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## **Original Article**

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#### Author for correspondence:

Hideharu Oka, Department of Pediatrics, Asahikawa Medical University, 2-1-1-1, Midorigaoka-Higashi, Asahikawa, 078-8510, Japan. Tel: +81-16668-2481; Fax: +81-16668-2489. E-mail: oka5p@asahikawa-med.acjp

# breath-holding and free-breathing

Hideharu Oka<sup>1</sup><sup>®</sup>, Kouichi Nakau<sup>1</sup>, Sadahiro Nakagawa<sup>2</sup>, Yuki Kobayashi<sup>3</sup>, Rina Imanishi<sup>1</sup>, Kunihiro Iwata<sup>2</sup> and Hiroshi Azuma<sup>1</sup>

<sup>1</sup>Department of Pediatrics, Asahikawa Medical University, Hokkaido, Japan; <sup>2</sup>Section of Radiological Technology, Department of Medical Technology, Asahikawa Medical University Hospital, Hokkaido, Japan and <sup>3</sup>Asahikawa Medical University, Hokkaido, Japan

Comparison of myocardial T1 mapping during

## Abstract

Background: T1 mapping is a recently developed imaging analysis method that allows quantitative assessment of myocardial T1 values obtained using MRI. In children, MRI is performed under free-breathing. Thus, it is important to know the changes in T1 values between free-breathing and breath-holding. This study aimed to compare the myocardial T1 mapping during breath-holding and free-breathing. Methods: Thirteen patients and eight healthy volunteers underwent cardiac MRI, and T1 values obtained during breath-holding and free-breathing were examined and compared. Statistical differences were determined using the paired t-test. Results: The mean T1 values during breath-holding were 1211.1 ± 39.0 ms, 1209.7  $\pm$  37.4 ms, and 1228.9  $\pm$  52.5 ms in the basal, mid, and apical regions, respectively, while the mean T1 values during free-breathing were 1165.1  $\pm$  69.0 ms, 1103.7  $\pm$  55.8 ms, and 1112.0 ± 81.5 ms in the basal, mid, and apical regions, respectively. The T1 values were lower during free-breathing than during breath-holding in almost all segments (basal: p = 0.008, mid: p < 0.001, apical: p < 0.001). The mean T1 values in each cross section were 3.1, 7.8, and 7.7% lower during free-breathing than during breath-holding in the basal, mid, and apical regions, respectively. Conclusions: We found that myocardial T1 values during free-breathing were about 3-8% lower in all cross sections than those during breath-holding. In free-breathing, it may be difficult to assess myocardial T1 values, except in the basal region, because of underestimation; thus, the findings should be interpreted with caution, especially in children.

Cardiac magnetic resonance has been widely used for myocardial scar and perfusion assessment and evaluation. T1 mapping is a recently developed imaging analysis method that allows quantitative assessment of myocardial T1 values obtained using MRI.<sup>1</sup> In the past, delayed-contrast MRI was commonly used to evaluate myocardial damage, but it allowed assessment of local lesions alone and not the entire myocardium. The advantages of T1 mapping are that it measures T1 values across the entire myocardium, which is ideal for the evaluation of diffuse lesions, and that it can be used in patients with renal dysfunction because it does not involve the use of a contrast medium. It is reported to be useful in evaluating myocardial damage in patients with cardiomyopathy or after chemotherapy.<sup>2–4</sup>

However, T1 mapping requires a breath-holding sequence, and children who are examined under sedation cannot hold their breath. Hence, the imaging is performed under free-breathing. It is possible to prevent blurring of images secondary to the heart's movement with motion correction; however, free-breathing causes more significant distortion than breath-holding due to the respiratory motion.<sup>5</sup> Since paediatric cardiac patients, such as those with CHD or cardiomyopathy, may develop myocardial damage from an early age, it would be useful if the myocardial properties could be assessed under free-breathing. Therefore, it is important to know the differences in T1 values between free-breathing and breath-holding images. In



Figure 1. Analysis methods. The images required for T1 mapping were taken in three short-axis slices (basal, mid, and apical) and measured T1 values.

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 Table 1. The demographic characteristics

	Cases (n = 21)
Age	21.6 ± 7.1
Sex (male:female)	10:11
Height (cm)	163.6 ± 9.1
Body weight (kg)	58.5 ± 13.7
Heart rate (bpm)	71.7 ± 14.8
Patients	
CHD	11
Haematological disease	2
Healthy volunteers	8

Data are given as mean and standard deviation.

#### Table 2. T1 mapping data

	Breath-holding	Free-breathing	р
Basal			
Anterior	1196.2 ± 49.6	1142.0 ± 78.1	0.003
Anteroseptal	1227.1 ± 45.2	1177.1 ± 68.3	0.002
Inferoseptal	1223.6 ± 38.2	1187.3 ± 70.2	0.043
Inferior	1247.7 ± 60.4	1195.8 ± 71.3	0.010
Inferolateral	1197.7 ± 47.5	1146.3 ± 68.3	0.005
Anterolateral	1174.2 ± 45.7	1142.3 ± 77.3	0.113
Mean	1211.1 ± 39.0	1165.1 ± 69.0	0.008
Mid			
Anterior	1192.6 ± 60.7	1075.8 ± 64.2	<0.001
Anteroseptal	1228.4 ± 43.9	1124.7 ± 57.3	<0.001
Inferoseptal	1238.4 ± 36.7	1142.1 ± 64.6	<0.001
Inferior	1223.2 ± 34.5	1112.9 ± 60.5	<0.001
Inferolateral	1181.7 ± 42.4	1085.1 ± 63.4	<0.001
Anterolateral	1193.7 ± 44.8	1081.6 ± 59.5	<0.001
Mean	1209.7 ± 37.4	1103.7 ± 55.8	<0.001
Apical			
Anterior	1228.9 ± 71.8	1100.8 ± 77.9	<0.001
Septal	1239.6 ± 59.9	1126.3 ± 86.6	<0.001
Inferior	1213.2 ± 64.8	1106.6 ± 92.2	0.001
Lateral	1234.1 ± 69.5	1114.3 ± 97.2	<0.001
Mean	1228.9 ± 52.5	1112.0 ± 81.5	<0.001

Data are given as mean and standard deviation.

the present study, we investigated the free-breathing effect by comparing myocardial T1 values obtained under breath-holding and free-breathing.

#### **Materials and methods**

Among the patients who had undergone cardiac MRI at the Asahikawa Medical University hospital since April 2020, 22 were included in the study after T1 mapping of the myocardium was



Figure 2. T1 mapping for each section. Images of T1 mapping for each section. The upper row is a breath-holding image, and the lower row is a free-breathing image.

carried out during breath-holding and free-breathing. Twelve patients had heart disease, two had undergone chemotherapy for haematological disease, and eight were healthy subjects. The breakdown of patients with heart disease was as follows: 3 patients were post-operative for tetralogy of Fallot, 1 patient was postoperative for double outlet right ventricle, 1 patient was post-operative for transposition of the great artery, 1 patient was post-operative for total anomalous pulmonary venous return, 1 patient was post-operative for ventricular septal defect, 1 patient was post-operative for tricuspid atresia, 1 patient had ventricular septal defect, 1 patient had post-myocarditis, and 1 patient had WPW syndrome. The patients had to be old enough to hold their breath during the test; therefore, we targeted students from middle school and above. The volunteers performed an echocardiogram to confirm that there were no abnormalities in heart function or intra-cardiac structures. Vital signs were monitored during the examination. Patients underwent cardiac MRI in a 3.0 Tesla whole-body scanner (MAGNETOM Vida 3.0 T; Siemens Healthcare, Erlangen, Germany) equipped with dual-source, parallel radiofrequency transmission, and 18-element cardiac phasedarray coils for radiofrequency reception. The Modified Look-Locker Inversion recovery sequence was used for T1 mapping. Basal, mid, and apical-ventricular short-axis slices were scanned, and myocardial T1 values were obtained.<sup>6</sup> Other scan parameters were field of view – 360 mm, matrix size –  $256 \times 144$ , acceleration factor - 2, repetition time - 2.53 ms, echo time - 1.06 ms, slice thickness - 8 mm, flip angle - 35°, and shot mode - true fast imaging with steady-state precession pulse sequence, using the 5(3)3 scheme. The images required for T1 mapping were taken in three short-axis slices (basal, mid, and apex) and measured during breath-holding and free-breathing (Figs 1 and 2). After visual inspection of the images, motion correction was implemented for cardiac movement and cross sections were compared for significant deviations between the two different breathing modes. The workstation (cvi42; ENTORESS, Mie, Japan) was used for analysis, and the region of interest settings for T1 measurements were carefully performed to exclude the endocardial blood pool, myocardium, or adjacent tissues. The T1 values were classified according to the American Heart Association 16-segment model.<sup>7</sup>

All parameters are expressed as mean  $\pm$  standard deviation (SD) values. The Shapiro–Wilk normality test was used to check the normal distribution of the data. Statistical differences were determined using the paired-samples t-test. A p-value < 0.05 indicates statistical significance. Statistical analysis was performed using the Statistical Package for the Social Sciences version 24.0 (IBM Corp., Armonk, New York, United States of America).



**Figure 3.** (*a*-*c*) T1 values for each section. Graphs of T1 values at each cross section for breath-holding and free-breathing. Free-breathing T1 values showed a lower than those of breath-holding in almost all segments. Comparing the mean values of each cross section, T1 values were lower by 3.1% for basal, by 7.8% for mid, and by 7.7% for apical as a result of free-breathing than those of breath-holding. \* p < 0.05, † p < 0.01.

This study was performed according to the 2013 Declaration of Helsinki and the current ethical guidelines. It was approved by our institutional ethics board (Approval No. 19,250). Informed consent was obtained from all the patients or their legally authorised representatives.

## Results

Table 1 shows the demographic data of the patients. One patient with post-operative total anomalous pulmonary venous return was excluded because deep breathing caused large artefacts in the images taken during free-breathing that could not be analysed. The mean age of the patients was  $21.6 \pm 7.1$  years, and 47.6% were males. The mean height and weight were  $163.6 \pm 9.1$  cm and  $58.5 \pm 13.7$  kg, respectively. The mean heart rate was  $71.7 \pm 14.8$  bpm.

The T1 values for breath-holding and free-breathing are shown in Table 2. The mean T1 values during breath-holding were  $1211.1 \pm 39.0$  ms,  $1209.7 \pm 37.4$  ms, and  $1228.9 \pm 52.5$  ms in the basal, mid, and apical regions, respectively, and the mean T1 values during free-breathing were 1165.1 ± 69.0 ms, 1103.7  $\pm$  55.8 ms, and 1112.0  $\pm$  81.5 ms in the basal, mid, and apical regions, respectively. The Shapiro-Wilk normality test confirmed that each sample followed a normal distribution. The lower T1 values were observed during free-breathing than during breathholding in almost all segments (basal: p = 0.008, mid: p < 0.001, apical: p < 0.001). The mean change in T1 values during free-breathing and breath-holding in each cross section was 3.1, 7.8, and 7.7% for the basal, mid, and apical regions, respectively (Fig 3a-c). Differences by sex were also examined. In both sexes, the lower T1 values were observed during free-breathing than during breath-holding in almost all segments. In the Bland-Altman analysis, the measurements in the basal region were randomly arranged around 0, indicating only random error. The measurements in the mid and apical regions often showed values higher than 0, indicating a fixed error. This indicated the presence of a higher degree of bias for T1 values during breath-holding than during free-breathing. (Fig 4a-c).

To ensure our methods' reproducibility, we examined the intra-observer and inter-observer differences in the T1 values in



Figure 4. (a-c) Bland-Altman Curve between breath-holding and free-breathing images. (a) Basal region, (b) mid region, and (c) apical region.

the LV basal, mid, and apical walls. Strong correlation was observed between intra-observer and inter-observer values (r = 0. 95 and r = 0. 96, respectively).

#### Discussion

In the present study, we found that free-breathing T1 values in all cross sections of the basal, mid, and apical regions were lower by approximately 3–8% than those of breath-holding T1 values.

T1 mapping provides a quantitative estimate of the myocardial properties and is useful for evaluating myocardial fibrosis and oedema.<sup>1</sup> Echocardiography and CT cannot quantitatively and objectively assess myocardial characteristics. In addition, although conventional delayed-contrast MRI is used for determining localised myocardial fibrosis, it is difficult to assess the entire myocardium. Thus, we think that T1 mapping can assess the entire myocardium and any potential damage quantitatively and will help monitor the progression of myocardial damage. In children, breath-holding cardiac MRI is difficult to perform unless manually ventilated under tracheal intubation and the examination is

performed during free-breathing under sedation with intravenous anaesthesia. Therefore, it is essential to understand the changes in the T1 values due to respiration. In the present study, we found that free-breathing T1 values in all cross sections were 3–8% lower than breath-holding T1 values. Figure 2 shows clear changes in T1 images in the mid and apical regions between breath-holding and free-breathing. This indicates that T1 values in the basal region may be clinically assessable.

Myocardial T1 values are known to be affected by magnetic fields, heart rate, and respiratory motion.<sup>8</sup> In this study, we found that respiration reduced the myocardial T1 values. Respiratory motion is known to cause errors in the pixel-by-pixel estimation of T1, degrading the final map.<sup>9</sup> Moreover, the motion artefact of the heart reduces T1 values.<sup>10,11</sup> Since the movement of the diaphragm affects the movement of the myocardial T1 values. Our results suggest that the mid and apical regions are strongly affected by the artefacts caused by respiration, resulting in lower T1 values, whereas the basal region is less affected by respiration. Hence, myocardial T1 values in the basal region can be evaluated even

in children. Bush et al reported that using the prospective motion correction strategy to measure T1 values during free-breathing produced results that were comparable to those during breathholding to lessen the impact of breathing. It may be important to consider this method in children.<sup>12</sup>

The first limitation of this study is the small number of cases. In order to generalise this result, we need to accumulate more cases by performing large prospective studies. Second, the mean age of our sample population is more reflective of adolescent/adult patients, and there were no cases of ventilated children. Thus, it is unclear whether these results apply to infants and young children. Third, because free-breathing inevitably causes positional deviations in the measurement area, we used motion correction for visual checks. We tried to perform measurements in the same crosssectional area; however, the results can be affected by even slight positional deviations.

In our study, we found that free-breathing T1 values in all cross sections were approximately 3–8% lower than breathholding T1 values. It may be difficult to accurately assess myocardial T1 values with free-breathing, except in the basal region, because of underestimation. This should be taken into consideration when interpreting myocardial T1 values in children with CHD.

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**Conflicts of interest.** The authors declare that they have no conflicts of interest.

**Ethical standards.** This study was approved by the Asahikawa Medical University Institutional Review Board. All study participants provided informed consent.

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