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Equal clinical performance of a novel point-of-care cardiac troponin I (cTnI) assay with a commonly used high-sensitivity cTnI assay

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ABSTRACT

Background: Efficient rule-out of acute myocardial infarction (MI) facilitates early disposition of chest pain patients in emergency departments (ED). Point-of-care (POC) cardiac troponin (cTn) may improve patient throughput. We compared the diagnostic accuracy of a novel cTnI test (Minicare cTnI, Philips), with current POC cTnI (I-Stat, Abbott) and high-sensitivity central laboratory cTnI (hs-cTnI; Architect, Abbott) assays.

Methods: The clinical performance of the assays were compared in samples from 450 patients from a previous clinical evaluation of Minicare cTnI.

Results: Minicare cTnI correlated with Architect hs-cTnI (r² = 0.85, p < 0.0001) and I-Stat cTnI (r² = 0.93, p < 0.0001). Areas under the receiver operating characteristics curves were 0.87–0.91 at admission (p = ns) and 0.96–0.97 3 h after admission (p = ns). The negative predictive values (NPV) at admission were 92–97% (95% CI) for Minicare cTnI and increased to 99% (97–100%) at 2–4 h, and similar to Architect hs-cTnI (98%, 96–100%), but higher than I-Stat cTnI (95%, 92–97%; p < 0.01). Negative likelihood ratios (LR−) after 2–4 h were 0.06 (0.02–0.17, 95% CI) for Minicare cTnI, 0.11 (0.05–0.24) for Architect hs-cTnI (p = 0.02) and 0.28 (0.18–0.43) for I-Stat cTnI (p < 0.0001). The clinical concordances between Minicare cTnI and Architect hs-cTnI were 92% (admission) and 95% (2–4 h), with lower concordances between Minicare cTnI and I-Stat cTnI (83% and 78%, respectively; p = 0.007).

Conclusions: The Minicare cTnI POC assay may become useful for prompt and safe ruling-out of AMI in ED patients with suspected AMI using a guideline supported 0/3 h sampling protocol.

1. Introduction

Achieving a prompt and accurate diagnosis of acute myocardial infarction (MI) is important, because early intervention saves lives. The diagnosis of MI is based on several criteria, including observation of circulating cardiac troponin I or T (cTnI or cTnT) above the upper reference limit (URL) [1–3]. Efficient rule-out of MI in patients presenting with symptoms of cardiac ischemia facilitates early disposition of the patient to outpatient care, and helps to relieve the increasing crowding of the emergency...
departments (ED) in most hospitals [4]. Sensitive cTn assays support such a decision based on results at admission and 3 h after admission, according to current guidelines (earlier rule-out protocols are based on the use of high-sensitive (hs) cTnT or cTnI assays) [3]. The time needed for results to become available from the hospital central laboratory often delays the disposition decision for another one or two hours, however [4]. Point-of-care (POC) assays have the potential to improve patient throughput in the ED, by providing earlier test results of cTn, but currently display a lower diagnostic accuracy than central laboratory cTnl/T assays [4–6]. A novel cTnI POC test, the Minicare cTnI (Philips), provides a cTnI result within 10 min.

The primary aim of this post-hoc analysis of a large, prospective multicentre diagnostic study [7] was to compare the clinical performance of the Philips Minicare cTnl assay with regard to rule-in or rule-out of MI in comparison with two widely used cTnl assays: the Abbott Architect® hs-cTnI and the POC Abbott I-Stat® cTnl.

2. Materials and methods

2.1. Patients

Patients admitted with suspected MI were enrolled prospectively at EDs or coronary care units/chest pain units of seven hospitals in Germany, France, Austria and the Netherlands as described previously [7]. Patients eligible for enrolment were adults (age ≥ 18 years) who had presented with symptoms suggestive of MI. Only patients who presented for the first time and < 12 h from onset of symptom onset were included. The study was approved by local ethical committees and all patients provided written, informed consent.

2.2. Study outcome and definitions

The primary outcome of the study was MI, based on evidence of myocardial necrosis in a clinical setting consistent with myocardial ischemia, defined according to the Third universal definition of myocardial infarction [1]. MI secondary to coronary procedures (MI type 4a or 5) or patients who died prior to biomarkers being obtained (MI type 3) were not included. Cases of MI were further sub-classified according to ST-segment changes to ST-segment elevation MI (STEMI) and non-ST-segment elevation MI (NSTEMI).

Serial blood samples for cTnl analyses were drawn at three different time points: admission, 2–4 h, and 6–24 h after the first blood draw/admission. The time window of blood sampling was set according to local hospital standard procedures.

2.3. Adjudication process

The pre-specified primary outcome was documented and reported by the Minicare cTnl study investigators, and adjudicated by an independent clinical events committee (CEC). The CEC adjudication process and precision of adjudicated diagnoses has been presented previously [8]. Only the data from patients with at least one central laboratory cTnT result above the 99th percentile upper reference limit (URL) using the hospital hs-cTnT test were adjudicated. Patients without elevated hs-cTnT levels were not considered as having an MI and were not adjudicated.

The final diagnosis of MI was based on the adjudication performed independently by two CEC cardiologist reviewers (Phase I review), who were blinded to the outcome of the Minicare cTnl test. Each reviewer assessed the information in the event package (e.g. ECG, clinical records) and the results from the hospital standard cTnl assay. If the two reviewers disagreed on the diagnosis or time of event in Phase I, the event was assessed at a second review in a CEC committee meeting (Phase II review). At least three cardiologist reviewers adjudicated the event and reached a consensus decision on the final diagnosis.

2.4. Measurement of cTnl

The concentrations of cTnT in lithium-heparin plasma were measured at the local hospital laboratories using a hs-cTnT assay (Roche); cTnl was measured in fresh lithium-heparinised whole blood and plasma samples using the Minicare cTnl assay at admission and 2–4 h after admission. Lithium-heparin plasma samples were frozen and shipped to Department of Clinical Chemistry, Uppsala University Hospital, Uppsala, Sweden, for the analysis by the Abbott Architect® hs-cTnI and the Abbott I-Stat® cTnl assays. Analysis of plasma samples occurred within 4 h of thawing.

The 99th percentile URLs provided by manufacturers were 43 ng/L for the Minicare cTnl assay, 26.2 ng/L for the Architect hs-cTnl assay, and 80 ng/L for the I-Stat cTnl assay. The analytical specifications of the Minicare cTnl were LoB (level of blank) 6.5 and 8.5 ng/L, LoD (level of detection) 18 and 17 ng/L for two different lots, respectively and LoQ (level of quantitation) at 20% CV was 38 ng/L (95% CI 28.3–47.7 ng/L). The CV at 99th percentile URL was calculated to be 18.6% [7].

2.5. Statistics

Non-parametric statistics were used for comparisons between groups and comparisons between assays were made using Passing-Bablok regression analysis. The clinical performance was evaluated by receiver operating characteristics (ROC) curve analysis, which in addition to area under the ROC curve (AuROC) provides information on sensitivity, specificity, positive predictive value (PPV), negative predictive value (NPV), and positive (LR+) and negative (LR-) likelihood ratios. The cut-off points used to calculate these parameters were the URLs (26.2 ng/L for Architect hs-cTnl, 80 ng/L for I-Stat cTnl and 43 ng/L for Minicare cTnl). Differences between AuROC were calculated according to DeLong et al. [10] and differences between proportions by Chi²-testing. The statistical programmes MedCalc (MedCalc Statistical Software version 16.8.4, MedCalc Software bvba, Ostend, Belgium) and SAS™ were used.

3. Results

3.1. Patients

Table 1 shows clinical characteristics of all 450 patients included in the study; a more detailed table of clinical characteristics is reported in Supplementary Table 1. Results from all three cTnl assays were obtained from 417 patients at admission and from 353 patients 2–4 h after admission. The comparative results given below are based on these cohorts. Supplementary Fig. 1 shows the number of patients included at admission and 2–4 h after admission as well as the numbers of patients given the diagnosis of MI by the adjudication panel.

3.2. Correlations and correspondences between cTnl analysers

Significant (p < 0.0001) correlations were observed using Passing-Bablok regression analysis for cTnl concentrations measured on the Minicare cTnl, either as whole blood or plasma, compared with the Architect hs-cTnl and I-Stat cTnl (Fig. 1). There was evidence of bias for both comparisons: 0.59 (whole blood) and 0.48 (plasma) for the Architect hs-cTnl and 0.48 (whole blood) and 0.43 (plasma) for the I-Stat cTnl. The correspondences relative to the cut-offs of the respective assay at admission were 92% between the Minicare cTnl and the Architect hs-cTnl, 83% between the Minicare cTnl and the I-Stat cTnl, and 81% between Architect hs-cTnl and I-Stat cTnl. The 8% discordant results among MI patients between Minicare cTnl and Architect hs-cTnl were almost equally distributed with 3% showing elevated concentrations with Minicare cTnl and 5% showing elevated results with Architect hs-cTnl, but non-elevated results with the other. The 17%
discordant results among MI patients between Minicare cTnI and I-Stat cTnI showed a much more skewed distribution since all discordant results had elevated Minicare cTnI results but non-elevated I-Stat cTnI results. Similarly, all 19% discordant results among MI patients between Minicare cTnI and I-Stat cTnI were due to elevated results. Between Architect hs-cTnI and I-Stat cTnI were due to elevated results of the assays. Sixteen patients with MI showed a much more skewed distribution since all discordant results had elevated Minicare cTnI results but non-elevated I-Stat cTnI results. Similarly, all 19% discordant results among MI patients between Minicare cTnI and I-Stat cTnI were due to elevated concentrations of cTnI in the MI patients are highly significant between all three assays (–normal (false negative) cTnI 2 respectively. The proportions of patients with adjudicated MI who had discordant results between Minicare cTnI and I-Stat cTnI of 83% and 78% (–hs-cTnI at admission and reduced to 3 patients (5%) after 2 h (p = 0.003 vs. admission; see Table 2). The overall concordance (Table 2) between Minicare cTnI whole blood and Architect hs-cTnI was 92% (admission) and 95% (2–4 h), with a lower concordance between Minicare cTnI and I-Stat cTnI of 83% and 78% (p = 0.007), respectively. The proportions of patients with adjudicated MI who had normal (false negative) cTnI 2–4 h after admission were 28% for the I-Stat cTnI and 5% for Minicare cTnI whole blood (p = 0.009). Similar results were obtained in the comparison with Minicare cTnI plasma and Architect hs-cTnI. False negative cTnI results were obtained in the comparison with Minicare cTnI plasma and Architect hs-cTnI. False negative cTnI at admission were 27–30% for Minicare cTnI and Architect hs-cTnI and 47% for I-Stat cTnI (p < 0.05). The clinical concordances between the assays relative to delay from onset of symptoms to admission are given in the supplementary file, Tables 2a and b. The concordances between the Minicare cTnI and Architect hs-cTnI were similar to the above results and with lower concordances to I-Stat cTnI when the patients were divided into those admitted < 4 h from onset and those admitted > 4 h from onset of symptoms.

### 3.3. Clinical performance

The comparisons between the clinical performances of the three assays are illustrated in Fig. 1 in which the results of patients with and without MI are compared by the findings with the Minicare cTnI whole blood assay and the Architect hs-cTnI and I-Stat cTnI assays at admission and 2–4 h after admission, respectively. The correlations between concentrations of cTnI in the MI patients are highly significant between all three assays (p < 0.0001). However, discrepancies were seen relative to the cut-offs of the assays. Sixteen patients with MI (25%) had normal cTnI according to either Minicare cTnI or Architect hs-cTnI at admission and reduced to 3 patients (5%) after 2–4 h (p = 0.003 vs. admission; see Table 2). The overall concordance (Table 2) between Minicare cTnI whole blood and Architect hs-cTnI was 92% (admission) and 95% (2–4 h), with a lower concordance between Minicare cTnI and I-Stat cTnI of 83% and 78% (p = 0.007), respectively. The proportions of patients with adjudicated MI who had normal (false negative) cTnI 2–4 h after admission were 28% for the I-Stat cTnI and 5% for Minicare cTnI whole blood (p = 0.009). Similar results were obtained in the comparison with Minicare cTnI plasma and Architect hs-cTnI. False negative cTnI results were obtained in the comparison with Minicare cTnI plasma and Architect hs-cTnI and 47% for I-Stat cTnI (p < 0.05). The clinical concordances between the assays relative to delay from onset of symptoms to admission are given in the supplementary file, Tables 2a and b. The concordances between the Minicare cTnI and Architect hs-cTnI were similar to the above results and with lower concordances to I-Stat cTnI when the patients were divided into those admitted < 4 h from onset and those admitted > 4 h from onset of symptoms.

The clinical performances of the Minicare cTnI whole blood and plasma assays were further evaluated in comparison to the two plasma cTnI assays by ROC curve analysis (Table 3, Fig. 2). The AuROC (95%CI) for the Minicare cTnI whole blood was 0.88 (0.83–0.94) at admission and 0.97 (0.95–0.99) 2–4 h after admission. The AuROC for Minicare cTnI plasma was 0.87 (0.81–0.93) and 0.97 (0.95–0.99), respectively. Corresponding figures were 0.91 (0.87–0.95) and 0.97 (0.95–0.99) for the Architect hs-cTnI and 0.88 (0.82–0.94) and 0.96 (0.92–0.99), respectively, for the I-Stat cTnI. AuROCs were not statistically different between the three assays at the two different time points. Significant differences were observed in terms of clinical performance in relation to their respective cut-offs (43 ng/L for Minicare cTnI, 26.2 ng/L for Architect hs-cTnI and 80 ng/L for I-Stat cTnI), as shown in Table 3. Thus, the sensitivities of the Minicare cTnI and Architect hs-cTnI assays were significantly higher at admission or 2–4 h after admission, compared with the sensitivity of I-Stat cTnI (p < 0.0001) whereas the specificity of the I-Stat cTnI assay was higher (p < 0.0001). The NPV at admission was 95% for the Minicare cTnI and Architect hs-cTnI, which was not significantly different from that of the I-Stat cTnI assay (92%). NPV increased to 99% for Minicare cTnI in whole blood, to 98% for Minicare cTnI plasma and to 98% for Architect hs-cTnI at 2–4 h, whereas the NPV of the I-Stat cTnI was lower, 95% (p = 0.002 and p = 0.03, respectively). The corresponding LR- after 2–4 h was lower for the Minicare cTnI whole blood (0.06) than for Architect hs-cTnI (0.11) (p = 0.02) and for I-Stat cTnI (0.28)
Minicare cTnI plasma (0.08) was not different from Architect hs-cTnI, but significantly different from I-Stat cTnI ($p < 0.0001$).

The time between onset of symptoms and admission varied considerably among the patients and it was therefore of interest to relate cTnI concentrations to the duration of symptoms before admission (Supplementary file, Tables 3–6). The majority of patients with MI were admitted within 1–4 h after start of symptoms. Concentrations of cTnI increased in line with the duration of symptoms for all assays (Fig. 3). Non-elevated cTnI levels were seen with either Minicare cTnI or Architect hs-cTnI in 4–8% of patients with MI admitted > 4 h after start of symptoms, in contrast to 21% with I-Stat cTnI. The proportions of patients without a diagnosis of MI who had elevated cTnI concentrations at admission were about 1% for the I-Stat cTnI, 7–8% for the Minicare cTnI and Architect hs-cTnI respectively (Supplementary Fig. 2). Elevation of cTnI in the absence of MI was seen in 10% of patients with a prior history of coronary artery disease. Clinical performance characteristics in relation to onset of symptoms i.e. $< 4$ h or $> 4$ h, are further given in Supplementary Tables 7a and b.

### 4. Discussion

The present study is the first prospective evaluation using fresh whole blood POC assays that demonstrated equal clinical performance with a hs-cTnI laboratory assay for cTnI measurements made at 0 h or 3 h after admission. The clinical performance of the POC Minicare cTnI assay suggests that this assay, whether used as a whole blood assay or as a plasma assay, has comparable clinical performance to the Architect hs-cTnI assay, in terms of diagnosis of MI. This is remarkable in the light of the modest analytical sensitivity of Minicare cTnI, which can measure cTnI in about 5% of a healthy population [7], compared with the Architect hs-cTnI assay which can measure cTnI in 95% of a healthy population.

### Table 2

Clinical concordance of the three assays: numbers of MI diagnoses in relation to assay cut-offs.

<table>
<thead>
<tr>
<th>Test</th>
<th>Admission</th>
<th>2–4 h after admission</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>2 h after admission</td>
<td>2 h after admission</td>
</tr>
<tr>
<td>No. with Minicare cTnI $\leq 43$ ng/L $^a$</td>
<td>16</td>
<td>2</td>
</tr>
<tr>
<td>No. with Minicare cTnI $&gt; 43$ ng/L $^a$</td>
<td>3</td>
<td>43</td>
</tr>
<tr>
<td>No. with I-Stat cTnI $\leq 80$ ng/L $^a$</td>
<td>19</td>
<td>11</td>
</tr>
<tr>
<td>No. with I-Stat cTnI $&gt; 80$ ng/L $^a$</td>
<td>0</td>
<td>34</td>
</tr>
</tbody>
</table>

We also compared the clinical performance of the Minicare cTnI and the I-Stat cTnI, a POC assay used worldwide in the ED setting. The performance of I-Stat cTnI was inferior to the Minicare cTnI when adopting the cut-off given by the manufacturer. Moreover, the Minicare cTnI assay can be used equally with whole blood, capillary blood or plasma [7].

The analytical and clinical sensitivities of the Minicare cTnI appear difficult to reconcile with the reported imprecision of the assay at its clinical cut-off, which is < 20% CV as required in guidelines, and above the 10% CV required for a hs-cTn assay [7]. One explanation could be overestimation of the imprecision, which was determined by the Clinical & Laboratory Standards Institute protocol and not by the

### Table 3
Clinical performance characteristics of the three cTnI assays based on measurements made at admission and 2–4 h after admission.

<table>
<thead>
<tr>
<th>Assay</th>
<th>AuROC</th>
<th>Sensitivity (%)</th>
<th>Specificity (%)</th>
<th>PPV (%)</th>
<th>NPV (%)</th>
<th>LR+</th>
<th>LR-</th>
</tr>
</thead>
<tbody>
<tr>
<td>Minicare TnI, whole blood, cut-off 43 ng/L</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Admission</td>
<td>0.88</td>
<td>(0.83–0.94)</td>
<td>70 (58–81)</td>
<td>93 (89–95)</td>
<td>63 (51–75)</td>
<td>95 (92–97)</td>
<td>10 (6–15)</td>
</tr>
<tr>
<td>2–4 h</td>
<td>0.97</td>
<td>(0.95–0.99)</td>
<td>95 (86–99)</td>
<td>91 (87–94)</td>
<td>68 (57–78)</td>
<td>99 (97–100)</td>
<td>11 (7–17)</td>
</tr>
<tr>
<td>Minicare TnI, Plasma, cut-off 43 ng/L</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Admission</td>
<td>0.87</td>
<td>(0.81–0.93)</td>
<td>73 (61–84)</td>
<td>92 (89–95)</td>
<td>64 (52–74)</td>
<td>95 (92–97)</td>
<td>9 (6–15)</td>
</tr>
<tr>
<td>2–4 h</td>
<td>0.97</td>
<td>(0.95–0.99)</td>
<td>93 (83–98)</td>
<td>90 (86–93)</td>
<td>66 (55–76)</td>
<td>98 (96–100)</td>
<td>10 (6–15)</td>
</tr>
<tr>
<td>Architect® hs-cTnI, Plasma, cut-off 26.2 ng/L</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Admission</td>
<td>0.91</td>
<td>(0.87–0.95)</td>
<td>72 (59–82)</td>
<td>92 (88–94)</td>
<td>61 (49–72)</td>
<td>95 (92–97)</td>
<td>9 (6–13)</td>
</tr>
<tr>
<td>2–4 h</td>
<td>0.97</td>
<td>(0.95–0.99)</td>
<td>90 (79–96)</td>
<td>91 (87–94)</td>
<td>67 (55–77)</td>
<td>98 (95–99)</td>
<td>10 (7–16)</td>
</tr>
<tr>
<td>I-Stat® POC cTnI, cut-off plasma, 80 ng/L</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Admission</td>
<td>0.88</td>
<td>(0.82–0.94)</td>
<td>52 (40–66)***</td>
<td>99 (97–100)***</td>
<td>89 (75–97)***</td>
<td>92 (89–95)</td>
<td>47 (17–128)***</td>
</tr>
<tr>
<td>2–4 h</td>
<td>0.96</td>
<td>(0.92–0.99)</td>
<td>72 (59–83)***</td>
<td>98 (96–99)***</td>
<td>89 (77–96)***</td>
<td>95 (92–97)**</td>
<td>43 (17–106)***</td>
</tr>
</tbody>
</table>

AuROC: area under the receiver operating characteristics curve; PPV, positive predictive value; NPV, negative predictive value; LR+, positive likelihood ratio; LR-, negative likelihood ratio. Figures in parentheses are 95% CI. *-values as compared to Minicare cTnI whole blood; *** = p < 0.001, ** = p < 0.01, * = p < 0.05.

Fig. 2. Receiver operating characteristics curves for diagnosis of acute myocardial infarction with the three assays at admission and at 2–4 h after admission.
construction of an imprecision profile based on duplicates of a large number of patient samples. We argued previously that the latter procedure probably better reflects the actual imprecision of the day-to-day use of the assay [12]. Indeed in recent evaluations, after the completion of the present study, of the imprecision of the Minicare cTnI assay, lower imprecision was reached and reported by the manufacturer (Philips). The unique antibody configuration of the assay, which includes an antibody against troponin C, may also be relevant to the discussion of the sensitivity of the assay, as we showed earlier that the clinical performance of another cTnI assay was dramatically improved by such measures [13]. For example, the Vidas cTnI assay was judged not clinically useful due to its inability to measure cTnI in the blood of any of a population of healthy subjects [14]. Nevertheless, its clinical performance matched contemporary cTnI assays. Thus, the inclusion of an anti-TnC antibody in the Minicare cTnI assay may detect processes in the ischemic myocardium that increase the expression and release of troponin C [15]. Alternatively, TnC antibodies may increase the epitope recognition of cTnI-cTnC complexes in the assay.

Prompt and accurate rule-out of MI for a patient presenting with symptoms suggestive of MI is a major goal in the crowded emergency room. The introduction of laboratory hs-cTn assays supports rule-out of MI within 3 h [3], or as little as 1 h, after admission [16]. The use of laboratory assays often incurs a delay in response time, typically of 1–2 h after blood draw; the availability of a sufficiently sensitive POC cTnI assay could support earlier discharge by providing a rapid cTnI measurement [4]. This has not been achieved to date, apparently due to the seemingly insurmountable challenge of achieving the analytical sensitivity of the hs-cTnI laboratory assays in a POC test. The clinical performance of the Minicare cTnI POC assay therefore seems unique, having achieved a negative predictive value of 99% and clinical diagnostic performance equivalent to one of the most sensitive cTnI laboratory assays currently available. As discussed above, the clinical performance of the Minicare cTnI arose from its clinical specificity, rather than its analytical sensitivity in measuring very low cTnI levels. The Minicare cTnI assay may be the first POC assay suitable for safe rule-out of patients with suspicion of MI 3 h after admission [3]. However, until the clinical performance of the Minicare cTnI assay has been confirmed in further studies, the diagnosis of AMI should be based on the results of this assay only when the results of sensitive laboratory assays are not available.

Most patients (60%) were admitted to the hospital < 4 h after start of symptoms, consistent with the lower negative predictive value at admission. It was interesting that, in the group of patients with MI and longer (> 6 h) duration of symptoms pre-admission all patients had Architect hs-cTnI or Minicare cTnI plasma concentrations above their respective clinical cut-offs except one patient at the cut-off as measured by Minicare cTnI in whole blood. Several patients without an MI diagnosis had elevated cTnI concentrations i.e. “false positives” (7.2%), in a diagnostic sense. As might be expected about 50% of these patients had a history of coronary artery disease and the raised cTnI therefore were consistent with a degree of myocardial ischaemia.

Our study focussed on the clinical performance of the respective assays, in terms of their ability to identify a level of cTnI that would support a diagnosis of MI (i.e. above the 99th percentile cut-off from a healthy population) in patients with an adjudicated diagnosis of MI, independently of the assays discussed here. Current guidelines for the diagnosis of non–ST elevation MI also require observation of dynamic changes (rise and/or fall) in cTnI [1,3]. This aspect of MI diagnosis is beyond the scope of our study, which represents a limitation of the current analysis. However, our observations are directly relevant to the clinical use of these cTnI tests: clinical performance depends on the identification of cTnI levels above the diagnostic cut-off, irrespective of the precise diagnostic protocol used with respect to dynamic changes in cTnI. Other limitations of the study were the moderate size of the cohort and the post-hoc analysis of the prospective trial, which also precluded any conclusions as to gender-specific cut-offs. However, these limitations were counterbalanced by the input of very high quality data of the patients and by the state-of-the art diagnostic process.

We conclude that the novel Minicare cTnI POC assay equals the clinical performance in our patient cohort of a hs-cTnI assay. The diagnostic performance of the assay may arise from the inclusion of a monoclonal antibody directed against troponin C. If confirmed by future clinical studies our data suggest that the clinical performance of the Philips Minicare cTnI assay, with a very high negative predictive value, may become suitable for prompt and safe rule-out of MI for patients with symptoms suggestive of MI in the emergency setting using a 0/3 h sampling protocol.

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Appendix A. Supplementary data

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References
