

# Role of Diffusion-Weighted Imaging and Proton MR Spectroscopy in Distinguishing between Pyogenic Brain Abscess and Necrotic Brain Tumor

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**Abstract-** Brain abscesses and brain tumors may have similar clinical presentations. For example, only 50% brain abscess patients have fever, which could be masked by corticosteroid therapy. Also, the differential diagnosis of brain abscesses versus cystic or necrotic tumors may be difficult based on computed tomography (CT) or magnetic resonance (MR) imaging findings. However, the strategies of management for abscess and neoplasm are very different, and it is especially imperative to have a correct diagnosis before any surgical intervention of cystic brain lesions. The MR special techniques, e.g. diffusion-weighted imaging (DWI) and proton (1H) MR spectroscopy, are useful as additional diagnostic modalities for differentiating brain abscesses from cystic or necrotic brain tumors. DWI shows high signal intensity in most cases of pyogenic abscesses and low signal intensity in most cases of cystic or necrotic tumors. MR spectroscopy shows characteristic metabolites in pyogenic abscesses, distinct from those in cystic or necrotic tumors.

**Key Words:** Brain abscess, Brain tumor, Diffusion MR imaging, MR spectroscopy

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## INTRODUCTION

Differentiation between brain abscesses and cystic or necrotic tumors with computed tomography (CT) or magnetic resonance (MR) imaging may be difficult. Difficulties in the diagnosis of intracranial abscess are mainly due to the combination of nonspecific clinical findings and possible similarities in the radiological appearances of cystic gliomas, metastases, and brain abscesses<sup>(1)</sup>. With conventional magnetic resonance imaging (MRI), typical brain abscess has low signal

intensity on T1-weighted images, high signal intensity on T2-weighted images, and ring enhancement after contrast medium administration. However, cystic or necrotic brain tumors may have very similar pictures, making the differential diagnosis difficult.

Diffusion-weighted imaging (DWI) and proton (1H) MR spectroscopy (MRS) are results of recent progress in MR technique, and could serve as useful diagnostic modalities for between brain abscesses and cystic or necrotic brain tumors. We would divide the following discussion into three parts: (a) DWI of Pyogenic Brain

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Abscesses and Necrotic Brain Tumors; (b) 1H MRS of Pyogenic Brain Abscesses and Necrotic Brain Tumors; and (c) DWI vs. 1H MRS of Pyogenic Brain Abscesses and Necrotic Brain Tumor.

### DWI of pyogenic brain abscesses and necrotic brain tumors

DWI provides a unique way to evaluate the diffusion properties of water molecules in tissue (Brownian motion), and has been demonstrated to be useful in the diagnosis and treatment of ischemia, tumors, epilepsy and white matter disorders<sup>(2)</sup>. Application of DWI in the differential diagnosis between brain abscess and cystic or necrotic brain tumors has been well documented in the literature<sup>(3-10)</sup>. In 1996, Ebisu et al.<sup>(3)</sup> first reported a case of brain abscess with high signal intensity on DWI and a low ADC value. The pus itself (verified by aspiration) could account for the restricted diffusion and therefore high DWI signal intensity (Fig. 1). The other studies showed very similar or the same results<sup>(4-12)</sup>.

The low ADC values for pyogenic abscesses varied<sup>(5-10)</sup>. The variation might be ascribable to differences in the concentration of inflammatory cells, the necrotic tissue debris and bacteria, and the viscosity of the abscess fluid. These differences may be in turn related to the age of the abscess, the causative organism, and the host's immune responses<sup>(5-10)</sup>.

The signal intensity of pyogenic brain abscess on DWI is not always high, and the ADC value could be higher than that of the normal brain parenchyma. Krabbe et al.<sup>(13)</sup> reported a cerebral abscess with a high ADC value, but did not mention the signal intensity on DWI. The explanation for this might be the partial volume effect proposed by Desprechins et al.<sup>(6)</sup>. Guo et al.<sup>(14)</sup> reported two small cerebral abscesses (mean area of 90 mm<sup>2</sup>) with high signal on DWI but ADC values similar to that of normal parenchyma (the ADC ratios were 1.01 and 1.03). This similarity was postulated to be due to a partial volume effect or to intrinsic differences in diffusion properties of the smaller abscesses. However, signal intensity on DWI was high and ADC value was low regardless of the lesion size, and these results were supposed to have improved in-plane resolution<sup>(8)</sup>.

All tumors showed low signal intensities on DWI in

the cystic or necrotic area, and their cystic or necrotic areas had high ADC values [<sup>(2-10,13,15)</sup>-Fig. 2]. However, contradictory DWI findings of cystic or necrotic tumors do exist in the literature<sup>(7,8,16)</sup>. Park et al. reported two cases of cystic or necrotic brain metastasis with markedly high signal intensity on DWI, with surgical findings showing that the cyst had a thick and creamy necrotic content similar to pus<sup>(7)</sup>. Chang et al. reported a ring-enhanced fibrillary low-grade astrocytoma with high signal intensity on DWI and a low ADC value, mimicking that of abscesses, and viscous creamy fluid within the tumor was found at surgery<sup>(8)</sup>. Holtas et al. reported a ring-enhanced brain metastasis with hyperintensity on DWI and a low ADC value in the necrotic part of the tumor. The reason for restricted diffusion was possibly due to early necrosis with intracellular edema of the lesion<sup>(16)</sup>.

In summary, DWI may be a much more valuable diagnostic tool in the differential diagnosis between brain abscesses and cystic or necrotic brain tumors than conventional MRI. However, there are some conflicting DWI reports in the literature, and we still need to be cautious about making the diagnosis with DWI findings.

### 1H MRS of pyogenic brain abscess and necrotic brain tumors

Findings from several studies have suggested that in vivo proton magnetic resonance spectroscopy (1H MRS), a non-invasive examination, might contribute to the establishment of the differential diagnosis between brain tumors and abscesses<sup>(17-29)</sup>. The predominant resonance lines (N-acetyl-aspartate, choline, and creatine/phosphocreatine) that are usually observed in the parenchyma of the normal brain were hardly detectable in either tumor or abscess necrosis. The main characteristic features of pyogenic abscesses were the resonances of amino acids (valine, leucine, and isoleucine) (0.9 ppm), acetate (1.9 ppm), alanine (1.5 ppm), lactate (1.3 ppm), and succinate (2.4 ppm) (Fig. 1). In contrast, Spectra in patients with cystic or necrotic tumor showed only the peak attributed to lactate (1.3 ppm) (Fig. 2). Lipid (0.8-1.3 ppm) may also be found in patients with a tumor.

Lactate and lipids are nonspecific metabolites pro-

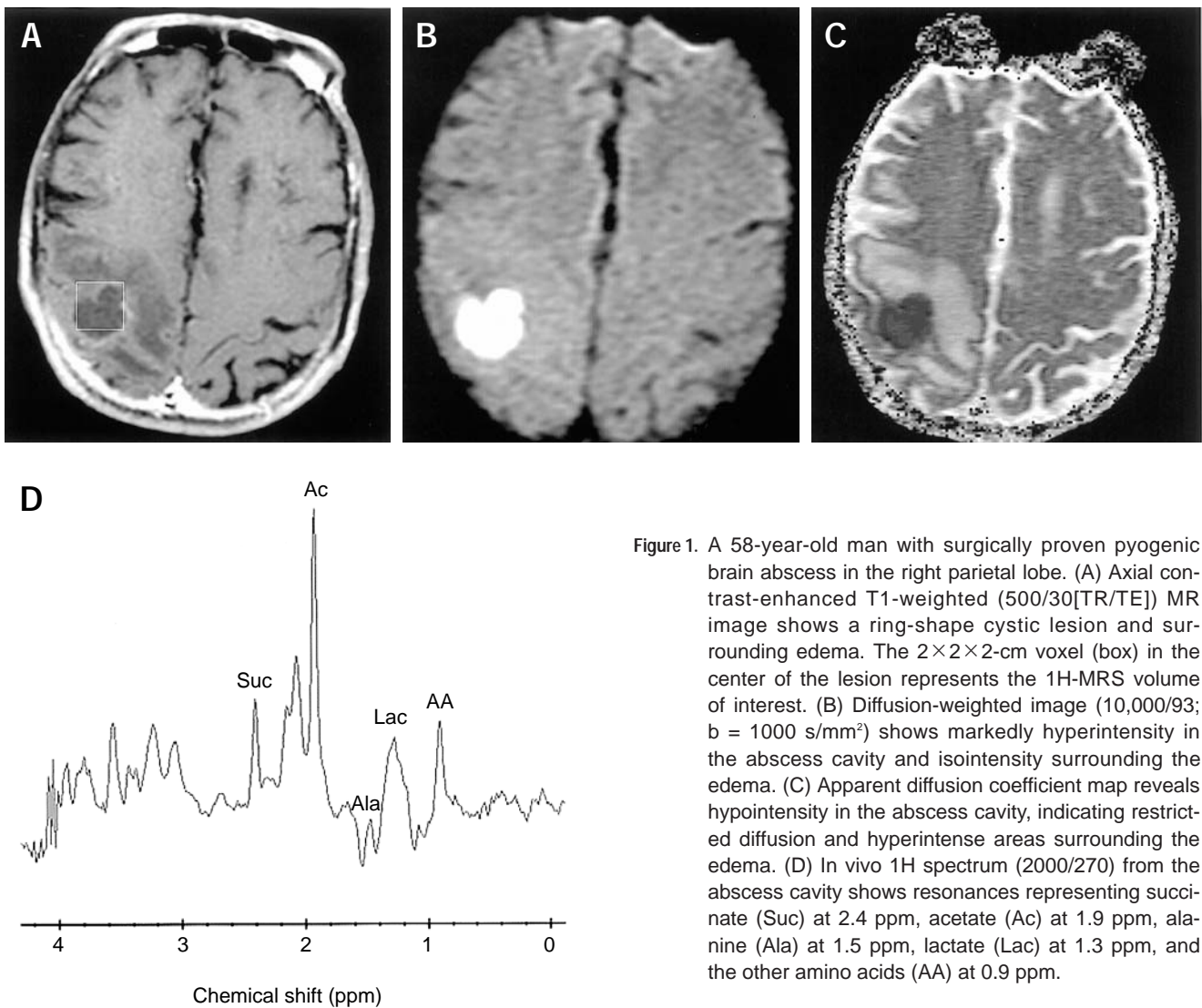


Figure 1. A 58-year-old man with surgically proven pyogenic brain abscess in the right parietal lobe. (A) Axial contrast-enhanced T1-weighted (500/30[TR/TE]) MR image shows a ring-shape cystic lesion and surrounding edema. The  $2 \times 2 \times 2$ -cm voxel (box) in the center of the lesion represents the  $^1\text{H}$ -MRS volume of interest. (B) Diffusion-weighted image (10,000/93;  $b = 1000 \text{ s/mm}^2$ ) shows markedly hyperintensity in the abscess cavity and isointensity surrounding the edema. (C) Apparent diffusion coefficient map reveals hypointensity in the abscess cavity, indicating restricted diffusion and hyperintense areas surrounding the edema. (D) In vivo  $^1\text{H}$  spectrum (2000/270) from the abscess cavity shows resonances representing succinate (Suc) at 2.4 ppm, acetate (Ac) at 1.9 ppm, alanine (Ala) at 1.5 ppm, lactate (Lac) at 1.3 ppm, and the other amino acids (AA) at 0.9 ppm.

duced by anaerobic glycolysis and necrotic brain tissue in brain abscesses. Both lactate and lipids peaks can also be observed in necrotic tumors<sup>(19,20,22-25)</sup>. Increases in acetate and succinate presumably originate from the enhanced glycolysis and fermentation by the infecting microorganisms<sup>(19-26)</sup>. Amino acids such as valine and leucine are probably end products of proteolysis by enzymes released from neutrophils in pus<sup>(19-26)</sup>.

Discrimination between amino acids (i.e., valine, leucine, and isoleucine at 0.9 ppm) and lipid (at 0.8 to 1.2 ppm) is important. Lipid signals may exist in both brain tumors and abscesses, whereas amino acids are not seen in proton MR spectra of brain tumors in vivo and

are only detectable in vitro<sup>(21)</sup>. It is known that with an echo time of 135, phase inversion occurs as a result of J-coupling between lactate and amino acids, but not in lipid, a feature which may be helpful in differential diagnosis between brain abscess and tumor<sup>(19,22)</sup>.

The findings of lactate and lipid peaks after long term treatment with antibiotics may become nonspecific and thus can be easily missed in patients with cystic or necrotic tumors<sup>(25,27-29)</sup>. Dev et al. demonstrated a decline of the acetate and pyruvate in 5 patients after 1 week of aspiration and medical treatment. The authors proposed that the disappearance of metabolites of bacterial origin might suggest a positive response to therapy<sup>(25)</sup>.

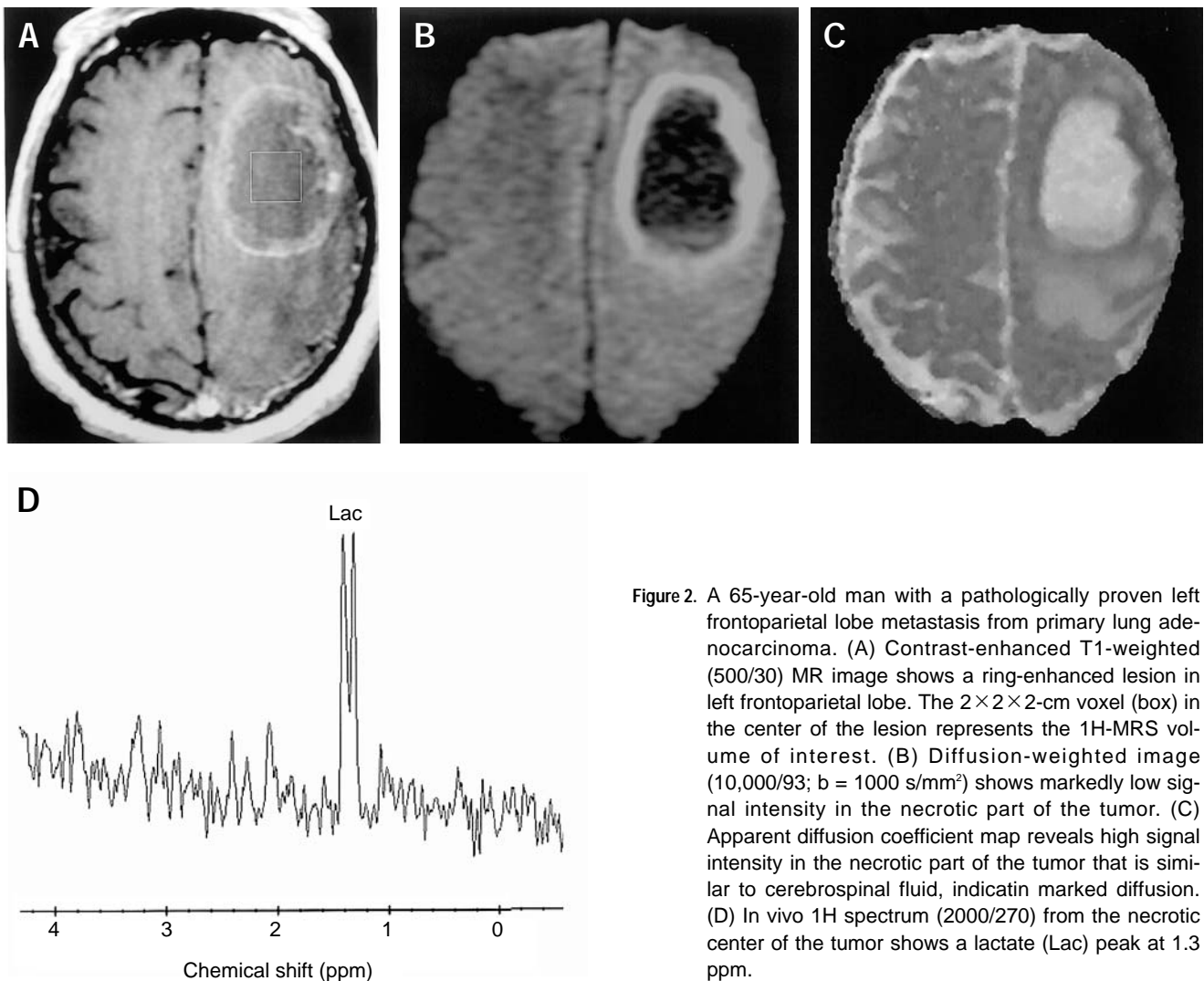


Figure 2. A 65-year-old man with a pathologically proven left frontoparietal lobe metastasis from primary lung adenocarcinoma. (A) Contrast-enhanced T1-weighted (500/30) MR image shows a ring-enhanced lesion in left frontoparietal lobe. The  $2 \times 2 \times 2$ -cm voxel (box) in the center of the lesion represents the  $^1\text{H}$ -MRS volume of interest. (B) Diffusion-weighted image (10,000/93;  $b = 1000 \text{ s/mm}^2$ ) shows markedly low signal intensity in the necrotic part of the tumor. (C) Apparent diffusion coefficient map reveals high signal intensity in the necrotic part of the tumor that is similar to cerebrospinal fluid, indicating marked diffusion. (D) In vivo  $^1\text{H}$  spectrum (2000/270) from the necrotic center of the tumor shows a lactate (Lac) peak at 1.3 ppm.

Burtscher et al. also showed dramatic changes in the follow-up MR spectroscopy of abscesses 29 and 113 days after the beginning antibiotic treatment. All resonances (representing succinate, acetate, alanine, and amino acids) except that of lactate disappeared<sup>(27)</sup>.

The fact that these spectral changes disappear with effective antibiotic treatment, as shown in previous studies<sup>(25,27-29)</sup>, underlines the need for 1) the neuroradiologist or spectroscopist interpreting the spectra to be aware of the case history and 2) the MR spectroscopic examinations to be performed as early as possible, preferably before the start of antibiotic therapy. Knowledge of the expected spectral changes after treatment also offers the

possibility of evaluating the efficacy of nonsurgical treatment of brain abscesses. Further prospective studies with a large number of cases will be necessary for thorough evaluation of particular spectral peaks and time courses of the spectral changes with antibiotic treatment.

In recent studies, it is possible to differentiate anaerobic from aerobic brain abscesses on the basis of metabolite patterns observed on in vivo  $^1\text{H}$  MRS<sup>(30,31)</sup>. The spectral metabolite pattern of the lactate, amino acids, alanine, acetate, and succinate would suggest an obligate anaerobic or a mixture of obligate and facultative anaerobic abscess. On the other hand, the lack of acetate and succinate signal would suggest an obligate

aerobic or a facultative anaerobic abscess. This information may be useful for a timely and appropriate treatment of the abscesses.

Compared with the *in vivo* spectra, a number of additional resonance peaks of lysine at 1.73 and 3.0 ppm, of glutamate/glutamine at 2.09-2.36 ppm, of taurine at 3.24 and 3.42 ppm, of glycine at 3.55 ppm; and of the other amino acids at 3.75 ppm could be observed on the *in vitro* one-dimensional single-pulse or spin echo spectrum, and in the two-dimensional correlation spectroscopy 1H MR spectra<sup>(19,23,30,31)</sup>. It is noteworthy that the *in vitro* measurements may offer complementary information that cannot be extracted from *in vivo* MRS spectra.

#### **DWI vs. 1H MRS of pyogenic brain abscess and necrotic brain tumor**

DWI is a more practical and accurate method compared with 1H MRS<sup>(28)</sup>. There are several reasons for this conclusion. First, DWI scan time is very short (about 40 sec), whereas MR spectroscopy takes a much longer scan time. Single-voxel 1H MRS takes around several minutes and 2D chemical shift imaging takes about 10 minutes according to the repetition time, excitation number, etc. DWI thus is more practical in clinical use, especially in emergent conditions. Second, single-voxel 1H MRS is more limited in voxel size (roughly  $1.5 \times 1.5 \times 1.5$ -cm to  $2 \times 2 \times 2$ -cm) than DWI, and the smaller lesion in MRS will be more affected by the partial volume effect. This is true even in chemical shift imaging, although 2D chemical shift imaging may have a smaller voxel than the single-voxel technique in the case of small lesions. In any case it is likely that 2D chemical shift imaging would still be more time consuming. If the lesion is located peripherally, images of the lesion produced by either single-voxel MRS or 2D chemical shift imaging might be contaminated by the signals from the fat close-by. Third, the treated pyogenic abscess may have high signal on DWI, whereas MR spectroscopic examination shows only lactate/lipid peaks (no amino acid peak [end-products of bacterial breakdown]), a spectrum the same as what is found in necrotic tumor.

Nevertheless, there are some advantages of 1H MRS. First, 1H MRS may provide valuable information

on the changes of metabolites after nonsurgical treatment of brain abscesses, and may thus be helpful for the evaluation of the efficacy of antimicrobial therapy<sup>(25,27-29)</sup>. Second, 1H MRS may differentiate anaerobic from aerobic brain abscesses on the basis of metabolite patterns<sup>(30,31)</sup>. This would be an important information for prompt and appropriate treatment of patients with brain abscesses.

In summary, DWI and 1H MRS are useful tools for the differential diagnosis between brain abscesses and cystic or necrotic brain tumors. The combination of DWI and 1H MRS modalities may provide a more accurate diagnosis of brain abscesses and cystic or necrotic brain tumors than the use of only a single modality.

#### **CONCLUSION**

The MR special techniques, e.g. DWI and 1H MRS, are useful tools for the differential diagnosis between brain abscesses and cystic or necrotic brain tumors. Moreover, it is possible to differentiate anaerobic from aerobic brain abscesses based on the metabolite patterns on the *in vivo* 1H MRS. Further studies should evaluate the possibility of the detection of the overlapped metabolites that cannot be differentiated with one-dimensional localized spectroscopy. Probably one may be able to subcategorize these abscesses in the future with the application of two-dimensional or three-dimensional localized *in vivo* spectroscopy.

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