
Yun Luo,1 Zhen-Ni Guo,1 Peng-Peng Niu,1 Yang Liu,2 Hong-Wei Zhou,2 Hang Jin,1 Yi Yang1

ABSTRACT

Objective: We aimed to investigate the value of three-dimensional (3D) T1 volumetric isotropic turbo spin echo acquisition (VISTA) in the diagnosis of cervical artery dissection (CAD).

Methods: We prospectively included patients who were suspected as having a CAD within 1 month of onset. For T1 VISTA, the diagnosis of the dissection was based on the presence of intramural high-signal, intimal flap, double lumen and aneurysmal dilatation. The final diagnosis of dissection was based on the clinical history, physical examination, and all of the imaging tests.

Results: A total of 46 patients were included in this study. The final diagnosis of CAD was made for 21 patients. Diagnosis of dissection was made for 20 of the 21 patients after assessing T1 VISTA. A definitive diagnosis of dissection was not made for 5 patients (including 3 patients with digital subtraction angiography) before the T1 VISTA examination. The sensitivity and specificity for T1 VISTA were 95.2% (95% CI, 76.2% to 99.9%) and 100% (95% CI, 86.3% to 100%), respectively. The agreement between the two researchers for T1 VISTA for diagnosis of CAD was very good (k=0.91). For patients without acute artery occlusion, all of them had a definite conclusion with or without dissection by T1 VISTA (n=29). However, for 17 patients with acute artery occlusion, the possibility of dissection could not be excluded for 6 of them by T1 VISTA (p=0.001).

Conclusions: 3D T1 VISTA at 3.0 Tesla was useful in the diagnosis of acute CAD. However, for some patients with total occlusion of the artery without typical imaging features of dissection, the unequivocal distinction between intramural haematoma and intraluminal thrombus may be not adequate by T1 VISTA alone. Future studies should investigate whether a follow-up scan, a contrast-enhanced imaging or an optimal VISTA technique could be useful.

INTRODUCTION

Spontaneous cervical artery dissection (CAD) accounts for only ~2% of all ischaemic strokes, but it accounts for 10–25% of ischaemic strokes in young adult patients.3 CAD can cause ischaemic symptoms typically through two mechanisms, which are thromboembolic and haemodynamic compromises.2 3 The early and reliable diagnosis of CAD is highly important for treatment decision-making in the era of precision medicine.4 5

Conventional angiography has long been the gold standard for the diagnosis of CAD.5 However, this method is invasive and does not have the advantage of demonstrating mural haematoma.7 With the advent and development of MRI techniques, like MR angiography (MRA) combined with an axial two-dimensional (2D) fat-saturated spin echo T1-weighted sequence has shown an evident advantage in the diagnosis of CAD.8–10 However, the axial 2D T1-weighted sequence has several limitations. It is time-consuming to cover all arteries and shows poor performance for the arteries with a tortuous course.11 12

The three-dimensional (3D) isotropic T1-weighted spin echo sequence can overcome the limitations of a 2D sequence.13 14 Some researchers have found that a 3D black blood T1-weighted sequence of volumetric isotropic turbo spin echo acquisition (VISTA) offers similar or more information than a 2D T1-weighted spin echo sequence.15–17 We aimed to investigate the value of a 3D T1 VISTA sequence at 3.0 T for the diagnosis of CAD. We also investigated the limitations of 3D T1 VISTA in the diagnosis of CAD.

METHODS

In reporting the current study, the standards for reporting diagnostic accuracy studies were followed.18 This study was not publicly registered.

Participants

Between September 2014 and February 2016, we prospectively included consecutive patients who were suspected as having an acute CAD with the following symptoms or

signs: Horner’s syndrome, unusual neck pain and/or headache, cranial nerve palsy and tinnitus. The CAD was considered acute if the duration of symptoms was ≤30 days. Patients who had the above signs or symptoms, particularly if in combination and/or associated with a cerebral or retinal ischaemia, were highly suspected as having a CAD. Patients who were suspected as CAD by imaging tools of MRA or ultrasound were also included. This study was approved by the institutional review board of the first hospital of Jilin University. Written consent forms were obtained from all patients.

**Imaging protocols**

A 3D T1 VISTA examination was performed on all patients who were included in this study. At least one examination of angiography including MRA, CT angiography (CTA), and digital subtraction angiography (DSA) was performed on each patient. A 3D T1 VISTA protocol was performed using a 3.0 Tesla scanner (Philips Ingenia, Eindhoven, The Netherlands) with a standard 8-channel head/neck coil. The following parameters were used for the 3D T1 VISTA sequence: an oblique coronal plane acquisition, spectral adiabatic inversion recovery fat saturation mode, repetition time (TR)/echo time (TE)=350 ms/19 ms, field of view=280×199×120 cm³, 400×284 matrix, variable refocusing flip angle, slice interval=0; voxel size=0.7×0.7×0.7 cm³, over-sample factor=1.5, and number of excitations=2. The acquisition time was 3 min and 38 s.

**Data analysis**

The analysis of 3D T1 VISTA results was performed by two experienced neuroradiologists who were blinded to all of the patient information and the final diagnosis. Images along the short and long axes of the arteries could be reconstructed at the workstation. For 3D T1 VISTA, the diagnosis of the dissection was based on the presence of the following features: intramural high signal, particularly if it is a semilunar hyperintense signal; intimal flap and/or double lumen; and aneurysmal dilation. The final diagnosis of the dissection was based on the clinical history, physical examination, and all of the imaging tests by two experienced neurologists. For conventional imaging tests, the imaging signs for diagnosis of CAD including double lumen, intimal flap, pearl and string sign, string sign, and tapered occlusion. Disagreement between the two observers were resolved be consensus.

SPSS V.19.0 (IBM, West Grove, Pennsylvania, USA) was used to perform the analysis. Since no gold standard for the diagnosis of CAD was available, the final diagnosis results were chosen to be the reference standard to calculate the sensitivity and specificity (including the corresponding 95% CI) for 3D T1 VISTA. Interobserver agreement for 3D T1 VISTA was examined by using the k-coefficient of agreement. Fisher’s exact test was used for count data. The level of statistical significance was set at p<0.05.

**RESULTS**

A total of 46 patients were included in this study. 3D T1 VISTA examination was performed on each of them. The final diagnosis of the dissection was made for 21 patients (6 females) (table 1). There were eight patients with single carotid artery dissection and nine patients with single vertebral artery dissection. One patient had bilateral carotid artery dissections and bilateral vertebral artery dissections. Three patients had bilateral vertebral artery dissections. One patient with vertebral artery dissection presented with neck pain accompanied by rightsided weakness and leftsided numbness (Brown-Séquard syndrome) (patient number 11). The median age of patients with dissection was 38 years (range 27–74 years old).

The median time between disease onset and the examination with 3D T1 VISTA was 11 days with a range of 3–26 days for patients with dissection. Diagnosis of dissection was made for 20 patients after assessing the 3D T1 VISTA data alone by two researchers. Among these 20 patients, the diagnosis of dissection was not made for five (including 3 patients with DSA) of them before the examination with T1 VISTA. The sensitivity and specificity for 3D T1 VISTA were 95.2% (95% CI, 76.2% to 99.9%) and 100% (95% CI, 86.3% to 100%), respectively. The agreement between the two researches for T1 VISTA for diagnosis of CAD was very good (k=0.91).

Patient number 1 presented with headache and transient motor weakness on the right side (table 1). Duplex ultrasonography and a subsequently CTA showed stenosis and possible dissection of the right vertebral artery, which was confirmed by 3D T1 VISTA (figure 1). T1 VISTA images of the other 18 patients all showed typical crescent-shaped hyperintense signals surrounding the lumen (patients number 2–19).

Patient number 20 presented with aphasia and motor weakness on the right side. Duplex ultrasonography of the left internal carotid artery at 15 days of onset showed a double lumen and intima, which suggested the aetiology was dissection. However, there was no typical crescent-shaped hyperintense signal on VISTA at 26 days of onset, one of the two researchers insisted that the definite diagnosis of dissection could not be made by T1 VISTA alone (figure 2).

Patient number 21 presented with dizziness, headache, and hemiplegia. 3D T1 VISTA at 5 days of onset showed high-signal intensity of the left vertebral artery (figure 3). One of the reconstructed images along the short-axis of left vertebral artery showed a non-typical crescent-shaped hyperintense signal. One researcher suggested that it was unreliable to make a diagnosis of dissection based on the images of T1 VISTA. However, a repeat scan of T1 VISTA at 29 days of onset showed the recanalisation of occlusion and a residual intramural haematoma, which confirmed the diagnosis of dissection.

For patients without artery occlusion, all of them had a definite conclusion with or without dissection by VISTA (n=29). However, for 17 patients with artery occlusion,
the possibility of dissection could not be excluded for six of them by VISTA (0/29 vs 6/17; p=0.001). For these six patients, the aetiology has not been determined for five of them (table 2), even after assessing all of the data, including T1 VISTA and DSA. 3D T1 VISTA showed hyperintense signals in the occluded artery segment without typical features of dissection. One of the patients was <30 years old without any risk factors of cerebrovascular diseases (patient number 25). The follow-up T1 VISTA for two of these five patients showed no recanalisation of the occluded arteries.

**DISCUSSION**

This study showed that the 3D T1 VISTA sequence at 3.0 T is useful in the diagnosis of spontaneous CAD. It can show intramural haematoma of the cervical artery clearly with a single acquisition and acceptable scan time. However, for some patients with totally acute occlusion of the artery without typical crescent-shaped hyperintense signals, a diagnosis of the aetiology of artery occlusion may be difficult by VISTA alone.

Several studies have investigated the efficiency of 3D black blood T1 sequences using variable refocusing
flip-angle turbo-spin-echo imaging in the diagnosis of CAD. All of them showed that a 3D black blood T1 sequence is more useful than conventional imaging tools. Takemoto et al. first reported the value of T1 VISTA sequence at 1.5 T in the diagnosis of CAD. They concluded that a 3D black blood T1 sequence can improve the assessment of intramural haematoma in vertebral artery dissection compared with 2D spin-echo T1-weighted images and time-of-flight MRA (TOF-MRA). Another study showed that abnormal vessel enhancement was recognised in 15 of 15 patients with vertebral artery dissection on contrast-enhanced T1 VISTA images. There are two other similar 3D black blood T1 sequences that are commercially available (T1 CUBE and T1 SPACE from GE Healthcare, Milwaukee, Wisconsin and Siemens, Erlangen, Germany, respectively). Studies showed that these two 3D black blood T1 sequences at 1.5 T or 3.0 T may also be a substitute...
for 2D T1 sequences in the diagnosis of CAD.\textsuperscript{11} \textsuperscript{20} The inter-rater and intrarater agreements were good for 3D black blood T1 sequences.\textsuperscript{11} \textsuperscript{14} There are several advantages of 3D black blood T1 sequences in the diagnosis of CAD compared with traditional imaging tools. First, this method can reveal the vessel wall of intracranial and extracranial arteries in a single acquisition with good image quality, good dark blood contrast, isotropic voxels, and a relatively short scan time. Second, the method can show the mural haematoma as high-signal intensity clearly. The mural haematoma is not easily identified from the source image of MRA and CTA and cannot be revealed by vascular images such as MRA, CTA, and DSA. Third, because of its isotropic volume acquisition, it can obtain the images of multiplanar reformation. Multiplanar reformation images can show the features of dissection at different angles and planes, which is especially helpful for an artery with tortuous course.

A recently published study showed that 3D simultaneous non-contrast angiography and intraplaque haemorrhage (SNAP) imaging can provide non-contrast MRA and vessel wall images simultaneously in a single acquisition with a shorter scanning time.\textsuperscript{21} This study demonstrated excellent agreement with multisequence MRI in evaluating luminal stenosis and intramural haematoma in patients with craniocervical artery dissection. It seems that a 3D SNAP sequence may be a better choice than the aforementioned 3D black blood sequences because it can provide MRA and vessel wall images simultaneously in a single scan. However, this technique has not been widely investigated and is not commercially available.

We also investigated the limitations of 3D T1 VISTA in the diagnosis of CAD by analysing the patients for whom

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**Figure 3** Patient number 21 with a left vertebral artery dissection. (A–C) Coronal of three-dimensional (3D) T1 volumetric isotropic turbo spin echo acquisition (VISTA) image, curved planar reconstruction of 3D T1 VISTA images and short-axis view of the left vertebral artery at 5 days of onset show high-signal intensity of the left vertebral artery. (D–F) A repeat scan at 23 days of onset shows the recanalisation of the occlusion and the residual intramural haematoma.

**Table 2** Characteristics of patients without a definitive diagnosis of with or without dissection

<table>
<thead>
<tr>
<th>Patient</th>
<th>Age</th>
<th>Symptoms</th>
<th>Acute infarction</th>
<th>Location of lesion</th>
<th>3D T1 VISTA</th>
<th>Occlusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>22</td>
<td>40s</td>
<td>Dizziness and hemiplegia</td>
<td>Yes</td>
<td>Bilateral VAs</td>
<td>At 18 days of onset</td>
<td>Yes</td>
</tr>
<tr>
<td>23</td>
<td>50s</td>
<td>Hemiplegia</td>
<td>Yes</td>
<td>Left ICA</td>
<td>At 25 and 48 days of onset</td>
<td>Yes</td>
</tr>
<tr>
<td>24</td>
<td>60s</td>
<td>Dizziness and double vision</td>
<td>Yes</td>
<td>Right VA</td>
<td>At 9 days of onset</td>
<td>Yes</td>
</tr>
<tr>
<td>25</td>
<td>20s</td>
<td>Dizziness and hemiplegia</td>
<td>Yes</td>
<td>Left VA</td>
<td>At 19 days of onset</td>
<td>Yes</td>
</tr>
<tr>
<td>26</td>
<td>60s</td>
<td>Hemiplegia</td>
<td>Yes</td>
<td>Right VA</td>
<td>At 15 days of onset</td>
<td>Yes</td>
</tr>
</tbody>
</table>

3D, three-dimensional; ICA, internal carotid artery; VA, vertebral artery; VISTA, volumetric isotropic turbo spin echo acquisition.
a definite conclusion was not reached, with or without dissection by VISTA. Disagreement between the two readers for T1 VISTA existed for two patients who had acute artery occlusion. For one patient, the diagnosis of dissection was made by comprehensive analysis of ultrasound scan, T1 VISTA, and the other information. For the other patient, the diagnosis of dissection was made after a second scan with T1 VISTA. For five patients with acute artery occlusion, the aetiology has not been determined even after assessing all of the data including VISTA and DSA. All of this suggested that, for some patients with acute artery occlusion in whom even the aetiology may be dissection, the diagnosis of dissection cannot be made by T1 VISTA if there was no typical imaging features of dissection.

In this study, patients without artery occlusion all had a definite conclusion with or without dissection. However, for patients with artery occlusion, the possibility of dissection may not be excluded, especially for patients with occlusion of the vertebral artery because of its smaller size. This suggested that the difference between intramural haematoma and intraluminal thrombus may be difficult with 3D T1 VISTA even at 3.0 T for some patients. For this group of patients, a follow-up scan, a contrast-enhanced scan, or an optimal VISTA technique may be helpful (patient number 16).

There were several limitations in this study. First, the sample size was small and the contrast-enhanced imaging was not performed. Contrast-enhanced 3D T1 VISTA might be useful in manifesting subtle structure abnormalities, assessing vessel inflammatory reaction, and distinguishing intramural haematoma from intraluminal thrombus. Second, because previous studies already showed that a 3D black blood sequence can provide similar information and may be a substitute for 2D sequence, the 2D T1 black blood sequence was not performed for the patients. Third, we did not perform a follow-up 3D T1 VISTA scan for certain patients. Finally, all of the information including T1 VISTA were used to make the reference standard, which may lead to inaccurate results of the sensitivity and specificity for T1 VISTA.

CONCLUSIONS

In conclusion, this study showed that a 3D T1 VISTA sequence at 3.0 T is useful in the diagnosis of spontaneous CAD. This sequence can show intramural haematoma of the cervical artery clearly with a single acquisition and acceptable scan time. However, for some patients with totally acute occlusion of the artery without typical features of dissection, the unequivocal distinction between intramural haematoma and intraluminal thrombus may still be difficult with T1 VISTA alone. Future studies should investigate whether a repeat scan or an optimal VISTA technique would be useful for making a definitive diagnosis.

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