Original Article Clinical application of radiofrequency ablation negative pressure automatic biopsy gun in liver space-occupying lesions patients

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Abstract: Objective: With the rapid developments in hepatic lesions detection technologies, the incidence rate of liver space-occupying lesions (SOLs) has accelerated dramatically. This investigation is aimed to evaluate the safety and clinical efficacy of radio frequency ablation negative pressure automatic biopsy gun (RFANPABG) for treatment in liver space-occupying lesions (SOLs) patients. Methods: A comparison study of puncture biopsy with 348 cases of diagnosed liver SOLs patients enrolled from January 2013 to August 2014. RFANPABG were performed on the average random distribution 174 patients, and radiofrequency ablation conventional puncture biopsy gun (RFACPBG) were regularly conducted on the other 174 patients to evaluate the technical success rate of puncture efficiency, the needle tract bleeding rate, the needle tract tumor growing rate and so on. Results: For the RFANPABG, the technical success rate of puncture efficiency was 97.70% (170/174), higher than that of RFACPBG with 6.32% (148/174) (P<0.05); the needle tract bleeding rate was 1.15% (2/174), lower than that of RFACPBG with 6.32% (148/174) (P<0.05); and the needle tract tumor growing rate was 0.58% (1/174), also lower than that of RFACPBG, which is 4.60% (8/174) (P<0.05). Conclusion: The higher puncture efficiency, lower needle tract bleeding rate and needle tract tumor growing rate of RFANPABG, in conjunction with the clinical data findings, make RFANPABG a promising useful diagnostic modality in SOLs.

Keywords: Hepatic space-occupying lesions, radiofrequency ablation negative pressure automatic biopsy gun, puncture efficiency, needle tract bleeding rate, needle tract tumor growing rate

Introduction

With the wide spreading infection of hepatovirus, especially the hepatitis C infection [1, 2], the global incidence of liver cancer is rising sharply year by year in worldwide, especially in China. For example, WHO estimate about 47,000 deaths caused by liver cancer per year in Europe [3]; while in China, the mortality rate was estimated about 26.26 per 100,000, taking about 19.33% of all sites of cancers [4]. Hepatic resection and liver transplantation are known as the curative treatment options for malignant liver tumor [5], however, just 10-54% malignant liver tumor patients are suitable to surgery [6]. In addition, with low respectability rate (20-37%) and shortage of the grafts, the majority of malignant liver tumor patients are not received surgical treatment [7, 8]. Different loco regional nonsurgical therapies, such as transarterial chemoembolization (TACE), percutaneous ethanol injection(PEI), cry therapy, interstitial laser therapy, microwave coagulation, radiofrequency ablation (RFA) and so on, have been developed for treatment of malignant liver tumor patients [5, 9]. Several local ablative techniques have been used for several decades [10], and the radiofrequency ablation (RFA), as the most popular local ablative technique with easily to operate and low morbidity, have been widely used in malignant liver tumor treatments [11-13]. The space-occupying lesions (SOLs) of liver mean a unique and distinct spectrum when compared with those normal livers. Although many pathological factors, such as hepatic hemangioma, hepatoblastoma, focal nodular hyperplasia, hepatic adenoma, hepatocellular carcinoma and so on may lead to the

RFANPABG treatment	RFACPBG treatment	P-value
15-72 (49.3±12.7)	14-73 (49.1±13.1)	NS
93/81	93/81	NS
73/101	74/100	NS
96/78	94/80	NS
138/36	137/37	NS
	15-72 (49.3±12.7) 93/81 73/101 96/78	15-72 (49.3±12.7) 14-73 (49.1±13.1) 93/81 93/81 73/101 74/100 96/78 94/80

Table 1. General Data of studied liver SOLs patients before treatment by RFANPABG

Abbreviations: NS, Not significant.

performance of liver SOLs [14-17], clinical presentations of liver SOLs have played an important role for malignant liver tumor judgment [18, 19]. As the radiofrequency ablation (RFA) have been widely applied in malignant liver tumor treatments [11-13], these techniques should also be applied in liver SOLs treatment.

Radiofrequency ablation, known as "radiofrequency thermal ablation", is a recently developed thermo ablative technique, with combination of energy absorption, heating and conduction to generate areas of coagulative necrosis and tissue desiccation [20-22]. With low morbidity and mortality rates, effective tumor ablation, producing larger volume of coagulation necrosis and so on, RFA gained a tremendous enthusiasm in modern management of malignant liver tumors in recent years [11-13, 20]. However, the risks and probability of RFA operation failure should not be ignored, one of the most important, the use of RFA should be in the target lesions and be justified by critically appraising. In addition, the lesions should be accurately diagnosed as malignant liver tumors before received the RFA. As the main basis and means for clinical pathologic diagnosis, puncture pathology biopsy has been in clinical application for more than 50 years, and biopsy gun also has been applied for more than 10 years. However, some application flaws and complications has hindered the clinical application of biopsy gun to a certain extent, such as the puncture efficiency were 75 to 95%, the needle tract bleeding rate were 1.7 to 7.5%, and the needle tract tumor growing rate were 2.1 to 8.0%.

Hence, we performed a radiofrequency ablation negative pressure automatic biopsy gun (RFANPABG) for treatment the liver space-occupying lesions (SOLs) patients. This study was aim to evaluate the safety and clinical efficacy of using RFANPABG) for liver SOLs treatment, and also to investigate the possible clinical application prospect.

Materials and methods

General data

All the 348 patients (general data were shown in Table 1) were consecutive studied and average random distributed from Chinese diagnosed and/or treated with liver SOLs in our hospital, during January 2013 to August 2014. This study was conducted in accordance with the declaration of Helsinki. This study was approved by the local ethics committee, and the written informed consent form was obtained by each subject. All participants' general data were obtained from questionnaires, including living environment conditions, occupation, ages and so on. All patients were diagnosed with liver SOLs and the corresponding diseases by B ultrasound, CT, and pathological examination, and they accepted treatment by RFANPABG or RFACPBG with the recommend operational procedures.

Establishment the 3D-CT image of target liver SOLs

The 3D-CT image of target liver SOLs was obtained by using INNOVA4100 IQ angiography machine with 3D-CT navigation function. The puncture point and route to the target lesions were determined avoiding ribs. In addition, the puncture angle and probe insertion depth were determined under bull's-eye view.

Local negative pressure biopsy and radiofrequency ablation

The puncture of target lesions with negative pressure biopsy gun needle and RFA needle were conducted under bull's-eye view. When reached the proper position, operate the nega-

Index	RFANPABG	RFACPBG	χ^2 and		
	treatment	treatment	P-value		
Puncture efficiency					
Success	170 (97.7%)	148 (85.1%)	χ ² = 17.655		
Fail	4 (2.3%)	26 (14.9%)	P<0.001		
Needle tract bleeding rate					
Bleeding	2 (1.1%)	11 (6.3%)	$\chi^2 = 6.473$		
No bleeding	172 (98.9%)	163 (93.7%)	P = 0.011		
Needle tract tumor growing rate					
Growing	1 (0.6%)	8 (4.6%)	χ ² = 5.589		
No growing	173 (99.4%)	166 (97.4%)	P = 0.018		

Table 2. Puncture efficiency, needle tract bleeding rate and needletract tumor growing rate between the RFANPABG treatment groupand the RFACPBG treatment group

and the RFACPBG treatment group, whether been statisticated by lesion site, lesion maximum diameter distribution or lesion number.

Technical success rate of puncture efficiency and post procedure complications

All patients could tolerate the combined treatment, and both the RFANPABG needles and the RFACPBG needles are successfully entered the target lesions for each puncture. After the puncture by nega-

tive pressure biopsy gun needle to cut tissue for biopsy, then pull out the biopsy needle and puncture the insulated coating RFA needle. The multi-polar radiofrequency was opened at the right target lesion position, and all the RFA parameters were determinate according to the lesion location, size and shape. After two or three times RFA, the solidification was conducted to avoid bleeding and tumor implantation and metastasis in probe tract, and the cutting tissue samples were sent for pathological examination.

Statistical analysis

All statistical analyses were performed by using the Statistical Package for Social Sciences software (SPSS, Windows version release 17.0; SPSS Inc.; Chicago, IL, USA). The chi-squared (χ^2) test was utilized to evaluate the Hardy-Weinberg equilibrium in treatment efficacy. A level of P<0.05 was considered statistically significant.

Results

Patient characteristics

The demographic and clinical characteristics of the studied diagnosed liver SOLs patients in the RFANPABG treatment group and the RFACPBG treatment group are summarized in **Table 1.** There are no significant statistical different of age and gender between the RFANPABG treatment group and the RFACPBG treatment group. In addition, there are also no significant statistical different of patients number between the RFANPABG treatment group tive pressure automatic biopsy gun or conventional puncture biopsy gun, each patient successfully received RFA treatment. The success of puncture is defined as obtaining the right expected tissue samples, and the location of lesions are desiccated and coagulative necrosis under medical examination; the needle tract bleeding is assessed by serum hemoglobin level, prothrombin time, platelet count and the treatment response by a CT scan of the liver from 3-5 days after RFA; and the needle tract tumor growing is assessed with reexamination once for every 2 to 3 months by CT or B ultrasound, and blood, liver and renal function detection. There are significant statistical differences of the puncture efficiency, the needle tract bleeding rate and the needle tract tumor growing rate between the RFANPABG treatment group and the RFACPBG treatment group. As shown in Table 2, the technical success rate of puncture efficiency in the RFANPABG treatment group is 97.7%, about 1.15 fold than that (85.1%) in the RFACPBG treatment group (χ^2 = 17.655, P<0.001). At the same time, the needle tract bleeding rate in the RFANPABG treatment group is 1.1%, while 6.3% in the RFACPBG treatment group, which is significant much lower with treated by RFANPABG (χ^2 = 6.473, P = 0.011). After treatment by RFANPABG or RFACPBG, the post procedure complications of all patients are been studied by follow-up surveys, and the corresponding needle tract bleeding rates are been statistically analyzed. 173 patients are free of needle tract tumor growing after received the RFANPABG treatment, and only 166 patients are free of needle tract tumor growing after received the RFACPBG treatment,

the needle tract tumor growing rate is significantly lower in the RFANPABG treatment group ($\chi^2 = 5.589$, P = 0.018).

Discussion

With the increasing incidence rates of malignant liver tumor, more and more alternative treatments to control or potentially cure liver disease have been develop [23-25]. As a new multidisciplinary technique with many key advantages including low morbidity and mortality rates, effective tumor ablation [11-13, 20], radiofrequency ablation (RFA) technique has been used as one effective loco regional nonsurgical treatment modality in modern management of malignant liver tumors in recent years [11, 12, 26]. However, many factors should be taken into consideration before received the RFA, for an example, the lesions should be accurately diagnosed before RFA treatment. Accordingly, many techniques for clinical pathologic diagnosis also get remarkable progress [23, 24, 27].

As the liver SOLs work as the very possible symptoms for malignant liver tumors, such as hepatocellular carcinoma, hepatic adenoma and hepatoblastoma, accurate diagnosis of the liver SOLs may show very important role in timely diagnosis and treatment for malignant liver tumors [18, 19]. Liver SOLs have unique and distinct spectrums different from normal liver, many classic and modern techniques have been developed for these diagnose, for example, B ultrasound, CT and fine needle aspiration biopsy [17]. Puncture pathology biopsy works as the main basis and means for lesions diagnose in clinical application for a long history, and the biopsy gun also has been used in clinical diagnose for 10 years. Although the puncture efficiency could up to 95%, the needle tract bleeding rate could be minimized to 1.7%, and the needle tract tumor growing rate also could be minimized to 2.1%, these deficiencies still have hindered the clinical application of biopsy gun in some extent.

In this study, we invented the radiofrequency ablation negative pressure automatic biopsy gun (RFANPABG), then preliminary applied in diagnosis and treatment of liver SOLs, and the corresponding puncture efficiency, needle tract bleeding rate and needle tract tumor growing rate were also compared with treatment the ablation conventional puncture biopsy gun (RFACPBG). Results showed a very higher puncture efficiency, lower needle tract bleeding rate and needle tract tumor growing rate of treatment by RFANPABG, which realized the fast in situ diagnosis and treatment of SOLs, moreover, applying this technique may significantly reduce the risk of possible complications. With these innovations and advantages, the RFANPABG may become into a promising useful clinical diagnostic modality in liver SOLs in the future.

Disclosure of conflict of interest

None.

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References

- [1] Lim EJ and Torresi J. Prevention of hepatitis C virus infection and liver cancer. Recent Results Cancer Res 2014; 193: 113-133.
- [2] Chang MH. Prevention of hepatitis B virus infection and liver cancer. Recent Results Cancer Res 2014; 193: 75-95.
- [3] Blachier M, Leleu H, Peck-Radosavljevic M, Valla DC and Roudot-Thoraval F. The burden of liver disease in Europe: A review of available epidemiological data. J Hepatol 2013; 58: 593-608.
- [4] Chen JG and Zhang SW. Liver cancer epidemic in China: Past, present and future. Semin Cancer Biol 2011; 21: 59-69.
- [5] Ng KK and Poon RT. Radiofrequency ablation for malignant liver tumor. Surg Oncol 2005; 14: 41-52.
- [6] Forner A, Llovet JM and Bruix J. Hepatocellular carcinoma. Lancet 2012; 379: 1245-1255.
- [7] Poon RT, Fan ST, Lo CM, Ng IO, Liu CL, Lam CM and Wong J. Improving survival results after resection of hepatocellular carcinoma: a prospective study of 377 patients over 10 years. Ann Surg 2001; 234: 63-70.
- [8] Fong Y, Fortner J, Sun RL, Brennan MF and Blumgart LH. Clinical score for predicting recurrence after hepatic resection for metastatic colorectal cancer: analysis of 1001 consecutive cases. Ann Surg 1999; 230: 309-318.
- [9] Sutherland LM, Williams JA, Padbury RT, Gotley DC, Stokes B and Maddern GJ. Radiofrequency ablation of liver tumors: a systematic review. Arch Surg 2006; 141: 181-190.
- [10] Joosten J and Ruers T. Local radiofrequency ablation techniques for liver metastases of

colorectal cancer. Crit Rev Oncol Hematol 2006; 62: 153-163.

- [11] Stoltz A, Gagnière J, Dupré A and Rivoire M. Radiofrequency ablation for colorectal liver metastases. J Viscl Surg 2014; 151: S33-S44.
- [12] Meijerink MR, van den Tol P, van Tilborg AA, van Waesberghe JH, Meijer S and van Kuijk C. Radiofrequency ablation of large size liver tumours using novel plan-parallel expandable bipolar electrodes: Initial clinical experience. Eur J Radiol 2011; 77: 167-171.
- [13] Gravante G, Ong SL, Metcalfe MS, Bhardwaj N, Lloyd DM and Dennison AR. The effects of radiofrequency ablation on the hepatic parenchyma: Histological bases for tumor recurrences. Surg Oncol 2011; 20: 237-245.
- [14] Mostafa MG, Dalquen P, Kunze D and Terracciano L. Telecytological diagnosis of space-occupying lesions of the liver. Acta Cytol 2014; 58: 174-181.
- [15] Tremper J, Aulmann S, Willig F and Zechmann CM. Multiple space-occupying lesions of the liver: a rare differential diagnosis. Radiologe 2009; 49: 1058-1062.
- [16] Eickhoff A, Spiethoff A, Hartmann D, Jakobs R, Weickert U, Schilling D, Eickhoff JC, Bohrer MH and Riemann JF. Space-occupying lesions in the liver: incidence of adenocarcinoma metastases of unknown primary site. Dtsch Med Wochenschr 2006; 132: 369-374.
- [17] Bakshi P, Srinivasan R, Rao KL, Marwaha RK, Gupta N, Das A, Nijhawan R and Rajwanshi A. Fine needle aspiration biopsy in pediatric space-occupying lesions of liver: a retrospective study evaluating its role and diagnostic efficacy. J Pediatric Surg 2006; 41: 1903-1908.
- [18] D'Errico A, Fiorentino M, Corti B, Altimari A, Gabusi E, Gruppioni E, Pirini MG and Grigioni WF. Space-occupying (malignant) lesions of the liver. Pathologica 2003; 95: 253-254.
- [19] Beissert M, Jenett M and Kessler C. Diagnosis of space-occupying lesions of the liver. What is the value of diagnostic imaging?. MMW Fortschr Med 2001; 143: 29-33.

- [20] Pandya GJ and Shelat VG. Radiofrequency ablation of pancreatic ductal adenocarcinoma: The past, the present and the future. World J Gastrointest Oncol 2015; 7: 6-11.
- [21] Fuller CW, Nguyen SA, Lohia S and Gillespie MB. Radiofrequency ablation for treatment of benign thyroid nodules: systematic review. Laryngoscope 2014; 124: 346-353.
- [22] Minami Y and Kudo M. Radiofrequency ablation of hepatocellular carcinoma: a literature review. Int J Hepatol 2011; 2011: 104685.
- [23] de Sio I, ladevaia MD, Vitale LM, Niosi M, Del Prete A, de Sio C, Romano L, Funaro A, Meucci R, Federico A, Loguercio C and Romano M. Optimized contrast-enhanced ultrasonography for characterization of focal liver lesions in cirrhosis: A single-center retrospective study. United European Gastroenterol J 2014; 2: 279-287.
- [24] Yazgan C, Akata D, Ozmen M and Karcaaltincaba M. Precision imaging of focal liver lesions: comparison with conventional sonography in terms of image quality. J Ultrasound Med 2013; 32: 1405-1410.
- [25] Jang JY, Kim MY, Jeong SW, Kim TY, Kim SU, Lee SH, Suk KT, Park SY, Woo HY, Kim SG, Heo J, Baik SK, Kim HS and Tak WY. Current consensus and guidelines of contrast enhanced ultrasound for the characterization of focal liver lesions. Clin Mol Hepatol 2013; 19: 1-16.
- [26] Lee JM, Han JK, Chang JM, Chung SY, Kim SH, Lee JY, Lee MW and Choi BI. Radiofrequency Ablation of the Porcine Liver In Vivo: Increased Coagulation with an Internally Cooled Perfusion Electrode. Acad Radiol 2006; 13: 343-352.
- [27] Smith AB, Filby A and Carr LM. Heterogeneity in patient diagnostic pathways: an example from contrast-enhanced ultrasound diagnostic scans for focal liver lesions. BMC Res Notes 2014; 7: 199.