ORIGINAL RESEARCH

Fast hemostasis: a win–win strategy for ultrasound and microwave ablation

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Introduction: Hemorrhage is a serious complication following percutaneous biopsy requiring detecting and immediate treatment of active bleeding. This study aimed to explore the potential benefits of ultrasound (US)-guided microwave ablation (MWA) to treat acute hemorrhage in risky locations.

Materials and methods: We present seven patients (four males and three females) aged 19–69 years with solid-organ arterial hemorrhage treated by US-guided MWA and followed up with contrast-enhanced US (CEUS).

Results: All seven cases successfully underwent MWA for hemostasis, and their vital signs subsequently stabilized. During the follow-up from 13 to 36 days, the ablation area decreased slowly in five patients and was still stable in two cases. There were no complications observed in this study after MWA treatment. We also reviewed a total of 12 publications in the past 10 years.

Conclusion: This study suggested that US-guided MWA may be an effective and safe strategy for acute hemorrhage in the emergency setting. To confirm this method and benefit more patients, more prospective studies with larger samples and longer follow-ups are recommended.

Keywords: hemorrhage, hemostasis, microwave, ablation, sonography, contrast-enhanced US, biopsy

Background

Percutaneous biopsy is the main diagnostic method of tissue tumors. Although more and larger size biopsy specimens have enabled more reliable pathology results, it has increased the risk of hemorrhage from arterial sources,^{1,2} which requires detecting and immediate action against massive active bleeding. Traditional hemostasis methods, such as surgical practice with clamping, suturing,^{3,4} or removing the bleeding parts of the organs, have been performed, which occasionally fail due to central artery rupture, suture-induced tissue infarction, and pseudoaneurysm formation⁵ and thus add to the total expenditure. Another common technology is transcatheter angiographic embolization (TAE),⁶ although the radiation may be hazardous and cause severe complications7 and might not be easily accessible and available. Compared to these approaches, thermal ablation - particularly microwave ablation (MWA) - would ablate bleeding vessels under color Doppler flow imaging (CDFI) or contrast-enhanced ultrasound (CEUS), reaching the target tissue and causing faster and larger-scale irreversible tissue coagulation and necrosis in the treated region. MWA has been mainly used in the treatment of active hemorrhage of livers, and its feasibility has been confirmed in both animal studies and clinical practice.8-10 However, few publications have reported MWA for bleeding in greater risk locations such as the thyroid, spleen, and thoracic wall. This study aimed to summarize the potential benefits of US-guided MWA to treat acute hemorrhage and to preliminarily evaluate its therapeutic effect.

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Case series

We prospectively included seven patients (four males and three females) aged 19-69 years, whose main complication was solid-organ hemorrhage caused by biopsy, interventional operation, or trauma from December 2013 to September 2016. This clinical trial was registered in Clinical Trials.gov ID: NCT03026452, and the procedure was approved by the ethics committee of The First Affiliated Hospital of Zhejiang University. All patients signed written informed consents and were informed of the risk and advantages of MWA for hemostasis. The baseline of these cases in the study is summarized in Table 1. Eligible patients met the following criteria: (1) ultrasound (US) clearly found arterial bleeding, bleeding spots with velocity >100 cm/s; (2) US detected hematoma, along with continuous fall of hemoglobin (Hb) and blood pressure levels; (3) the bleeding position was deep, and the 5-min surface pressure for hemostasis was invalid if bleeding velocity was <100 cm/s. Among these seven cases, hemorrhages appeared in two cases caused by spleen biopsy (Figure 1A and B), liver biopsy (Figure 2A-C), or percutaneous transhepatic cholangial drainage (PTCD). In two cases, hemorrhages had been induced by thyroid biopsy (Figure 3A-C), and one resulted from trauma in the thoracic wall (Figure 4A and B). The mean of Hb, heart rate (HR), and velocity in these cases was 111.7 ± 35.0 g/L, 98.9 ± 16.3 beats/min, and 88 ± 18.5 cm/s, respectively. Furthermore, CEUS successfully detected the hemorrhages, and free fluid was available in all cases, which contributed to the location of the bleeding with US and hematoma.

All patients received conservative therapies such as realtime monitoring under US, local pressing with US probe, or US-guided injection of hemostatic drugs into hemorrhage regions. However, improvement was not observed. One patient, whose Hb declined continuously following spleen biopsy, was in shock, and her vital signs were unstable. Two other patients with thyroid biopsies to obtain pathologic diagnosis experienced uncontrollable bleeding even after pressing the puncture point for 30 min.

Under sonographic guidance, we selected CDFI patterns to observe active bleeding and to estimate its velocity by pulse wave (PW). When active bleeding velocity was faster than 100 cm/s, MWA was considered. CEUS was performed to locate the target area before MWA. Then a 15-gauge microwave electrode was inserted into the bleeding sites, which were ablated by the output power of 50-100 W with 2,450 MHz for 6-8 min using ECO (ECO Medical, Nanjing, China). During the procedure, the electrode was retracted by 1.5-2 cm to ablate small leaking vessels during ablation. The

Tabl	e I The	basic cha	Table I The basic characteristics of patient with hemorrhage	th hemorrhag(0									
Case	Case Sex	Age	Primary disease	Bleeding	Cause	Puncture	Hb (g/L)	HP	HR	Sonography Contrast- Velocity Free Hematoma	Contrast-	Velocity	Free	Hematoma
		(years)		organ		needle type		(mmHg)	(mmHg) (beats/min)		enhanced (cm/s)	(cm/s)	fluid	
_	Male	61	Thrombocytopenia	Spleen	Biopsy	18G	100	09/06	115	Ι	+	105	Yes	Yes
2	Male	53	Spleen multiple tumors	Spleen	Biopsy	18G	120	126/75	92	I	+	001	Yes	Yes
m	Male	67	Obstruction of the biliary Liver	Liver	PTCD	18G	131	97/56	103	I	+	113	Yes	No
			tract											
4	Male	47	Hepatic tumor	Liver	Biopsy	18G	149	147/86	88	I	+	78	Yes	Yes
ъ	Female	60	Thyroid node	Thyroid	Biopsy	23G	115	112/90	96	+	+	82	Yes	Yes
9	Female	58	Thyroid node	Thyroid	Biopsy	23G	127	130/85	75	+	+	77	Yes	Yes
7	Female 69	69	Bilateral pleural effusion	Thoracic wall	Thoracic wall Thoracocentesis	18G	40	75/40	123	+	+	61	Yes	No
Abbre	viations:	Hb, hemog	Abbreviations: Hb, hemoglobin; HP, blood pressure; HR, heart rate; PTCD, percutaneous transhepatic cholangial drainage.	ieart rate; PTCD,	percutaneous transhep	vatic cholangial dra	tinage.							



Figure 1 (A) Crescent-shaped hematoma around the spleen as detected by sonography (arrowheads). (B) Enhanced hemorrhage of puncture tract on contrast-enhanced ultrasound (CEUS). (C) Ultrasound (US)-guided microwave electrode was set into the hemorrhage region (arrowhead). (D) After ablation, enhancement was not detected in the ablation zone (arrowheads).



Figure 2 (A and B) Active bleeding originated from the puncture tract with a velocity of 1.13 m/s (arrowheads). (C) A 1.3 cm crescent-shaped free fluid was detected by sonography around the liver (arrowheads). (D) After microwave ablation, contrast-enhanced sonography did not show any microbubble extravasation inside the liver (arrowheads).



Figure 3 (A) Active bleeding lesion was detected along with small bleeding vessels leaking to the thyroid surface (arrowhead). (B) Contrast-enhanced sonography showed an irregular hematoma measuring 2.6×1.6 cm around the right thyroid (arrowheads). (C and D) Irregular hematoma was detected around the right thyroid, and ablation region showed no enhancement on contrast-enhanced sonography (arrowheads).



Figure 4 (A and B) Active bleeding originated from a branch of thoracic wall artery (arrowheads). (C) Ultrasound (US)-guided microwave ablation (MWA) (arrowhead). (D) Visible microbubbles extravasation was not observed in this section of hemostasis.

results depended on the microwave frequency and whether treated tissues were coagulated. The hemostatic effect gradually increased with increasing power until carbonized tissues prevented heat conduction.

To estimate the effects of MWA for hemostasis, CEUS was performed immediately using SonoVue (Bracco SpA, Milan, Italy) after interventional therapy. Successful hemostatic criteria were as follows: first, the hyperechoic area would overlay the entire lesions, and enhancement was not observed in the ablation zones under CEUS. Extravasation of the contrast agent was also not detected around the target organs or tissue spaces. Finally, all vital signs including decreased HR that was within the normal range and elevated blood pressure subsequently stabilized. Hemostasis failure was considered when CEUS showed microbubble extravasation in the trauma regions after ablation, or when the electrocardiogram (ECG) monitor showed unstable blood pressure and HR. When hemostasis was unsuccessful after the first treatment, a second ablation was performed immediately. When interventional hemostasis therapy had not been available after three attempts, open surgery was required immediately.

Results

Table 2 presents the detailed procedures and outcomes of US-guided ablation for each case. During the procedures, increased temperature made the bleeding coagulated, and the bleeding speed reduced. Then tissues around the electrodes were coagulated slowly, and there was no abnormally fast blood flow around the bleeding spot, indicating that hemostasis was successfully available. After MWA, enhancement was not detected in the ablation zone (Figures 1C and D, 2D, 3C and D, and 4C and D).

Patients' blood pressure rose gradually to the normal after MWA and became stable. During the follow-up of 13 to 36 days, the ablation area decreased slowly in five patients and was still stable in two cases. There were no complications observed in this study after MWA treatment.

Table 2 The detailed description in the procedures of microwave ablation (MWA) for hemostasis

Case	Parenchymal/	Power	Time	Follow-up	Ablation
	vessel injury	(W)	(min)	(days)	area
I	Spleen parenchymal	100	4	28	Decreased
2	Spleen parenchymal	80	6	36	Decreased
3	Liver parenchymal	100	4	22	Stable
4	Liver parenchymal	100	5	18	Decreased
5	Thyroid vessel	40	2	26	Decreased
6	Thyroid vessel	40	2	30	Decreased
7	Thoracic wall vessel	40	I	13	Stable

In addition, we reviewed a total of 12 publications from PubMed, Scopus, and Web of Science before April 2017 documenting thermal ablation for hemostasis following biopsy or surgical resection (Table 3).^{11–22} In the literature review, nine studies reported the efficacy of MWA/radiofrequency ablation (RFA)/percutaneous ethanol injection (PEI) on 22 patients (aged 29–83 years, five males and 17 females) with solid-organ bleeding from uterine, liver, and spleen. They suggested 90.9% efficiency rate without major complications.

Discussion

Solid-organ hemorrhage deriving from trauma is a serious condition requiring immediate diagnosis and treatment. In our hospital, US-guided tissue biopsy, which involves various solid organs such as thyroid, breast, prostate, liver and spleen, was performed in ~3,000 cases every year. The number of thermal ablation treatments for solid-organ tumors increased to 720 last year. With such a huge amount of interventional operations, the occurrence of various complications like hemorrhage is inevitable, amounting to ~0.7% (0.4% slight bleed-ing using surface pressure for hemostasis and 0.3% severe bleeding by MWA against solid-organ hemorrhage).

Baseline sonography is not sensitive to guide the insertion of ablation needles in the treatment of abdominal solid organ injury. Sonography is a reliable screening tool for detecting active bleeding in combination with CDFI pattern, particularly with the presence of continuous free fluid in the abdominal cavity or hematoma in the tissue space. In particular, CEUS could show real-time and contrast-related images via microbubbles produced, and CEUS was clearer in detecting hemorrhagic spots and assessing the degree of damage.^{23–25} Several studies showed that CEUS is a potential tool that is nearly as sensitive as contrast-enhanced-computed tomography (ceCT) for the detection of traumatic abdominal organ injuries.^{26,27} Moreover, the proposed method has no radiation hazard and is a safer contrast agent compared to ceCT.

Previous studies showed that RFA decreased blood loss in hepatectomy both in animal and clinical experiences.²⁸ The temperature in RFA increased slowly, and heat was easily taken away by vessels. Thus, it could not reach the target temperature, and the limited coagulation zone size was an important issue in RFA.¹¹ Compared with RFA, MWA depended less on tissue impedance and has a larger zone of active heating.^{19,29} MWA enabled the microwave antenna in the targeting tissue to locally obtain an average temperature of 65°C–100°C in seconds, resulting in its coagulation necrosis, which made it possible to quickly

Author	Year	Country	Year Country Object of study	Characteristics of patients	Treatment method	Patients (no of tumors)	Male/ female	Mean age (range)	Prognosis	Prognosis Complication	Ablation time (min)	Energy (kJ)
Yekuo	2009	China	Porcine	10 liver trauma	CEUS-guided RFA	01	AN	AN	Success		8.5 ± 4.5	86 ± 51 (16–150)
Sun et al ¹² 2013 China	2013	China	00 D	21 renal bleeding	CEUS-guided percutaneous MWA	21	۲ ۲	A	Success	۲ ۲	 (a) subcapsular artery: 2.1 ± 0.3; (b) interlobar arteries: 3.1 ± 0.6; (c) section artervex. 5 2 + 0.7 	80 ×
Bin et al ¹³	2014	China	Porcine	8 spleen bleeding	US-guided MWA	ω	٩N	AN	Success	٩N	ar ter y. J.z. – U./ 10	80 W
Aytekin	2010		Human	l massive	Angiographic PEI	_	0/1	55	Success	Left flank pain	AN	AN
et al ¹⁴				multifocal bleeding from subcapsular						from subcapsular hematoma		
				hematoma								
Chapa	2010	America	Human	2 acute uterine	Thermal balloon ablation	2	0/2	43	Success	No	NA	NA
et al ¹⁵				hemorrhage								
Nakayama	2011	Japan	Human	7 uterine	US-guided MEA	7	0/7	45.1 ± 5.4	Success	No	$\textbf{46.6} \pm \textbf{24.5} \; \textbf{(0.83)}$	70 W
et al ¹⁶				hemorrhage								
Maroulis et al ¹⁷	2013	Greece	Human	2 liver bleeding during laparotomy	RFA	2	2/0	29	Success	No	35	180–200 W
Weigt	2014	Germany	Human	I bleeding through	Endoscopic intraductal	_	1/0	83	Success	No	NA	NA
et al ¹⁸				the ampulla of	RFA							
71-12												101 00 0E
et al ¹⁹		CIIIIa	Пиплап	bleeding (5 in liver	Percutaneous MWA, 2	٥	c/c	0.0 ⊥ 10.0	4 success, 2 failure	0	107 ± 20.1	AA 02-07
				and I in spleen)	US-guided percutaneous RFA							
Ogura et al ²⁰	2016	Japan	Human	I bleeding from the ampulla of Vater	Endoscopic intraductal RFA	_	0/1	80	Success	No	1.5	10 M
Wai et al ²¹	2016	China	Human	I biopsy liver hemorrhage	US-guided percutaneous MWA	_	0/1	61	Success	AN	6	140
Pope et al ²²	2016	America	Human	I liver hemorrhage following 2 failed	US-guided PEI	_	0/1	42	Success	AN	AN	٩N
				embolizations								

achieve hemostasis and lower the cost without any radiation hazard and major complications from TAE.⁷ Additionally, contrast agent in angiography may increase the metabolic burden of liver and kidney while contrast agent in CEUS not due to its pulmonary pathway for discharge. Some studies also demonstrated that modified MWA has a larger ablative range and higher top temperatures than the previous one because of their longer wavelength, deeper penetration depth, and lower energy attenuation.^{30,31} Thus, MWA was useful for controlling life-threatening hemorrhages even when active bleeding occurred in a larger diameter vessel.

The main limitation of this study was its small sample size and the short-term follow-up, which may have impact on the delayed bleeding if possible. The existing heat-sink in MWA may limit tissue heating and tissue ablation.

Conclusion

US-guided percutaneous MWA may be an effective and safe strategy for hemostasis in patients with hemorrhage in the emergency setting. To verify this method and benefit more patients, more prospective studies with larger samples and longer follow-ups are needed.

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Disclosure

The authors report no conflicts of interest in this work.

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