Diagnostic Accuracy of 3.0-T Magnetic Resonance T1 and T2 Mapping and T2-Weighted Dark-Blood Imaging for the Infarct-Related Coronary Artery in Non–ST-Segment Elevation Myocardial Infarction

Jamie Layland, PhD; Samuli Rauhalammi, MSc; Matthew M. Y. Lee, MBChB; Nadeem Ahmed, MBChB; Jaclyn Carberry, BMedSci; Vannesa Teng Yue May, BMedSci; Stuart Watkins, MD; Christie McComb, PhD; Kenneth Mangion, MBChB; John D. McClure, PhD; David Carrick, PhD; Anna O’Donnell, BN; Avind Sood, MD; Margaret McEntegart, PhD; Keith G. Oldroyd, MD (Hons); Aleksandra Radjenovic, PhD; Colin Berry, PhD

Background—Patients with recent non–ST-segment elevation myocardial infarction commonly have heterogeneous characteristics that may be challenging to assess clinically.

Methods and Results—We prospectively studied the diagnostic accuracy of 2 novel (T1, T2 mapping) and 1 established (T2-weighted short tau inversion recovery [T2W-STIR]) magnetic resonance imaging methods for imaging the ischemic area at risk and myocardial salvage in 73 patients with non–ST-segment elevation myocardial infarction (mean age 57±10 years, 78% male) at 3.0-T magnetic resonance imaging within 6.5±3.5 days of invasive management. The infarct-related territory was identified independently using a combination of angiographic, ECG, and clinical findings. The presence and extent of infarction was assessed with late gadolinium enhancement imaging (gadobutrol, 0.1 mmol/kg). The extent of acutely injured myocardium was independently assessed with native T1, T2, and T2W-STIR methods. The mean infarct size was 5.9±8.0% of left ventricular mass. The infarct zone T1 and T2 times were 1323±68 and 57±5 ms, respectively. The diagnostic accuracies of T1 and T2 mapping for identification of the infarct-related artery were similar (P=0.125), and both were superior to T2W-STIR (P<0.001). The extent of myocardial injury (percentage of left ventricular volume) estimated with T1 (15.8±10.6%) and T2 maps (16.0±11.8%) was similar (P=0.838) and moderately well correlated (r=0.82, P<0.001). Mean extent of acute injury estimated with T2W-STIR (7.8±11.6%) was lower than that estimated with T1 (P<0.001) or T2 maps (P<0.001).

Conclusions—In patients with non–ST-segment elevation myocardial infarction, T1 and T2 magnetic resonance imaging mapping have higher diagnostic performance than T2W-STIR for identifying the infarct-related artery. Compared with conventional STIR, T1 and T2 maps have superior value to inform diagnosis and revascularization planning in non–ST-segment elevation myocardial infarction.

Clinical Trial Registration—URL: http://www.clinicaltrials.gov. Unique identifier: NCT02073422. (J Am Heart Assoc. 2017;6:e004759. DOI: 10.1161/JAHA.116.004759.)

Key Words: acute coronary syndrome • area at risk • edema • mapping • noninvasive imaging • non–ST-segment elevation acute coronary syndrome

Followings acute myocardial infarction (MI), MRI techniques have been shown to retrospectively estimate the ischemic area at risk (AAR), that is, the myocardial perfusion bed directly affected by ischemia because of acute occlusion of a coronary artery.1 Furthermore, as a result of even transient ischemia producing differences in the myocardial longitudinal (T1) and transverse (T2) relaxation times, newer techniques have shown improved sensitivity in identifying...
myocardial injury. Dark-blood fat-suppressed (short tau inversion recovery [STIR]) T2-weighted MRI methods (T2W-STIR) are widely used for clinical and research purposes. T2W-STIR, however, is prone to artifacts from motion, blood–tissue borders, and coil sensitivity issues that cause signal loss with depth of field and may lead to diagnostic uncertainty. Novel fast T1 and T2 parametric methods for mapping myocardial injury have been described in patients with acute coronary syndrome—predominantly in patients with ST-segment elevation MI (STEMI). In these cohorts, both T1 and T2 mapping techniques demonstrated superior accuracy compared with T2W-STIR. Given that edema may persist for days or weeks after acute ischemic injury, STEMI patients have generally been a focus for research with MRI following revascularization. Non-STEMI (NSTEMI) is a condition typified by smaller degrees of myocardial injury and nonocclusive culprit lesions. Compared with STEMI, the clinical utility of identifying the territory of acute injury in NSTEMI patients may be enhanced because NSTEMI is a more heterogeneous condition associated with diagnostic uncertainty, and MRI can be performed before a decision to proceed to an invasive management strategy. Nevertheless, data allowing direct comparison of T1 and T2 mapping sequences and T2W-STIR in NSTEMI, particularly at 3.0 T, are lacking and may not necessarily be congruent with findings in the STEMI cohorts.

We performed a prospective study of medically stabilized NSTEMI patients referred for coronary angiography to compare the diagnostic accuracy of T1 and T2 mapping compared with T2-weighted dark-blood edema imaging (T2W-STIR) for detection of the infarct-related artery (IRA). We also compared these MRI methods for estimation of the angiographic ischemic AAR.

Methods

Patient Population

Patients with a diagnosis of type 1 NSTEMI with at least 1 cardiac risk factor scheduled for early coronary angiography were prospectively recruited from 2 cardiac institutions. Exclusion criteria represented standard contraindications to MRI, including metallic devices and severe kidney disease (an estimated glomerular filtration rate <30 mL/min per 1.73 m²). The participants in this study were enrolled in the Fractional Flow Reserve Versus Angiographically Guided Management to Optimise Outcomes in Unstable Coronary Syndromes (FAMOUS NSTEMI) clinical trial, a multicenter study examining the utility of fractional flow reserve use in this population (ClinicalTrials.gov identifier NCT02073422). Only patients enrolled in centers with access to cardiac MRI (CMR) were approached to participate. All patients provided written informed consent. The local ethics board approved the study. Pharmacological management of patients reflected contemporary guidelines including treatment with dual-antiplatelet therapy, an HMG-CoA enzyme inhibitor, and a beta blocker. Troponin checks were routinely performed before and 12 hours after the procedure to assess for procedure-related (type 4a) MI.

MRI Acquisition

MRI was performed on a Siemens MAGNETOM Verio 3.0-T scanner with an 8-element phased-array cardiac surface coil. The MRI protocol included steady-state free procession cine MRI, T1 and T2 parametric maps, and T2W-STIR before gadolinium contrast administration. Cine steady-state free procession images with 2-fold accelerated parallel imaging (GRAPPA [generalized autocalibrating partially parallel acquisition]) were acquired in a stack of short-axis views of the left ventricle. Shimming and a radio frequency localizer were used at the outset of the scan to optimize data quality. Imaging parameters were repetition time 3.4 ms, echo time 1.51 ms, flip angle 50°, typical field of view 340×286 mm², matrix 256×216, slice thickness 7 mm, slice gap 3 mm, receiver bandwidth 977 Hz/pixel, 25 cardiac phases. Late gadolinium enhancement (LGE) CMR was performed with a T1-weighted, segmented, gradient echo, phase-sensitive inversion recovery sequence, with following parameters: echo time 760 ms, repetition time 1.56 ms, flip angle 20°. The inversion time was adjusted for optimal suppression of signal from normal myocardium (inversion time ≈340 ms). Typical field of view was 350×262 mm², matrix 256×192, slice thickness 7 mm, slice gap 2.8 mm, and bandwidth 465 Hz/pixel. Images were collected 10 to 15 minutes after the administration of 0.1 mmol/kg of contrast agent (Gadovist; Bayer).

The ECG-gated single-shot modified Look-Locker inversion recovery (MOLLI) method (3[3][3][5] protocol) was used to measure myocardial longitudinal relaxation times (T1). The MOLLI parameters included a T1 start of 100 ms, an increment of 80 ms, and a trigger delay of 160 ms. The protocol included a GRAPPA acceleration factor of 2. Following pixel-wise T1 fitting, T1 (ms) was displayed on a quantitative, color, scaled map.

Myocardial transverse (T2) relaxation times were estimated using a T2 mapping technique that involved a T2 prepared True Fast Imaging with Steady state Precession (TrueFISP) pulse sequence to produce single-shot T2 prepared images, each with different T2 preparation times. The T2 prepared TrueFISP images were acquired at intervals of at least 2 R–R intervals to allow for sufficient magnetization recovery in between acquisitions. Motion correction was enabled to prevent misregistration between images. T2 was estimated by
pixel-wise fitting assuming monoexponential signal decay, and a color, scaled, motion-corrected myocardial T2 map was then generated. For both T1 and T2 maps, an algorithm for motion correction based on previous work was applied before curve fitting.17

The breath-hold black-blood triple inversion recovery sequence (T2W-STIR) involves a pair of selective and nonselective 180° inversion pulses to null the blood pool signal followed by a third inversion pulse (STIR) to null the fat signal.4 Surface coil intensity correction was routinely performed with prescan normalization and slice-related shimming, as appropriate. Typical imaging parameters were echo time of 2 R-R intervals, repetition time of 47 ms, and flip angle of 180°, with 310 Hz/pixel bandwidth and turbo factor 15. Inplane spatial resolution was 2.2×1.4 mm (with acquisition matrix of 151×256, GRAPPA=2), and slice thickness was 6 mm. Short-axis left ventricular (LV) views were sequentially acquired with T1/T2 parametric maps and T2W-STIR matched to the same slice position. We obtained 3 slices: basal, mid, and apical. Sample images are shown in Figure 1.

**MRI Analysis**

Anonymized images were analyzed in random order on a Siemens workstation by 2 MRI-trained cardiologists with >3 years experience with imaging myocardial edema. An image analyst coordinated data management to ensure blinded analysis. A third MRI-trained cardiologist and an MRI-trained technician performed quantitative assessments of infarct area and AAR as determined by both T1 and T2 maps.

Two MRI-trained cardiologists who were blind to the patients’ clinical presentation reviewed hyperintense zones on T1- and T2-weighted MRI images independently. Specific edema imaging sequences were analyzed randomly and independently from each other to avoid observer bias. Each observer assessed for the presence or absence of edema in each short-axis slice and ascribed it to a segment in accordance with the 16-segment model of the American Heart Association. A third observer resolved discordance between cardiologists. Each observer was also asked to assign a culprit coronary arterial territory for each patient.

Figure 1. Magnetic resonance imaging findings in a 62-year-old male patient 4 days after hospitalization because of an acute non–ST-segment elevation myocardial infarction: (A) T2 map, (B) T2-weighted short tau inversion recovery (T2W-STIR), (C) T1 map, (D) late gadolinium enhancement (LGE). Inferior subendocardial infarction, as revealed by LGE, corresponds with the transmural extent of myocardial injury revealed with T1 and T2 maps, but this is not seen with T2W-STIR. Conversely, the anterior wall is hyperintense with T2W-STIR.
Detecting Infarct-Related Territory With 3.0-T MRI

Layland et al

V.T.

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Statistical analyses were carried out using IBM SPSS Statistics software, version 21.0. Normality was tested with the Shapiro–Wilk test. All results are given as mean±SD unless otherwise stated. Correlations between AAR were quantified with T1/T2 maps and T2W-STIR. APPROACH lesion scores were tested by Pearson or Spearman methods, as appropriate. Comparisons of normally distributed continuous data between T1/T2 maps and T2W-STIR were undertaken using a Student t test. Between-group comparisons of nonnormally distributed data were performed with a Mann–Whitney test. The level of agreement between the myocardial territory of acute injury was quantified with T1/T2 maps, and T2W-STIR was assessed using Bland–Altman plots and 95% limits of agreement. The 95% limits of agreement were calculated using the mean difference between areas of acute injury quantified by the 2 imaging modalities ±2 SD of these differences and contained ≈95% of all such differences. The McNemar exact test was used to compare accuracy of the T1, T2, and T2W-STIR sequences in identifying the IRA compared with the current gold standard method of coronary angiography. In addition, binomial tests (with 95% CIs calculated using the likelihood ratio) were also used to determine how well the novel mapping techniques correctly identified the IRA compared with other methods such as ECG, wall motion abnormalities on MRI, and LGE. To compare the degree of overlap between remote and injured zones for each mapping technique, effect sizes (in the form of Cohen’s d) were calculated, and these values were converted to overlapping coefficients using a formula described previously.

An interobserver agreement reliability analysis, using the κ statistic, was performed to determine the consistency of IRA diagnosis among observers. To assess reproducibility, quantitative assessment of the area of acute injury on T1 and T2 maps was performed by a second observer in a subset of 18 patients. The level of agreement between the 2 observers assessing the AAR on T1 and T2 maps was assessed using Bland–Altman plots and 95% limits of agreement. P<0.05 was considered statistically significant.

The study is publicly registered (ClinicalTrials.gov identifier NCT02073422).

Results

Patient Characteristics

Between February 2012 and May 2013, we recruited 73 NSTEMI patients (mean age 57±10 years, 78% male) who underwent CMR within 6.5±3.5 days of invasive management. All patients fulfilled the diagnostic criteria for acute NSTEMI, and the infarct-related coronary arteries were identified by clinical criteria including ECG and coronary angiography. Patient demographics and clinical characteristics are provided in Table 1. The mean Global Registry of Acute Coronary Events score at hospital was 156±37. Overall, 27.4% of patients were imaged before and 72.6% were imaged after coronary angiography. Angiographic characteristics, in particular, the IRA characteristics and assignments, are provided in Table 1. Overall, 63% of the study population underwent percutaneous coronary intervention. All patients had evidence of coronary artery disease at angiography.

Angiographic Analysis

The attending interventional cardiologist used a combination of clinical history, ECG, and coronary angiographic findings to identify the culprit coronary artery. Culprit artery assignment was independently analyzed and verified by an accredited interventional cardiologist (M.M.C.). The Alberta Provincial Project for outcome Assessment in Coronary Heart Disease (APPROACH) lesion score was used to provide estimates of the AAR. In addition, all angiograms were independently analyzed to assess plaque characteristics by a core lab (J.C., V.T.).

Statistical Analyses

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The jeopardized myocardial territory of acute injury was defined as the percentage of LV area delineated by the hyperintense zone on T1 and T2 images. The signal intensity threshold indicating injured myocardium was set at 2 SD above the mean intensity of reference (region of interest) placed in remote unaffected myocardium (180° from the affected zone with no visible evidence of infarction, edema, or wall motion abnormalities). Myocardial salvage was calculated by subtraction of the percentage of infarct size from the percentage of the area of acute injury. The extent of infarct scar was delineated as an area of myocardial enhancement (cm²) mapped on LGE images, using a signal intensity threshold of >5 SD above remote region and expressed as a percentage of total LV mass. Microvascular obstruction (MVO) was defined as a dark zone present within an area of gadolinium enhancement. Infarct regions with evidence of MVO were included within the infarct area, and MVO area was also recorded individually as a percentage of total LV mass.

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Cardiac MRI Findings

The CMR findings are summarized in Tables 2 and 3. Of 26 patients (35.6%) without evidence of LGE, 16 (61.5%) had evidence of acute myocardial injury revealed by T1 and T2 mapping, whereas fewer participants (7, 27%) had evidence of myocardial injury using T2W-STIR. The mean infarct size as a percentage of LV volume was 5.9 ± 8.0%. There was no evidence of acute myocardial injury using 2 of the 3 imaging methods in 10 patients (13.6%).

Microvascular obstruction was detected in 10 patients (14%). The troponin concentration in patients with MVO was numerically higher (7.43 ± 9.9) than in patients without MVO (4.37 ± 8.01; \( P = 0.57 \)). The mean LV ejection fraction was 57 ± 12%.

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age, y</td>
<td>57±10</td>
</tr>
<tr>
<td>Male, n (%)</td>
<td>57 (78)</td>
</tr>
<tr>
<td>Body mass index, kg/m²</td>
<td>28.7±4.8</td>
</tr>
<tr>
<td>Smoker, n (%)</td>
<td>52 (72.2)</td>
</tr>
<tr>
<td>Hypertension, n (%)</td>
<td>25 (34.7)</td>
</tr>
<tr>
<td>Hyperlipidemia, n (%)</td>
<td>23 (32.4)</td>
</tr>
<tr>
<td>Diabetes mellitus, n (%)</td>
<td>9 (12.3)</td>
</tr>
<tr>
<td>Previous PCI, n (%)</td>
<td>6 (8.2)</td>
</tr>
<tr>
<td>GRACE score</td>
<td>156±37.25</td>
</tr>
<tr>
<td>Troponin, mg/dL</td>
<td>0.92 (0.2, 5.3)</td>
</tr>
</tbody>
</table>

Infarct-related artery

<table>
<thead>
<tr>
<th>Category</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Left anterior descending, n (%)</td>
<td>33 (45)</td>
</tr>
<tr>
<td>Circumflex, n (%)</td>
<td>23 (32)</td>
</tr>
<tr>
<td>Right coronary, n (%)</td>
<td>17 (23)</td>
</tr>
</tbody>
</table>

Angiographic characteristics

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Thrombus</td>
<td>17.6%</td>
</tr>
<tr>
<td>Ulcerated</td>
<td>58.8%</td>
</tr>
<tr>
<td>Irregular</td>
<td>64.7%</td>
</tr>
<tr>
<td>Calcified</td>
<td>17.6%</td>
</tr>
<tr>
<td>TIMI flow &lt;3</td>
<td>19.6%</td>
</tr>
<tr>
<td>Stenosis, %</td>
<td>84.79±15.7%</td>
</tr>
<tr>
<td>Multivessel disease, n (%)</td>
<td>36 (49)</td>
</tr>
</tbody>
</table>

Treatment strategies

<table>
<thead>
<tr>
<th>Category</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Medical therapy only, n (%)</td>
<td>21 (28.7)</td>
</tr>
<tr>
<td>Coronary artery bypass grafting, n (%)</td>
<td>6 (8.2)</td>
</tr>
<tr>
<td>PCI, n (%)</td>
<td>46 (63)</td>
</tr>
<tr>
<td>APPROACH lesion score, %LV mass</td>
<td>23.45±12.6</td>
</tr>
</tbody>
</table>

GRACE indicates Global Registry of Acute Coronary Events; LV, left ventricular; PCI, percutaneous coronary intervention; TIMI, Thrombolysis in Myocardial Infarction.

Detection of Acute Myocardial Injury by T1 and T2 Maps and T2-Weighted Short Tau Inversion Recovery

There was no significant difference between T1 and T2 maps for classification of acute myocardial injury (T1 versus T2 maps: 64 [88%] versus 63 [86%] patients, respectively); however, there were significant differences when comparing T2W-STIR (42, 58%) with T1 maps (\( P = 0.019 \)) and T2 maps (\( P = 0.012 \)).

Detection of Acute Myocardial Injury by T1 and T2 Maps and T2W-STIR Images

On average, 3 short-axis slices were available for native T1 and T2 maps and T2W-STIR images. Overall, 83.3% of patients had either excellent or good image quality. No patient was excluded because of poor image quality. There were significantly more artifacts with T2W-STIR (57.1% of images) compared with T1 maps (12.7%) and T2 maps (3.7%; \( P < 0.0001 \)). A detailed breakdown of artifacts is provided in Table 4. When comparing the 2 parametric mapping techniques directly, there were significantly more artifacts with T1 maps compared with T2 maps (\( P < 0.001 \)).

Table 3. Comparisons of the presence and extent of myocardial injury and salvage as revealed by T1 and T2 mapping and T2W-STIR.

<table>
<thead>
<tr>
<th></th>
<th>T1</th>
<th>T2</th>
<th>T2W-STIR</th>
<th>( P ) Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patients with evidence of myocardial injury</td>
<td>64 (88)</td>
<td>63 (86)</td>
<td>42 (58)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Area of myocardial injury (%LV volume)</td>
<td>15.8±10.6</td>
<td>16.0±11.8</td>
<td>7.8±11.6</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Myocardial salvage (%LV volume)</td>
<td>10.1±8.9</td>
<td>10.5±9.5</td>
<td>4.7±9.7</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

LV indicates left ventricular.
Regarding patient characteristics among those with any myocardial injury (any) compared with those without, patients with injury were significantly more likely to have LGE (44 of 66 versus 1 of 7, \( \chi^2 = 7.34, P = 0.007 \)) and higher troponin (mean 5.26 ± 8.54 versus 0.24 ± 0.31, \( P = 0.001 \)). There were no significant differences in TIMI (Thrombolysis in Myocardial Infarction) flow (2.71 ± 0.7 versus 2.86 ± 0.38, \( P = 0.59 \)), fractional flow reserve values (0.78 ± 0.19 versus 0.82 ± 0.17, \( P = 0.57 \)), ECG evidence of ischemia (37 of 66 versus 2 of 7, \( P = 0.16 \)), or MVO (10 of 66 versus 0 of 7, \( P = 0.26 \)).

**Diagnostic Accuracy of T1 and T2 Maps and Dark-Blood T2-Weighted Short Tau Inversion Recovery**

The diagnostic accuracy of each imaging method for identification of the IRA, as defined by clinical data including ECG and coronary angiogram, are shown in Table 5. The IRA was correctly identified more often with T2 maps compared with other imaging methods (77% T2 maps, 71% T1 maps, 44% T2W-STIR); however, when assessing diagnostic accuracy, there was no difference when comparing T1 and T2 maps (\( P = 0.125 \)). In contrast, differences in diagnostic accuracy were shown between T1 maps and T2W-STIR (\( P < 0.001 \)) and between T2 maps and T2W-STIR (\( P < 0.001 \)). For IRA identification, a high level of interobserver agreement was found with T1 maps (\( \kappa = 0.790, P < 0.001 \)) and T2 maps (\( \kappa = 0.794, P < 0.001 \)), whereas the level of agreement was moderate (\( \kappa = 0.555, P = 0.001 \)) with T2W-STIR. Moreover, there was a lower level of discordance between observers for individual patients using T1 maps (discordant findings in 13.69%) and T2 maps (15.06%) compared with dark-blood T2W-STIR (30.14% cases, \( P < 0.001 \)). We also compared each technique with other methods of identifying the IRA, such as ECG, presence of LGE, and regional wall motion abnormalities. The proportions of correctly identified infarct-related arteries for T1 (0.84, 95% CI 0.74–0.91, \( P < 0.001 \)), T2 (0.86, 95% CI 0.77–0.93, \( P < 0.001 \)), and regional wall motion abnormalities (0.64, 95% CI 0.53–0.75, \( P = 0.019 \)) were significantly higher than chance alone. For several other techniques, however, the proportions of accurate results did not significantly differ from 0.50: ECG 0.53 (95% CI 0.42–0.65, \( P = 0.640 \)), LGE 0.62 (95% CI 0.50–0.72, \( P = 0.061 \)), T2W-STIR 0.51 (95% CI 0.39–0.62, \( P = 0.999 \)).

When assessing the accuracy of these techniques in specific subgroups, there were no differences in the numbers of patients with correctly identified infarct-related arteries, depending on whether they had multivessel disease, using T1 maps (29 of 36 versus 32 of 37, \( \chi^2 = 0.47, P = 0.494 \)), T2 maps (31 of 36 versus 31 of 36, \( \chi^2 = 0.00, P = 1.00 \)), and T2W-STIR (18 of 36 versus 19 of 37, \( \chi^2 = 0.01, P = 0.908 \)).

### Table 4. Detailed Breakdown of Artifacts

<table>
<thead>
<tr>
<th></th>
<th>IRA Territory Identified</th>
<th>T2 Map</th>
<th>T2-W-STIR Map</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Correct (n=56)</td>
<td>Incorrect (n=17)</td>
<td>All</td>
</tr>
<tr>
<td>T1 map</td>
<td>52</td>
<td>0</td>
<td>52 (71%)</td>
</tr>
<tr>
<td>Incorrect (n=21)</td>
<td>4</td>
<td>17</td>
<td>21 (29%)</td>
</tr>
<tr>
<td>T2 map</td>
<td>56 (77%)</td>
<td>17 (23%)</td>
<td>( P = 0.13 )</td>
</tr>
</tbody>
</table>

### Table 5. Diagnostic Accuracies of Each Imaging Method for Identification of the IRA

<table>
<thead>
<tr>
<th>IRA Territory Identified</th>
<th>T2 Map</th>
<th>T2-W-STIR Map</th>
</tr>
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<tbody>
<tr>
<td>Correct (n=56)</td>
<td>Incorrect (n=17)</td>
<td>All</td>
</tr>
<tr>
<td>T1 map</td>
<td>52</td>
<td>0</td>
</tr>
<tr>
<td>Incorrect (n=21)</td>
<td>4</td>
<td>17</td>
</tr>
<tr>
<td>T2 map</td>
<td>56 (77%)</td>
<td>17 (23%)</td>
</tr>
</tbody>
</table>

**Table 4. Detailed Breakdown of Artifacts**

<table>
<thead>
<tr>
<th></th>
<th>T1 (219)</th>
<th>T2 (219)</th>
<th>T2W-STIR (219)</th>
</tr>
</thead>
<tbody>
<tr>
<td>MOCO</td>
<td>18</td>
<td>12</td>
<td></td>
</tr>
<tr>
<td>M</td>
<td>5</td>
<td>4</td>
<td>79</td>
</tr>
<tr>
<td>RF</td>
<td>28</td>
<td></td>
<td></td>
</tr>
<tr>
<td>SB</td>
<td>10</td>
<td></td>
<td></td>
</tr>
<tr>
<td>G</td>
<td>4</td>
<td></td>
<td></td>
</tr>
<tr>
<td>FE</td>
<td>1</td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>W</td>
<td>4</td>
<td>2</td>
<td>1</td>
</tr>
<tr>
<td>SSFP</td>
<td>1</td>
<td>1</td>
<td></td>
</tr>
</tbody>
</table>

F indicates flow; FE, field effects; G, gating; M, motion (eg, breathing); MOCO, motion correction; RF, radiofrequency field inhomogeneity; S, susceptibility; SB, stagnant blood; SSFP, steady-state free precession off-resonance bands; T2W-STIR, T2-weighted short tau inversion recovery; W, Wrap.

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45±3 ms, P<0.001; T2W-STIR, 146±55 versus 90±25, P<0.001); however, there was some overlap between distributions. For T1 maps, there was a large effect size (d=3.1, P<0.001) for the difference between infarct zone and remote scores with a small amount of overlap (12.5%). For T2 maps, there was an equally large effect (d=2.9) for the difference between infarct zone and remote scores with a small amount of overlap (14.9%). For T2W-STIR, there was a somewhat smaller effect size (d=1.3) for the difference between infarct zone and remote scores with much greater overlap (51.0%) (Figure 2).

For the extent of acute myocardial injury (percentage of LV volume) estimated by 2 independent observers using T1 mapping, there was good correlation between observers 1 and 2 (r²=99.7%). The 95% limits of agreement were −1.76 and 1.63 (percentage of LV volume), respectively, and there was no evidence of bias (0.25). For T2 mapping between these observers, there was good correlation (r²=98.9%), with 95% limits of agreement of −2.98 and 3.36 (percentage of LV volume) and no evidence of bias (0.19).

**Variable Quantification of the Extent of Acute Myocardial Injury by T1 and T2 Maps and T2-Weighted Short Tau Inversion Recovery**

There was good correlation between T1 and T2 maps for estimation of myocardial injury (r=0.82, P<0.001; T1 maps versus T2 maps: 15.8±10.6% versus 16.0±11.8% of LV volume, P=0.838). The 95% limits of agreement for mean myocardial injury estimated with T1 versus T2 maps were −13% and 13%, respectively, of LV volume with a minimal bias (0.2±6.8%) (Figures 3 and 4).

There were moderate correlations between areas of acute injury estimated with T1 maps versus T2W-STIR (r=0.54, P<0.001) and injury estimated with T2 maps versus T2W-STIR (r=0.46, P<0.001). Mean area of acute injury estimated with T2W-STIR (7.8±11.6% of LV volume) was smaller than that estimated with T1 (P<0.001) or T2 (P<0.001). The 95% limits of agreement for mean area of acute injury estimated with T1 maps versus T2W-STIR and T2 maps versus T2W-STIR were wider with a higher level of bias compared with the comparison for T1 versus T2 maps (T1 versus T2W-STIR: −28% and 12% of LV volume with a bias of −8.0±10.6%; T2 versus T2W-STIR: −32% and 16% with a bias of −8.1±12.1%).

The average amount of myocardial salvage (percentage of LV myocardial volume) estimated with T1 maps (10.1±8.9%) compared with T2 maps (10.5±9.5%, P=0.616) was similar. However, the mean myocardial salvage estimated with T2W-STIR (4.7±9.7% of LV volume) was significantly lower than estimates with T1 maps (P=0.027) or T2 maps (P=0.009).

**Correlation of the Extent of Acute Myocardial Injury Estimated by T1- and T2-Weighted MRI Versus APPROACH Lesion Score**

The mean APPROACH lesion score was 23.4±12.6 of the LV mass. CMR estimates of extent of acute myocardial injury did not correlate with the APPROACH lesion score. Myocardial

![Figure 2. T1 and T2 values (m/s) in remote territory and in the injury zone (IZ).](https://example.com/figure2.png)
injury estimates were compared for T1 maps versus APPROACH ($r=0.13$, $P=0.29$), T2 maps versus APPROACH ($r=0.11$, $P=0.37$), and T2W-STIR versus APPROACH ($r=0.08$, $P=0.49$).

The main results of our study are as follows. First, using 3.0-T MRI, native T1 and T2 maps have higher diagnostic accuracy than dark-blood T2W-STIR and other techniques, including ECG, in detecting the IRA in patients with recent NSTEMI. Second, T1 and T2 maps produced highly congruent results when estimating the extent of myocardial injury in contrast to dark-blood T2W-STIR imaging that tended to underestimate the extent of injury compared with these methods. Third, reflecting the heterogeneity of NSTEMI patients (including interindividual variations in natural history, presence of multivessel disease, and typical preservation of culprit artery blood flow), T2-weighted CMR does not reliably quantify the ischemic area of acute injury. Fourth, there was a high level of interobserver agreement for identification of the IRA with T1 and T2 maps compared with dark-blood T2W-STIR. Finally, artifacts were significantly more prevalent with dark-blood imaging compared with T1 and T2 maps.

Discussion

The main results of our study are as follows. First, using 3.0-T MRI, native T1 and T2 maps have higher diagnostic accuracy than dark-blood T2W-STIR and other techniques, including ECG, in detecting the IRA in patients with recent NSTEMI. Second, T1 and T2 maps produced highly congruent results when estimating the extent of myocardial injury in contrast to dark-blood T2W-STIR imaging that tended to underestimate the extent of injury compared with these methods. Third, reflecting the heterogeneity of NSTEMI patients (including interindividual variations in natural history, presence of multivessel disease, and typical preservation of culprit artery blood flow), T2-weighted CMR does not reliably quantify the ischemic area of acute injury. Fourth, there was a high level of interobserver agreement for identification of the IRA with T1 and T2 maps compared with dark-blood T2W-STIR. Finally, artifacts were significantly more prevalent with dark-blood imaging compared with T1 and T2 maps.
Unlike patients with STEMI, identification of the IRA in patients with the IRA in patients with NSTEMI for several reasons including a lack of ECG changes and the presence of multivessel disease. Furthermore, a significant proportion of patients who present with NSTEMI have normal coronary arteries at the time of coronary angiography. This number could be expected to increase with the increasing use of high-sensitivity troponin. Our data demonstrate that novel T1 and T2 maps have higher accuracy for predicting the IRA in patients with NSTEMI compared with standard diagnostic techniques such as ECG and location of regional wall motion abnormalities. Consequently, the use of CMR in the diagnostic workup of patients with NSTEMI may enable selection of appropriate patients for invasive treatment and direct treatment to the appropriate coronary territory.

Imaging Acute Myocardial Injury

The detection of acute myocardial injury using a dark-blood turbo spin-echo technique has been shown to facilitate prompt diagnosis of acute coronary syndromes and to allow identification of both the AAR and the amount of myocardial salvage after reperfusion. Yet, despite great theoretical promise, the clinical reality of using dark-blood T2W-STIR imaging has been far from optimal because of methodological problems inherent in the technique. Recently, a noncontrast quantitative T2 mapping sequence using T2 steady-state free procession has been proposed and shown to be more accurate than conventional T2W-STIR for predicting myocardial injury in animal and human models.

Previous experimental studies have also shown that T1 values increase with increasing myocardial water content and thus with ischemia and infarction. By directly quantifying T1 values for each voxel in the myocardium, a parametric map can be generated representing the T1 relaxation times of any region of the heart. The most widely adopted T1 mapping sequence is based on the MOLLI technique. The use of T1 maps has also been shown to be more accurate than conventionally T2W-STIR for predicting myocardial injury in animal and human models.

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Dark-Blood Imaging

Following its development by Simonetti and colleagues, dark-blood T2W-STIR has been widely used to assess for the presence of myocardial edema. Several investigators have shown prognostic information using STIR imaging. In a prospective study involving 88 patients with NSTEMI, for example, Raman and colleagues highlighted the potential for T2W-STIR imaging in identifying acute myocardial injury and distinguishing patients who required coronary revascularization from those who did not. Moreover, the presence of an increase in regional signal intensity was associated with a higher hazard of a cardiovascular event or death within 6 months after a non-ST-segment elevation acute coronary syndrome.

Recent studies have shown improved accuracy in the detection of acute myocardial injury with both T1 mapping and T2 mapping techniques compared with dark-blood imaging at 1.5 T. These findings were echoed by McAlindon and colleagues, who compared the accuracy of these techniques in 40 patients with reperfused STEMI. These investigators demonstrated that T2 mapping had less variability and improved detection of the IRA compared with dark-blood T2-weighted imaging.

NSTEMI is the most common form of acute coronary syndrome, yet to date, no study has directly compared T1 and T2 maps with dark-blood T2W-STIR in NSTEMI. Our study showed—for the first time in a large NSTEMI population—that at 3.0 T, the diagnostic performance of dark-blood imaging is inferior to newer novel mapping sequences, with underestimation of the area of acute injury as well as myocardial salvage. The identification of acute myocardial injury and the diagnostic accuracy of dark-blood imaging were also limited, which is important from a clinical perspective. Previous studies comparing dark-blood imaging with newer T2 or T1 methods have included only small numbers of patients with NSTEMI. Overall, the mean infarct size in these subpopulations has been moderate—16.7% of LV mass reported by Payne et al—and in the study by Verhaert and colleagues, the mean peak troponin concentration was 50 mg/dL. In our NSTEMI population, the mean infarct size was 5.9% and the troponin value was 0.92 mg/dL—a much lower infarct burden. The poor performance of STIR in this population calls into question the use of dark-blood edema MRI for imaging NSTEMI patients, either for clinical or research purposes. This point becomes all the more relevant because the sensitivity of the biochemical detection of NSTEMI increases with the adoption of high-sensitivity troponin assays. The higher sensitivity of T1 and T2 mapping for detecting subtle alterations in signal intensity that can occur in NSTEMI patients suggests that these methods may be more useful in clinical practice.

T1 and T2 Maps in Non-ST-Segment Elevation Myocardial Infarction

Dall’Armellina and colleagues assessed T1 with T2-weighted bright-blood imaging in 41 patients presenting with MI at 3.0 T. Overall, 73% of the study cohort presented with STEMI. T1- and T2-weighted imaging had similar diagnostic performance for detecting edematous injured myocardium. In a subgroup analysis performed in only 9 patients with...
Detecting Infarct-Related Territory With 3.0-T MRI

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NSTEMI, the authors demonstrated superior diagnostic performance of T1 maps over T2 imaging and less variability in T1 maps. In a much larger cohort of NSTEMI patients and using a different T1 mapping method, our results indicate that T1 and T2 maps had similar accuracy for estimating the myocardial AAR in patients with NSTEMI. In contrast to the findings of Dall’Armellina and colleagues, although there was no advantage in terms of accuracy, there were significantly fewer artifacts with T2 maps—suggesting an advantage for this mapping sequence—compared with T1 maps. Importantly, and despite this lower degree of injury, the utility of T1 and T2 mapping was preserved.

**Angiographic Area at Risk**

Several studies have clearly demonstrated the accuracy of both T1 and T2 maps in the assessment of the angiographic AAR in patients with acute coronary syndromes; however, the patient populations have largely consisted of STEMI patients. The APPROACH score has been shown to correlate with the myocardium at risk derived through CMR, and the magnitude of the correlation increased with the transmural extent of infarction. Typically, NSTEMI patients have smaller, subendocardial infarcts; therefore, it is not surprising to find that the AAR, as measured with edema imaging sequences, in our population did not correlate with the APPROACH score.

**Clinical Application**

T1 and T2 maps may prove useful to help identify the artery that should be treated before an invasive percutaneous coronary intervention is performed. CMR could potentially provide useful information to optimize the percutaneous coronary intervention plan by targeting the intervention to the culprit artery and by providing clinically relevant information on regional LV systolic function and myocardial viability. Our data support the potential role of CMR imaging as an adjunctive test to facilitate the invasive management of NSTEMI patients. Further research is warranted.

Retrospective imaging of the ischemic AAR is clinically relevant because the amount of salvageable myocardium is a predictor of prognosis and potential response to therapy. This study demonstrates the greater accuracy of T1 and T2 maps in identifying ischemically jeopardized myocardium compared with T2W-STIR imaging in patients with NSTEMI. Patients who experience an NSTEMI tend to be older and have more concomitant health problems and multivessel coronary disease. Using a more sensitive marker of myocardial injury than dark-blood T2W-STIR in this heterogeneous population may allow for improved risk stratification for patients who would benefit from early invasive assessment and may accurately identify the culprit territory when faced with a patient who has multivessel coronary disease.

**Study Limitations**

The timing of MRI after MI was influenced by the clinical course of the patients (eg, recurrent ischemia, the timing of invasive angiography). Although this variation in timing may have introduced some measurement heterogeneity, we think the MRI data became more representative of real-life clinical practice, reflecting the timing of when MRI might actually be performed in relation to acute presentation and/or invasive management. The participants were also enrolled in a clinical trial, but the intervention in this trial was not relevant to the MRI study, and the characteristics of our patients were similar to those of the trial participants and registry patients.

Despite higher accuracy for predicting the IRA, novel mapping sequences still failed to identify the IRA in ≈20% of cases. This may limit the clinical usefulness of the technique in a nonselected NSTEMI population.

MOLLI is sensitive to heart rate and this method may lead to modest inaccuracies in T1 (ms) in patients with a rapid heart rate, an irregular heart rate, or frequent ectopic beats. T2 mapping is also heart rate dependent; however, the majority of patients were on beta blockers before the MRI scan, with a controlled heart rate to minimize this effect.

We obtained only 3 slices to cover the left ventricle, and thus a theoretical concern exists that small NSTEMIs could have been missed using the MRI mapping technique.

Using the angiogram as the gold standard to identify the IRA is not without its flaws and could lead to a situation in which the mapping technique is correct but the IRA is incorrectly identified.

We used 6-mm slice thickness for STIR imaging in the study, and this may have enhanced imaging artifacts; however, this is within the range implemented by other researchers.

From an anatomical perspective, interindividual variation in coronary artery distribution may confound IRA assignment. In NSTEMI, multivessel coronary artery disease is common, and there may be >1 culprit lesion. Therefore, it could be that an IRA that was missed on angiography was correctly identified with the MRI maps. Because patients underwent CMR after invasive management with percutaneous coronary intervention, it is possible that the myocardial injury detected with T1 maps, T2 maps, and T2W-STIR actually originated from the percutaneous coronary intervention treatment rather than the initial MI. However, according to the current Universal Definition of Myocardial Infarction, none of the participants in this study experienced periprocedural MI.

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Conclusion

T1 and T2 maps are novel quantitative imaging methods that have high diagnostic accuracy in detecting the IRA in NSTEMI. They produce congruent results when estimating AAR, whereas T2W-STIR imaging yielded less accurate results with more artifacts. Findings suggest that T1 and T2 mapping represents a clinically valuable novel imaging method, with particular utility for the assessment of reversible myocardial injury and salvaged myocardium. Based on our findings, the utility of T2W-STIR imaging at 3.0-T for the assessment of edema in patients with NSTEMI is limited.

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Disclosures

None.

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