

Quantitative assessment of right ventricular function and pulmonary regurgitation in surgically repaired tetralogy of Fallot using 256-slice CT : comparison with 3-Tesla MRI

山崎, 誘三

<https://doi.org/10.15017/1500556>

出版情報 : 九州大学, 2014, 博士 (医学), 課程博士
バージョン :
権利関係 : やむを得ない事由により本文ファイル非公開 (2)



**Quantitative Assessment of Right Ventricular Function and Pulmonary
Regurgitation in Surgically Repaired Tetralogy of Fallot Using 256-slice CT:
Comparison with 3 Tesla MRI**

Yuzo Yamasaki¹, Michinobu Nagao², Kenichiro Yamamura³, Masato Yonezawa¹,
Yoshio Matsuo¹, Satoshi Kawanami², Takeshi Kamitani¹, Ko Higuchi¹,
Ichiro Sakamoto⁴, Yuichi Shiokawa⁵, Hidetake Yabuuchi⁶, Hiroshi Honda¹

Departments of ¹Clinical Radiology, ²Molecular Imaging & Diagnosis, ³Pediatrics,

⁴Cardiovascular Medicine, ⁵Cardiovascular Surgery, and ⁶Health Sciences,

Graduate School of Medical Sciences, Kyushu University

Abstract

Objectives: To compare 256-slice cardiac computed tomography (CCT) with cardiac magnetic resonance (CMR) imaging to assess right ventricular (RV) function and pulmonary regurgitant fraction (PRF) in patients with repaired tetralogy of Fallot (TOF)

Methods: 33 consecutive patients with repaired TOF underwent retrospective ECG-gated CCT and 3-Tesla CMR. RV and left ventricular (LV) end-diastolic volume (EDV), end-systolic volume (ESV), stroke volume (SV), and ejection fraction (EF) were measured using CCT and CMR. PRF-CCT (%) was defined as: $(RVSV - LVSV) / RVSV$. PRF-CMR (%) was measured by the phase-contrast method. Repeated measurements were performed to determine intraobserver and interobserver variability.

Results: CCT measurements, including PRF correlated highly with the CMR reference ($r=0.71$ to 0.96). CCT overestimated RVEDV (mean difference, 17.1 ± 2.9 ml), RVESV (12.9 ± 2.1 ml) and RVSV (4.2 ± 2.0 ml), and underestimated EF (-2.6 ± 1.0 %) and PRF (-9.1 ± 2.0 %) compared with CMR. The limits of agreement between CCT and CMR were in a good range for all measurements. The variability in CCT measurements was lower than those in CMR. The estimated effective radiation dose was 7.6 ± 2.6 mSv.

Conclusions: 256-slice CCT can assess RV function and PRF with relatively low-dose radiation exposure in patients with repaired TOF, but overestimate RV volume and underestimate PRF.

Key points (3 to 5 sentences, up to 12 words)

256-slice CT assessment of RV function is highly reproducible in repaired TOF.

Pulmonary regurgitation can be evaluated by biventricular systolic volume difference.

CT overestimates RV volume and underestimates pulmonary regurgitation, compared with MRI.

Key words (5 words)

Tetralogy of Fallot; Right ventricular function; Pulmonary regurgitation; Multislice computed tomography; Magnetic resonance imaging.

Abbreviations and acronyms

TOF = tetralogy of Fallot

PR / PRF = pulmonary regurgitation / pulmonary regurgitant fraction

RV = right ventricle / ventricular

LV = left ventricle / ventricular

CCT = cardiac computed tomography

CMR = cardiac magnetic resonance

EDV = end-diastolic volume

ESV = end-systolic volume

SV = stroke volume

EF = ejection fraction

ECG = electrocardiogram

ROI = region of interest

HU = Hounsfield units

CHD = congenital heart disease

Introduction

In patients with surgical repaired tetralogy of Fallot (TOF), residual pulmonary regurgitation (PR) is an important determinant of outcome, as it may contribute to right ventricular (RV) enlargement and dysfunction. Furthermore, these changes may result in exercise intolerance, a propensity for arrhythmias and an increased risk of sudden cardiac death. Previous studies reported that RV volume or ejection fraction can be used as an index of RV dysfunction, and that severe PR was the most common indication for reoperation in patients with repaired TOF [1-4].

Cardiac magnetic resonance (CMR) is the current reference standard to evaluate RV performance and PR, due to its ability to quantify ventricular volumes and pulmonary arterial flow [5-8]. CMR is noninvasive and does not expose the patient to ionizing radiation. Nevertheless, CMR is contraindicated in patients with an implanted pacemaker/defibrillator, patients with claustrophobia or any clinical condition that prohibits a long CMR examination. Therefore, echocardiography and cardiac computed tomography (CCT) are used in these patients [9]. However, Echocardiographic assessment of RV function is also limited due to complex RV geometry [5]. Furthermore,

Doppler echocardiographic assessment of PR is dependent on the observer's experience, and is limited by post-operative structural changes. CCT has recently undergone major technological developments that currently allow quantification of cardiac function with low dose radiation exposure. The unprecedented quality of CCT images, which offer superb endocardial definition optimal for boundary detection, suggests that this modality could potentially constitute a more reproducible reference technique. Moreover, the incremental value of concomitant assessment of cardiac function in addition to coronary artery morphology by CCT has been recognized [10].

A previous study demonstrated good correlations between CCT and CMR measurements of right ventricular functional parameters in patients with ischemic cardiomyopathy [11; 12]. However, in patients with repaired TOF, the accuracy and reproducibility of assessing RV function with CCT has not been examined. Furthermore, the accurate assessment of PR is important in managing patients with repaired TOF, but CCT cannot directly evaluate PRF. In the present study, we attempted to quantify PR using biventricular stroke volume data obtained from CCT. Thus, the purpose of this study was to investigate the accuracy and reproducibility of RV function and PR

measurements with 256-slice CCT in patients with repaired TOF using CMR measurements as the reference standard.

Materials and Methods

Study population

Between June 2010 and November 2013, we prospectively enrolled 33 consecutive patients with surgically repaired TOF. All patients underwent both CCT and CMR for the assessment of cardiac function within 1 week at our hospital. CCT and CMR were performed on the same day in 21 patients and within 1 week in 12 patients. Patients' characteristics are shown in Table 1. General exclusion criteria for contrast CT studies were renal insufficiency, severe arrhythmia (e.g., atrial fibrillation), pregnancy and history of contrast media reaction. Exclusion criteria for CMR were an implanted cardiac pacemaker or defibrillator or other MR-unsafe ferromagnetic objects and claustrophobia. The study was approved by our institutional review board and written informed consent was obtained from each patient prior to imaging.

256-slice CCT

All CCT examinations were performed using a 256-MSCT scanner (Brilliance iCT: Philips Healthcare, Cleveland, OH, USA). All patients underwent retrospective electrocardiogram (ECG)-gated helical scans with ECG tube current modulation. The detector collimation was $2 \times 128 \times 0.625$ mm with a dynamic z-focal spot, resulting in a sample collimation of 256 simultaneous slices of 0.625 mm thickness. The tube voltage was 80 or 100 kV. To improve image quality, reduced-dose scan series were reconstructed using an iterative reconstructive technique (iDose4, Philips Healthcare, Cleveland, OH, USA). Images were reconstructed with an individually adapted field of view encompassing the heart in a matrix of 512×512 pixels, the cardiac standard convolution kernel and a section thickness of 2 mm with an increment of 1 mm. No additional beta-blocker or nitroglycerin was administered to avoid influencing cardiac function. For all patients, an iodinated contrast agent (Iopamidol, 370 mgI/ml; Bayer Healthcare, Osaka, Japan) was administered at a volume based on the patient's weight. The contrast agent was injected over 13 seconds and was followed by the injection of contrast agent diluted 1:1 with saline for 10 seconds as a chaser. The contrast agent and

chaser were injected at a rate of 0.7 ml/kg/sec into the antecubital vein via a 20-gauge catheter using a dual-head injector. Automatic bolus tracking was performed with the region of interest (ROI) placed in the aortic root. All CCT scans were initiated 5 s after the mean ROI contrast reached a pre-determined threshold of 200 Hounsfield units (HU) at a tube voltage of 100 kV or 230 HU at a tube voltage of 80 kV.

Cardiac MRI

All patients underwent 3 Tesla MR imaging (Achieva 3.0T Quasar Dual; Philips Healthcare, Best, The Netherlands) equipped with dual-source parallel radiofrequency transmission, 32-element cardiac phased-array coils used for radiofrequency reception and a 4-lead vectorcardiogram used for cardiac gating. Cine-balanced turbo field-echo sequences in axial view images and short axis view images acquired in parallel to the atrioventricular groove from the base to apex were performed with the following imaging parameters: repetition time 2.8 ms, echo time 1.4 ms, flip angle 45°, slice thickness 8 mm, field of view 380 mm, matrix size 176 × 193, SENSE factor 2, 20 cardiac phases/RR intervals of the ECG. Phase contrast velocity

mapping with a flow-sensitive, gradient-echo sequence was performed in the main pulmonary artery to assess the pulmonary regurgitant fraction (PRF) (Fig. 1). Imaging parameters were as follow: repetition time 6.2 ms, echo time 3.9 ms, flip angle 10°, velocity encoded value (VENC) set to 100–250 cm/s, slice thickness 3mm, field of view 320×300 , matrix size 128×256 , 30 cardiac phases/RR intervals of the ECG.

CCT and MRI data analysis

Image reconstruction of CCT was retrospectively gated to the ECG; 10 phases were reconstructed throughout the cardiac cycle, with the RR interval divided into 10% increments. One experienced radiologist (over 5 years' clinical experience in cardiac radiology) who was blinded to the patient's clinical information evaluated the CCT and CMR data sets and determined RV and LV end-diastolic volume (EDV), end-systolic volume (ESV), stroke volume (SV) and ejection fraction (EF). All volumes were indexed to body surface area. End-diastolic and end-systolic phases were identified visually on those images showing the largest and smallest left and right ventricular cavity areas, respectively. Ventricular volumes were measured on CCT based on axial

images using a workstation (Extended Brilliance Workspace, Philips Healthcare, Cleveland, OH, USA). CCT images were analyzed semiautomatically, followed by manual correction (Fig. 2). CMR images were measured using a workstation (Extended Workspace, Philips Healthcare, Cleveland, OH, USA). CMR images were analyzed semiautomatically, followed by manual correction. RV volumes were measured based on axial images (Fig. 3A, 3B), as previously reported [13]. LV volumes were measured based on short-axis images (Fig. 3C, 3D). Papillary muscles, moderator bands and trabeculations were assigned to the intracavitary lumen of the ventricles.

PRF-CCT (%) was defined as the difference between RV and LV stroke volume divided by RV stroke volume using CCT volumetric data. When PRF-CCT was a negative value, it was assigned a value of zero. Patients with more than trivial regurgitation of the atrioventricular or aortic valves, or significant residual shunt were excluded from the calculation of pulmonary regurgitant fraction by stroke volume difference.

Reproducibility

In 15 randomly selected patients, image analysis was repeated at least 1 month later by the same primary reader and by an additional reader (with over 5 years' clinical experience in cardiac radiology) who was blinded to the results of the initial study to determine the reproducibility of measurements for each imaging modality. The reproducibility of the CCT- and CMR-derived measurements was evaluated by calculating the inter- and intra-observer variability, defined as the absolute value of the difference between each pair of measurements (by the same observer or different observers) divided by the mean of the measurement pair (expressed as a %).

Statistical analysis

Data are presented as means \pm SDs. A paired t test was applied to assess the differences between CCT and CMR parameters. The relationship between CCT and CMR was tested by two-variable linear regression analysis, with Pearson's correlation coefficient. Bland-Altman analysis was used to further determine the agreement between CCT and CMR values by calculating the bias (mean difference) and the 95% limits of agreement ($\pm 1.96 \times$ SD of the mean difference). All statistical analyses were

performed using JMP software (Version 9.0.2), with p-values <0.05 considered statistically significant.

Results

During acquisition of CCT data, all patients had regular sinus rhythm, and the mean heart rate was 70 ± 11 beats per minute (bpm) (range, 54 to 96 bpm). The tube voltage was 80 kV for 14 patients and 100 kV for 19 patients. The estimated effective radiation dose was 7.6 ± 2.6 mSv (range, 3.7–11.4 mSv). The image noise for LV and RV cavities, determined as the SD of the attenuation value in a single round ROI was 27.7 ± 9.3 HU (range, 11.9–41.3 HU) and 27.7 ± 10.2 HU (range, 11.1–46.7 HU), respectively.

Correlations between CCT and CMR measurements

Right ventricular function

The mean RVEDV, RVESV, RVSV and RVEF measured by CCT and CMR are summarized in Table 2. Mean RVEDV and RVESV measured on CCT were

significantly higher compared to CMR (CCT vs. CMR: RVEDV 164.4 ± 59.1 vs. 147.3 ± 53.8 ml, RVESV 93.0 ± 39.7 vs. 80.0 ± 37.3 ml; $p < 0.001$). Mean RVSV and RVEF derived from CCT images were slightly but nevertheless significantly lower compared to CMR (CCT vs. CMR: RVSV 71.5 ± 25.0 vs. 67.3 ± 21.9 ml, RVEF 44.2 ± 7.7 vs. $46.8 \pm 7.5\%$; $P < 0.05$). The Pearson correlation coefficients between CMR and CCT measurements of RV function were good ($r = 0.71$ to 0.96) (Table 2, Fig. 4). The limits of agreement between CCT and CMR were in a good range for all measurements (Fig. 5).

Left ventricular function

The mean LVEDV, LVESV, LVSV and LVEF measured by CCT and CMR are summarized in Table 2. Mean LVEDV and LVSV measured on CCT were significantly higher compared to CMR (CCT vs. CMR: LVEDV 97.4 ± 31.9 vs. 92.9 ± 30.1 ml, LVSV 51.6 ± 15.5 vs. 47.9 ± 14.8 ml; $p < 0.05$). Mean LVESV and LVEF measured on CCT were slightly higher compared to CMR, as reflected by nonsignificant biases of 1.3 ml and 1.3 %, respectively (CCT vs. CMR: LVESV 45.7 ± 21.3 vs. 44.5 ± 18.9 ml, LVEF

53.9±8.7 vs. 52.6±6.8 %; $p>0.05$). The Pearson correlation coefficients between CMR and CCT measurements of LV function were good ($r = 0.73$ to 0.95) (Table 2, Fig. 6). The limits of agreement between CCT and CMR were in a good range for all measurements (Fig. 7).

Pulmonary Regurgitation

In 3 patients, PRF analysis using phase-contrast MRI could not be performed because of susceptibility artifacts from sternal wires or implants. 3 patients had moderate or severe valvular insufficiency of aortic or mitral valve, 1 patient had a partial anomalous pulmonary venous return, and 1 patient had large major aortopulmonary collateral arteries. Total of the 8 patients were excluded from the pulmonary regurgitation analysis.

The PRF measured by both MRI and CT in 25 patients are shown in Table 3. There was an excellent correlation between PRF-CCT and PRF-CMR ($r=0.90$, $p<0.0001$) (Table 3, Fig. 8A). CCT significantly underestimated PRF compared with MRI (mean difference, -9.1 ± 2.0 %) (Fig. 8B). The limits of agreement ($\pm 1.96 \times$ SD of

the mean difference) between the two modalities were 9.7 to −27.9 %.

Intra-observer and inter-observer variability

Inter- and intra-observer variability data for the cardiac function measurements obtained by CCT and CMR are summarized in Table 4. Both the inter- and intra-observer variability values for the functional parameters were lower with CCT than with CMR.

Discussion

Advances in medical treatment, cardiac surgery, intensive care and non-invasive diagnosis over the last 50 years have led to enormous worldwide growth in the number of adults with congenital heart disease (CHD) [14]. The prevalence of CHD in adults and the median age of patients with severe CHD have increased in the general population. In 2000, there were nearly equal numbers of adults and children with severe CHD [15]. Patients with TOF following surgical repair represent a growing population with congenital heart disease, as they now survive into adulthood.

Non-invasive modalities to assess cardiac function in these patients will become more important.

Our results confirmed the linear relationship between CCT and CMR, as reflected by the high correlation coefficients obtained for RV and LV functions (Table 2; Figs. 4, 6).

CCT measurements were found to be highly reproducible, as reflected by lower interobserver and intraobserver variability in all measurements. Meanwhile, CCT significantly overestimated RV volumes compared with the CMR values, resulting in a small but significant negative bias in the calculated RVEF. The patients in our cohort did not receive beta-blockers, because the use of beta-blockers is associated with significant reductions in heart rate, SV and EF, and significant increases in ESV due to negative inotropic effects [16]. Therefore, the overestimations of EDV and SV for both LV and RV in our study might have been caused by transient increases in preload due to rapid inflow of contrast medium. In addition, 10% steps for the reconstruction phase of CCT and interventricular dyssynchrony (RV contraction delay) often seen in repaired TOF patients might have missed the actual end-systolic point using CCT [17]. This might be considered a reason for the overestimation of RVESV and underestimation of

RVEF. Alternative explanations could be the inter-modality differences in the ability to visualize endocardial boundary details and to include or exclude trabeculae that are prominent, especially in RVs from the ventricular cavity. Finally, these discrepancies between CCT and CMR measurements contributed to the lower regurgitant fraction derived by the CCT volumetric method compared with the CMR flow method, although methodological errors in either type of measurement could have also contributed to the difference. RV volume and the severity of PR are often used as criteria for reoperation for pulmonary valve replacement (PVR) [4; 18-20], and the tendency for CCT to overestimate RV volume and underestimate PRF must be recognized. However, this tendency can be corrected by applying a fitting equation on input CCT-variable; 1) $RVEDV-CMR (ml) = 0.87 \times RVEDV-CCT (ml) + 3.6$, 2) $RVESV-CMR (ml) = 0.89 \times RVESV-CCT (ml) - 3.1$, and 3) $PRF-CMR (\%) = 0.91 \times PRF-CCT (\%) + 11.8$. The use of these fitting formulas obtained from well-correlated two data sets allows predicting RV functional parameters and PRF.

PRF-CCT obtained from bi-ventricular stroke volumes was excellently correlated with PRF-CMR obtained from phase-contrast MRI. The measurement of

PRF by phase-contrast MRI has the limitation of susceptibility artifacts due to metallic implants in the RV outflow tract or pulmonary valve in patients with repaired TOF. In the present study, PRF-CMR could not be measured in three patients due to post-operative implants, and CT may circumvent this limitation of MRI. In addition, CCT is more useful than CMR in patients with intellectual impairment or congestive heart failure who cannot withstand long examination time in a supine position and repeated breath-hold in the examinational periods, because of a shorter scan time with CT. The patients with PRF-CMR > 40% were considered to have severe PR. The results of the present study demonstrated that a PRF-CMR of 40% corresponds to a PRF-CCT of 31%. In our study, 12 patients had PVR after CCT and CMR examination, and 2 of the 12 patients had cardiac function re-evaluated by CCT after PVR. Both patients had improved RV function, and PRF-CCT was reduced to nearly 0%, as shown in Table 5.

As previously reported, CMR measurements of RV and LV function are highly reproducible in patients with both normal and dilated right ventricles [21]. In our study, the assessment of the reproducibility of the RV and LV function measurements showed that the CCT measurements had lower intra- and inter-observer variability than

CMR measurements. These findings indicate that CCT measurements of RV and LV function are highly reproducible in patients with repaired TOF. Lower variability values of CCT measurements compared with CMR measurements may be related to better definition of contours obtained from high spatial resolution of CT. In repaired TOF patients who have large RV volumes, the RV cavity becomes more spherical and this makes it easier to define the boundary of the intracavitary lumen; however, the shape of LV cavity may become distorted because of exclusion by the large RV volume. These changes might have helped to reduce the intra- and inter-observer variability in the RV functional parameters in our patient cohort.

The disadvantages of CCT compared with CMR include radiation exposure and the need to use contrast material, which may lead to allergic reactions or kidney damage. Most CCT protocols currently used for the evaluation of ventricular function utilize retrospective ECG-gating, which can result in a substantial radiation exposure of about 14.8–21.1 mSv [22–24]. Recently, retrospective ECG-gated CCT performed with iterative reconstruction and low tube voltage allowed a reduction in the radiation dose while maintaining image quality and the accuracy of functional analysis [25; 26]. Our

CT protocol in combination with iterative reconstruction and low tube voltage could reduce the effective radiation dose (mean, 7.6 mSv) by 50% of the amount typically required for retrospective ECG-gating. Patients with repaired TOF are younger than patients with ischemic cardiomyopathy, and they must receive repetitive cardiac examination; therefore, low dose radiation exposure is necessary.

Furthermore, we used the split-bolus injection with diluted contrast media to obtain an accurate contour of the inner margin of the RV cavity [27; 28]. It provided sufficient attenuation for visualization of the right heart, and the right heart structures could be seen clearly. This probably reduced the time required for ventricular volume measurements by CCT.

Our study has several limitations. First, the size of the study population was rather small. Second, our study did not compare volumetric data and pulmonary regurgitation with measurements obtained by cardiac catheterization. Finally, using 10% steps for the reconstruction phase and acquisition during 10-90% of RR interval on CCT measurements could potentially affect the determination of the actual end-diastolic and end-systolic phases.

In conclusion, ECG-gated 256-slice CCT can assess the RV and LV function, and PRF with high reproducibility in patients with repaired TOF; and the combination of low tube voltage and an iterative reconstruction technique lowers that radiation dose exposure by as much as 50%. RV and LV functional parameters, and PRF obtained from CCT are excellently correlated with those from CMR in patients with repaired TOF. Although the tendency for CCT to overestimate RV volume and underestimate PRF was observed, CCT measurements can be corrected by applying the fitting formulas from excellent correlations between CCT and CMR parameters.

References

- 1 van Straten A, Vliegen HW, Hazekamp MG et al (2004) Right ventricular function after pulmonary valve replacement in patients with tetralogy of Fallot. Radiology 233:824-829
- 2 Oechslin EN, Harrison DA, Harris L et al (1999) Reoperation in adults with repair of tetralogy of fallot: indications and outcomes. J Thorac Cardiovasc Surg 118:245-251
- 3 Ammash NM, Dearani JA, Burkhart HM, Connolly HM (2007) Pulmonary regurgitation after tetralogy of Fallot repair: clinical features, sequelae, and timing of pulmonary valve replacement. Congenit Heart Dis 2:386-403
- 4 Lee C, Kim YM, Lee CH et al (2012) Outcomes of pulmonary valve replacement in 170 patients with chronic pulmonary regurgitation after relief of right ventricular outflow tract obstruction: implications for optimal timing of pulmonary valve replacement. J Am Coll Cardiol 60:1005-1014
- 5 Mercer-Rosa L, Yang W, Kutty S, Rychik J, Fogel M, Goldmuntz E (2012) Quantifying pulmonary regurgitation and right ventricular function in surgically

repaired tetralogy of Fallot: a comparative analysis of echocardiography and magnetic resonance imaging. *Circ Cardiovasc Imaging* 5:637-643

- 6 Michaely HJ, Nael K, Schoenberg SO et al (2006) Analysis of cardiac function--comparison between 1.5 Tesla and 3.0 Tesla cardiac cine magnetic resonance imaging: preliminary experience. *Invest Radiol* 41:133-140
- 7 Lotz J, Doker R, Noeske R et al (2005) In vitro validation of phase-contrast flow measurements at 3 T in comparison to 1.5 T: precision, accuracy, and signal-to-noise ratios. *J Magn Reson Imaging* 21:604-610
- 8 Gutberlet M, Noeske R, Schwinge K, Freyhardt P, Felix R, Niendorf T (2006) Comprehensive cardiac magnetic resonance imaging at 3.0 Tesla: feasibility and implications for clinical applications. *Invest Radiol* 41:154-167
- 9 Takx RA, Moscariello A, Schoepf UJ et al (2012) Quantification of left and right ventricular function and myocardial mass: comparison of low-radiation dose 2nd generation dual-source CT and cardiac MRI. *Eur J Radiol* 81:e598-604
- 10 Seneviratne SK, Truong QA, Bamberg F et al (2010) Incremental diagnostic value of regional left ventricular function over coronary assessment by cardiac

computed tomography for the detection of acute coronary syndrome in patients with acute chest pain: from the ROMICAT trial. *Circ Cardiovasc Imaging* 3:375-383

- 11 Maffei E, Messalli G, Martini C et al (2012) Left and right ventricle assessment with Cardiac CT: validation study vs. Cardiac MR. *Eur Radiol* 22:1041-1049
- 12 Guo YK, Gao HL, Zhang XC, Wang QL, Yang ZG, Ma ES (2010) Accuracy and reproducibility of assessing right ventricular function with 64-section multi-detector row CT: comparison with magnetic resonance imaging. *Int J Cardiol* 139:254-262
- 13 Alfakih K, Plein S, Bloomer T, Jones T, Ridgway J, Sivananthan M (2003) Comparison of right ventricular volume measurements between axial and short axis orientation using steady-state free precession magnetic resonance imaging. *J Magn Reson Imaging* 18:25-32
- 14 Ochiai R, Yao A, Kinugawa K, Nagai R, Shiraishi I, Niwa K (2011) Status and Future Needs of Regional Adult Congenital Heart Disease Centers in Japan. *Circulation Journal* 75:2220-2227

- 15 Marelli AJ, Mackie AS, Ionescu-Ittu R, Rahme E, Pilote L (2007) Congenital heart disease in the general population: changing prevalence and age distribution. *Circulation* 115:163-172
- 16 Jensen CJ, Jochims M, Hunold P et al (2010) Assessment of left ventricular function and mass in dual-source computed tomography coronary angiography: influence of beta-blockers on left ventricular function: comparison to magnetic resonance imaging. *Eur J Radiol* 74:484-491
- 17 Mueller M, Rentzsch A, Hoetzer K et al (2010) Assessment of interventricular and right-intraventricular dyssynchrony in patients with surgically repaired tetralogy of Fallot by two-dimensional speckle tracking. *Eur J Echocardiogr* 11:786-792
- 18 Geva T (2011) Repaired tetralogy of Fallot: the roles of cardiovascular magnetic resonance in evaluating pathophysiology and for pulmonary valve replacement decision support. *J Cardiovasc Magn Reson* 13:9
- 19 Warnes CA, Williams RG, Bashore TM et al (2008) ACC/AHA 2008 Guidelines for the Management of Adults with Congenital Heart Disease: a report of the

- American College of Cardiology/American Heart Association Task Force on Practice Guidelines (writing committee to develop guidelines on the management of adults with congenital heart disease). *Circulation* 118:e714-833
- 20 Baumgartner H, Bonhoeffer P, De Groot NM et al (2010) ESC Guidelines for the management of grown-up congenital heart disease (new version 2010). *Eur Heart J* 31:2915-2957
- 21 Mooij CF, de Wit CJ, Graham DA, Powell AJ, Geva T (2008) Reproducibility of MRI measurements of right ventricular size and function in patients with normal and dilated ventricles. *J Magn Reson Imaging* 28:67-73
- 22 Earls JP, Berman EL, Urban BA et al (2008) Prospectively gated transverse coronary CT angiography versus retrospectively gated helical technique: improved image quality and reduced radiation dose. *Radiology* 246:742-753
- 23 Maruyama T, Takada M, Hasuike T, Yoshikawa A, Namimatsu E, Yoshizumi T (2008) Radiation dose reduction and coronary assessability of prospective electrocardiogram-gated computed tomography coronary angiography: comparison with retrospective electrocardiogram-gated helical scan. *J Am Coll*

Cardiol 52:1450-1455

- 24 Hausleiter J, Meyer T, Hadamitzky M et al (2006) Radiation dose estimates from cardiac multislice computed tomography in daily practice: impact of different scanning protocols on effective dose estimates. *Circulation* 113:1305-1310
- 25 Hou Y, Liu X, Xv S, Guo W, Guo Q (2012) Comparisons of image quality and radiation dose between iterative reconstruction and filtered back projection reconstruction algorithms in 256-MDCT coronary angiography. *AJR Am J Roentgenol* 199:588-594
- 26 Oda S, Utsunomiya D, Funama Y et al (2011) A low tube voltage technique reduces the radiation dose at retrospective ECG-gated cardiac computed tomography for anatomical and functional analyses. *Acad Radiol* 18:991-999
- 27 Kerl JM, Ravenel JG, Nguyen SA et al (2008) Right heart: split-bolus injection of diluted contrast medium for visualization at coronary CT angiography. *Radiology* 247:356-364
- 28 Kondo M, Nagao M, Yonezawa M et al (2014) Improvement of Automated Right Ventricular Segmentation Using Dual-bolus Contrast Media Injection with

256-slice Coronary CT Angiography. Acad Radiol 21:648-653

Figure 1

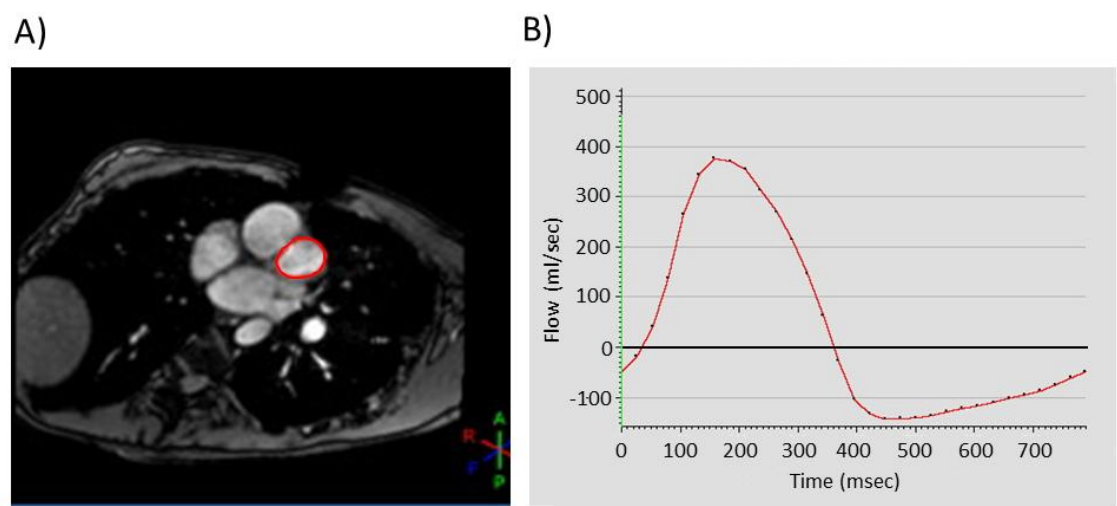


Figure 2

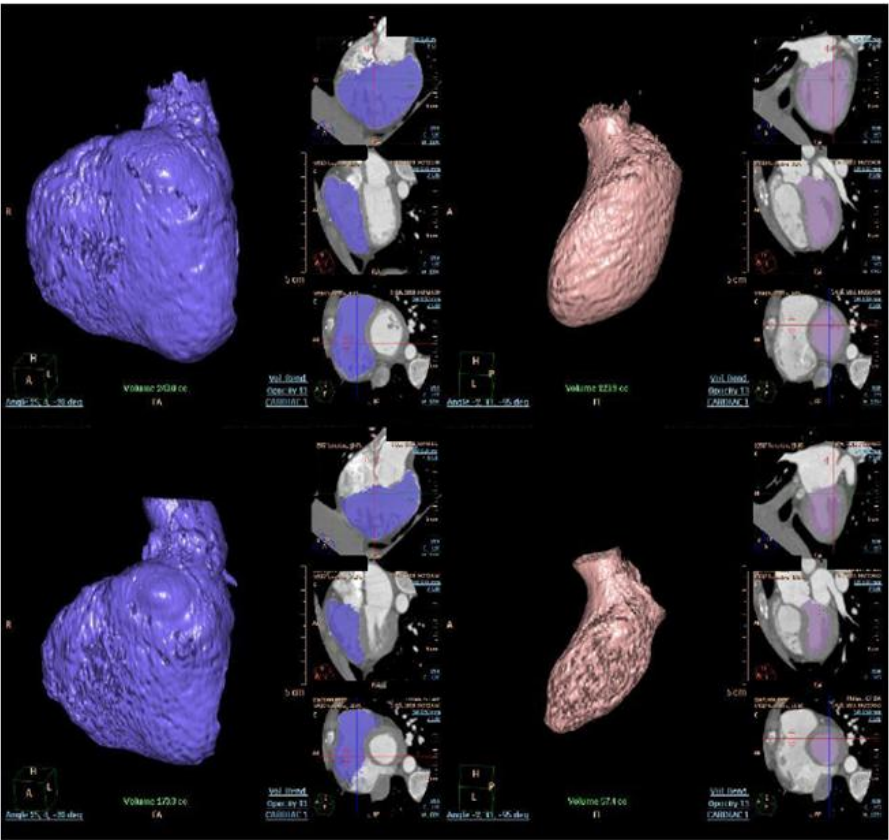


Figure 3

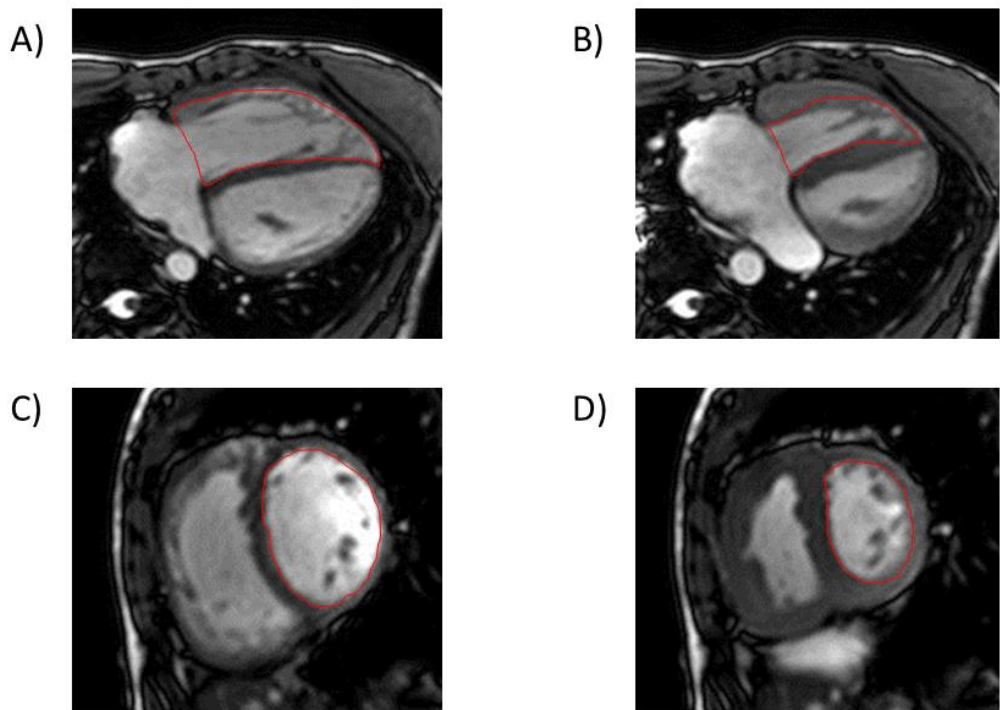


Figure 4

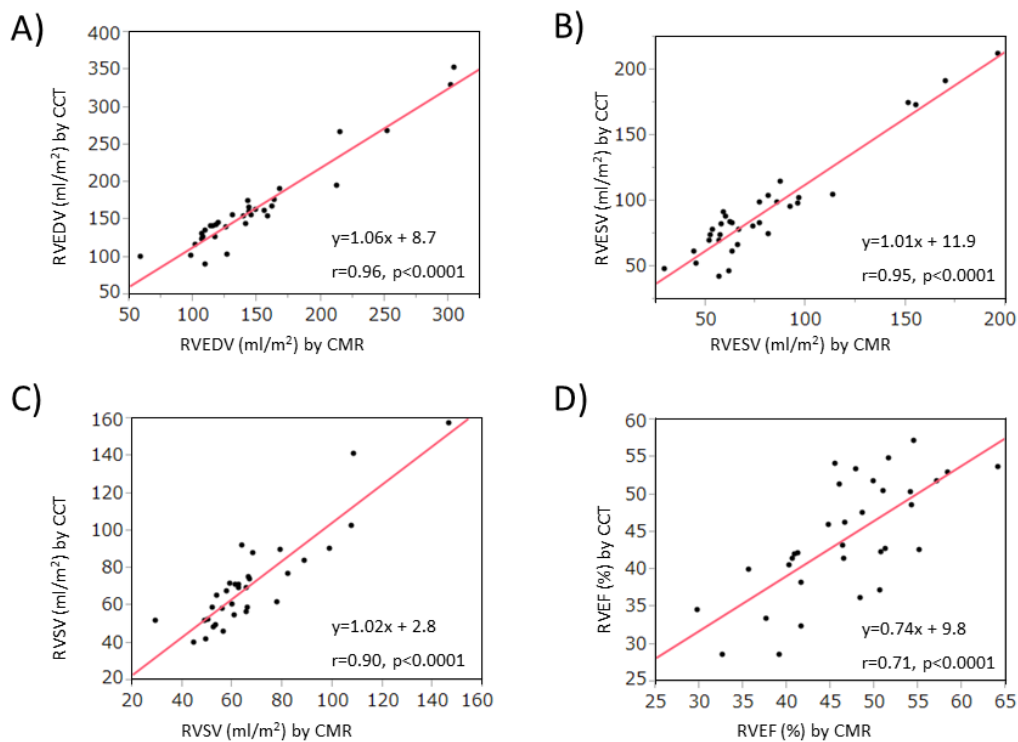


Figure 5

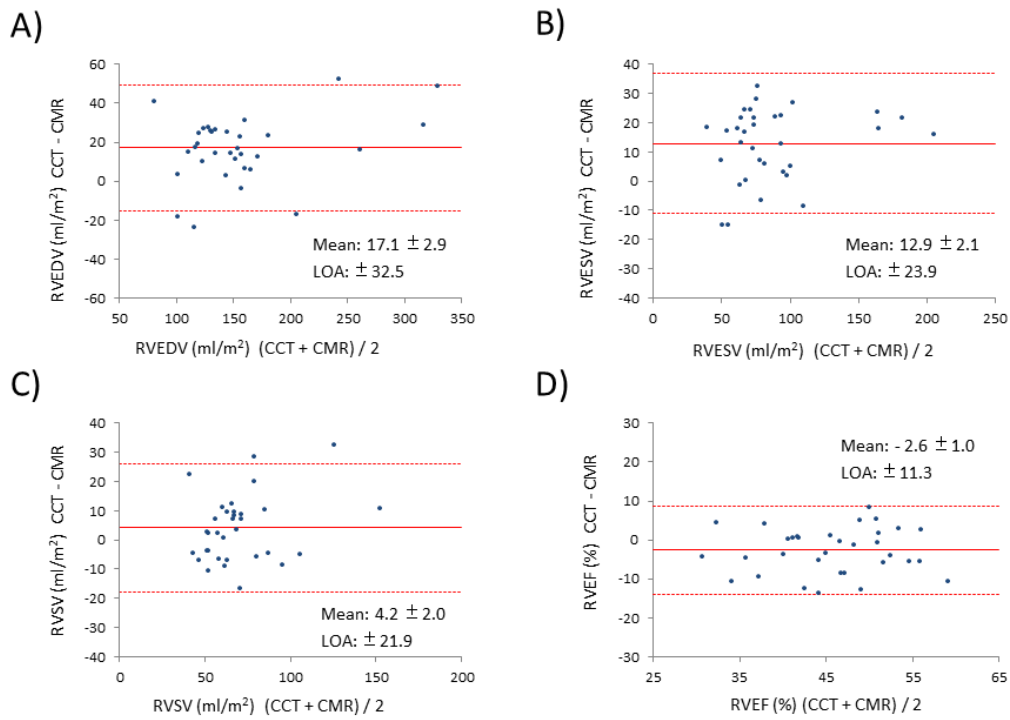


Figure 6

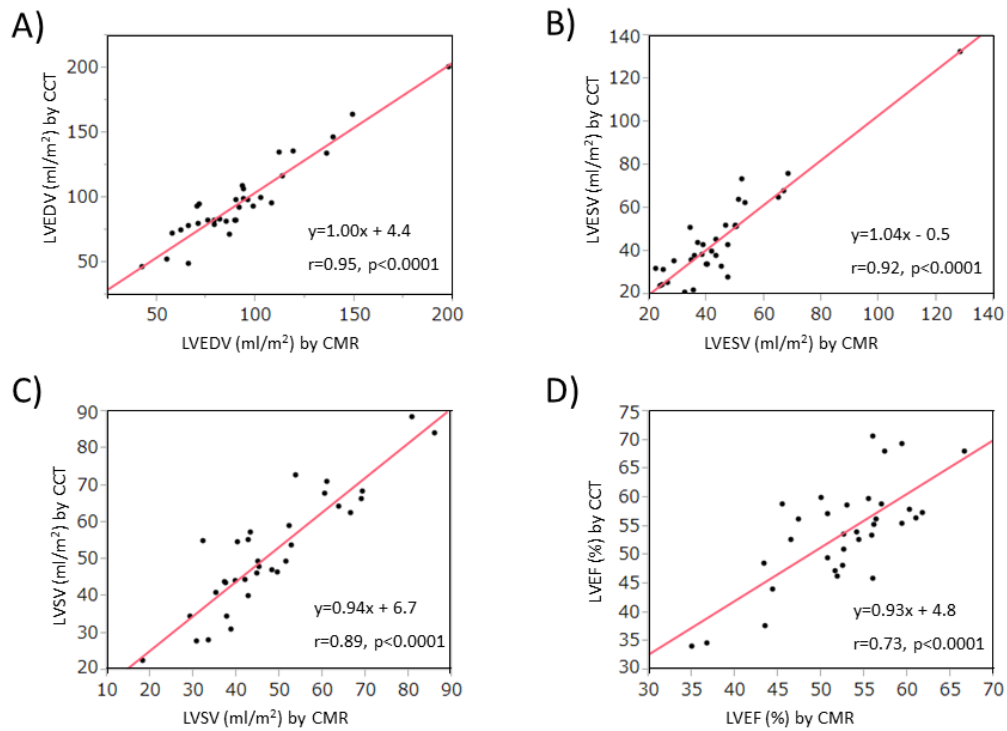


Figure 7

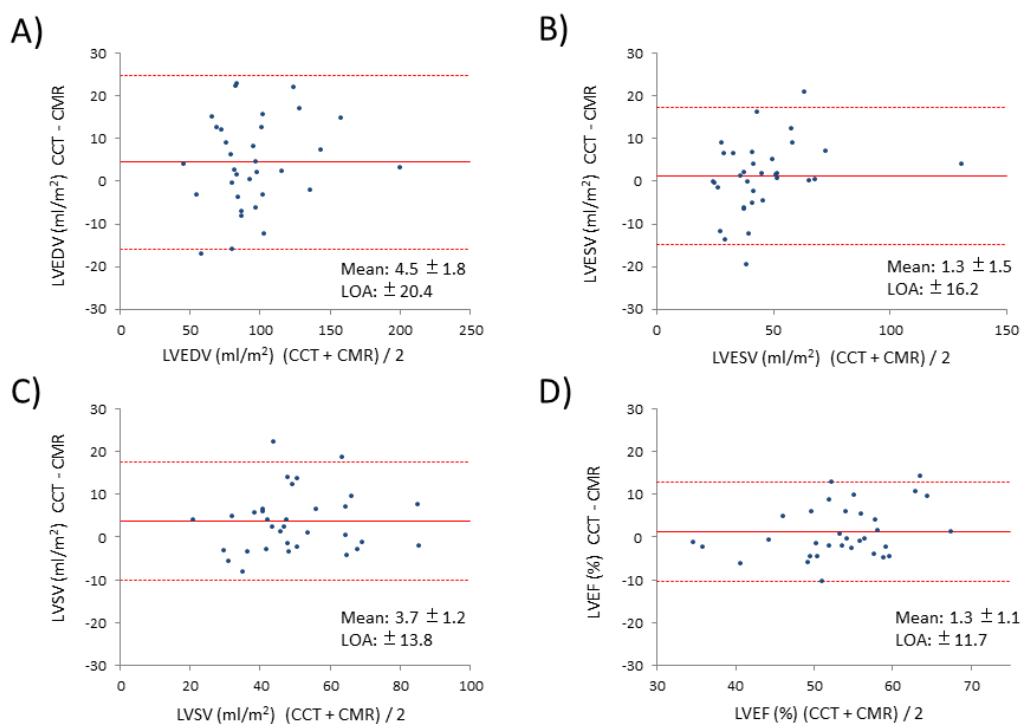


Figure 8

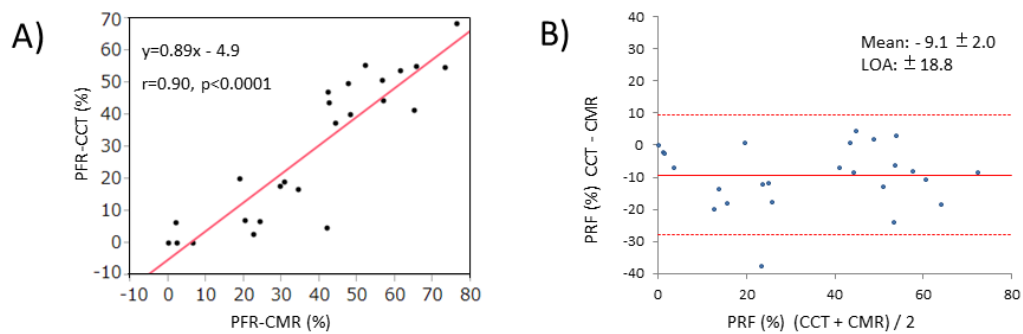


Table 1. Patient Characteristics

Characteristics	value
Total	33
Age	28.9 ± 13.1
Male / Female	19 / 14
BSA(m ²)	1.5 ± 0.3
Age at corrective surgery	5.7 ± 6.8
NYHA Functional Classification I / II / III / IV	25 / 8 / 0 / 0
BNP	56.1 ± 42.7

Data are presented as mean ± standard deviation or number of patients .

BSA, body surface area; NYHA, New York Heart Association; BNP, brain natriuretic peptide.

Table 2. Cardiac function by CCT compared to CMR (n=33)

	CCT	CMR	Paired-t test (p)	Pearson's coefficient (r)
RVEDV (ml/m ²)	164.4±59.1	147.3±53.8	< 0.001	0.96*
RVESV (ml/m ²)	93.0±39.7	80.0±37.3	< 0.001	0.95*
RVSV (ml/m ²)	71.5±25.0	67.3±21.9	0.04	0.90*
RVEF (%)	44.2±7.7	46.8±7.5	0.02	0.71*
LVEDV (ml/m ²)	97.4±31.9	92.9±30.1	0.02	0.95*
LVESV (ml/m ²)	45.7±21.3	44.5±18.9	0.40	0.92*
LVSV (ml/m ²)	51.6±15.5	47.9±14.8	0.01	0.89*
LVEF (%)	53.9±8.7	52.6±6.8	0.23	0.73*

Note-Data are mean ± standard deviation; n=number of patients; *p<0.0001.

CCT = Cardiac computed tomography; CMR = Cardiac magnetic resonance; RV = right ventricle; LV = left ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction.

Table 3. PFR by CCT compared to CMR (n=25)

	PRF-CT	PRF-MRI	Paired-t test (p)	Pearson's coefficient (r)
PRF (%)	29.4±22.2	38.7±22.5	< 0.001	0.90*

Note-Data are mean ± standard deviation; n=number of patients; *p<0.0001. PRF = pulmonary regurgitant fraction

Table 4. Inter- and intra- observer variability of RV and LV function obtained from repeated measurements by CCT and CMR in a subset of 15 randomly selected patients.

	Interobserver variability (%)		Intraobserver variability (%)	
	CCT	CMR	CCT	CMR
RVEDV	3.5 ± 2.8	5.3 ± 6.2	3.4 ± 3.1	6.2 ± 4.9
RVESV	5.1 ± 5.0	11.0 ± 11.0	3.7 ± 2.5	10.6 ± 10.9
RVSV	7.5 ± 5.6	8.2 ± 7.3	5.5 ± 5.2	8.5 ± 10.5
RVEF	5.6 ± 5.3	8.5 ± 8.3	3.9 ± 2.6	9.7 ± 9.7
LVEDV	5.4 ± 4.1	6.8 ± 6.0	4.9 ± 3.4	6.8 ± 6.3
LVESV	7.3 ± 6.8	9.6 ± 7.2	7.6 ± 5.4	9.5 ± 6.7
LVSV	7.5 ± 6.6	10.2 ± 8.6	5.5 ± 4.4	9.7 ± 9.9
LVEF	4.5 ± 3.9	5.2 ± 5.8	5.1 ± 2.7	6.8 ± 4.3

Note-Data are mean ± standard deviation.

CCT = Cardiac computed tomography; CMR = Cardiac magnetic resonance; RV = right ventricle; LV = left ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction.

**Table 5. Comparison RV function between pre- and post- PVR
by CCT**

	Patient 1		Patient 2	
	Pre-operation	Post-operation	Pre-operation	Post-operation
RVEDV (ml/m ²)	269.0	154.4	174.9	125.1
RVESV (ml/m ²)	192.1	104.0	99.2	76.2
RVSV (ml/m ²)	76.9	50.5	75.7	48.9
RVEF (%)	28.6	32.7	43.3	39.1
PRF-CT (%)	55.0	0	47.0	3.6

RV = right ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction;

PRF = pulmonary regurgitant fraction

Figure Legends

Figure 1.

A) Phase-contrast imaging in a patient with repaired TOF and PR. Cine image from a phase-contrast sequence shows the pulmonary artery (rounded red line) in cross section.

B) Graph illustrates flow (ml/sec) versus time (msec) during phase-contrast imaging.

PR is the ratio of retrograde flow volume (regurgitation) to antegrade flow volume (ml).

Figure 2.

Post-processing screen illustrates the single step used to measure RV end-diastolic volume (upper-left), RV end-systolic volume (lower-left), LV end-diastolic volume (upper-right) and LV end-systolic volume (lower-right) from a CCT data set.

Figure 3.

A) Cine axial image at the end-diastolic phase to measure RVEDV

B) Cine axial image at the end-systolic phase to measure RVESV

C) Cine short-axis image at the end-diastolic phase to measure LVEDV

D) Cine short-axis image at the end-systolic phase to measure LVESV

Figure 4.

Scattergram shows the results of linear regression analysis between CCT and CMR measurement of RVEDV (A), RVESV (B), RVSV (C) and RVEF (D) for all patients. The regression equation and Pearson's correlation coefficient are provided for each plot.

Figure 5.

Bland-Altman plots show degree of agreement between CCT and CMR of RVEDV (A), RVESV (B), RVSV (C) and RVEF (D) for all patients. The solid line represents mean difference. The broken line represents the 95% limits of agreement (LOA) ($\pm 1.96 \times \text{SD}$). The Bland-Altman analysis show overestimation of RVEDV and RVESV and close agreement of the two methods for RVSV and RVEF.

Figure 6.

Scattergram shows results of linear regression analysis between CCT and CMR

measurement of LVEDV (A), LVESV (B), LVSV (C) and LVEF (D) for all patients.

The regression equation and Pearson's correlation coefficient are provided for each plot.

Figure 7.

Bland-Altman plots show the degree of agreement between CCT and CMR

measurement of RVEDV (A), RVESV (B), RVSV (C) and RVEF (D) for all patients.

The solid line represents mean difference. The broken line represents the 95% limits of

agreement (LOA) ($\pm 1.96 \times \text{SD}$). The Bland-Altman analysis shows close agreement

with little bias between the two methods for the measurement of LV function.

Figure 8.

A) Scattergram shows the results of linear regression analysis between PRF-CCT and

PRF-CCT.

B) Bland-Altman plots show the degree of agreement between PRF-CCT and

PRF-CMR. The solid line represents the mean difference. The broken line represents the

95% limits of agreement (LOA) ($\pm 1.96 \times \text{SD}$). The Bland-Altman analysis shows that

CCT underestimated PRF compared with CMR.

Table Legends

Table 1. Patient Characteristics

Data are presented as mean \pm standard deviation or number of patients.

BSA, body surface area; NYHA, New York Heart Association; BNP, brain natriuretic peptide.

Table 2. Cardiac function by CCT compared to CMR (n=33)

Note-Data are mean \pm standard deviation; n=number of patients; *p<0.0001.

CCT = Cardiac computed tomography; CMR = Cardiac magnetic resonance; RV = right ventricle; LV = left ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction.

Table 3. PFR by CCT compared to CMR (n=25)

Note-Data are mean \pm standard deviation; n=number of patients; *p<0.0001. PRF = pulmonary regurgitant fraction

Table 4. Inter- and intra- observer variability of RV and LV function obtained from repeated measurements by CCT and CMR in a subset of 15 randomly selected patients.

Note-Data are mean \pm standard deviation.

CCT = Cardiac computed tomography; CMR = Cardiac magnetic resonance; RV = right ventricle; LV = left ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction.

Table 5. Comparison RV function between pre- and post- PVR by CCT

RV = right ventricle; EDV = end-diastolic volume; ESV = end-systolic volume; SV = stroke volume; EF = ejection fraction; PRF = pulmonary regurgitant fraction