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## Accuracy of emergency medical services (EMS) telephone triage in identifying acute coronary syndrome (ACS) for chest pain patients: A systematic literature review

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# Accuracy of emergency medical services (EMS) telephone triage in identifying acute coronary syndrome (ACS) for chest pain patients: A systematic literature review

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#### Abstract;

#### **Objective:**

To systematically appraise the available evidence to determine the diagnostic accuracy of the

EMS telephone triage for patients with chest pain suspected to be caused by ACS or life-

threatening conditions.

#### **Design:**

Systematic review.

#### **Data sources:**

Electronic searches were performed in Embase, Medline, and Cinahl databases for relevant papers.

#### **Eligible criteria:**

The review included all types of original studies with adult patients (>18 years with a primary complaint of chest pain who called EMS for an ambulance and evaluated dispatch triage priority, and linked to final diagnosis of ACS, acute myocardial infarction (AMI), or other life-threatening conditions.

#### Data extraction and synthesis:

standardised data extraction form was used to extract and collect data as study design, population, study period, and outcome. Also, all data required were extracted for diagnostic accuracy assessment including 2x2 tables when available for dispatch priority and prediction models priority for suspected ACS, or life-threatening conditions. Two authors independently extracted the data. Risk of bias was assessed using (QUDAS-2) assessment tool.

#### **Result:**

In total, 553 studies were identified from the literature search and cross-referencing. After excluding 550 studies, three were eligible for inclusion. Among those 3 studies, there are different prediction models developed by authors with variation in variables used to detect ACS. The overall result showed that dispatch triage tools have good sensitivity to detect ACS and life-threatening conditions even though they are used to triage sign and symptoms rather than diagnosing the patients. On the other hand, prediction models were built to detect ACS and life-threatening conditions and therefore it showed better sensitivity and NPV.

#### **Conclusion:**

EMS dispatch systems accuracy for ACS and life-threatening conditions associated with chest pain is good. Since the dispatch tools were built to triage ambulance response priority based on sign and symptoms, this lead to over triage among non life-threatining chest pain patients. Over triage were slightly reduced by deriving different prediction models and showed better sensitivity.

#### Strength and limitations:

- Up to our knowledge, this is the first systematic review looking into the accuracy of pre-hospital chest pain telephone triage.
- It is possible that some relevant papers were missed because we only included studies that have adult participants who are >18 years old and we excluded papers not published in English language.
- Unfortunately, we couldn't conduct a meta-analysis as there was variations between dispatch system tools and priority types.
- Also, there was a difference in the definition of life-threatening conditions definition between included studies.

#### **Abbreviations list:**

ED: Emergency Department; ACS : Acute Coronary Syndrome; AMI: Acute Myocardial Infarction; ECG: Electrocardiogram; NPV: Negative Productive Value; QUADAS: Quality Assessment of Diagnostic Accuracy Studies; PPV: Positive Predictive Value;

E.

STEMI: ST-Elevation Myocardial Infarction; ACS: Acute Coronary Syndrome;

## **Keywords:**

Pre-hospital, Chest pain, telephone triage, ACS

#### **Background:**

Chest pain, which is one of the main symptoms of ACS, is one of the most common reasons for ambulance callouts and presentations to EDs (1, 2). Patients with ACS typically present with chest pain. However, the majority patients with chest pain transported by ambulance are ultimately diagnosed with self-limiting, non-cardiac disease. Thus, chest pain leads to over-triage patients and increases ambulance resource consumption, and crowed EDs (2, 3).

Two EMS dispatch systems are currently used in the UK: the Advanced Medical Prioritizing Dispatching System (AMPDS), and the National Health Service (NHS 999) System (4). Both systems use computerized telephone triage software that identify life-threatening conditions by a series of questions at the beginning of the call. For patients with a primary complaint of chest pain, they also include a specific series of questions to identify ACS and triage the case using the patient's sign and symptoms (3, 5). However, systematic reviews have shown that the accuracy of medical triage systems are backed by a low level of evidence (6, 7). There is also insufficient data on the dispatch protocol efficiency (7, 8). There is no consensus on the definitions used in Criteria Based Dispatch (CBD), which is program used to patients by signs and symptoms, and AMPDS (9, 10). Also, there is no consensus on the accepted level for over-triage, under-triage for medical emergency dispatch (6, 9). Therefore, systematic over-triage for chest pain patients is used as safe method by dispatch system to avoid potential harm to patients. As a result, it caused an over use of EMS resources (6) and showed difference between clinical findings and dispatch priority (8, 9). Therefore, with the current clinical guidelines for emergency medical services (EMS), it is hard to rule in or rule out ACS by telephone-triage (11).

We aimed systematically to appraise the available evidence to determine the diagnostic accuracy of the EMS telephone triage for patients with chest pain suspected to be caused by ACS or life-threatening conditions.

#### **Methods:**

The systematic review was conducted in accordance with PRISMA guidelines and following Cochrane methodology for diagnostic test accuracy reviews (12). PRISMA checklist provided in supplementary appendix. This systematic review was pre-registered on the PROSPERO database (reference CRD42020171184)

#### Search strategy and eligibility criteria:

We searched the Embase, Medline, and Cinahl databases on 3/03/2020 for the terms "chest pain, telephone or dispatch, triage, ACS or AMI, life-threatening conditions, and EMS or prehospital". Retrospective and prospective cohort studies written in English and investigating EMS telephone triage for chest pain patients linked with final diagnosis of ACS, or life-threatening conditions were eligible.

#### Studies included:

Based on the inclusion criteria, titles and abstracts were independently screened and shortlisted by two reviewers (AbA and ChR). The inclusion criteria were: (1) the study included adult patients (>18 years); (2) the study included patients with a primary complaint of chest pain who called EMS for an ambulance; (3) the studied evaluated dispatch triage priority; (4) the study provided data for a linked final diagnosis of ACS, acute myocardial

infarction (AMI), or other life-threatening conditions. Two reviewers independently reviewed and screened potentially relevant full text papers to identify those fulfilling the inclusion criteria. Any discrepancies between the reviewers were resolved by a third reviewer (AhA).

#### **Outcome measures:**

The primary outcome was a diagnosis of ACS, including ST elevation myocardial infarction (STEMI), non-ST elevation myocardial infarction (NSTEMI) and unstable angina. AMI (incorporating both STEMI and NSTEMI) is defined according to the universal definition of AMI (13). Unstable Angina is defined as sudden prolonged chest pain at rest >20 minutes, new onset severe angina, or previous angina that is increasing in severity of pain, duration of pain, and/or frequency (14).

The secondary outcome was the diagnosis of a life-threatening condition associated with chest pain. Since there is no universal definition for life-threatening conditions associated with chest pain, we included all relevant data from the studies, regardless of the precise definition used for 'life-threatening conditions'.

#### Methodological quality assessment:

The quality assessment of eligible articles was independently assessed by two reviewers (AhA and AbA), using modified Quality Assessment of Diagnostic Accuracy Studies (QUADAS-2) tool to suit the purpose of this systematic review (15). Discrepancies between reviewers were solved by discussion.

#### Data extraction:

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After selecting the eligible studies, two investigators (AhA and AbA) used a standardised data extraction form to extract and collect data as study design, population, study period, and outcome. Also, both investigators extracted all data required for diagnostic accuracy assessment including 2x2 tables when available for dispatch priority and prediction models priority for suspected ACS, or life-threatening conditions.

# Statistical analysis:

In line with our protocol, we considered pooling sensitivities, specificities, negative and positive predictive values (NPV and PPV, respectively) by meta-analysis. However, after reviewing the final list of studies meeting inclusion criteria, the reviewers unanimously concluded that meta-analysis would have been inappropriate due to clear evidence of clinical heterogeneity between studies, and missing data. We therefore undertook a narrative synthesis of the existing evidence.

#### **Results:**

In total, our literature search identified 553 potentially relevant articles of which 26 articles were considered eligible for inclusion after titles and abstract screening. After independent full text review, 23 articles were excluded and 3 articles met the inclusion criteria for this review (Figure 1). Excluded studies with reason of exclusion is shown in the supplementary appendix. (16-18).

#### **Study Characteristics and Quality Assessments:**

General study characteristics are summarized in Table 1. Patient characteristics and study weaknesses are summarised in Table 2. The details of the prediction models identified, together with the test characteristics (including sensitivity, specificity, NPV, and PPV) for the diagnosis of ACS and life-threatening conditions are shown in Tables 3 and 4. The modified QUADAS-2 methodological quality assessment tool was used to assess all eligible studies as shown in the supplementary appendix 2. A summary of the quality assessment for each eligible study is shown in Figure 2. Two out of the three studies were retrospective cohort studies and compared the dispatch triage for ambulance response priority to the prediction model (16, 17) and one was prospective cohort study that only testing prediction model accuracy (18). ACS was adjudicated by experts in only one study (18) while the other two studies used the hospital final diagnosis of ACS, AMI, or life-threatening conditions. All eligible studies utilized the appropriate ACS or AMI definitions at the time of ieu conducting the study.

#### Summary of the existing evidence

Gellerstedt et al derived a prediction model to detect AMI and life-threatening conditions patients calling for an emergency ambulance with chest pain. They aimed to effectively triage calls to determine the appropriate level of response (capable of providing advanced life support vs only basic life support). The final model used eight variables (age, gender, strong pain, dyspnoea, cold sweat, nausea, vertigo, and syncope) to calculate the probability of AMI or a life-threatening condition. Setting an arbitrary cut-off, the authors found that the model had greater sensitivity than and similar specificity to the judgement of the call handlers, which could potentially improve triage. However, performance of the model was only evaluated in the derivation set, with no validation presented (16).

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In 2006, the same group derived a new prediction model in another study. They aimed to determine whether a computer-based decision support system could have superior accuracy for identifying life threating conditions or ACS in patients who called for an emergency ambulance with chest pain. Gellerstedt et al collected data from patients who called 112 complaining of chest pain using ten questions related to ACS with yes or no answers (17). They derived two prediction models based on those answers. The first prediction model (full model) incorporated data from all ten questions that dispatchers had asked, including: age, gender, central chest pain, high intensity pain, pain duration between 15 minutes and 24 hours, aggravated pain when breathing or moving, abnormal breathing, cold sweat, diabetes or previous CVD, previous ACS, and belief that symptoms are heart-related. The authors also derived a 'limited model', which only included variables that had a significant association with the presence of ACS or a life-threatening condition on bivariate analysis. These included: gender, age, central chest pain, high intensity pain, previous ACS, and a high priority based on dispatcher judgement. There was no significant difference in performance between the full model and limited model. However, both the limited and full models had superior sensitivity to the priority assigned by dispatchers in practice, both for ACS (Table 3) and life-threatening conditions (Table 4), with similar specificity (17). The model was derived in 70% of the available cohort and validated in the remaining 30%. The authors state that performance was 'similar' in the validation set, although no further details were provided.

*Reuter et al* used backward logistic regression to derive two separate prediction models to detect ACS among men and women, respectively (18). The male prediction model to detect ACS consisted of 8 variables, which are: age, smoking, severe pain, permanent pain,

breathing non-related pain, retrosternal pain, radiating pain, and additional symptoms. The female prediction model to detect ACS consisted of four variables, which are: age >60 years, history of coronary artery disease, breathing non-related pain, and radiating pain.

The male prediction model had an area under the receiver operating characteristic curve (AUC) of 0.76 (95% CI 0.73 – 0.79) in the derivation set (70% of the sample, which was randomly selected), which was maintained at 0.76 (95% CI 0.73 – 0.80) in the internal validation set (30% of the cohort). However, while the female prediction model had a similar AUC in the derivation set (0.79, 95% CI 0.75 – 0.83), it had lower accuracy in the validation set (AUC 0.67, 95% CI 0.60 – 0.74). No data were presented to enable calculation of sensitivity, specificity, PPV or NPV, and the output of the model was not compared to standard care or clinical judgement.

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#### **Discussion:**

In this systematic review, we have identified three studies that presented the derivation of clinical prediction models to enhance the telephone triage of patients calling for an emergency ambulance with chest pain. None of the models presented had sufficiently high sensitivity to avoid the need for ambulance dispatch. The model derived by Gellerstedt *et al* (2006) used an outdated definition of AMI (using creatine kinase-MB as the reference standard biomarker), meaning that the results could not be applied to modern clinical practice. The second model derived by Gellerstedt *et al* (2016) underwent internal validation, but full details that validation were not published. Finally, Reuter *et al* derived two prediction models (for men and women, respectively) but did not present sufficient data to compare their performance to standard care. The prediction model for females had a poor AUC (0.67)

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on internal validation. We identified no prediction models that have undergone external validation.

It is an extremely challenging task for dispatchers to identify ACS over the telephone as there are so many different causes of chest pain (11, 19, 20), and there is very limited information available at the time of the initial phone call. It has previously been reported that only 1 out of 18 patients who request an ambulance for a complaint of chest pain has ACS(3). While a validated prediction model would have clear benefits, this systematic review has identified no validated tools that could be used in current practice.

It has also been reported that the AMPDS triage system is not an accurate tool for identifying ACS patients (3). Moreover, in Denmark, Criteria-based Danish index is used to triage emergency calls and it resulted in over triage for 51% of chest pain calls as they were discharged without specific diagnosis and AMI were only found in 11% of chest pain calls (2). Dispatch triage systems were constructed to prioritize patient's condition and ambulance response grade and it were not developed for diagnosis. Therefore, it important to highlight that the use of prediction models would enhance ACS detection by looking into different symptoms associated with ACS.

It is clear that chest pain is a main symptom of ACS (2, 11, 19, 21-23). However, previous studies have shown that some ACS patients has been given lower priority during triage due to not having chest pain (3, 24). Deakin *et al* found that 13% of patients who called for an ambulance with confirmed ACS presented with no chest pain A study was conducted in Australia by Linda *et al* (25) to identify the sex differences in symptoms related to MI reported to EMS dispatch agrees with Deakin *et al* (3) and our findings that chest pain alone

is not a solid predictor of MI. Among female patients who called for an ambulance and were finally diagnosed with MI only 54.4% presented with chest pain, while among male patients 68.7% had chest pain. As a result, priority 1 allocation for women was 67% and for men 81% (25). Also, several studies have identified differences of symptoms reported by ACS patients. Interestingly, as we found in this systematic review, symptoms of ACS differ between male and female patients (18, 19, 25). One of the studies included in the search developed two different prediction models for men and women. However, the female prediction model failed to stand up to internal validation and should not be implemented (18). The most common symptom with MI following chest pain is Shortness of Breath (SOB) which appeared in 28.3% women vs 25.8% men. In patients presenting with no chest pain, SOB was the most common symptom, occurring in 38% women vs 32% men. These findings agree with what we have found as two studies that nausea, SOB and cold sweat were the most common symptoms reported after chest pain (16, 17). Women with ACS are less likely to present with chest pain when compared to men. Therefore, these differences might lead to the fact that dispatch triage systems can't accurately detect ACS.

### **Future Research:**

Future work should focus on deriving and validating for chest pain triage to increase the sensitivity and specificity for detecting ACS and life-threatening conditions. Prediction models have shown that it has more accuracy than dispatch system tools. Prediction models could help in reducing the over triage and increasing the accuracy of detecting ACS and other life-threatening conditions associated with chest pain which could eventually lead to reduce over-triage for non life-threatining conditions, and EDs crowding.

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#### **Patient and Public Involvement:**

No Patient involved.

### Funds:

The publication fee is funded by the cardiovascular department, University of Manchester,

UK. No grant or award number.

#### Limitations:

It is possible that some relevant papers were missed because we only included studies that have adult participants who are >18 years old and we excluded papers not published in English language. Unfortunately, we couldn't conduct a meta-analysis as there was variations between dispatch system tools and priority types. Also, there was a difference in the definition of life-threatening conditions definition between included studies.

#### **Conclusion:**

EMS dispatch systems accuracy for ACS and life-threatening conditions associated with chest pain is good. Since the dispatch tools were built to triage ambulance response priority based on sign and symptoms, this led to over triage among non life-threatining chest pain patients. Over triage were slightly reduced by deriving different prediction models and showed better sensitivity.

# Table 1: Characteristics of included studies.

Study ID	Year	Country	Study design	Sites	Study period
Gellerstedt, et al.	2006	Municipality	Prospective cohort study	1	3 months
(16)		of Go <sup>~</sup> teborg			
Reuter, et al. (18)	2019	France	Follow-up prospective	1	18 months May 2010-
			cohort study		Nov 2011
Gellerstedt, et al.	2016	Sweden	Retrospective cohort study	1	7 months 1 may
(17)		0	6		2009- 28 Feb 2010

# Table 2: Study and patient characteristics of all studies included in the systematic

review.

12							
14 15	Study ID	N	Population	Triage	Exclusion criteria	Target	Study weaknesses
16 17 18				criteria		condition	
19 20	Gellerstedt	503	Patients who called	Dispatcher	NA	AMI, Life-	Retrospective, no
21 22 23	, et al. (16)		for an ambulance	judgement		Threatening	sample size
23 24 25			and who were	vs		Conditions	calculation, small
26 27			assessed by the	prediction			simple size, out of
28 29			dispatcher as having	model			date AMI
30 31 32			chest pain	(derived in			reference standard
33 34				this cohort)			(based on creatine
35 36							kinase-MB rather
37 38					2		than cardiac
39 40 41					0		troponin)
42 43	Reuter, et	3727	Adult 18 old called	Prediction	Difficult	ACS	Information bias,
44 45 46	al. (18)		for ambulance chest	model	communication,		unclear how AMI
40 47 48			pain with final		language barrier		was adjudicated.
49 50			diagnosis.		inability to speak		
51 52 53					with patient		
53 54 55	Gellerstedt	1942	Patients who called	Dispatcher	Lost diagnosis, or	ACS, Life-	95% CIs for test
56 57	, et al.		112 for an	judgement	follow up.	Threatening	characteristics
58 59	(17)		emergency	VS		Conditions	were not
60	L		1	1	L	1	

1 2						
2 3 4		ambulance with a	prediction			presented. Only
5 6		complaint of chest	model			internal validation
7 8		pain				completed, and
9 10 11						the full results of
12 13						that validation
14 15						were not
16 17 18						presented
19 20	ACS: Acute Coron	ary Syndrome; AMI: A	cute Myocardia	al Infraction; CI: Co	onfidence Interva	l; NA: Not
21 22	Available.					
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50 51 52 53 54 55 56 57 58	
58 59 60	

# Table 3: Diagnostic accuracy of dispatch tools and prediction models for ACS.

ID	N	Triage	Sensitivity	Specificity	NPV	PPV	TP	FP	FN	TN
		criteria	%	%	%	%	%	%	%	%
Gellerste	503	Dispatch	85.7%	26.9%	87.7	32.0	90	291	15	107
dt, et al.		system				2				
(16)		(standard								
		care)								
	503	Prediction	92.4%	28.6%	93.4	25.5	97	284	8	114
		model	Ô,							
Reuter,	3727	Male	NA	NA	NA	NA	NA	NA	NA	NA
et al.		Prediction		0,						
(18)		model		4						
	1824	Female	NA	NA	NA	NA	NA	NA	NA	NA
		Prediction			4					
		model			C					
Gellerste	1942	Dispatch	82.6	39.9	94.3	15.9	194	1026	41	681
dt, et al.		system								
(17)	1942	Full	90.2	40.9	96.8	17.4	212	1008	23	699
		Prediction								
		model								
	1942	Limited	91.1	41.1	96.9	17.5	214	1006	22	701
		Prediction			6					
		model								

# Table 4: Diagnostic accuracy of dispatch tools and prediction models for life-

# threatening conditions.

ID	N	Triage	Sensitivity	Specificity	NPV	PPV	TP %	FP %	FN	TN
		criteria	%	%	%	%			%	%
Gellerste	493	Dispatch	80.9	37.5	64.1	47.3	178	198	42	75
dt, et al.		system								
(16)	493	Prediction	86.4	31.8	74.4	50.5	190	186	30	87
		model		~						
Gellerste	1944	Dispatch	76.5	39.8	89.9	19.5	238	983	73	650
dt, et al.		system		4	•					
(17)	1944	Full	88.4	42.1	95	22.5	275	946	36	687
		Prediction			2					
		model				2				
	1944	Limited	88.7	42.1	95.1	22.6	276	945	35	688
		Prediction								
		model								

# Figure 1: Flow diagram of study selection

# Figure 2: QUADAS-2 assessment of eligible studies

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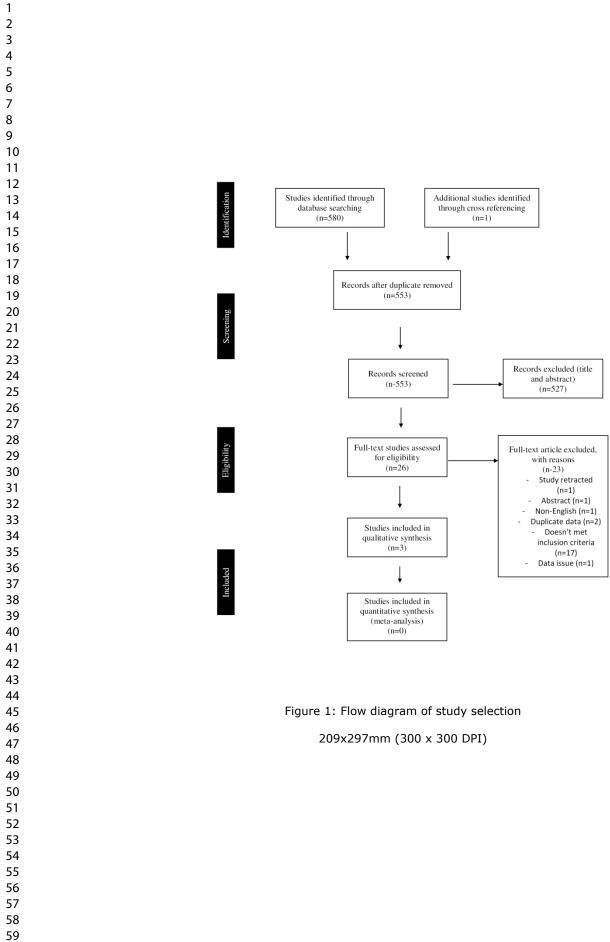
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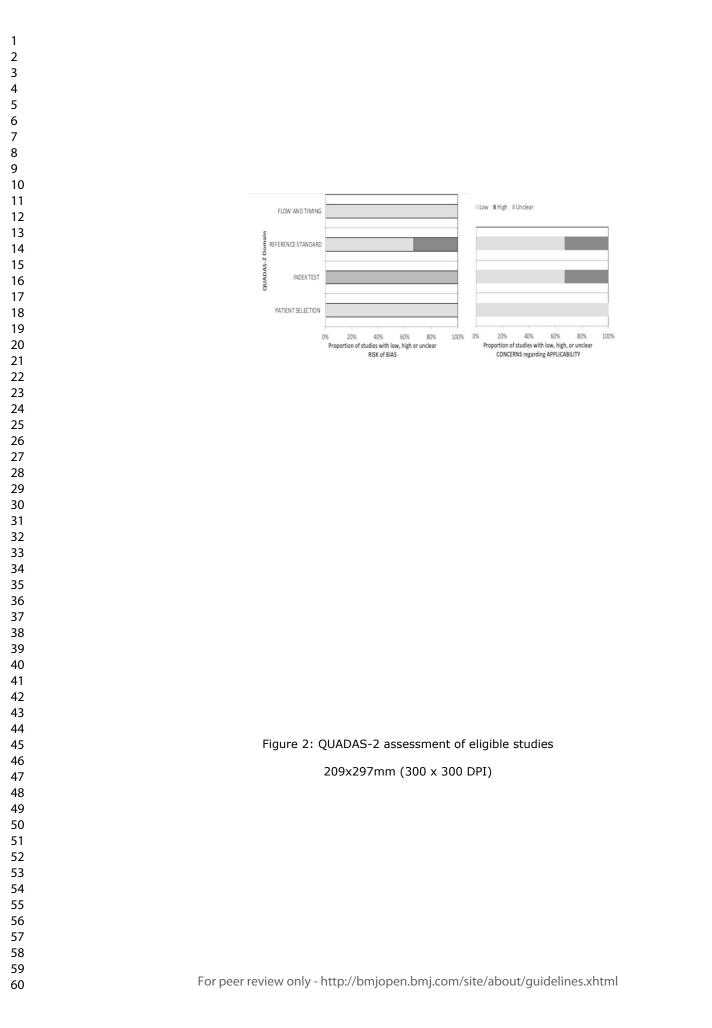
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Study	Reason for exclusion
(Deakin et al. 2006)	Included all medical (non-chest pain) calls as TN.
(Scott et al. 2017)	The authors didn't provide details on dispatch priority or sensitivity of triage accuracy.
(Herlitz et al. 2002)	Population Included based on any symptom that raise suspicion of AMI.
(Rawshani et al. 2017)	Duplicate data.
(Grzybowski et al. 2000)	No dispatch triage priority or sensitivity of triage accuracy.
(Plat et al. 2018)	Included all medical calls not only chest pain.
(Ball et al. 2016)	Included all medical and trauma calls.
(Sporer et al. 2008)	Included all patients transported by ambulance.
(Andersen et al. 2016)	included chronic diseases patients.
(Wouters et al. 2020)	Included patients called out of hours primary care no the emergency medical services.
(Rawshani et al. 2016)	Duplicate data.
(Sørensen et al. 2013)	Included patients based on ECG findings
(Thakore, McGugan, and Morrison 2002; Braunwald et al. 2002)	Included medical and trauma conditions.
(Higgins et al. 1993)	Included patients who arrived to ED only.
(Nehme, Andrew, and Smith 2016)	Measuring response time to time critical emergencies with no final diagnosis.
(Clawson et al. 2018)	Included AMI confirmed cases only and didn't provide non-cardiac chest pain to measure accuracy.
(Sporer and Wilson 2013)	Measured triage for accurately sending ALS or BLS team for drug administration.
(Adams et al. 2010)	No dispatch triage. Prioritizing patients based on ECG findings.
(Manzo-Silberman et al. 2015)	Calls collected from general practitioner and EMS dispatch with no linkage of priority to final diagnosis.
(Pedersen et al. 2019)	No dispatch priority assigned to the final diagnosis.
(Carmen Martín-Castro 2001)	Non-English.
(Pandey and Khandekar 2009)	Article retracted due to data issue.
(Bhargava et al. 2012)	Abstract.

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# Supplementary Appendix

# Customised QUADAS-2 tool for telephone triage in chest pain systematic review

# Domain 1: Patient selection

10 11	A. Risk of bias	
12	a. Was a consecutive or random sample of patients enrolled?	Yes / No / Unclear
13 14	b. Did the study avoid inappropriate exclusions?	Yes / No / Unclear
14 15 16	Could the selection of patients have introduced bias?	Low / High / Unclear
17	B. Concerns regarding applicability	
18 19	Is there concern that included patients do not match the review que	estion? Low / High / Unclear
20 21		
22	OVERALL:	
23		
24 25		
25 26	LOW: Patients who call 999, 911, 112, 108 or another number for emergence	y medical attention with a primary complaint of chest pain
27	HIGH: Selection of a specific high- or low-risk population: setting is not an e	mergency telephone consultation; convenience sampling with clear potential for
28		nergency telephone consultation, convenience sampling with clear potential for
29	systematic selection bias	
30 31	UNCLEAR: Insufficient information to determine the risk of bias.	
32		
33		
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Domain 2: Telephone triage intervention		
<ul> <li>A. Risk of bias         <ul> <li>a. Was the outcome of telephone triage</li> </ul> </li> </ul>	interpreted without knowledge of the outcome?	Yes / No / Unclear
b. Is it unlikely that the telephone triage	introduced biased risk group allocation?	Yes / No / Unclear
Could the scoring or interpretation of the tele	phone triage have introduced bias?	Low / High / Unclear
B. Concerns regarding applicability		
Is there concern that the telephone triage, its differ from the review question?	conduct, or interpretation	Low / High / Unclear
OVERALL:		
LOW: Telephone triage outcome calculated prospective	vely by treating clinicians, blinded to patient outcome.	
HIGH: Telephone triage outcome calculated without b	linding to outcome. Retrospective calculation of telepho	one triage outcome.
UNCLEAR: Insufficient information.		
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1				
2 3				
4	Do	mai	n 3: AMI/serious adverse event allocation	
5 6		A.	Risk of bias	
7			a. Was the universal definition of AMI used to define the reference standard?	Yes / No / Unclear
8 9 10			b. Was the reference standard adjudicated without knowledge of the telephone triage?	Yes / No / Unclear
10 11 12 13			Could the reference standard, its conduct, or its interpretation have introduced bias?	Low / High / Unclear
14		В.	Concerns regarding applicability	
15			Is there concern that the target condition as defined by the reference standard	Low / High / Unclear
16 17			does not match the review question?	
18				
19				
20 21		0	'ERALL:	
21		0		
23				
24		LO	W: All patients underwent reference standard investigations for AMI including troponin sampling	g. The diagnosis of AMI was adjudicated by at least two
25 26		inv	estigators who were blinded to the telephone triage outcome. Serious adverse events include d	eath (all cause), aortic dissection, pulmonary embolism and
20 27 28			nsion pneumothorax	
20 29		ні	GH: Outdated definition for AMI or definition inconsistent with the universal definition. The defin	nition of serious adverse events does not include one of the
30			e components specified above. Follow up procedure raises significant concerns about the possil	
31			hout some assurance that this method would capture all relevant events).	
32 33				
33 34		UN	ICLEAR: Insufficient information.	
35				
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44 45			For peer review only - http://bmjopen.bmj.com/site/about/g	uidelines.xhtml
45 46				
47				

# Domain 4: Flow and timing

# A. Risk of bias

	a.	Did all included patients receive an appropriate reference standard?	Yes / No / Unclear			
	b.	Did patients receive the same reference standard?	Yes / No / Unclear			
	с.	Follow-up procedure was sufficiently long to not miss relevant adverse events?	Yes / No / Unclear			
	d.	Did no significant loss to follow up/exclusion due to incomplete records occur?	Yes / No / Unclear			
	Could t	he patient flow have introduced bias?	Low / High / Unclear			
LOV	V: All pa	atients underwent reference standard investigations for AMI including troponin sampling	. Patients were followed up through their subsequent			
inpa	atient c	ourse or for at least 7 days.				

HIGH: Not all patients were subjected to appropriate reference standard investigations including troponin sampling.

UNCLEAR: Insufficient information. This option includes lack of detail about troponin testing and the follow up procedure.

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# Table 1: QUADAS-2 assessment of included studies ( / Low Risk, green, X High Risk, red, ? Unclear Risk, amber)

Study		RISK OF BIAS			APPLICABILITY CONCERNS				OVERALL	
	PATIENT SELECTION	TELEPHONE TRIAGE	AMI /SAE ALLOCATION	FLOW AND TIMING	PATIENT SELECTION	TELEPHONE TRIAGE	AMI /SAE ALLOCATION	PATIENT SELECTION	TELEPHONE TRIAGE	AMI /SAE ALLOCATION
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## PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #		
TITLE					
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1		
ABSTRACT					
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	2-3		
INTRODUCTION					
Rationale	3	Describe the rationale for the review in the context of what is already known.	5-6		
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	7		
METHODS					
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration including registration number.			
Eligibility criteria	criteria 6 Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered language, publication status) used as criteria for eligibility, giving rationale.				
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.			
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.			
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	6-7		
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	7-8		
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	7		
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	7,8		
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	7		
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I <sup>2</sup> ) for each meta-analysis.	8		
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## **PRISMA 2009 Checklist**

Section/topic	#	Checklist item	Reported on page #			
Risk of bias across studies	15 Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).					
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	8			
RESULTS						
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	Figure 1			
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	Table 1,2			
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	9			
Results of individual studies	sults of individual studies 20 For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.					
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.				
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).				
Additional analysis	23 Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).					
DISCUSSION	4					
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	11-13			
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	14			
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	14			
FUNDING	1					
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	14			
<i>From:</i> Moher D, Liberati A, Tetzlaff doi:10.1371/journal.pmed1000097	J, Altm	an DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med For more information, visit: <u>www.prisma-statement.org</u> .	6(7): e1000097			
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#### Accuracy of emergency medical services (EMS) telephone triage in identifying acute coronary syndrome (ACS) for chest pain patients: A systematic literature review

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Secondary Subject Heading:	Cardiovascular medicine
Keywords:	Myocardial infarction < CARDIOLOGY, Coronary heart disease < CARDIOLOGY, ACCIDENT & EMERGENCY MEDICINE

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## Accuracy of emergency medical services (EMS) telephone triage in identifying acute coronary syndrome (ACS) for chest pain patients: A systematic literature review

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- D) Pre-hospital, Chest pain, telephone triage, ACS, Life-threatening conditions,

 E) Words count: 3170

#### Abstract;

#### **Objective:**

To systematically appraise the available evidence to determine the accuracy of decision aids for Emergency Medical Services (EMS) telephone triage of patients with chest pain suspected to be caused by acute coronary syndrome (ACS) or life-threatening conditions.

#### **Design:**

Systematic review.

#### **Data sources:**

Electronic searches were performed in Embase 1974, Medline 1946, and Cinahl 1937 databases from 03/03/2020 to 04/03/2020.

#### **Eligibility criteria:**

The review included all types of original studies that included adult patients (>18 years) who called EMS with a primary complaint of chest pain and evaluated dispatch triage priority by telephone. Outcomes of interest were a final diagnosis of ACS, acute myocardial infarction (AMI), or other life-threatening conditions.

#### Data extraction and synthesis:

Two authors independently extracted data on study design, population, study period, outcome and all data for assessment of accuracy, including cross-tabulation of triage priority against the outcomes of interest. Risk of bias was assessed using the QUADAS-2 assessment tool.

#### **Results:**

Searches identified 553 papers, of which three were eligible for inclusion. Those reports described the evaluation of three different prediction models with variation in the variables

used to detect ACS. The overall results showed that dispatch triage tools have good sensitivity to detect ACS and life-threatening conditions, even though they are used to triage signs and symptoms rather than diagnosing the patients. On the other hand, prediction models were built to detect ACS and life-threatening conditions and therefore prediction models showed better sensitivity and Negative Predictive Value (NPV) than dispatch triage tools.

#### **Conclusion:**

We have identified three prediction models for telephone triage of patients with chest pain. While they have been found to have greater accuracy than standard EMS dispatch systems, prospective external validation is essential before clinical use is considered.

#### Strength and limitations:

- To our knowledge, this is the first systematic review looking into the accuracy of prehospital chest pain telephone triage.
- It is possible that some relevant papers were missed because researchers only included studies that have adult participants who are >18 years old and excluded papers not published in English language.
- Unfortunately, because of the paucity of data for each decision aid, the data were not suitable for meta-analysis.
- Also, there was a difference in the definition of life-threatening conditions definition between included studies.

#### **Funding:**

The publication fee is funded by the cardiovascular department, University of Manchester,

UK. No grant or award number.

#### **Registration:**

This systematic review was pre-registered on the International prospective register of systematic reviews (PROSPERO) database (reference CRD42020171184)

#### **Abbreviations list:**

ED: Emergency Department; ACS : Acute Coronary Syndrome; AMI: Acute Myocardial Infarction; ECG: Electrocardiogram; NPV: Negative Predictive Value; QUADAS: Quality Assessment of Diagnostic Accuracy Studies; PPV: Positive Predictive Value; STEMI: ST-Elevation Myocardial Infarction; ACS: Acute Coronary Syndrome; EMS:

**Emergency Medical Services** 

#### Keywords:

Pre-hospital, Chest pain, telephone triage, Acute Coronary Syndrome.

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#### **Background:**

Chest pain, which is one of the main symptoms of acute coronary syndromes (ACS), is one of the most common reasons for ambulance callouts and presentations to emergency departments (EDs) (1, 2). However, most patients with chest pain who are transported by ambulance are ultimately diagnosed with self-limiting, non-cardiac disease. Thus, patients with chest pain tend to be systematically over-triaged, increasing ambulance resource utilisation, and contributing to ED crowding (2, 3).

Two EMS dispatch systems are currently used in the UK: the Advanced Medical Prioritizing Dispatching System (AMPDS), and the National Health Service (NHS 999) System (4). Both systems use computerized telephone triage software that identify life-threatening conditions by a series of questions at the beginning of the call. For patients with a primary complaint of chest pain, they also include a specific series of questions to identify ACS and triage the case using the patient's sign and symptoms (3, 5). However, systematic reviews have shown that there is only a low level of evidence to support the use of current triage systems (6, 7). There is also insufficient data on the dispatch protocol efficiency (7, 8). There is no consensus on the definitions used in Criteria Based Dispatch (CBD), which is the program used to triage patients by signs and symptoms, and AMPDS (9, 10). Also, there is no consensus on the accepted level for over- or under-triage for medical emergency dispatch (6, 9). Therefore, systematic over-triage for chest pain patients is used by dispatch systems to avoid potential harm to patients. EMS resource utilisation is increased as a result (6), and there are important differences between patients clinical findings when examined and dispatch priority (8, 9). Therefore, with the current clinical guidelines for emergency medical services (EMS), it is hard to rule in or rule out ACS by telephone-triage (11).

We aimed systematically to appraise the available evidence to determine the accuracy of decision aids that have been used for EMS telephone triage of patients with chest pain suspected to be caused by ACS or life-threatening conditions.

#### **Methods:**

 The systematic review was conducted in accordance with Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines and following Cochrane methodology for diagnostic test accuracy reviews (12). A completed PRISMA checklist is provided in supplementary appendix 1. This systematic review was pre-registered on the International prospective register of systematic reviews (PROSPERO) database (reference elien CRD42020171184)

#### Search strategy and eligibility criteria:

We searched the Embase 1974, Medline 1946, and Cinahl 1937 databases from 3/03/2020 to 04/03/2020 for the terms "chest pain, telephone or dispatch, triage, ACS or AMI, lifethreatening conditions, and EMS or prehospital". Retrospective and prospective cohort studies written in English were eligible. Search strategy table is provided in supplementary appendix 2.

#### Studies included:

Based on the inclusion criteria, titles and abstracts were independently screened and shortlisted by two reviewers (AbA and ChR). The inclusion criteria were: (1) the study

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included adult patients (>18 years); (2) the study included patients with a primary complaint of chest pain who called EMS for an ambulance; (3) the studied evaluated dispatch triage priority; (4) the study provided data for a linked final diagnosis of ACS, acute myocardial infarction (AMI), or other life-threatening conditions. Two reviewers independently reviewed and screened potentially relevant full text papers to identify those fulfilling the inclusion criteria. Any discrepancies between the reviewers were resolved by a third reviewer (AhA).

#### **Outcome measures:**

The primary outcome was a diagnosis of ACS upon hospital admission, including ST elevation myocardial infarction (STEMI), non-ST elevation myocardial infarction (NSTEMI) and unstable angina. AMI (incorporating both STEMI and NSTEMI) is defined according to the universal definition of AMI (13). Unstable Angina is defined as sudden prolonged chest pain at rest >20 minutes, new onset severe angina, or previous angina that is increasing in severity of pain, duration of pain, and/or frequency (14).

The secondary outcome was the diagnosis of a life-threatening condition associated with chest pain. Since there is no universal definition for life-threatening conditions associated with chest pain, we included all relevant data from the studies, regardless of the precise definition used for 'life-threatening conditions'.

#### Methodological quality assessment:

The quality assessment of eligible articles was independently assessed by two reviewers (AhA and AbA), using a modified Quality Assessment of Diagnostic Accuracy Studies (QUADAS-2) tool to suit the purpose of this systematic review (15). Discrepancies between reviewers were solved by discussion.

#### Data extraction:

After selecting the eligible studies, two investigators (AhA and AbA) used a standardised data extraction form to extract and collect data as study design, population, study period, and outcome. Also, both investigators extracted all data required for diagnostic accuracy assessment including 2x2 tables when available for dispatch priority and prediction models priority for suspected ACS, or life-threatening conditions.

#### Statistical analysis:

In line with our protocol, we considered pooling sensitivities, specificities, negative and positive predictive values (NPV and PPV, respectively) by meta-analysis. However, after reviewing the final list of studies meeting inclusion criteria, the reviewers unanimously concluded that meta-analysis would have been inappropriate due to clear evidence of clinical heterogeneity between studies, and missing data. We therefore undertook a narrative synthesis of the existing evidence.

#### **Results:**

In total, our literature search identified 553 potentially relevant articles of which 26 were considered potentially eligible for inclusion after screening titles and abstracts. Following independent full text review, 23 articles were excluded and 3 articles met the inclusion

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criteria for this review (Figure 1). Excluded studies with reason of exclusion is shown in the supplementary appendix 3 (16-18).

#### **Study Characteristics and Quality Assessments:**

General study characteristics are summarized in Table 1. Patient characteristics and study weaknesses are summarised in Table 2. The number patients in each study with each component of the outcome measures studied is summarised in Supplementary appendix 4 Table 1. The details of the prediction models identified, together with the test characteristics (including sensitivity, specificity, NPV, and PPV) for the diagnosis of ACS and lifethreatening conditions are shown in Tables 3, 4.

The modified QUADAS-2 methodological quality assessment tool was used to assess all eligible studies as shown in the supplementary appendix 5. A summary of the quality assessment for each eligible study is shown in Figure 2. Two out of the three studies were retrospective cohort studies and compared the dispatch triage for ambulance response priority to the prediction model (16, 17) and one was prospective cohort study that only testing prediction model accuracy (18). ACS was adjudicated by experts in only one study (18) while the other two studies used the hospital final diagnosis of ACS, AMI, or life-threatening conditions. All eligible studies utilized the appropriate ACS or AMI definitions at the time of conducting the study.

#### Summary of the existing evidence

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Gellerstedt *et al* derived a prediction model to detect AMI and life-threatening conditions patients calling for an emergency ambulance with chest pain. They aimed to effectively triage calls to determine the appropriate level of response (capable of providing advanced life support vs only basic life support). The final model used eight variables (age, gender, strong pain, dyspnoea, cold sweat, nausea, vertigo, and syncope) to calculate the probability of AMI or a life-threatening condition. Setting an arbitrary cut-off, the authors found that the model had greater sensitivity than and similar specificity to the judgement of the call handlers, which could potentially improve triage. However, performance of the model was only evaluated in the derivation set, with no validation presented (16).

In 2016, the same group derived a new prediction model in another study. They aimed to determine whether a computer-based decision support system could have superior accuracy for identifying life threating conditions or ACS in patients who called for an emergency ambulance with chest pain. *Gellerstedt et al* collected data from patients who called 112 complaining of chest pain using ten questions related to ACS with yes or no answers (17). They derived two prediction models based on those answers. The first prediction model (full model) incorporated data from all ten questions that dispatchers had asked, including: age, gender, central chest pain, high intensity pain, pain duration between 15 minutes and 24 hours, aggravated pain when breathing or moving, abnormal breathing, cold sweat, diabetes or previous cardiovascular disease (CVD), previous ACS, and belief that symptoms are heart-related. The authors also derived a 'limited model', which only included variables that had a significant association with the presence of ACS or a life-threatening condition on bivariate analysis. These included: gender, age, central chest pain, high intensity pain, pain, previous ACS, and a high priority based on dispatcher judgement. There was no significant difference in performance between the full model and limited model. However, both the limited and full

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models had superior sensitivity to the priority assigned by dispatchers in practice, both for ACS (Table 3) and life-threatening conditions (Table 4), with similar specificity (17). The model was derived in 70% of the available cohort and validated in the remaining 30%. The authors state that performance was 'similar' in the validation set, although no further details were provided.

Reuter *et al* used backward logistic regression to derive two separate prediction models to detect ACS among men and women, respectively (18). The male prediction model to detect ACS consisted of 8 variables, which are: age, smoking, severe pain, permanent pain, breathing non-related pain, retrosternal pain, radiating pain, and additional symptoms. The female prediction model to detect ACS consisted of four variables, which are: age >60 years, history of coronary artery disease, breathing non-related pain, and radiating pain.

The male prediction model had an area under the receiver operating characteristic curve (AUC) of 0.76 (95% CI 0.73 – 0.79) in the derivation set (70% of the sample, which was randomly selected), which was maintained at 0.76 (95% CI 0.73 – 0.80) in the internal validation set (30% of the cohort). However, while the female prediction model had a similar AUC in the derivation set (0.79, 95% CI 0.75 – 0.83), it had lower accuracy in the validation set (AUC 0.67, 95% CI 0.60 – 0.74). No data were presented to enable calculation of sensitivity, specificity, PPV or NPV, and the output of the model was not compared to standard care or clinical judgement.

#### Discussion

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In this systematic review, we have identified three studies that presented the derivation of clinical prediction models to enhance the telephone triage of patients calling for an emergency ambulance with chest pain. None of the models presented had sufficiently high sensitivity to avoid the need for ambulance dispatch. While two of the prediction models were shown to have greater accuracy than standard dispatch systems, unfortunately the current evidence does not support the use of these prediction models in practice without further validation. However, the evidence to date does suggest that clinical prediction models for this use case could feasibly improve upon the accuracy of triage offered by standard EMS dispatch systems.

The original model derived by Gellerstedt *et al* (2006) used a now outdated definition of AMI (using creatine kinase-MB as the reference standard biomarker), meaning that the results could not be applied to modern clinical practice without further validation. The second model derived by Gellerstedt *et al* (2016) underwent internal validation, but full details that validation were not published. Finally, Reuter *et al* derived two prediction models (for men and women, respectively) but did not present sufficient data to compare their performance to standard care. The AUC for the male prediction model was 0.76, showing good overall accuracy. The prediction model for females had a poor AUC (0.67) on internal validation. We identified no prediction models that have undergone external validation.

It is an extremely challenging task for dispatchers to identify ACS over the telephone as there are so many different causes of chest pain (11, 19, 20), and there is very limited information available at the time of the initial phone call. It has previously been reported that only 1 out of 18 patients who request an ambulance for a complaint of chest pain has ACS(3). While a

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validated prediction model would have clear benefits, this systematic review has identified no validated tools that could be used in current practice.

It has also been reported that the AMPDS triage system is not an accurate tool for identifying ACS (3). Moreover, in Denmark, Criteria-based Danish index is used to triage emergency calls and it resulted in over-triage for 51% of chest pain calls as they were discharged without a specific diagnosis and AMI was only found in 11% of patients (2). Dispatch triage systems were constructed to prioritize the pre-hospital response based on a patient's condition. They were not intended to make diagnoses. Therefore, it important to highlight that the use of prediction models for telephone triage may not be expected to 'rule out' ACS but to identify patients who may safely receive a lower priority response, while correctly identifying the vast majority of those who require an urgent response.

It is clear that chest pain is a main symptom of ACS (2, 11, 19, 21-23). However, previous studies have shown that some ACS patients has been given lower priority during triage due to not having chest pain (3, 24). Deakin *et al* found that 13% of patients who called for an ambulance with confirmed ACS presented with no chest pain A study was conducted in Australia by Linda *et al* (25) to identify the sex differences in symptoms related to MI reported to EMS dispatch agrees with Deakin *et al* (3) and our findings that chest pain alone is not a solid predictor of MI. Among female patients who called for an ambulance and were finally diagnosed with MI only 54.4% presented with chest pain, while among male patients 68.7% had chest pain. As a result, priority 1 allocation for women was 67% and for men 81% (25). Also, several studies have identified differences of symptoms reported by ACS patients. Interestingly, as we found in this systematic review, symptoms of ACS differ between male and female patients (18, 19, 25). One of the studies included in the search developed two

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different prediction models for men and women. However, the female prediction model failed to stand up to internal validation and should not be implemented (18). The most common symptom with MI following chest pain is Shortness of Breath (SOB) which appeared in 28.3% women vs 25.8% men. In patients presenting with no chest pain, SOB was the most common symptom, occurring in 38% women vs 32% men. These findings agree with what we have found as two studies that nausea, SOB and cold sweat were the most common symptoms reported after chest pain (16, 17). Women with ACS are less likely to present with chest pain when compared to men. This may explain the relative underperformance of the female prediction model derived by Reuter *et al.* 

#### **Future Research:**

This systematic review has highlighted two potential areas for future research. First, the three prediction models identified may add value over existing telephone triage algorithms but this would require prospective external validation. Second, the existing evidence suggests that it is feasible to derive a prediction model to enhance telephone triage for patients with suspected ACS. The derivation and validation of a new, contemporary prediction model could reduce the systematic over-triage of patients in future, while ensuring that those with life-threatening conditions consistently receive an urgent response. However, given the risks of delayed responses (e.g. 'time is muscle for patients with STEMI), a higher sensitivity than has been observed in the studies identified is likely to be required.

#### **Patient and Public Involvement:**

No Patient involved.

#### **Ethical approval:**

Ethical approval is not required for this systematic review.

#### Limitations:

It is possible that some relevant papers were missed because we only included studies that have adult participants who are >18 years old and we excluded papers not published in English language. Unfortunately, we couldn't conduct a meta-analysis as there was variations between dispatch system tools and priority types. Also, there was a difference in the definition of life-threatening conditions definition between included studies.

#### **Conclusion:**

e e We have identified three clinical prediction models for telephone triage of patients with chest pain. While the models have been found to have greater accuracy than standard EMS dispatch systems, the level of evidence to support their use is currently low. Prospective external validation is therefore required. However, our findings do support the feasibility of deriving and validating clinical prediction models to reduce over-triage while ensuring urgent responses for those who need them most.

#### **Funding:**

The publication fee has been funded by the Division of Cardiovascular Sciences, University of Manchester, UK. No grant or award number.

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#### **Tables and Figures**

## Table 1: Characteristics of included studies.

Study ID	Year	Country	Study design	Sites	Study period
Gellerstedt, et al.	2006	Sweden	Prospective cohort study	1	3 months
(16)					
Reuter, et al. (18)	2019	France	Follow-up prospective	1	18 months May 2010-
			cohort study		Nov 2011
Gellerstedt, et al.	2016	Sweden	Retrospective cohort study	1	7 months 1 may
(17)		O C			2009- 28 Feb 2010

## Table 2: Study and patient characteristics of all studies included in the systematic

review.

1 2 3 4							
5 6	Study ID	N	Population	Triage	Exclusion criteria	Target	Study weaknesses
7 8				criteria		condition	
9 10 11	Gellerstedt	503	Patients who called	Dispatcher	NA	AMI, Life-	Retrospective, no
12 13	, et al. (16)		for an ambulance	judgement		Threatening	sample size
14 15			and who were	vs		Conditions	calculation, small
16 17 18			assessed by the	prediction			simple size, out of
19 20			dispatcher as having	model			date AMI
21 22 23			chest pain	(derived in			reference standard
23 24 25				this cohort)			(based on creatine
26 27				Ő.			kinase-MB rather
28 29 30							than cardiac
31 32				0			troponin)
33 34	Reuter, et	3727	Adult 18 old called	Prediction	Difficult	ACS	Information bias,
35 36	al. (18)		for ambulance chest	model	communication,		unclear how AMI
37 38 39			pain with final		language barrier		was adjudicated.
40 41			diagnosis.		inability to speak		
42 43					with patient		
44 45 46	Gellerstedt	1942	Patients who called	Dispatcher	Lost diagnosis, or	ACS, Life-	95% CIs for test
40 47 48	, et al.		112 for an	judgement	follow up.	Threatening	characteristics
49 50	(17)		emergency	VS		Conditions	were not
51 52			ambulance with a	prediction			presented. Only
53 54 55			complaint of chest	model			internal validation
56 57			pain				completed, and
58 59							the full results of
60				<u> </u>	<u> </u>	<u> </u>	

						that validation
						were not
						presented
ACS: Acute (	Coronary Syn	drome; AMI	I: Acute Myoca	rdial Infraction; Cl	: Confidence In	terval; NA: Not
Available.						

Table 3: Diagnostic accuracy of dispatch tools and prediction n	nodels for ACS.
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ID	N	Triage	Sensitivity	Specificity	NPV	PPV	TP	FP	FN	TN
		criteria	%	%	%	%				
Gellerste	503	Dispatch	85.7%	26.9%	87.7	32.0	90	291	15	107
dt, et al.		system				2				
(16)		(standard								
		care)								
	503	Prediction	92.4%	28.6%	93.4	25.5	97	284	8	114
		model								
Reuter,	2,363	Male	NA	NA	NA	NA	NA	NA	NA	NA
et al.		Prediction	Ő'							
(18)		model		~						
	1,824	Female	NA	NA	NA	NA	NA	NA	NA	NA
		Prediction		4.						
		model		0						
Gellerste		Dispatch	82.6	39.9	94.3	15.9	194	1026	41	681
dt, et al.		system			C					
(17)		Full	90.2	40.9	96.8	17.4	212	1008	23	699
	1,942	Prediction								
		model								
		Limited	91.1	41.1	96.9	17.5	214	1006	22	701
		Prediction			6					
		model								
Abbreviati	ions: NP	V: Negative Pre	edictive Valu	e ; PPV: Posi	tive Pre	dictive	Value	 : TP: Tr	ue Posi	tive;

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## Table 4: Diagnostic accuracy of dispatch tools and prediction models for life-

## threatening conditions.

ID	N	Triage	Sensitivity	Specificity	NPV	PPV	TP	FP	FN	TN
		criteria	%	%	%	%				
Gellerste	493	Dispatch	80.9	37.5	64.1	47.3	178	198	42	75
dt, et al.		system								
(16)	493	Prediction	86.4	31.8	74.4	50.5	190	186	30	87
		model								
Gellerste	1944	Dispatch	76.5	39.8	89.9	19.5	238	983	73	650
dt, et al.		system								
(17)	1944	Full	88.4	42.1	95	22.5	275	946	36	687
		Prediction		14	•					
		model		C	2					
	1944	Limited	88.7	42.1	95.1	22.6	276	945	35	688
		Prediction								
		model				2				

Supplementary appendix:

Table 1: Number of patients with each individual outcome by study

Legends to figures:

Figure 1: Flow diagram of study selection

Figure 2: QUADAS-2 assessment of eligible studies. Green = low risk; yellow = unclear;

red = high risk

#### A. Contributorship statement:

Ahmed Alotaibi: writing protocol, data search, data extraction, quality assessment, and writing.

Abdulrahman Alghamdi: data search, data extraction, quality assessment.

Charles Renard: data search, data extraction, quality assessment.

Richard Body: protocol planning, writing, proofreading.

#### **B.** competing interests:

The authors declare that they have **no** conflict of **interest** 

#### C. Funding:

The publication fee has been funded by the Division of Cardiovascular Sciences, University

of Manchester, UK. No grant or award number.

#### D. Data sharing statement:

All the data included were obtained from published articles.

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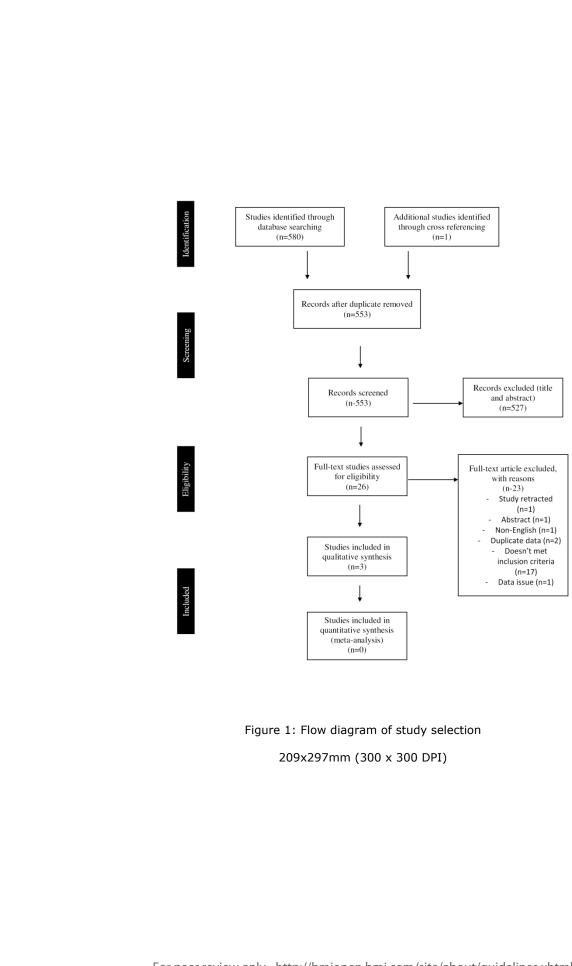
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Low High Unclear	Risk of bias			Concerns regarding applicability				
	Gellerstedt et al, 2016	Reuter et al, 2019	Gellerstedt et al, 2006	Gellerstedt et al, 2016	Reuter et al, 2019	Gellerstedt et al, 2006		
Flow and timing				N/A				
Reference standard								
Index test								
Patient selection								

Figure 2: QUADAS-2 assessment of eligible studies.

338x190mm (200 x 200 DPI)

#### The PRISMA for Abstracts Checklist

TITLE	CHECKLIST ITEM	REPORTED ON PAGE #
1. Title:	Identify the report as a systematic review, meta-analysis, or both.	1
BACKGROUND		
2. Objectives:	The research question including components such as participants, interventions, comparators, and outcomes.	2 (objectives)
METHODS		
3. Eligibility criteria:	Study and report characteristics used as criteria for inclusion.	2 (eligibility criteria)
4. Information sources:	Key databases searched and search dates.	2 (data sources)
5. Risk of bias:	Methods of assessing risk of bias.	2 (data extraction 8 synthesis)
RESULTS		
6. Included studies:	Number and type of included studies and participants and relevant characteristics of studies.	3 (results)
7. Synthesis of results:	Results for main outcomes (benefits and harms), preferably indicating the number of studies and participants for each. If meta-analysis was done, include summary measures and confidence intervals.	3 (results)
8. Description of the effect:	Direction of the effect (i.e. which group is favoured) and size of the effect in terms meaningful to clinicians and patients.	N/A (no meta- analysis)
DISCUSSION		
9. Strengths and Limitations of evidence:	Brief summary of strengths and limitations of evidence (e.g. inconsistency, imprecision, indirectness, or risk of bias, other supporting or conflicting evidence)	3 (conclusions
	For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml	

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10. Interpretation:	General interpretation of the results and important implications	3
		(conclusions
OTHER		
11. Funding:	Primary source of funding for the review.	4 (funding)
12. Registration:	Registration number and registry name.	4
		(registratior

For peer review only

#### **Search Strategy**

**EMBASE** database: Date of Search: 03/03/2020 Final Result: 98 Articles.

Search Number	Keywords	Result
1	chest pain.mp. or exp thorax pain/	100365
2	dispatch.mp. or exp emergency medical dispatch/ or	104949
	exp emergency health service/ or exp ambulance/	
3	telephone.mp. or exp telephone/	87404
4	phone.mp.	46215
5	exp heart infarction/ or myocardial infraction.mp. or	8159567
	exp acute heart infarction/ or exp adult/	
6	3 or 4	127453
7	prehospital.mp.	16529
8	2 or 7	114149
9	1 and 5 and 6 and 8 🔜	98

9	1 and 5 and 6 and 8 📈	98
MEDLINE database	2:	
Date of search: 04,	/03/2020	
Final result: 464		
	4	
Search Number	Keywords	Result
1	chest pain.mp. or exp Chest Pain/	70473
2	acute coronary syndrome.mp. or exp Myocardial	189638
	Infarction/ or exp Acute Coronary Syndrome/	
3	telephone.mp. or exp telephone/	64080
4	phone.mp.	21109
5	exp Emergency Medical Services/ or prehospital.mp.	139582
6	triage.mp. or exp Triage/	18912
7	emergency health services.mp. or Emergency Medical	42024
	Services/	
8	3 or 4 or 6	91939
9	5 or 7	139622
10	1 and 2 and 8 and 9	464

Study	Reason for exclusion	
(Deakin et al. 2006)	Included all medical (non-chest pain) calls as TN.	
(Scott et al. 2017)	The authors didn't provide details on dispatch priority or sensitivity of triage accuracy.	
(Herlitz et al. 2002)	Population Included based on any symptom that raise suspicion of AMI.	
(Rawshani et al. 2017)	Duplicate data.	
(Grzybowski et al. 2000)	No dispatch triage priority or sensitivity of triage accuracy.	
(Plat et al. 2018)	Included all medical calls not only chest pain.	
(Ball et al. 2016)	Included all medical and trauma calls.	
(Sporer et al. 2008)	Included all patients transported by ambulance.	
(Andersen et al. 2016)	included chronic diseases patients.	
(Wouters et al. 2020)	Included patients called out of hours primary care no the emergency medical services.	
(Rawshani et al. 2016)	Duplicate data.	
(Sørensen et al. 2013)	Included patients based on ECG findings	
(Thakore, McGugan, and Morrison 2002; Braunwald et al. 2002)	Included medical and trauma conditions.	
(Higgins et al. 1993)	Included patients who arrived to ED only.	
(Nehme, Andrew, and Smith 2016)	Measuring response time to time critical emergencie with no final diagnosis.	
(Clawson et al. 2018)	Included AMI confirmed cases only and didn't provident non-cardiac chest pain to measure accuracy.	
(Sporer and Wilson 2013)	Measured triage for accurately sending ALS or BLS team for drug administration.	
(Adams et al. 2010)	No dispatch triage. Prioritizing patients based on ECG findings.	
(Manzo-Silberman et al. 2015)	Calls collected from general practitioner and EMS dispatch with no linkage of priority to final diagnosis.	
(Pedersen et al. 2019)	No dispatch priority assigned to the final diagnosis.	
(Carmen Martín-Castro 2001)	Non-English.	
(Pandey and Khandekar 2009)	Article retracted due to data issue.	
(Bhargava et al. 2012)	Abstract.	

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# Supplementary appendix:

# Table 1: Number of patients with each individual outcome by study

Study	Total number	ACS (N) %	AMI (N) %	STEMI	NSTEMI (N)	LTC (N) %
	of			(N) %	%	
	participants					
Gellerstedt,	503	NA	(105) 20.3%	NA	NA	(220) 43.7%
et al. (16)		O,				
Reuter, et al.	2363	(508) 21.5%	NA	(215) 9.1%	(244) 10.3%	NA
(18)	Male					
	1824 Female	(139) 7.6%	NA	(46) 2.5%	(81) 4.4%	NA
Gellerstedt,	1942	(235) 12.1%	NA	NA	NA	NA
et al.						
(17)	1944	NA	NA	NA	NA	(311) 16%
				2		1

# Supplementary Appendix

# Customised QUADAS-2 tool for telephone triage in chest pain systematic review

# **Domain 1: Patient selection**

10 11	Α.	Risk of bias		
12		a. Was a consecutive or random s	sample of patients enrolled?	Yes / No / Unclear
13		b. Did the study avoid inappropria	ate exclusions?	Yes / No / Unclear
14			U h	
15		Could the selection of patients have int	roduced bias?	Low / High / Unclear
16		····		
17	В.	Concerns regarding applicability		
18		Is there concern that included patients	do not match the review question?	Low / High / Unclear
19		is there concern that meraded patients	do not match the review question.	
20				
21	OVERA	11.		
22 23				
24				
25		Dationts who call 000 011 112 108 or ar	acthor number for emergency modical attention	with a primary complaint of chast pain
26	LOVV. P	atients who can 999, 911, 112, 108 01 an	nother number for emergency medical attention	r with a primary complaint of chest pain
27	нсн	Selection of a specific high- or low-risk pr	opulation: setting is not an emergency telephon	e consultation; convenience sampling with clear potential for
28				e consultation, convenience sampling with clear potentiar for
29	system	atic selection bias		
30		AR: Insufficient information to determine	a tha rick of high	
31	UNCLE	AR. Insumcient information to determine	e the fisk of blas.	
32 33				
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1 2			
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5			
6	Domain 2: Telephone triage intervention		
7	bomain 2. relephone mage intervention		
8	A. Risk of bias		
9		and the state of t	
10		e triage interpreted without knowledge of the outcome?	Yes / No / Unclear
11	<li>b. Is it unlikely that the telephon</li>	e triage introduced biased risk group allocation?	Yes / No / Unclear
12			
13	Could the scoring or interpretation of	the telephone triage have introduced bias?	Low / High / Unclear
14			
15			
16	B. Concerns regarding applicability		
17	Is there concern that the telephone tri	iage, its conduct, or interpretation	Low / High / Unclear
18	differ from the review question?		
19 20	unter from the review question:		
20 21	OVERALL:		
22	OVERALE.		
23			
24	LOW/ Talashana triaga autooma calculated ar	association by two time aliginizes blinded to extigat a strange	
25	LOW: Telephone triage outcome calculated pr	ospectively by treating clinicians, blinded to patient outcome	
26		ithe ut blinding to outcome. Detugon active relation of tale	
27	HIGH: Telephone thage outcome calculated w	ithout blinding to outcome. Retrospective calculation of tele	phone triage outcome.
28	UNCLEAR: Insufficient information.		
29	UNCLEAR: Insumcient information.		
30			
31			
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34 25			
35 36			
30 37			
38			
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46			
47			

А.	Risk of bias	
	a. Was the universal definition of AMI used to define the reference standard?	Yes / No / Unclear
	b. Was the reference standard adjudicated without knowledge of the telephone triage?	Yes / No / Unclear
	Could the reference standard, its conduct, or its interpretation have introduced bias?	Low / High / Unclear
в.	Concerns regarding applicability	
	Is there concern that the target condition as defined by the reference standard does not match the review question?	Low / High / Unclear
<u>0\</u>	<u>/ERALL:</u>	
	W: All patients underwent reference standard investigations for AMI including troponin sampling	g. The diagnosis of AMI was adjudicated by at least two eath (all cause), aortic dissection, pulmonary embolism and
	nsion pneumothorax	
te HI co		
te HI co wi	nsion pneumothorax GH: Outdated definition for AMI or definition inconsistent with the universal definition. The defin re components specified above. Follow up procedure raises significant concerns about the possi	
te HI co wi	nsion pneumothorax GH: Outdated definition for AMI or definition inconsistent with the universal definition. The defin re components specified above. Follow up procedure raises significant concerns about the possil thout some assurance that this method would capture all relevant events).	
te HI co wi	nsion pneumothorax GH: Outdated definition for AMI or definition inconsistent with the universal definition. The defin re components specified above. Follow up procedure raises significant concerns about the possil thout some assurance that this method would capture all relevant events).	

1 2 3		
5	Domain 4: Flow and timing	
6	A. Risk of bias	
7 8	a. Did all included patients receive an appropriate reference standard?	Yes / No / Unclear
9	b. Did patients receive the same reference standard?	Yes / No / Unclear
10	c. Follow-up procedure was sufficiently long to not miss relevant adverse events?	Yes / No / Unclear
11 12	d. Did no significant loss to follow up/exclusion due to incomplete records occur?	Yes / No / Unclear
13 14 15	Could the patient flow have introduced bias?	Low / High / Unclear
16 17	LOW: All patients underwent reference standard investigations for AMI including troponin samp	ling. Patients were followed up through their subsequent
18 19	inpatient course or for at least 7 days.	
20 21	HIGH: Not all patients were subjected to appropriate reference standard investigations including	troponin sampling.
22 23	UNCLEAR: Insufficient information. This option includes lack of detail about troponin testing and	
24		
25		
26 27		
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## Table 1: QUADAS-2 assessment of included studies ( ✓ Low Risk, green, ✗ High Risk, red, ? Unclear Risk, amber)

Study RISK OF BI		RISK OF BIAS			APPLICABILITY CONCERNS			OVERALL		
	PATIENT	TELEPHONE	AMI /SAE	FLOW AND	PATIENT	TELEPHONE	AMI /SAE	PATIENT	TELEPHONE	AMI /SAE
9	SELECTION	TRIAGE	ALLOCATION	TIMING	SELECTION	TRIAGE	ALLOCATION	SELECTION	TRIAGE	ALLOCATION
1										
2										
3										
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### The PRISMA for Abstracts Checklist

TITLE	CHECKLIST ITEM	REPORTED ON PAGE #
1. Title:	Identify the report as a systematic review, meta-analysis, or both.	1
BACKGROUND		
2. Objectives:	The research question including components such as participants, interventions, comparators, and outcomes.	2 (objectives)
METHODS		
3. Eligibility criteria:	Study and report characteristics used as criteria for inclusion.	2 (eligibility criteria)
4. Information sources:	Key databases searched and search dates.	2 (data sources)
5. Risk of bias:	Methods of assessing risk of bias.	2 (data extraction & synthesis)
RESULTS		
6. Included studies:	Number and type of included studies and participants and relevant characteristics of studies.	3 (results)
7. Synthesis of results:	Results for main outcomes (benefits and harms), preferably indicating the number of studies and participants for each. If meta-analysis was done, include summary measures and confidence intervals.	3 (results)
8. Description of the effect:	Direction of the effect (i.e. which group is favoured) and size of the effect in terms meaningful to clinicians and patients.	N/A (no meta- analysis)
DISCUSSION		
9. Strengths and Limitations of evidence:	Brief summary of strengths and limitations of evidence (e.g. inconsistency, imprecision, indirectness, or risk of bias, other supporting or conflicting evidence)	3 (conclusions
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10. Interpretation:	General interpretation of the results and important implications	3 (conclusic
OTHER		
11. Funding:	Primary source of funding for the review.	4 (funding
12. Registration:	Registration number and registry name.	4 (registrati

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