

TIMI Reliance in a General Emergency Department Chest Pain Unit (TRIAGED CPU)

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ABSTRACT

Objective

Several studies have attempted to demonstrate that the Thrombolysis in Myocardial Infarction (TIMI) risk score has the ability to risk stratify Emergency Department (ED) patients with potential acute coronary syndromes (ACS). Most of the studies we reviewed relied on trained research investigators to determine TIMI risk scores rather than ED Providers functioning in their normal work capacity. We assessed whether TIMI risk scores obtained by ED providers in the setting of a busy ED differed from those obtained by trained research investigators.

Methods

This was an ED-based prospective observational cohort study comparing TIMI scores obtained by 49 ED providers admitting patients to an ED chest pain unit (CPU) to scores generated by a team of trained research investigators. Provider type, patient gender, and TIMI elements were examined for their effects on TIMI risk score discrepancy.

Results

Of the 501 adult patients enrolled in the study, 29% of TIMI risk scores determined by ED providers and trained research investigators were generated using identical TIMI risk score variables. In our low risk population the majority of TIMI risk score differences were small, however 12% of TIMI risk scores differed by two or more points.

Conclusion

TIMI risk scores determined by ED providers in the setting of a busy ED frequently differ from scores generated by trained research investigators who complete them while not under the same pressure of an ED provider. ED providers should not be expected to produce TIMI scores identical to those of trained research investigators.

Keywords: acute coronary syndrome; standard of care; cardiology; TIMI score; chest pain unit

INTRODUCTION

Chest pain is the second most common complaint of patients presenting to emergency departments (ED) in the United States, accounting for approximately seven million visits annually.¹ Early determination of whether a patient's chest pain origin is cardiac versus noncardiac is imperative. Patients diagnosed early with acute coronary diseases (ACS) may benefit from early interventions.²⁻⁶ A missed diagnosis of ACS may result in wrongful discharge, myocardial infarction and sudden death. Despite utilization of ECG results, biomarker assays, patient history and clinical acumen, 0.4-5% of patients with acute myocardial infarction are inadvertently discharged from the ED.⁷⁻¹⁴ Absence of ECG changes, biomarker assays, or history of heart disease does not entirely exclude the diagnosis of Non-ST-elevation ACS.

46 In an effort to improve outcomes in patients with acute coronary syndromes, researchers have
47 developed numerous risk stratification tools.¹⁵⁻⁵⁷ Of all the risk stratification systems developed,
48 the Thrombolysis in Myocardial Infarction (TIMI) risk score is the most studied, supported and
49 utilized.^{3,7,58,59}

50
51 A patient's TIMI risk score is determined by assigning a value of 1 for each of seven equally
52 weighted prognostic variables with the total score determining a patient's risk of adverse cardiac
53 outcome (death, MI, severe recurrent ischemia requiring revascularization) within 14 days of
54 presentation. TIMI risk score variables include a patient's age, presence of known coronary
55 artery stenosis, aspirin use in the past week, frequency of angina episodes, ECG and cardiac
56 marker changes, and risk factors for coronary artery disease (hypertension, diabetes, family
57 history of premature coronary artery disease, elevated cholesterol, and smoking).

58
59 The TIMI risk score was originally derived from a retrospective analysis of a relatively high-risk
60 population of patients with known unstable angina/NSTEMI.¹⁵ In this patient population the
61 TIMI risk score was associated with 4.7% to 40.9% (or greater) risk of adverse cardiac outcome
62 (Figure 1).¹⁵ Following the development of the TIMI risk score tool, several studies were
63 performed validating the tool's ability to stratify risk among patients with cardiac disease.^{16,60-62}

64
65 **Figure 1.** TIMI risk score.
66

TIMI risk score	Risk at 14 days of death, MI, or severe recurrent ischemia requiring urgent revascularization
0-1	4.7 %
2	8.3 %
3	13.2 %
4	19.9 %
5	26.2 %
6-7	at least 40.9 %

67 *(Table adapted from Antman et al.)¹⁵*
68

69 Though not originally designed for ED use, several additional studies have attempted to
70 demonstrate the TIMI risk score's ability to stratify risk among real-world ED populations.<sup>7,17-
71 21,63-68</sup> As a result of these studies, the TIMI risk score tool has made its way into the protocols of
72 Emergency Departments and hospitals around the world, often determining whether a patient is
73 admitted to a hospital, observation unit or discharged home.⁶⁴

74
75 The TIMI risk score is promoted for being simple to remember, easy to determine using data
76 readily available during an acute presentation, and for its ability to be applied early in a clinical
77 course.^{4,64,69} In addition to assisting with triage and disposition decisions, the TIMI risk score
78 improves the exchange of information between ED physicians and cardiologists.^{65,70}

79
80 **Importance**

81 For many reasons, complete and accurate TIMI risk scores can be difficult to obtain when
82 patients present with chest pain to a busy ED. Several studies have demonstrated how
83 interruptions, distractions, and workload affect an ED provider's ability to maintain thought flow

84 and increase the likelihood of errors occurring.⁷¹⁻⁷⁴ Pines et al⁷⁵ suggest that patients presenting
85 to the ED during times of increased ED crowding are at greater risk for adverse cardiovascular
86 outcomes. Inaccurate TIMI risk scores may result in inaccurate risk stratification, as well as
87 ineffectual or inappropriate management of patients with nonspecific chest pain.

88
89 Most studies validating the utility of the TIMI risk score among ED populations utilized trained
90 research investigators or a combination of trained researchers and ED providers to generate TIMI
91 risk scores.^{7,17,18,20,23,63} Trained research investigators do not work under the same time
92 constraints and in the same distracted environment as a working ED Provider. Trained research
93 investigators have the benefit of spending more time interviewing patients, reviewing medical
94 records, scrutinizing ECG patterns, and reviewing their own scores for errors and
95 clarification.^{7,17} Unfortunately, the ED provider does not usually have a trained research
96 investigator at his or her disposal to determine accurate TIMI risk scores. Our review of the
97 literature found very few prospective studies utilizing ED Providers exclusively as assessors for
98 the TIMI risk score. In the select studies where ED providers assessed TIMI risk scores, their
99 scores were not compared against those of trained study investigators for accuracy or
100 validity.^{64,65}

101
102 Current guidelines from the American College of Cardiology, American Heart Association, and
103 National Institute for Health and Clinical Excellence strongly encourage the use of early risk
104 stratification tools such as the TIMI risk score when patients present to health care providers
105 with chest pain.^{2-4,76} In addition, Gallegher et al⁷⁷ suggests the possibility of medicolegal pitfalls
106 by providers not utilizing risk stratifying tools when assessing patients for evidence of ACS. As a
107 result, the TIMI risk score tool is increasingly being used by ED providers as a basis for
108 therapeutic decision-making despite a lack of supporting studies using ED provider-obtained
109 data.

110 111 **Outcomes of Interest**

112 The primary goal of our study was to determine if TIMI risk scores obtained by ED providers in
113 the setting of a busy ED differ substantially from those obtained by trained research investigators
114 who complete them while not under the same pressure of a working ED provider. In addition, we
115 evaluated whether ED provider type or patient gender had any effect on TIMI risk score
116 discrepancy, which aspects of the TIMI risk score most frequently differ between assessors, and
117 whether lower TIMI risk scores (i.e., 0-3) or higher TIMI risk scores (i.e., >3) more frequently
118 match research investigator scores.

119
120 This is the first study we are aware of that evaluates how closely TIMI risk scores generated by
121 ED Providers obtained in the normal course of their work match those obtained by trained
122 research investigators, specifically when applied to patients admitted to a hospital's chest pain
123 unit (CPU).

124 125 **METHODS**

126 **Study Design**

127 This was a prospective observational cohort study comparing TIMI scores obtained by ED
128 providers admitting patients to the CPU at Lakeland Regional Medical Center (LRMC) to scores
129 generated by trained research investigators. The Lakeland Healthcare Institutional Review Board

130 approved the study without need for written informed consent because the data collected was
131 normal data already being obtained and charted during the normal course of an ED provider's
132 work, and could be collected anonymously.

133

134 **Study Setting and Population**

135 LRMC is an academic-based community hospital with an annual ED census of approximately
136 50,000 patients. The hospital's 6 bed CPU opened in 2010 and is situated adjacent to the ED.
137 The CPU is under the direct supervision of ED providers. All ED providers admitting patients to
138 the CPU from October 27, 2012 until July 28, 2013 were included in the study. Participating ED
139 providers included 18 Attending Physicians, 21 Resident Physicians and 10 Midlevel Providers
140 (Physicians Assistants and Nurse Practitioners). No ED providers were excluded from the study.
141 Patient inclusion criteria included all comers presenting to the ED with non-traumatic chest pain
142 suggestive of ACS who were admitted to our hospital's CPU, irrespective of age. At our
143 institution, ED providers independently determine who is to be placed in the CPU. TIMI risk
144 scores are not typically used in the decision to place patients in the CPU. Patient exclusion
145 criteria for study enrollment mirrored CPU exclusion criteria as set by the hospital's Chest Pain
146 Center Door-to-Balloon Committee in accordance with recommendations from the Society of
147 Cardiovascular Patient Care (Figure 2). The CPU is open 24 hours a day, 7 days a week and on
148 holidays, with research investigators available 24 hours a day to enroll patients.

149

150 **Figure 2.** LRMC Chest Pain Unit exclusion criteria.

151

- Patients with ST-elevation acute myocardial infarction (STEMI)
- Positive Cardiac Biomarkers suggestive of myocardial injury
- ECG changes
- Unrelenting chest pain
- Coronary Revascularization in the last 60 days
- Abnormal vital signs
- New dysrhythmia (any run of ventricular dysrhythmia is not a candidate for the CPU)
- Aortic dissection
- Pneumothorax
- Pneumonia
- Esophageal rupture
- Pulmonary Embolism
- Pericardial tamponade
- CHF
- Uncontrolled diabetes
- Electrolyte abnormalities that cannot be cared for with PO electrolyte replacement
- Psychiatrically unstable
- Patients unable to perform activities of daily living
- Pleural effusions
- Renal failure requiring dialysis during their time in the CPU
- Any diagnosis meeting admission criteria

152

153

154 **Study Protocol**

155 Research investigators consisted primarily of nurses already trained to care for CPU patients.
156 Prior to data collection, these research investigators received additional training on how to obtain
157 TIMI risk scores. Their standardized training involved handouts, Microsoft Office PowerPoint
158 presentations, and one-on-one training with clarification to ensure unambiguous collection of
159 data. Research investigators were instructed to use all resources available to them including a
160 patient's hospital record, accessible outside records, labs, prior cardiac catheterization reports,
161 cardiology notes, and patient reported responses. Research investigators routinely evaluated the
162 patient and assessed TIMI risk score variables within 24 hrs of a patient's presentation to the ED
163 (Figure 3). In situations where patients were unaware or unable to answer questions concerning
164 pertinent medical history (for example, an adopted patient unaware of his or her family history),
165 patients were not given any points for those variables.

166
167 Our goal for the research investigator was not to obtain 100% infallible TIMI scores, but rather
168 to generate scores as close as possible to scores assigned by research investigators performing
169 similar TIMI risk score validation studies.

170
171 Separately, ED providers assigned TIMI risk scores to all patients admitted to the CPU at the
172 time of CPU admission per hospital protocol and the ED provider's normal routine (typically
173 following the results of initial ECG and biomarker tests). No additional TIMI training or
174 education was provided to ED providers prior to data collection. ED provider TIMI scores were
175 recorded electronically in the patient's EpicCare electronic health record in a location research
176 investigators were told not to access. In addition, research investigators confirmed blindness by
177 recording whether or not they had prior knowledge of the ED provider's TIMI risk score for each
178 patient. Research investigator data was hand written on a standardized data collection form and
179 placed in a secure folder in the CPU area inaccessible to ED providers. In this way, research
180 investigators and ED Providers were blinded to one other's TIMI risk scores throughout the
181 study.

182
183 **Figure 3.** Variables assessed by research investigators.

184

- Age
- Presence of known coronary artery stenosis $\geq 50\%$ *
 - Prior cardiac catheterization with known disease
 - Prior MI, CABG, angioplasty, or stent
- Aspirin use in the preceding 7 days
- At least 2 episodes of severe chest pain within last 24 hrs
- ST changes $\geq 0.5\text{mm}$ on admission ECG
- Initial serum cardiac biomarker elevation (Troponin I above normal range)
- At least 3 of the following risk factors for Coronary Artery Disease (CAD):
 - High blood pressure ($\geq 140/90$ or on antihypertensive medicine)
 - Diabetes, prediabetes, or hyperglycemia
 - Family history of premature CAD or MI (CAD in male 1st-degree relative, or father < 55 , or female 1st-degree relative or mother < 65)
 - Elevated LDL (≥ 100), reduced HDL (≤ 40 for men, < 50 for women), elevated triglycerides (≥ 150)
 - Smoking in the past 5 years**
- ED provider type (Attending Physician, Resident Physician or Midlevel Provider)
- Confirmation of blinding to ED provider TIMI risk score

185
186 * Similar to Pollack et al⁶⁵, this parameter was expanded in our study because actual cardiac
187 catheterization reports were not always available in the ED.

188 ** 5 years was chosen as a cut-off because risk associated with smoking has been found to
189 diminish after 5 years.⁷⁸⁻⁸⁰

190
191 **Data Analysis**

192 Upon completion, the pertinent data was extracted from patient charts and data collection forms
193 and entered into a database using Microsoft Office Access 2007. The data was then exported into
194 a Microsoft Office Excel 2007 spreadsheet. We used SPSS software to make comparisons of
195 TIMI risk scores obtained by research investigators and ED providers. Where significance
196 testing was reported, variables were analyzed using the Pearson Chi-Square test.

197
198 **RESULTS**

199 The patient population consisted of 543 patients who presented to the ED with symptoms
200 suspicious for cardiac chest pain and were admitted to the CPU. Research investigators provided
201 all variables used to form the TIMI risk score for 543 patients. ED providers provided the
202 necessary variables for 501 patients. Because some ED providers did not record TIMI scores for
203 every patient, we only have complete data for 501 patients. Of these 501 patients, 277 were
204 female and 224 were male. The mean age of the patient study population was 59 (ages 18 to 94),
205 median age 57.

206
207 Though the frequency distributions for research investigators and ED providers were similar, the
208 two scores often did not match for a given patient (Table 1). In fact, of the 501 patients in the
209 study with complete data, ED provider and researcher TIMI risk scores matched for only 213
210 patients (42.5%). Of the 213 patients with the same TIMI scores, only 147 scores were
211 determined using identical TIMI variables. For example, one patient was given a TIMI score of 1
212 by both the research investigator and ED provider. On further analysis, however, the research

213 investigator gave a point for aspirin use over the past 7 days, while the ED provider gave a point
 214 for having 3 or more risk factors for CAD.

215
 216 **Table 1.** Research investigator and ED provider TIMI scores.
 217

TIMI Score	Researcher <i>n</i>	ED provider <i>n</i>	ED provider score matches researcher score
0	96	99	54 (56.3%)
1	130	121	48 (36.9%)
2	92	109	34 (37.0%)
3	89	88	33 (37.1%)
4	71	70	38 (53.5%)
5	22	12	5 (22.7%)
6	1	2	1 (100%)
7	0	0	0 (100%)
Total Patients	501	501	213 (42.5%)

218
 219 Further breakdown of TIMI scores revealed that scores differed by 1 point for 228 patients
 220 (45.5%), 2 points for 52 patients (10.4%), and 3 points for 8 patients (1.6%). No scores varied by
 221 more than 3 points (Table 2).

222
 223 **Table 2.** Discrepancy between ED Provider and Researcher TIMI scores.
 224

Range of TIMI Discrepancy	<i>n</i>	% of Total scores
-4	0	0
-3	4	0.8
-2	27	5.4
-1	125	25.0
0 (Matching)	213	42.5
+1	103	20.6
+2	25	5.0
+3	4	0.8
+4	0	0
Total	501	100

225
 226 Table 3 shows the incidence of TIMI variables as reported by research investigator and ED
 227 provider. The frequencies of several variables were similar, such as “Age ≥65”, “Aspirin use”,
 228 “ECG changes”, and “Elevated Troponin”. Research investigators reported a greater incidence
 229 of “Known CAD” and “Angina”, while ED providers reported a greater prevalence of “CAD
 230 Risk Factors”.

231
 232 **Table 3.** Incidence of TIMI variables.
 233

	Researcher <i>n</i> (%)	ED provider <i>n</i> (%)
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Age ≥65	166 (33.1%)	167 (33.3%)
Known CAD	149 (29.7%)	118 (23.6%)
ASA use	239 (47.7%)	254 (50.7%)
Angina	207 (41.3%)	126 (25.1%)
ECG changes	9 (1.8%)	7 (1.4%)
Elevated Trop	21 (4.2%)	10 (2.0%)
CAD risk factors	190 (37.9%)	274 (54.7%)

234
235 Our analysis shows that salient disagreements in TIMI variables exist between ED providers and
236 research investigators. For example, ED providers reported the incidence “Angina” in only 59 of
237 207 patients (28.5%) determined by research investigators to have had “Angina”. Additionally,
238 ED providers reported “Angina” as being present in 67 patients not reported by research
239 investigators. Table 4 shows how often research investigators and ED providers agreed on
240 reported variables, with their relative significance.

241
242 **Table 4.** TIMI variable agreement (ED provider variable matched research investigator variable
243 for the same patient).

	Positive <i>n</i> <i>(ED/R)</i>	Negative <i>n</i> <i>(ED/R)</i>	<i>p</i> value
Age ≥65	166/166 (100%)	334/335 (99.7%)	0.000
Known CAD	104/149 (69.8%)	338/352 (96.0%)	0.000
ASA use	181/239 (75.7%)	189/262 (72.1%)	0.000
Angina	59/207 (28.5%)	227/294 (77.2%)	0.147
ECG Changes	2/9 (22.2%)	487/492 (99.0%)	0.000
Elevated Trop	7/21 (33.3%)	477/480 (99.4%)	0.000
CAD Risk Factors	173/190 (91.1%)	210/311 (67.5%)	0.000

245 *(ED= ED provider, R=Research Investigator)*

246
247 Additional analysis was performed based on ED provider type assessing the TIMI score
248 (Attending Physician, Resident Physician or Midlevel Provider). Attending Physicians
249 determined the scores for 183 patients, Resident Physicians scored 225 patients, and Midlevel
250 Providers scored 93 patients. Overall TIMI risk score determinations were similar across all
251 provider types. TIMI scores matched 43.2% of researcher scores for Attending Physicians,
252 42.7% for Resident Physicians, and 40.9% for Midlevel Providers (Table 5). When discrepancies
253 occurred, Attending Physicians and Midlevel Providers reported slightly lower TIMI scores,
254 while Resident Physicians reported slightly higher TIMI scores (Figure 4).

255
256 **Table 5.** Range of TIMI score discrepancy from research investigator by ED provider type.

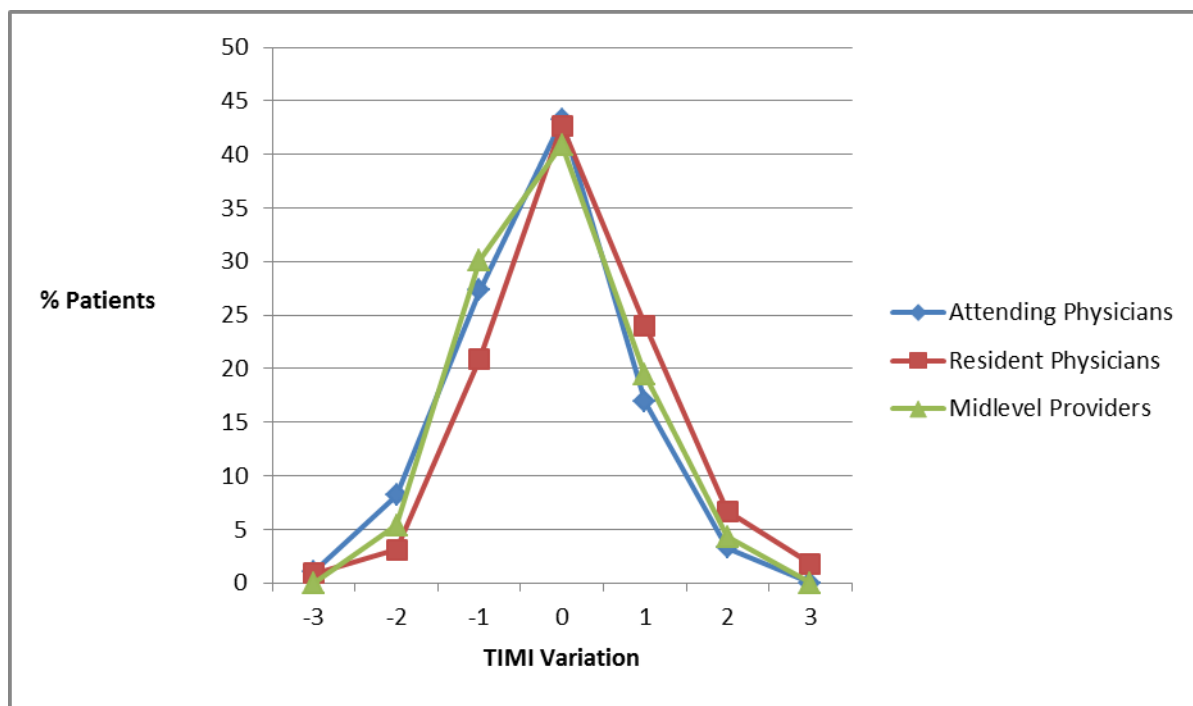
Range of TIMI Discrepancy	Attending Physician <i>n</i>	Resident Physician <i>n</i>	Midlevel Provider <i>n</i>
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257

from Researcher Score			
-3	2 (1.1%)	2 (0.9%)	0 (0.0%)
-2	15 (8.2%)	7 (3.1%)	5 (5.4%)
-1	50 (27.3%)	47 (20.9%)	28 (30.1%)
0 (Matching)	79 (43.2%)	96 (42.7%)	38 (40.9%)
+1	31 (16.9%)	54 (24.0%)	18 (19.4%)
+2	6 (3.3%)	15 (6.7%)	4 (4.3%)
+3	0 (0.0%)	4 (1.8%)	0 (0.0%)
Total Patients Scored	183	225	93

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260

Figure 4. Range of TIMI score discrepancy from research investigator by ED provider type.



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Further analysis shows that gender has little effect on TIMI score differences. ED provider scores agreed with research investigator scores for 112/277 female patients (40.4%) and for 103/224 male patients (46.0%).

Because the CPU at our institution is utilized to screen a population of patients at low-risk for ACS, far more low TIMI scores (TIMI 0-3) were generated. Based on the scores obtained by research investigators, 407 patients presenting to the CPU had TIMI scores 0-3, while only 94 had TIMI scores >3. There was no difference in the frequency of ED provider scores matching researcher scores on the basis of the number of variables involved (Table 6).

Table 6. TIMI risk score divergence by range.

TIMI risk score Range	Researcher n	ED provider matches researcher TIMI score	Matching TIMI score with identical variables
0 to 3	407	169 (41.5%)	116 (28.5%)
4 to 6	94	44 (46.8%)	31 (33.0%)
Total	501	213	147

275

276 **DISCUSSION**

277 This study demonstrated that a majority of TIMI scores as determined by ED providers in the
 278 setting of a busy ED differ from scores generated by trained research investigators who complete
 279 them while not under the same pressure of an ED provider. In our study only 29.3% of TIMI
 280 scores were calculated using identical TIMI risk score variables. The majority of TIMI risk score
 281 differences were either negligible (same TIMI risk score obtained despite differing TIMI
 282 variables used) or diverged by no more than 1 point in our low risk patient population; however,
 283 12% of patient scores differed by two or more points.

284

285 We have shown that ED provider type has little effect on the likelihood of TIMI risk scores
 286 matching TIMI scores obtained by trained research investigators. Neither the patient gender nor
 287 the quantity of positive variables had a significant effect on TIMI risk score differences.

288

289 Patient age was the variable most agreed upon by TIMI risk score assessors with only one
 290 instance of an ED Provider incorrectly giving a point to a 57 y/o for being ≥ 65 y/o. TIMI
 291 variables requiring more active investigation showed greater variation. Researchers reported
 292 greater incidence of known CAD, possibly due to having more time available to review patient
 293 records and interview the patient. ED providers were apt to report a greater incidence of ≥ 3 CAD
 294 risk factors. Confirmation bias (or myside bias) is one potential reason for this. For example, in
 295 ascertaining the presence of multiple CAD risk factors (a time consuming task), an ED provider
 296 might assume that when one or two risk factors are present, such as smoking and hypertension,
 297 other risk factors are likely present as well. Unfortunately, the TIMI risk score recorded in the
 298 electronic medical record by our ED providers simply shows when ≥ 3 CAD risk factors are
 299 present and does not further categorize which CAD risk factors were recognized by the ED
 300 provider.

301

302 ECG changes and biomarker elevations were seldom present in our study, likely reflecting the
 303 low-risk nature of our CPU study population. Research investigators reported a few more
 304 instances of ECG and biomarker changes than were reported by ED providers, though not
 305 statistically significant ($p = 0.000$).

306

307 Both ED providers and research investigators reported similar numbers of aspirin users among
 308 our population, however only 75.7% of these patients matched. 73 patients recognized by ED
 309 providers as having taken aspirin went unrecognized by our research investigators. Likewise,
 310 research investigators reported an additional 58 patients who ED providers said had not taken
 311 aspirin. Similar to aspirin, there was a discrepancy in the reporting of angina episodes.
 312 Researchers, who had the benefit of spending more time with patients, reported far more
 313 occurrences of angina than ED providers (207 to 126 occurrences). ED providers only

314 recognized 59 of the 207 patients (28.5%) designated as having had angina by research
315 investigators. Interestingly, ED providers reported angina as being present in 67 patients who
316 research investigators did not feel met criteria for angina.

317
318 There are many barriers to obtaining accurate histories from patients.⁸¹⁻⁸³ Patients who present to
319 the ED in chest pain often do so under great duress, likely compounding the already difficult
320 job of extracting accurate history. Studies have shown that patients in stressful situations have
321 impairments in cognition, memory and verbal recall.⁸⁴⁻⁸⁵ Many clinicians recognize the
322 phenomenon of the contradictory account, where the second person to interview a patient obtains
323 an entirely different story. Perhaps in recognition of this, Hess et al¹⁷ excluded patients with
324 unreliable history from his prospective study on TIMI score validity in the ED. The variability of
325 patient reported responses in the ED suggests a need for risk stratification tools which place
326 greater weight on objective variables which can be assessed independent of interviews with the
327 patient.

328
329 Many ED providers support the idea of utilizing a clinical prediction rule for the identification of
330 ACS among patients with chest discomfort in hopes of offering early discharge to low risk
331 patients.⁸⁶ A few recent studies have suggested that a rapid TIMI risk score protocol can be
332 employed to safely discharge low risk ED patients with chest discomfort home from the
333 ED.^{22,23,86} Though the TIMI risk score device has the potential to stratify risk among ED
334 populations, our study suggests that it may depend on how and by whom the TIMI risk score
335 data is obtained. It is important that these studies, as well as any study suggesting validity and
336 broad applicability of a risk stratification tool for regular use in the ED, be examined closely to
337 determine if the working data was obtained by ED providers while working in their normal
338 environment. We commend validation studies such as Chase et al⁶⁴ and Pollack et al⁶⁵ for using
339 ED providers to determine risk scores and call for more similar studies. We also question the
340 applicability of studies which rely on data largely obtained by trained research investigators in
341 place of ED providers.

342 343 **LIMITATIONS**

344 Some researchers have suggested that ECG and biomarker indices should carry greater weight in
345 risk stratification scores.^{17,40} Modified TIMI risk scoring tools have been developed which assign
346 more points to ECG and biomarker variables.^{17,40} Because so few ECG and biomarker changes
347 were present in our study it is difficult to make generalizations on the ED provider's ability to
348 recognize and assign a proper TIMI risk score for those variables. Though not significant, the
349 few ECG and biomarker changes recognized in our study were slightly underreported by ED
350 providers, which may reflect a degree of selection bias or simply differences in interpretation. It
351 is possible that ED providers under-report some aspects of the TIMI risk score (such as angina,
352 ECG and biomarker changes) since they have already deemed a patient low risk and not likely
353 suffering from true ACS by virtue of placing the patient in the CPU. In addition, ED providers
354 may be less likely than research investigators to report a Troponin I level at the very edge of the
355 cutoff as "positive", especially in a patient with known chronic renal insufficiency, for example.

356
357 We asked our research investigators to obtain scores within 24 hours of patient presentation. This
358 was done in order to improve the likelihood of obtaining complete data for the majority of

359 patients. We recognize that research investigators in other studies may have had additional time
360 to perform their investigations.

361
362 While CPU nurses are capable and trainable, most CPU nurses have minimal experience
363 participating in research and may not have performed to the same standard as professional
364 research investigators. Research investigator TIMI risk score ECG interpretation was performed
365 by our trained research investigators and not physicians well-versed in ECG interpretation.

366
367 Many TIMI risk score validation studies include a patient cut-off age for enrollment, such as ≥ 30
368 years old.¹⁵ Because we were performing a comparison of risk scores and not examining patient
369 outcomes, we did not feel that excluding patients by age was necessary.

370
371 Although pertinent patient history was occasionally obtained directly from a patient's
372 cardiologist by phone or when visiting the CPU, we did not routinely obtain data in this manner.
373 Most data was acquired using information readily available to the research investigator in the
374 CPU setting, which is similar to what is available to the ED provider. Midway through the
375 project some cardiologists released online access to their outpatient clinical electronic medical
376 records including catheterization lab reports, providing additional means of data acquisition to
377 researcher investigators. Prior to obtaining access to these records, data in question could
378 sometimes be obtained via fax or telephone during regular business hours.

379
380 As mentioned earlier, our researchers were not focused on obtaining infallible data. Where data
381 was unknown and could not easily be produced we gave no points for those variables.

382
383 Patient demographics may have also contributed to some study variation. Though predominantly
384 English-speaking, our geographic area does contain some non-English speaking individuals
385 which could have impeded an assessor's ability to obtain a reliable history.

386
387 Our study examined a specific cohort of low risk patients presenting to the ED with chest pain.
388 CPU patients do not make up the entirety of patients presenting to the ED complaining of chest
389 pain. Many times high risk patients with ACS are admitted directly to the hospital or cath lab,
390 and patients with noncardiac etiologies of chest pain (such as trauma or rash) are discharged
391 home. The results of our study may not be generalizable to all populations of patients presenting
392 with chest pain to the ED, however there clearly exists a discordance of TIMI risk scores
393 between ED providers and trained research investigators.

394 395 **CONCLUSION**

396 Several studies and guidelines have been published suggesting that TIMI scores obtained in ED
397 populations are valid.^{2-4,7,63,65,66,68,76} Our study demonstrates that there is discordance between
398 TIMI scores generated by trained research investigators and busy ED providers. Our study
399 questions the reliability, validity, and applicability of TIMI risk score validation studies where
400 scores were ascertained predominantly by trained research investigators.

401
402 ED providers should not be expected to produce TIMI scores identical to those of trained
403 research investigators and until more validation studies are available, should continue to use
404 sound clinical judgment in patients presenting to the ED with evidence of ACS.

405
406 Areas for future research may include comparing time spent by ED providers and research
407 assistants determining risk stratification scores, reliability of patient reported history in an ED
408 environment, difficulties associated with access to outside medical records, effects of ED
409 crowding and ED provider staffing on job efficiency, accuracy, and capacity for risk
410 stratification, further risk tool validation studies using ED provider-obtained data, and studies
411 evaluating all patients presenting to the ED with chest pain, not just CPU patients.

412
413 **Competing Interests**

414 None to declare.

415
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422 **Contributors**

423 BT and MM were responsible for conception and design of this study. BT conducted the
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425 data, trained research investigators, and wrote the manuscript. MM supervised the project,
426 reviewed the manuscript and approved the final version submitted for publication.

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