyroid Nodule Rupture after RFA

| No. of patients | Sex | Age, yr | Nodule site | Size, mm | Sonographic composition | No. of RFAs | Ablation time | Maximum ablation power, W |
|-----------------|-----|---------|-------------|----------|-------------------------|-------------|--------------|-------------------------|
| 1               | F   | 75      | RLP         | 8×9×5    | Predominantly solid     | 1           | NA           | NA                      |
| 2               | F   | 16      | RLP         | 24×32×44 | Predominantly solid     | 1           | 8 min 16 sec | 30                      |
| 3               | F   | 42      | RMP-RLP     | 25×34×55 | Predominantly solid     | 1           | 5 min 10 sec | 30                      |
| 4               | M   | 67      | Right lobe  | 38×59×64 | Predominantly solid     | 2           | 13 min (1st) | 50 (1st)     |
|                 |     |         |             |          |                         |             | 8 min 40 sec (2nd)| 50 (2nd)   |
| 5               | F   | 53      | LLP         | 10×21×23 | Predominantly cystic    | 1           | 4 min 40 sec | 30                      |
| 6               | F   | 41      | LLP         | 53×47×28 | Predominantly solid     | 1           | 6 min 50 sec | 110                     |
| 7               | F   | 44      | RLP         | 25×17×31 | Solid                   | 2           | 30 min (1st) | 70 (1st)     |
|                 |     |         |             |          |                         |             | 40 min (2nd) | 50 (2nd)     |
| 8               | M   | 52      | RLP         | 21×25×34 | Predominantly solid     | 1           | 15 min 4 sec | 60                      |
| 9               | F   | 30      | LMP         | 22×28×46 | Predominantly solid     | 2           | 16 min 3 sec (1st) | 60 (1st) |
|                 |     |         |             |          |                         |             | 15 min 24 sec (2nd) | 60 (2nd) |
| 10              | M   | 20      | LLP         | 15×24×33 | Predominantly solid     | 1           | 12 min 49 sec | 60                      |
| 11              | F   | 17      | LLP         | 17×31×40 | Predominantly solid     | 1           | 11 min 13 sec | 50                      |
| 12              | F   | 30      | LMP         | 10×15×15 | Solid                   | 3           | 6 min (1st)  | 50 (1st)     |
|                 |     |         |             |          |                         |             | 15 min 11 sec (2nd)| 80 (2nd) |
|                 |     |         |             |          |                         |             | 6 min (3rd)  | 50 (3rd)     |

RFA, radiofrequency ablation; RLP, right lower pole; NA, not applicable; RMP, right middle pole; LLP, left lower pole; LMP, left middle pole.

Table 2. Clinical and Imaging Manifestations in the 12 Patients with Thyroid Nodule Rupture after RFA

| Patient no. | Time to rupture from RFA, day | Symptoms                     | Type         | Initial treatment                     | Invasive procedure | Outcome       |
|-------------|-------------------------------|------------------------------|--------------|---------------------------------------|--------------------|---------------|
| 1           | 30                            | NA                           | Anterior     | NA                                    |                    | Complete recovery|
| 2           | 156                           | Sudden neck bulging and pain | Posterolateral| Compression                           |                    | Complete recovery|
| 3           | 65                            | Cough, fever, neck discomfort| Medial       | Antibiotics and observation            |                    | Complete recovery|
| 4           | 80                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            | Aspiration         | Complete recovery|
| 5           | 11                            | Sudden neck bulging and pain | Anterior     | Observation                           |                    | Complete recovery|
| 6           | 48                            | Sudden neck bulging and pain | Posterolateral| Observation                           |                    | Complete recovery|
| 7           | 30                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            | Aspiration         | No FU         |
| 8           | 40                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            |                    | Complete recovery|
| 9           | 57                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            | I&D                | Complete recovery|
| 10          | 21                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            | I&D                | Complete recovery|
| 11          | 20                            | Sudden neck bulging and pain | Anterior     | Antibiotics and observation            |                    | Complete recovery|
| 12          | 48                            | Sudden neck bulging and pain | Anterior     | Observation                           |                    | Complete recovery|

RFA, radiofrequency ablation; NA, not applicable; FU, follow-up; I&D, incision and drainage.

Ered completely with conservative treatment after mean follow-up period of 71 days (range, 11 to 202). The mean period of use of antibiotics was 16 days (range, 7 to 34). Four patients required invasive procedures, including incision and drainage or aspiration. The nature of aspirated fluid were hemorrhage in two patients and pus in two patients. All of these four patients had the anterior type of rupture. Fig. 1 shows the US images of one patient who underwent aspiration. Sonography showed breakdown of the anterior thyroid capsule and the communication between intra- and extrathyroidal lesions at the RF site. US
also showed a heterogeneous mass-like lesion extending from the thyroid nodule to the superficial aspect of the strap muscle, with Doppler US showing increased vascularity. US-guided core needle biopsy showed that the mass-like lesion was an organizing hematoma. Despite compression, this hematoma increased in size on follow-up US. US-guided aspiration resulted in the removal of dark blood material, with follow-up US showing that this patient recovered completely. Including this patient, all patients, except for the one lost to follow-up, experienced complete recovery.

**DISCUSSION**

This multicenter study, which describes 12 patients who experienced thyroid nodule rupture after RFA, is, to our knowledge, the first original study to classify types of post-RFA nodule rupture according to their localization. The anterior type is the most often, followed by posterolateral and medial type. Despite differences in their imaging features, 67% patients (eight of 12 patients) with thyroid nodule ruptures were successfully managed conservatively. Knowledge about the typical clinical symptoms (sudden neck bulging and pain) and imaging features of post-
RFA nodule rupture can ensure the proper diagnosis of patients and reduce unnecessary treatment.

RFA is a minimally invasive technique that may serve as a safe and effective alternative to surgery in managing benign thyroid nodules that cause symptoms and cosmetic problems [1-4,22-27]. A recent meta-analysis indicated that the mean volume change following RFA for benign thyroid nodules was 76.1% (95% confidence interval [CI], 70.1% to 82.1%), with an absolute mean change in volume of 8.9 mL (95% CI, 6.6 to 11.2) [27]. RFA is also reported to improve symptom and cosmetic scores, as well as having low complication rates [1-3,22-24]. Although RFA has a lower complication rate than surgery, complications can still occur [8-10,13,28], with nodule rupture being the second most common major complication [8,10]. Thyroid nodule rupture has firstly been reported after laser ablation [29]. In that study, three patients had pseudocysts with fasciitis owing to fluid leakage in the neck muscle fascia, followed by spontaneous reabsorption of these at 3 to 6 months. A subsequent report described six patients at four institutions who experienced thyroid nodule rupture after RFA [12], and there have been occasional case reports describing the occurrence of nodule rupture [13,14].

All previous ruptures that could be evaluated using imaging modalities were anterior type [8,12]. However, although our review of 12 patients at four institutions found that the anterior type was the most common, two patients had the posterolateral type and one had the medial type of rupture. The anterior type typically exhibits an anterior bulging of a mass that can extend to the strap muscle layer. The posterolateral type is a hyperechoic lesion that surrounds the thyroid capsule rather than a bulging anterior mass. The medial type shows a discontinuity of the medial thyroid capsule, with the internal contents bulging toward the tracheoesophageal (TE) groove, causing a mass effect on the trachea. Because the TE groove area is a relatively limited space, unlike the anterior or posterolateral area, the medial rupture is the most uncommon type and its causes remain unclear. Familiarity with these imaging characteristics is important in distinguishing among these types of nodule rupture after RFA and in avoiding additional unnecessary imaging workup or invasive procedures. In fact, the one patient with the posterolateral type and one with the medial type were referred to tertiary referral center from local clinics because of misdiagnosis.
Thyroid nodule rupture after RFA is thought to be caused by delayed bleeding, as shown by the high attenuation observed on CT with volume expansion of treated nodule, as well as aspiration of old blood in all patients who underwent invasive procedures. Bleeding may result from microvascular leakage within the tumor, leading to delayed volume expansion and rupture; tearing of the tumor wall and thyroid capsule at a weak point; or post-procedural massaging or moving of the neck. The differences in imaging features observed in the three types of post-RFA nodule rupture may arise from differences in the causes of rupture. The anterior type may be caused by a more acute expansion of the treated nodule with hemorrhage, whereas the posterolateral type may be caused by a more persistent oozing. All four patients who required invasive procedures had the anterior type of rupture, perhaps because this type is caused by a relatively large volume of hemorrhage. Because hemorrhage is basically aseptic and self-limiting, we recommend that these patients should be managed conservatively with observation and compression, with or without oral antibiotics, rather than undergo invasive procedures [12]. Eight of the 12 patients in this study completely recovered without any invasive procedure, such as aspiration or drainage. Invasive procedures should be performed only when conservative management is unsuccessful.

In conclusion, thyroid nodule rupture after RFA can be classified into three types based on its localization: anterior, posterolateral, and medial types. Because majority of thyroid nodule ruptures after RFA can be managed conservatively, familiarity with these imaging features is essential in avoiding unnecessary imaging workup or invasive procedures.

CONFLICTS OF INTEREST

No potential conflict of interest relevant to this article was reported.

AUTHOR CONTRIBUTIONS

Conception or design: J.H.B. Acquisition, analysis, or interpretation of data: S.R.C., J.H.B., J.Y.S., J.H.R., S.L.J. Drafting the work or revising: S.R.C., J.H.B., J.Y.S., J.H.R., S.L.J. Final approval of the manuscript: J.H.B.

ORCID

Sae Rom Chung https://orcid.org/0000-0003-4219-7166
Jung Hwan Baek http://orcid.org/0000-0003-0480-4754

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