INTRODUCTION

As a new clinical and pathological group, noncompaction of the ventricular myocardium (NVM) is categorized as primary cardiomyopathy by the American Heart Association (AHA) (1). It was first described in 1990 and is thought to be related to the arrest of myocardial development (2). It is characterized by prominent trabeculations on the luminal surface of the ventricle, deep intertrabecular recesses communicating with ventricular lumen and regional wall motion abnormalities. Though it usually affects the left ventricle (LV) alone, right ventricular (RV) noncompaction may also be involved (< 50% of cases) (3). NVM can be isolated (without other structural cardiac abnormalities) and or co-exist with other congenital cardiac malformations. This cardiac abnormality is increasingly detected by echocardiography, computed tomography (CT) and magnetic resonance imaging (MRI). However, there is no universally accepted definition and diagnostic criteria of NVM and many cases are easily overlooked or misdiagnosed in clinical practice. CMRI is an excellent modality for the noninvasive assessment of patients with NVM, for its high resolution, multiplane imaging and superior contrast between the myocardium and intracardiac blood-pool (3-5). Currently, there are few studies describing the MRI features of NVM. The purpose of our study was to summarize and...
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describe abnormal changes in CMRI in patients with isolated noncompaction of ventricular myocardium (INVM).

**MATERIALS AND METHODS**

**Patient Population**

Eleven patients were enrolled between January 2009 to April 2010. They were at first suspected cases of INVM by echocardiography, but then recommended to undergo CMRI and diagnosed with INVM by CMRI. Exclusion criteria for performing CMRI included general contradictions to MRI examination (claustrophobia, pacemaker, coronary stents, bypass grafts and aneurysm clamp). Detailed information about history of drug allergies and renal function were acquired before contrast administration. Our institution does not require Institutional Review Board (IRB) approval for this retrospective study and informed consent was waived.

**Clinical Data**

The clinical features of 11 patients were recorded by cardiologists when they first presented to the hospital. Electrocardiographic (ECG) findings were evaluated by two cardiologists (with more than 5 years and 10 years experience with ECG diagnosis, respectively). The clinical data of 11 patients were acquired by electronic medical record systems and then comprehensively reevaluated by a senior cardiologist.

**Cardiac MRI**

Cardiac MRI was performed on a 1.5-Tesla (T) clinical scanner (Magnetom Avanto; Siemens AG, Erlangen, Germany) equipped with quantum gradients (slope, 45 mT/m; slew rate, 200 mT/m/s) and a six-element coil for each patient. Three ECG leads were placed on the left anterior hemithorax (vector ECG) and readjusted to obtain reasonable amplitude and a clean signal trace in the monitoring display. The MRI sequences and parameters were as follows: Axial steady-state free precession (SSFP) (repetition time [TR]/echo time [TE], 292.1/1.22 ms; matrix, 136 x 256; spatial resolution, 2.0 x 1.3 x 6.0 mm) and a coronal single shot fast spin echo sequence (SS-FSE) (TR/TE, 959.0/26.0 ms; matrix, 104 x 256; spatial resolution, 2.3 x 1.3 x 6.0 mm) were initially obtained covering the whole heart. All the localization sections including the 2-chamber, 4-chamber, and short-axis sections were acquired using SSFP with the following parameters: TR/TE, 300/1.21 ms; spatial resolution, 2.1 x 1.3 x 8.0 mm. Dark-blood fast spin echo (SE) T1-weighted sequence (TR/TE, 842.1/30 ms; matrix, 125 x 256; spatial resolution, 2.2 x 1.3 x 5.0 mm) and fast SE T2-weighted sequence (TR/TE, 1725.5/77 ms; matrix, 160 x 256; spatial resolution, 1.7 x 1.3 x 5.0 mm) were subsequently performed in LV short-axis and long-axis views. All SE images were acquired in end-diastole and SSFP cine imaging (TR/TE, 39.75/1.12 ms; matrix, 156 x 192; spatial resolution, 1.8 x 1.8 x 6.0 mm) with GRAPPA (generalized autocalibrating partially parallel acquisition) parallel imaging at an acceleration factor of two were carried out in a 2-, 4-chamber and LV short-axis covering whole LV. T-SENSE parallel imaging was alternatively used for the patients with poor breath-hold behavior using an acceleration factor of three to speed up the acquisition. In nine patients, 0.2 mmol/kg body weight of Gadopentetate Dimeglumine (Magnevist; Schering AG, Berlin, Germany) was injected using a power injector at a rate of 2 ml/s, followed immediately by 20 ml of saline at the same rate. Ten to fifteen minutes after contrast administration, delayed-enhancement MRI was performed using the inversion recovery spoiled gradient echo (IR-SPGR) (TR/TE, 670/3.36 ms; matrix, 156 x 256; spatial resolution, 2.0 x 1.5 x 8.0 mm; time to inversion, 200-350 ms) with magnitude and phase sensitive reconstruction in ventricular long-axis and short-axis views.

**Cardiac MRI Diagnostic Criteria**

Diagnostic criteria of left INVM typically included the presence of prominent or numerous trabeculations, multiple deep intertrabecular recesses communicating with the ventricular cavity, and a two-layered appearance of the myocardium with an increased ratio of noncompacted to compacted myocardium (NC/C ratio > 2.3) in diastole (4). The RV apex often presents with hypertrophic trabeculae in a normal individual, so it is difficult to differentiate between variants of normal and pathologic patterns. In previous studies, no specific CMRI criteria have been proposed for the diagnosis of right INVM due to the small sample size described. In this study, echocardiographic criteria for right INVM, including prominent two-layered structure, deep intertrabecular spaces communicating with RV cavity and thickness of the noncompacted RV greater than 75%, were used (6).

**Cardiac MRI Data Analysis**

All images were evaluated by two senior radiologists (more than 10 years experience with cardiovascular imaging),
and disagreement of evaluation was settled by a consensus reading.

Cine images were analyzed with the Argus software (Siemens). The LV wall was divided into nine segments: the whole apex was one segment (apical segment); at the base and at the midventricular level, the LV was divided into four segments each (inferior, lateral, anterior and septal) (3). In LV, the ratio of NC/C was measured manually on the long-axis and short-axis view on the SSFP cine sequence in diastole. At the base and mid-ventricular level for the short-axis view, the location with the most pronounced trabeculations were chosen for measurement of the thickness of noncompacted and the compacted myocardium perpendicular to the compacted myocardium. The NC/C ratio of each segment was independently measured by two readers and their mean value was used for the final diagnosis. The apex was measured on long-axis of LV using the same method. Ventricular volume was measured as normal or dilated.

Delayed-enhancement segments were independently judged visually on magnitude and phase sensitive reconstruction images in ventricular long-axis and short-axis views by two readers, and disagreement was settled by a consensus reading.

RESULTS

Clinical Data

The patient series consisted of 11 patients; nine males and two females with a mean age of 35 years and an age range of 15-58 years old. Eight patients (8 of 11, 73%) were referred to our hospital with a history of chest tightness or palpitations. One (1 of 11, 9%) suffered from sudden syncop and one adolescent patient (1 of 11, 9%) from sudden edema in extremities. One (1 of 11, 9%) was asymptomatic and detected by echocardiographic examination by chance.

Electrocardiographic findings were summarized as followed: three patients (3 of 11, 27%) with normal ECG performance, one (1 of 11, 9%) with LV high voltage, one (1 of 11, 9%) with RV high voltage, left atrium hypertrophy and ST-T changes, one (1 of 11, 9%) with only ST-T changes, two (2 of 11, 18%) with pre-excitation syndrome, and three (3 of 11, 27%) with ventricular premature contraction. The patient clinical data were summarized in Table 1.

Cardiac MRI Data

Right INVM (1 of 11, 9%), left INVM (7 of 11, 64%), and bilateral INVM (3 of 11, 27%) were found. In SSFP combined with fast SE sequence, prominent trabecular recessations, deep intertrabecular recesses communicating with ventricle cavity, and a regional ventricular motion abnormality were found. Fast SE sequence showed myocardial wall structure with both a thin epicardial compacted zone and an extremely thickened endocardial noncompacted zone which mimicked hypertrophic cardiomyopathy (Fig. 1). Three patients showed high signals within the trabecular structures on black-blood fast SE T2-weighted sequence, providing obvious contrast with black blood in the cavity. The SSFP cine sequence displayed a two-layered structure of ventricular myocardium consisting of an outer thinning layer and inner spongiform layer. It also showed high signals within the trabecular recesses suggestive of blood flow communicating with the ventricular cavity (Fig. 2).

A total of 99 segments were analyzed in the LV and noncompacted segment was found in 36 segments (36 of 99, 36%). The left apex was the most frequently affected segment (10 of 99, 10%), in ten of eleven patients (10 of 11, 91%); seven mid-lateral, six mid-anterior, six mid-inferior, three basal-anterior and three basal-lateral segments were also found to be affected. The basal-inferior was the least affected segment with only one segment was involved. In this study, there was no basal-septal or mid-septal segment affected. The mean NC/C ratio of the 36 segments was 3.1, with a range of 2.3-9.2, and a maximum NC/C ratio happened on the apex.

Systolic function of the LV wall was impaired to some degree in nine patients and normal cardiac function was only measured in two patients (No. 1 and No. 10). The average LV ejection fraction in the 11 patients was 40% (range: 23% to 70%), the LV end-diastolic volume and LV end-systolic volume were expanded to a different degree.

Delayed contrast-enhancement was performed in nine patients, with six patients having positive results (6 of 9, 67%). Only one case of apical delayed enhancement was found. The delayed enhancement area was present in both compacted and noncompacted segments and the patterns of enhancement included subendocardial and transmural myocardial enhancement.

Eight patients had dilated ventricles (8 of 11, 73%). Among the 10 patients with left involved NVM (7 with left NVM and 3 with bilateral NVM), seven had a dilated LV (7 of 10, 70%). Among the four patients with right involved NVM
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(1 with right NVM and 3 with bilateral NVM), three (3 of 4, 75%) had a dilated RV.

One female patient (1 of 11, 9%) with left NVM had a ventricular aneurysm in the LV apex (Fig. 3). An apical thrombus was found in one adolescent patient (1 of 11, 9%) (Fig. 4), and thrombus of the right brachiocephalic vein and vena cava were subsequently found by other examinations.

**DISCUSSION**

Noncompaction of ventricular myocardium can co-exist with other congenital cardiac disorders such as an atrial septal defect, interventricular septal defect, pulmonic...
stenosis, anomalous pulmonary venous connection, coronary artery anatomical anomalies and Ebstein’s syndrome (6-9). In our study, all patients enrolled were isolated NVM and the majority were male. In previous studies, male preponderance has been suggested among adult INVM patients, but no proper evidence has been confirmed (10).

Cardiac MRI is increasingly recognized as an alternative modality for the noninvasive assessment of patients with INVM. Compared with traditional echocardiography, CMRI has less operator dependence, superior spatial resolution, higher contrast between blood and myocardium, which can provide better delineation of the abnormal trabeculations in INVM patients (11). It is particularly useful when the lesions are confined to the ventricular apex, which is difficult to be detected by echocardiography (4, 12). Unfortunately, we were not able to compare echocardiographic and MRI findings, because the affected segments could not be demonstrated in three patients for difficult echocardiographic windows and echocardiographic examination results were not available. Although echocardiographic examination showed hypertrabeculation in the other seven patients, this evidence was not enough to make a specific diagnosis of INVM.

In addition, contrast-enhanced CT is capable of displaying INVM and allow the evaluation of coronary arteries to exclude anomalies or significant stenosis, which is usually not possible with MRI or echocardiography. However, CT remains limited for the detection of myocardial viability, which can be obtained in CMRI by delayed enhancement imaging. To date, there is no consensus about MRI diagnostic criteria for INVM and the main cause of this is the threshold of NC/C ratio. The generally accepted criterion was proposed by Petersen et al. (4) in 2005. They concluded that the ratio of NC/C > 2.3 in the diastole distinguished pathological LV noncompaction with high specificity and sensitivity. Therefore, we used NC/C > 2.3 in diastole as a diagnostic cutoff value to differentiate between normal and pathologic patterns in our series.

Delayed enhancement was seen in six of the nine patients in our study. The patterns of enhancement included subendocardial enhancement and transmural myocardial enhancement. It has been previously reported that delayed enhancement could be found in 70% of INVM.
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patients, in both noncompacted compacted segments (3, 5), which was consistent with our findings. Regrettfully, the perfusion sequence was not performed in this study. However, hypoperfusion had been previously demonstrated by CMRI and ultrafast CT in NVM patients (13, 14). The histological findings support that delayed enhancement and hypoperfusion are closely related to the abnormalities of the myocardial microcirculation, which result in necrosis, fibrosis, and scar (15-18). Interestingly, the most frequently involved segment was the LV apex, while only one apex had delayed enhancement in our series. This may be due to thinned apical myocardium and the subsequent difficult observation in this area.

In this study, one ventricular aneurysm was observed in LV apex. Few cases of INVM associated with ventricular aneurysm have been reported (5). This may be due to regional ventricular wall thinning (19). Thromboembolic events were observed in one adolescent patient, including thrombus in the LV apex, the right brachiocephalic vein and vena cava. The deep recesses may worsen the risk of thrombus formation and be an additional factor for this serious complication (12).

Four of 11 cases were RV involving INVM in our study (1 right NVM and 3 bilateral NVM). Because it is usually difficult to differentiate pathologic noncompaction from normal trabeculations, the prevalence of RV involvement may have been underestimated in the past. Previously, right INVM has been reported in some cases by CMRI or echocardiography (20). Based on a pathohistologic study, Burke et al. (6) proposed a histological criterion for right NVM: transmural thickness of the noncompacted RV greater than 75%. This criterion was used in our series. It also has been suggested that a diagnosis of right NVM should be made associated with dilatation of the RV (21). In our series, three bilateral INVM had dilated ventricular lumen and only one right INVM had a normal volume. According to the data from our own experience, the dilatation of the RV may be a helpful supportive feature for the diagnosis of right NVM.

The clinical manifestation of INVM is highly variable, ranging from no symptoms to a progressive deterioration in cardiac function, arrhythmias, and systemic thromboembolic events. Eight of the 11 patients (73%) in our study had abnormalities in the resting ECG, which were nonspecific, including voltage signs of LV and RV hypertrophy, ST-T segment changes, ventricular premature beat, intraventricular block and pre-excitation syndrome. It has been reported that 94% of adult INVM patients had abnormal ECG manifestations, with ventricular arrhythmias being the most common (12). This may be due to myocardial ischemia and scar tissue induced by microcirculatory dysfunction. Moreover, the prominent trabecular meshwork on the luminal surface of the ventricle was presumed to increase the probability of an ectopic pacemaker and reentrant pathways (12).

The main limitation of our study was that we could not testify the accuracy of > 2.3 as a cutoff value in differentiating pathology myocardium from normal prominent myocardium because of small sample size and retrospective method used in our study. The cases enrolled were not confirmed by the gold standard, which was endomyocardial biopsy. A more comprehensive sequence such as perfusion imaging should be performed in a future study, which may help in the prediction of the prognosis and provide suggestions for treatment.

In conclusion, although the diagnosis of INVM is usually made by echocardiography, CMRI has been increasingly recognized to be particularly useful for assessing the INVM patient. Our findings support the clinical use of CMRI in diagnosing INVM, especially for those patients with suspected diagnoses for an echocardiographic examination. Cine SSFP sequence is a major sequence of CMRI that can represent the contractibility of the myocardium and the ventricular function (22). Characteristic CMRI features of INVM can be clearly revealed in the cine SSFP sequence and black-blood fast SE sequence, which are both important to MRI diagnosis.

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