Quantitative Assessment of Left Ventricular Function and Myocardial Mass: A Comparison of Coronary CT Angiography with Cardiac MRI and Echocardiography

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Summary

Background: The purpose of this study was to compare the left ventricular parameters obtained from multidetector row computed tomography (MDCT) studies with two-dimensional echocardiography (2DE), and magnetic resonance imaging (MRI), which is accepted as the gold standard in the evaluation of left ventricular functions. The study also aimed to evaluate whether or not there is a relationship between the MR-Argus and CMR tools software programs which are used in post-process calculations of data obtained by MRI.

Material/Methods: Forty patients with an average age of 51.4±14.9 years who had been scanned with cardiac MDCT were evaluated with cardiac MRI and 2DE. End-diastolic volume (EDV), end-systolic volume (ESV), stroke volume (SV), ejection fraction (EF), cardiac output (CO), and myocardial mass values calculated by MDCT, MRI, and 2DE were compared with each other. Two different MR software programs were used to compare left ventricular functions. The CMR tools LV tutorials method is accepted as the gold standard because it can be used in three-dimensional functional evaluation. The Pearson Correlation and Bland-Altman analysis were performed to compare the results from the two MR methods (MR-Argus and CMR tools) and the results from both the MDCT and the 2DE with the CMR tools results.

Results: Strong positive correlations for EF values were found between the MDCT and CMR tools (r=0.702 p<0.001), and between the MR-Argus and CMR tools (r=0.746 p<0.001). The correlation between the 2DE and CMR tools (r=0.449 p<0.004), however, was only moderate. Similar results were obtained for the other parameters. The strongest correlation for ESV, EDV, and EF was between the two MR software programs. The correlation coefficient between the MDCT and CMR tools is close to the correlation coefficient between the two software programs. While the correlation between 2DE and CMR tools was satisfactory for ESV, EDV, and CO values, it was at a moderate level for the other parameters.

Conclusions: Left ventricular functional analysis can be performed easily and reliably with cardiac MDCT used for coronary artery evaluation and it also gives more accurate results than 2DE.

MeSH Keywords: Echocardiography • Magnetic Resonance Imaging • Multidetector Computed Tomography • Ventricular Function, Left

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Background

Cardiovascular disease, including coronary artery disease (CAD) and cerebrovascular disease, is the primary cause of death globally, even when including low-income and middle-income countries [1–3]. Correct assessment of the left ventricular (LV) function and mass is important for diagnostic, therapeutic, and prognostic evaluation and risk assessment of patients with CAD [4]. CAD and left ventricular (LV) hypertrophy have been indicated as independent risk factors for premature death [5–8].

There are several non-invasive diagnostic methods for the evaluation of left ventricular function. Although 2DE is a widely used, inexpensive, and noninvasive method, it is operator-dependent and the acoustic window is restricted in some patients [9]. In the last decade, alternative imaging modalities have significantly improved in the evaluation of LV mass and function through technological developments [5]. Cardiac magnetic resonance imaging (CMRI) is a noninvasive imaging modality which does not expose the patient to ionizing radiation. Because of its high temporal and spatial resolution and consistency in the assessment of LV function and myocardial mass, it is considered the reference standard for these applications [5,10,11]. However, CMRI can be difficult to obtain compared to alternative imaging modalities, such as two-dimensional echocardiography (2DE) and multi-detector computed tomography (MDCT), in acutely ill or uncooperative patients. Moreover, contraindications to CMRI, such as pacemakers, claustrophobia, or clinical conditions that disallow long MRI examinations, can pose challenges in some patients [12,13]. Coronary CT angiography (CCTA) with MDCT is a new, noninvasive coronary artery imaging technique which is widespread and whose use has been increasing in recent years [8,14,15]. The evaluation of LV function, including end-diastolic volume (EDV), end-systolic volume (ESV), stroke volume (SV), ejection fraction (EF), and cardiac output, as well as myocardial mass with MDCT utilizes retrospective ECG-gating, which can be examined using only single-step breath holding [16,17].

The aim of this study was to compare the accuracy of measurements of LV function and myocardial mass using 64-Slice Coronary CT, 2DE, and CMRI with two different software programs, while using the CMR tools LV tutorials software, with three-dimensional results, as the reference standard.

Material and Methods

Study population

Forty patients (25 male and 15 female) with a mean age of 51.4±14.9 years and an age range of 16–80 years, with known or suspected CAD, were included in this study. The age, weight, and height of the patients were recorded. All patients underwent CCTA, CMRI (Figure 1), and 2DE on the same day. The exclusion criteria for the CCTA studies were a history of contrast medium allergy, renal failure (defined as serum creatinine >1.5 mg/dL), severe arrhythmia, pregnancy, acute hypotension (<100 mm Hg systolic), and clinically unstable disease. The exclusion criteria for CMRI were cardiac pacemaker, ferromagnetic objects or implants unsafe for MRI, and claustrophobia. The study was approved by the institutional review board, and written informed consent was obtained from each patient prior to the examinations.

CT image acquisition protocol

All CCTA studies were performed on a 64-Slice MDCT scanner (Somatom Sensation, Siemens Medical Systems, Erlangen, Germany). The MDCT acquisition parameters were as follows: detector configuration 64×0.6 mm; gantry rotation time 330 ms; and matrix 512×512, effective mAs: 870–900, kVp: 120. The pitch was set automatically by MDCT. Images were reconstructed with 0.6 mm slice thickness and 0.6 mm increments using a medium smooth tissue iterative reconstruction kernel (I26f).

Three days before the procedure, the patients’ heart rates were measured. In the case of a heart rate greater than 75 beats/min, oral beta-blocker therapy (Beloc 50–100 mg) was routinely administered. Before the procedure, heart rates and rhythms were checked again using ECG. Patients with heart rates above 75 beats/min were treated with up to three ascending doses of 5 mg/5 mL of a beta-blocker (5, 10, and 15 mg) as needed. The scanning area was determined to be between the level of the carina and the diaphragmatic surface of the heart. A total of 90–100 mL of non-ionic contrast matter, containing 400 mg of iodine (Iomeron®, Bracco, Italy), was given at a rate of 4.5–6 mL/s through an 18 G intravenous antecubital catheter using an automatic injector system (Ulrich, Medizintechnik, Germany). After the contrast was administered, flushing was carried out with 40 mL of NaCl at a rate of 6 mL/s using the automatic

Figure 1. Left ventricular analysis with (A) MR Argus, (B) CMR tools, and (C) MDCT.
injector. After performing a scenogram, one slice was taken at the level of the aortic root and bolus tracking (CARE Bolus, Siemens Medical Systems, Erlangen, Germany) was placed at that level. The scan trigger was adjusted to 100 HU (Hounsfield Unit). All patients were scanned in a craniocaudal direction. No complications occurred during the CT examinations.

CT data analysis

The CT images with retrospectively ECG-triggered acquisition windows were applied across an adjustable portion of the cardiac cycle (0–100% in this investigation). For analysis, a series of images in the axial plane was reconstructed from the raw data in 10% steps throughout 0–100% of the cardiac cycle interval with a slice thickness of 0.75 mm and an interval of 1.0 mm. Following that, all of the data were evaluated with Circulation software on a Leonardo workstation (Siemens Medical Solutions, Erlangen, Germany). Three-dimensional reformatted images and cardiac planes were obtained with the axial plane images. The end-systolic and end-diastolic phases, interventricular septum, and level of the mitral valve were detected at all of the three basic cardiac planes (horizontal long axis, vertical long axis, and short axis) with Left Ventricular Analysis (LVA) software. Then, the diastolic and systolic levels of the mitral and aortic valves and the apex of the heart were detected visually and marked manually. The endocardial and epicardial borders were traced semiautomatically in both series of images. The contours were checked visually for accuracy and, if necessary, manually adjusted using a software tool ("nudge tool"). The papillary muscles were included in the LV cavity. The images obtained were also reviewed visually and corrected manually if necessary. The ESV, EDV, SV, EF, and LV myocardial masses were determined. The elapsed time from inputting the data to obtaining the results was calculated for each patient. MDCT images were evaluated by one radiologist (Y.P.) with 10 years of experience in CCTA.

MR image acquisition protocol

All CMRI studies were performed on a 1.5-T MRI system (Magnetom Symphony, Siemens, Erlangen, Germany). Patients were examined with a combination of spine and body coils in the supine position. Following scout images in the axial, coronal, and sagittal planes, functional analysis was performed with cardiac-gated, multi-segmented, cine steady-state free precession (SSFP) sequence. The CMRI acquisition parameters were as follows: 10-mm slice-thickness with 2-mm slice gap, TR: 60.2 ms, TE: 1.89 ms, flip angle: 80°, field of view: 320×320 mm, matrix: 192×256.

To improve the quality of the images, patients were asked to hold their breath at the end of expiration. First, lower resolution scout images in the axial, coronal, and sagittal orientations were obtained. Then, long and short axis pseudoventricular images were created from the scout images. Horizontal long axis (HLA/four-chamber) images were planned according to the long and short axis pseudoventricular images. Vertical long axis (VLA/two-chamber) images were planned according to the horizontal long and short axis images. Short axis images were obtained from the mitral valve to the apex of the heart, and were planned parallel to the mitral valve. A total of 7–13 slice images were obtained for all patients including all of the left ventricle. The average examination time was 20–25 minutes. The obtained images were transferred to the workstation (Leonardo, Siemens Medical Solutions, Erlangen, Germany). For the functional analysis, two different commercially available software programs, Argus (Siemens Healthcare, Erlangen, Germany) and CMR tools LV tutorials (Cardiovascular Imaging Solutions, London, UK) were used.

CMRI data analysis

The endocardial and epicardial borders were traced manually using both software systems and functional analysis was performed.

Analysis using Argus (Siemens Healthcare, Erlangen, Germany) Software

For each study, the end-diastolic and end-systolic phases were determined. For the detection of each phase, the largest and narrowest diameters of the ventricular cavity at the middle of the ventricle were used. The endocardial and epicardial borders were traced manually in short axis images in both phases. The borders of the endocardium were traced by using the intensity difference between the chamber when filled with blood and the moderate intensity of the myocardium. The papillary muscles were included in the LV volumetric analysis. While the epicardial border was being detected, the interventricular septum was included in the LV volume. The most basal slice that was surrounded by at least 50% of the myocardium with filled blood was defined as the basal segment of the left ventricle. This was included in the LV chamber volume. The apex was defined as the last slice with a visible lumen throughout the entire cardiac cycle. The ESV, EDV, SV, EF, and LV myocardial masses were determined according to the Simpson’s rule. The elapsed time from inputting of the data to obtaining the results was calculated for each patient.

Analysis using CMR tools LV tutorials (Cardiovascular Imaging Solutions, London, UK) Software

The largest and narrowest volumes for each end-diastolic and end-systolic phases were determined in the same way as the MR Argus technique. Atrioventricular valve levels were detected at the end of the systolic and diastolic phases. All of the endocardial and epicardial borders were checked and corrected by checking the levels of the valves for all phases. Three-dimensional modeling was used and three-dimensional and movie images were obtained. The ESV, EDV, SV, EF, and LV myocardial masses were determined. The papillary muscles were included in the LV volumetric analysis. The elapsed time from inputting of the data to obtaining the results was calculated for each patient.

To eliminate all operator-related differences, all image analysis procedures were performed by the same experienced radiologist (Y.P.) using both the Argus and CMR tools programs.
Two-dimensional echocardiography (2DE) imaging acquisition protocol

All 2DE examinations were performed using an ultrasonography machine (Philips Ultrasound M2540A, Philips Medical Systems, Bothell, WA, USA) with a linear transducer of 2.5 MHz by the same cardiologist (E.E.G.) with 5 years of experience. Patients were evaluated with two-dimensional and M-mode Doppler echocardiographic examinations in the left lateral decubitus position. Conventional views were as follows: parasternal long and short axis, apical four- and two-chamber. The 2DE measurements were performed based on the recommended criteria of the American Society of Echocardiography. The M-mode was used for the assessment of LV dimensions (in systole and diastole), interventricular septum thickness at the level of the mitral valve, and posterior wall thickness with parasternal long axis images. Following that, the EF and LV myocardial masses were calculated using those data. Considering the 2DE data sets, contours were manually traced around the endocardial borders using apical four-chamber and two-chamber images. The papillary muscles were included in the myocardial mass. The ESV and EDV for the four-chamber and two-chamber images and the SV, CO, and EF were calculated according to the Simpson’s method.

Statistical analysis

For statistical analysis, the mean values of ESV, EDV, SV, and EF were used. LV functional values were obtained from three modalities with four different calculation systems (MDCT, 2DE, and MRI with the use of the two different software systems). The data of ESV, EDV, EF, SV, CO, and myocardial mass were obtained for evaluation. Minimum, maximum, and mean values with standard deviation were obtained from cardiac MDCT, CMR tools, MR Argus, and 2DE for determining the LV functional parameters. Continuous variables are presented as mean±standard deviation (SD).

For a linear correlation analysis, the Pearson correlation coefficient (r) and the Intraclass Correlation Coefficient (ICC) were determined. To assess the degree of agreement between each parameter, the Bland-Altman analysis (including 95% confidence intervals) was performed. All statistical analyses were performed using SPSS 17.0 software (SPSS; Chicago, Illinois, USA), with a p-value of <0.05 considered statistically significant.

Results

Patients' characteristics

Data were collected for all of the 40 patients (25 males and 15 females) enrolled in the study. The mean age of the patients was 51.4±14.9 years with a range of 16-80 years. The mean height and weight of the patients was 167.9±10.68 cm and 78.2±15.12 kg, respectively. The demographic characteristics of the patients are shown in Table 1. The preliminary diagnoses of patients were as follows: coronary artery anomaly in two patients, evaluation of the bypass graft in one patient, coronary stent patency control in one patient, aortic aneurysm in one patient, atypical chest pain in five patients, and CAD in the patients. All the patients were evaluated with CCTA. The definitive diagnoses of the patients were as follows: myocardial bridge in eight patients, coronary artery anomaly in five patients, aortic aneurysm in one patient, atheromatous plaques in a significantly stenotic coronary vessel in seven patients,
atheromatous plaques in a not significantly stenotic coronary vessel in seven patients, dysfunction in the left ventricle in seven patients, and normal coronary arteries in eleven patients.

**Ejection fraction**

The Pearson correlation coefficients between the MDCT and CMR tools measurements of LV EF were strong ($r=0.702$, $p<0.001$) and the ICC was $R_1=0.773$ (Figure 2). The Pearson correlation coefficients between the 2DE and CMR tools measurements of EF were moderate ($r=0.449$, $p<0.004$) and the ICC was $R_1=0.611$. The Pearson correlation coefficients between the MR Argus and CMR tools measurements of LV EF were excellent ($r=0.746$, $p<0.001$). The best correlation and reliability was between the MR Argus and CMR tools results (Figure 3) with an ICC of $R_1=0.854$. The EF values for MDCT, 2DE, MR Argus, and CMR tools are listed in Table 2.

**End-systolic volume and end-diastolic volume (ESV, EDV)**

The Pearson correlation coefficients between the MDCT and CMR tools measurements of LV ESV were very strong ($r=0.881$, $p<0.001$) and the ICC was $R_1=0.935$. The Pearson correlation coefficients between 2DE and CMR tools measurements of LV ESV were also very strong ($r=0.792$, $p<0.001$) and the ICC was $R_1=0.854$. The Pearson correlation coefficients between the MDCT and CMR tools measurements of LV EDV were excellent ($r=0.915$, $p<0.001$) and the ICC was $R_1=0.948$. The Pearson correlation coefficients between the 2DE and CMR tools measurements of LV EDV were also very strong ($r=0.783$, $p<0.001$) and the ICC was $R_1=0.849$. The Pearson correlation coefficients between the MR Argus and CMR tools measurements of LV ESV and EDV were excellent ($r=0.961$, $p<0.001$; $r=0.961$, $p<0.001$, respectively). The ICC were $R_1=0.927$ and $R_1=0.927$, respectively. Although the papillary muscles were included in the blood pool in the MR Argus analysis and the myocardial mass was included in the CMR tools analysis, the best correlation and reliability was found between the MR Argus and CMR tools.

**Stroke volume (SV) and cardiac output (CO)**

The Pearson correlation coefficients between the MDCT and CMR tools measurements of SV were good to excellent ($r=0.796$, $p<0.001$) and the ICC was $R_1=0.869$. The Pearson correlation coefficients between the 2DE and CMR tools measurements of SV were moderate ($r=0.435$, $p<0.005$) and the ICC was $R_1=0.559$. The Pearson correlation coefficients between the MDCT and CMR tools measurements of CO were good to excellent ($r=0.968$, $p<0.001$) and the ICC was $R_1=0.937$. The Pearson correlation coefficients between the MR Argus and CMR tools measurements of SV and CO were good ($r=0.608$, $p<0.001$; $r=0.619$, $p<0.001$, respectively). The ICCs were $R_1=0.727$ and $R_1=0.734$, respectively.

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**Table 2. Values of LV EF in MDCT, 2DE, MR Argus, and CMR tools.**

|          | N   | Minimum (%) | Maximum (%) | Mean ±SD (%) |
|----------|-----|-------------|-------------|--------------|
| MDCT     | 40  | 25.0        | 81.0        | 61.52±12.18  |
| 2DE      | 40  | 28.0        | 76.1        | 60.29±8.97   |
| MR Argus | 40  | 31.5        | 79.4        | 61.39±11.16  |
| CMR tools| 40  | 29.0        | 81.0        | 63.57±11.08  |

MDCT – multi-detector row computed tomography; 2DE – two-dimensional echocardiography.

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**Figure 3.** The Pearson correlations with Bland-Altman analysis between the MR Argus and CMR tools measurements of LV EF were excellent ($r=0.746$, $p<0.001$). The ICC was $R_1=0.854$. LV, left ventricle; EF, Ejection fraction; $r$, Pearson correlation coefficient; and ICC – intraclass correlation coefficient.

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The Pearson correlation coefficients between the MDCT and CMR tools measurements of myocardial mass were excellent \((r=0.884, p<0.001)\). The ICC was \(R_1=0.938\). LV, left ventricle; \(r\), Pearson correlation coefficient; and ICC – intraclass correlation coefficient.

The Pearson correlation coefficients between the 2DE and CMR tools measurements of myocardial mass were moderate \((r=0.414, p<0.008)\) and the ICC was \(R_1=0.559\). The Pearson correlation coefficients between the MR Argus and CMR tools measurements of myocardial mass were excellent \((r=0.885, p<0.001)\) and the ICC was \(R_1=0.932\) (Figure 5). The myocardial mass values for MDCT, 2DE, MR Argus, and CMR tools are listed in Table 3.

In the Bland-Altman analysis, systemic error and correlation of all functional parameters were determined for each modality with CMR tools and they were demonstrated graphically. The difference of each method from the CMR tools results and their means were calculated. The \(X\pm 2SS\) interval of the differences were accepted as the correlation limits. The methods in which the differences were close to zero, in which the correlation limits were narrowest,
and with homogenous distribution were accepted as the methods with high correlation and with the least errors. According to these parameters, the highest correlation and the least error was between the two MR analysis methods. The correlation between the MDCT and CMR tools and the limits for error were at least as good as the correlation between the MR Argus and CMR tools values. The difference between the 2DE and CMR tools was far from zero and the correlation limits were wider. The correlation between the 2DE and CMR tools was at a lower level when compared to the correlation between the MDCT and CMR tools.

**Left ventricular function computation time with MDCT and MRI**

The mean postprocessing time was 206±45 s, 250±55 s and 492±72 s for MDCT, MR Argus and CMR tools, respectively. According to these results, the postprocessing time was much longer for the CMR tools software than for the MDCT and MR Argus software.

**Discussion**

In this study, the data related to left ventricle function, which were obtained with all three modalities and two different MR software programs, were compared. The results of this study indicate three important findings. First, in the correlation and reliability analysis, the correlation between MR Argus and CMR tools’ data in EF, ESV, and myocardial mass values was excellent. The calculation of the same data with similar methods in both techniques could have an effect on this finding. Second, the correlation of MDCT with CMR tools was at least as good as the MR Argus method. For SV, CO, and EDV data, MDCT-CMR tools’ correlation was better when compared to the MR Argus-CMR tool correlation. The common characteristic of MDCT and CMR tools’ analyses is the use of three-dimensional modelling. We also believe that the reason for this good correlation is three-dimensional modelling. The degree of end-diastolic imaging quality in MDCT was at the highest level, and depending on this, the endocardial borders were clearer so the MDCT-CMR tools’ correlation in EDV values was better. Similarly for ESV data, the end-systolic imaging quality decreased and errors developed in the determination of endocardial borders. Depending on this, ESV MDCT-CMR tools’ correlation became lower according to MR Argus-CMR. Finally, 2DE-CMR tools’ correlations were worse than MDCT-CMR tools and MR Argus-CMR tools’ correlations. In particular, an intermediate degree of correlation was found in myocardial mass, SV, and CO values. The 2DE-CMR tools’ correlation was poorer and the reason for this was thought to be related to the sensitivity of 2DE and its dependency on the patient and user factors. Furthermore, it was thought that using the measurements that were obtained with M-mode 2DE in clinical practice might have had an effect on this.

In the study of Palazzuoli et al. [18], left ventricle function and volume assessment were performed with an MDCT segmental reconstruction algorithm in 93 patients and the results were compared with the 2DE results that were obtained within the last two months. The 2DE and MDCT results were evaluated with a double-blind study. As a result, the data obtained from MDCT, which is used to assess left ventricle function and size, were better in terms of repeatability and validity when compared to 2DE. In this study, although a good correlation was detected in ESV and EDV values between 2DE and MDCT data, an intermediate correlation was detected in EF, CO, SV, and myocardial mass values; 2DE-MR correlations were similar. In the study of Yamamuro et al. [19], MDCT and MR were conducted on 50 patients. Out of those patients, 2DE was conducted on 41 patients and SPECT was conducted on 27 patients. The EDV, ESV, EF, and left ventricle mass were calculated and the obtained values were compared to the MR results, which were accepted as standard. As a result, it was observed that the left ventricle parameters calculated with MDCT were well-correlated with MR results. Furthermore, it was found that the functional analysis that was done with MDCT was more accurate than 2D-2DE or SPECT. In the study of Juergens et al. [20], the results of left ventricle volumetric and functional analyses, which were obtained by using MDCT semi-automatic analysis software, were compared with MR findings in 30 patients with coronary artery disease. It was found that the results of left ventricle volumetric and functional analyses, which were carried out with MDCT, had a good correlation with MR findings.

In the current study, the best correlation values were found between MDCT and CMR tools, and between MR Argus and CMR tools. It was observed that MDCT had more accurate and reliable results when compared with 2DE. Finally, in the study of Akram et al. [21], left ventricle functions that were calculated with CMR and MDCT were compared in 20 patients and good and perfect degree correlations were obtained in all values. Furthermore, they reported that the duration of post-process procedures in calculations that were done with MDCT were 50% lower according to CMR. In that study, less time was expended for function assessment with MDCT, as well. Analysis with the CMR tools method was conducted over the longest period of time. As a result, when all three modalities were compared, it was observed that the best correlation was between the two MR softwares. However, the correlation of the MDCT results with CMR was better than 2DE, whereas 2DE-CMR correlations were lower than expected. Although the gold-standard noninvasive technique to evaluate left ventricle functions is MR, its use in clinical practice for this purpose is limited, as it is an expensive and long-lasting procedure. On the other hand, CCTA examination is a newer and rapidly developing noninvasive cardiac and coronary artery imaging method. As there is radiation exposure and contrast material used in this method, it is not rational to perform cardiac CCTA to assess only cardiac function. However, coronary CCTA, which is a noninvasive coronary artery imaging method, could be also used to evaluate left ventricle functions. Thus, by using the available raw data, anatomical and functional assessment could be done without leading to any increase in cost and dose of contrast material and radiation. Furthermore, it provides more reliable information compared to 2DE, which is the most commonly used method in routine practice.
Conclusions

In conclusion, with CCTA, it is possible to determine left ventricle functions together with the condition of coronary arteries in patients with coronary artery diseases more easily, and important information related to the prognosis could be obtained.

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Conflicts of interest

The authors declare that they have no conflicts of interest.