

Quality indicators for acute myocardial infarction: A position paper of the Acute Cardiovascular **Care Association**

European Heart Journal: Acute Cardiovascular Care 1-26 © The European Society of Cardiology 2016 Reprints and permissions: sagepub.co.uk/iournalsPermissions.nav DOI: 10.1177/2048872616643053 acc.sagepub.com



Francois Schiele¹, Chris P Gale², Eric Bonnefoy³, Frederic Capuano⁴, Marc J Claeys⁵, Nicolas Danchin⁶. Keith AA Fox⁷, Kurt Huber⁸, Zaza lakobishvili⁹, Maddalena Lettino¹⁰, Tom Quinn¹¹, Maria Rubini Gimenez¹², Hans E Bøtker¹³, Eva Swahn¹⁴, Adam Timmis¹⁵, Marco Tubaro¹⁶, Christiaan Vrints⁵, David Walker¹⁷, Doron Zahger¹⁸, Uwe Zeymer¹⁹ and Hector Bueno²⁰

Abstract

Evaluation of quality of care is an integral part of modern healthcare, and has become an indispensable tool for health authorities, the public, the press and patients. However, measuring quality of care is difficult, because it is a multifactorial and multidimensional concept that cannot be estimated solely on the basis of patients' clinical outcomes. Thus, measuring the process of care through quality indicators (QIs) has become a widely used practice in this context. Other professional societies have published QIs for the evaluation of quality of care in the context of acute myocardial infarction (AMI), but no such indicators exist in Europe. In this context, the European Society of Cardiology (ESC) Acute Cardiovascular Care Association (ACCA) has reflected on the measurement of quality of care in the context of AMI (ST segment elevation myocardial infarction (STEMI) and non-ST segment elevation myocardial infarction (NSTEMI)) and created a set of Qls, with a view to developing programmes to improve quality of care for the management of AMI across Europe. We present here the list of QIs defined by the ACCA, with explanations of the methodology used, scientific justification and reasons for the choice for each measure.

¹University Hospital of Besancon, EA3920 University of Franche-Comté, Besançon, France

- ²Leeds Institute of Cardiovascular and Metabolic Medicine, University of Leeds, Leeds; York Teaching Hospital NHS Foundation Trust, York, UK
- ³Unité de soins intensifs cardiologiques, Hôpital Cardiologique Louis-Pradel, Bron, France
- ⁴Haute Autorité de Santé, Saint Denis La Plaine, France
- ⁵Cardiology Department, University Hospital Antwerp, Edegem, Belgium
- ⁶Assistance Publique-Hôpitaux de Paris (AP-HP); Hôpital Européen Georges Pompidou (HEGP), Department of Cardiology, Paris, France; Université Paris-Descartes, Paris, France
- ⁷Centre for Cardiovascular Science, University of Edinburgh, Edinburgh, UK
- 83rd Medical Department, Cardiology and Intensive Care Medicine, Wilhelminenhospital, Vienna, Austria
- ⁹Department of Cardiology, Rabin Medical Center, Petah Tikva, Israel
- ¹⁰Humanitas Research Hospital, Rozzano (Milan), Italy
- ¹¹Faculty of Health and Medical Sciences, University of Surrey, Guildford, UK

¹²Department of Cardiology and Cardiovascular Research Institute Basel (CRIB), University Hospital Basel, Basel

- ¹³Department of Cardiology, Aarhus University Hospital, Aarhus, Denmark
- ¹⁴Department of Cardiology and Department of Medical and Health Sciences, Linköping University, Sweden
- ¹⁵National Institute for Health Research Biomedical Research Unit, Barts Heart Centre, London, UK
- ¹⁶Ospedale San Filippo Neri, Rome, Italy
- ¹⁷East Sussex Healthcare, Conquest Hospital, Hastings, UK
- ¹⁸Department of Cardiology, Soroka University Medical Center, Faculty of Health Sciences, Ben-Gurion University of the Negev, Beer-Sheva, Israel
- ¹⁹Klinikum Ludwigshafen and Institut für Herzinfarktforschung Ludwigshafen, Ludwigshafen, Germany
- ²⁰Centro Nacional de InvestigacionesCardiovasculares (CNIC), Cardiology Department, Hospital Universitario 12 de Octubre, and Universidad Complutense de Madrid, Madrid, Spain

Corresponding author:

Francois Schiele, Department of Cardiology, EA3920, University Hospital Jean Minjoz, Boulevard Fleming, 25000 Besançon, France. Email: francois.schiele@Univ-fcomte.fr

Keywords

Quality of care, acute cardiovascular care, quality indicators, myocardial infarction, centre organisation, reperfusion, risk assessment, anti-thrombotic treatment, patient satisfaction, discharge treatment, composite indicator

Date received: 15 March 2016; accepted: 15 March 2016

Introduction

In a report published in 2001, the Institute of Medicine stated that 'in its current form, habits and environment, American healthcare is incapable of providing the public with the quality health care it expects and deserves' (p.43).¹ Evaluation of the quality of care is an integral part of modern healthcare, and has become an indispensable tool, much sought after by health authorities, the general public, the press and even patients themselves. However, measuring the quality of care is difficult, because it cannot be estimated solely on the basis of patients' clinical outcomes, even if achieving a favourable outcome for the greatest number of patients possible is the ultimate goal of high quality care.

Thus, measuring the process of care through quality indicators (QIs) or performance measures (PMs) has become a widely used practice in this context. The American College of Cardiology (ACC) and the American Heart Association (AHA) have considerable experience in the measurement of quality of care in the setting of acute myocardial infarction (AMI). Indeed, these two organisations have jointly published several Task Force documents and position papers with precise definitions of QIs and PMs that can be used to describe and evaluate management and outcomes of patients with acute coronary syndrome (ACS).² Furthermore, they have published several documents on the optimal methodology for defining QIs and PMs,^{3,4} the statistics suitable for public reporting,5 and the use of composite indicators.6 The first set of PMs for AMI published by the ACC/AHA was released in 2006,7 and updated in 2008,8 and a position paper specific to coronary reperfusion published in 2008.9

The lack of standard definitions of QIs validated by the European Society of Cardiology (ESC) for clinical situations such as AMI or heart failure (HF) is in stark contrast with the professional societies of cardiology in the USA. Thus, the question arises as to whether it is time for the ESC to define QIs for AMI, which would be in line with the ESC's own recommendations for the management of these conditions? The ESC Acute Cardiovascular Care Association (ACCA) has reflected on the measurement of quality of care in the context of AMI and aimed to create QIs, with a view to developing programmes to improve quality of care for the management of AMI across Europe.

Objectives

The objective of the ACCA Quality of Care Working Group was to define suitable QIs for the management of AMI, with or without ST segment elevation.

Scope

Specifically, in an attempt to improve the management of patients hospitalised with AMI, the ACCA proposes standardisation of the evaluation of quality of care across all centres in Europe. Whilst the link between assessment and improvement of quality of care is a matter of debate, the hypothesis that improvement in quality partially relates to internal or external assessment is supported by observational data.¹⁰ Either way, the Quality of Care Working Group's scope was to select not only QIs based on existing guidelines, but also to incorporate measures whose implementation is deemed to be important, like centre facilities, or patient satisfaction. Lastly, a set of simple and reliable QIs can be used to evaluate the conformity of actual practices with the ESC guidelines for management in AMI.

The definition of QIs relies on the existence of treatments and strategies of proven efficacy, as well as on highgrade recommendations. These prerequisites are satisfied in several clinical situations, such as AMI, acute HF, pulmonary embolism, as well as chronic conditions including stable angina, coronary revascularisation, atrial fibrillation, cardiovascular prevention, peripheral arterial disease and cardiac rehabilitation.

Membership of the Quality of Care Working Group and selection of the domains of care

Under the supervision of the Board of the ACCA, a Quality of Care Working Group was formed and comprised international experts selected for their expertise in the management of patients with ACS. All members were invited to participate in the selection and definition of the QIs in the selected domains of management and were representative of the different European countries that comprise the ESC, and included members of the ACCA, members of the ESC Practice Guidelines Committee, and ad hoc members, experts in clinical practice, public health or statistics. The full list of the group leaders and members of the Quality of Care Working Group, with their respective area of expertise, is displayed in Table 1.

Seven different domains of care where quality should be assessed were defined, with one chairperson responsible for coordinating the discussions in each domain.

The aim of this selection was to extend the quality assessment beyond simply the process of care and its outcomes, by incorporating the full spectrum of the patient pathway, from organisation of care, to outcomes and the patient experience. The seven domains selected are

Members of the Country Area of expertise Quality WG		Area of expertise	Participation in QI domain		
Francois Schiele	France	ICCU, interventional cardiology, statistics, cardiovascular quality and outcomes	Chair of the Quality of Care WG Composite and outcomes (chair)		
Christopher Gale	UK	Statistics, cardiovascular quality and outcomes, public health	Secondary prevention, discharge treatme (chair)		
Eric Bonnefoy	France	iccu	Reperfusion-invasive strategies		
Marc Claeys	Belgium	ICCU, interventional cardiology	Composite and outcomes		
Frederic Capuano	France	Statistics, cardiovascular quality, public health	Composite and outcomes		
Nicolas Danchin	France	ICCU, statistics, cardiovascular quality and outcomes, public health	Reperfusion-invasive Strategies		
Keith AA Fox	UK	ICCU, statistics, cardiovascular quality and outcomes, public health	Reperfusion-invasive strategies		
Kurt Huber	Austria	ICCU, interventional cardiology	Centre organisation (chair)		
Zaza lakobishvili	Israel	ICCU, internal medicine	Centre organisation		
Maddalena Lettino	Italy	ICCU, clinical cardiology cardiovascular quality and outcomes	In-hospital risk assessment (chair)		
Tom Quinn	UK	Emergency cardiovascular care, nursing	Patient's satisfaction (chair)		
Maria Rubini Gimenez	Switzerland	Clinical cardiology	Secondary prevention, discharge treatment		
Hans E Bøtker	Denmark	ICCU, statistics, cardiovascular quality and outcomes, public health	Centre organisation		
Eva Swahn	Sweden	Clinical cardiology	Secondary prevention, discharge treatment		
Adam Timmis	UK	Clinical cardiology, editor, cardiovascular quality and outcomes	In-hospital risk assessment		
Marco Tubaro	Italy	ICCU, clinical cardiology cardiovascular quality and outcomes	Anti-thrombotic during hospitalisation (chair)		
Christiaan Vrints	Belgium	Clinical cardiology, editor, cardiovascular quality and outcomes	Centre organisation		
David Walker	Belgium	Emergency cardiovascular care	Centre organisation		
Doron Zahger	Israel	ICCU, cardiovascular quality and outcomes	Anti-thrombotic during hospitalisation		
Uwe Zeymer	Germany	ICCU, interventional cardiology	Reperfusion-invasive strategies (chair)		
Hector Bueno	Spain	ICCU, interventional cardiology clinical cardiology, editor, cardiovascular quality and outcomes	In-hospital risk assessment		

Table I. List, area of expertise and participation in quality indicator (QI) domains of the members of the Quality of Care Working Group (WG) (by order of authorship).

ICCU: Intensive Cardiac Care Units.

relevant to the clinical situation of AMI, namely: (a) centre organisation; (b) reperfusion/invasive strategy; (c) in-hospital risk assessment; (d) anti-thrombotic treatment during hospitalisation; (e) discharge treatments; (f) patient satisfaction; and (g) composite QIs (CQIs) and outcomes.

Methodology

Selection of candidate QIs

A set of 45 candidate QIs were identified, based on existing QIs and international guidelines for the management of ST segment elevation myocardial infarction (STEMI) and non-ST segment elevation myocardial infarction (NSTEMI). These covered all seven pre-selected dimensions. The 45 candidate QIs were selected by the Quality of Care Working Group using an online survey circulated to the whole group and completed by all members. Each candidate QI was

graded on a scale of 1–5 according to the following criteria: (a) supported by evidence/guidelines; (b) interpretability; (c) actionability and room for improvement; (d) feasibility of assessment; and (e) global fit.

Definition of QIs

For each domain, one or more 'main' QIs, as well as one or more 'secondary' QIs were finally retained. The grading of the QIs and the final selection of those to be retained was based on the feasibility and reliability of the assessment of the QI. The main QIs were selected because they were considered an essential element, mandatory for basic assessment. Conversely, the secondary QIs were considered as complementary measures that could be used to perform more advanced assessment, and/or may only be suitable for use in certain centres. The list of the main and secondary QIs for each domain is presented in Table 2, with details of **Table 2.** Summary of the quality indicators (QIs): definition, numerator and denominator, rationale, support from guidelines and method of reporting.

1.1 Centre organisation. Main QI: the centre should be part of a network organisation with written protocols for rapid and efficient management covering the following points:

Single emergency phone number for the patient to be connected with a medical system for triage. Pre-hospital interpretation of ECG for diagnosis and decision for immediate transfer to a centre with catheterisation laboratory facilities.

Pre-hospital activation of the catheterisation laboratory.

Numerator: all centres that are part of a network organisation.

Denominator: all centres.

Clinical rationale: to improve speed and efficiency of pre-hospital care and reperfusion for STEMI patients.

Sources of data: administrative data.

Corresponding guidelines:

- Network organisation: ESC STEMI GL class I, level B.
- Written protocol: ESC STEMI GL class I, level C.
- Single phone call: no recommendation.
- Pre hospital interpretation of ECG: ESC STEMI GL class I, level B.
- Pre hospital activation of the catheterisation laboratory: ESC STEMI GL class IIa, level B.

Method of reporting: qualitative measure per centre.

1.2 Centre organisation. Secondary QI (1): routine assessment of relevant times for the reperfusion process in STEMI patients (i.e. times from 'call to first medical contact', 'first medical contact to door', 'door to arterial access'; and 'door-in-door-out' for centres without a catheterisation laboratory on site).

Numerator: all centres with routine assessment of relevant intervals for the reperfusion process.

Denominator: all centres.

Clinical rationale: to identify system inefficiencies and steps where reduction in time for reperfusion for STEMI patients is possible. *Sources of data:* administrative data.

Corresponding guidelines:

- Routine assessment of time to reperfusion for STEMI patients (time 'call to first medical contact', first medical contact to door', door to arterial access'): ESC STEMI GL, class I, level C.
- All hospital must record and monitor delay times: ESC STEMI GL, class I, level B.
- Direct access to catheterization laboratory, bypassing the emergency department: ESC STEMI GL class IIa, level B. **Method of reporting:** qualitative measure (per centre).

1.3 Centre organisation. Secondary QI (2): the centre should participate in a regular registry or programme for quality assessment.

Numerator: centres participating in a registry.

Denominator: all centres.

Clinical rationale: to allow assessment of quality of care.

Sources of data: administrative data, registry data.

Corresponding guidelines:

- The centre should participate regularly in a registry for quality assessment: ESC STEMI GL, class I, level C.
- Development of regional or national programmes to measure performance indicators systematically and provide feedback to individual hospitals: proposed as PM by ESC GL NSTE-ACS.

Method of reporting: qualitative measure (per centre).

2.1 Reperfusion-invasive strategy. Main (STEMI 1): proportion of STEMI patients reperfused among those eligible (onset of symptoms to diagnosis <12 h).

Numerator: number of STEMI patients with onset of symptoms to diagnosis <12 h who receive reperfusion therapy. **Denominator:** all STEMI patients eligible for reperfusion (onset of symptoms to diagnosis <12 h, without contraindication or patient refusal).

Clinical rationale: all STEMI patients (within the first 12 h) should receive reperfusion therapy.

Sources of data: administrative data and medical records.

Corresponding guidelines: ESC STEMI GL: reperfusion <12 h: class I, level A.

Method of reporting: proportion (standard error).

- 2.2 Reperfusion-invasive strategy. Main QI (STEMI 2): proportion of patients with timely reperfusion. Timely is defined as:
- For patients treated with fibrinolysis: <30 min from Fist Medical Contact FMC to needle.
- For patients treated with primary PCI and admitted to centres with catheterisation laboratory facilities: <60 min from door-to-arterial access for reperfusion with PCI.
- For transferred patients: door-in-door-out time of <30 min.

Numerator: number of STEMI patients treated with primary PCI within the above delays.

Denominator: all STEMI patients eligible for reperfusion by primary PCI (onset of symptoms to diagnosis <12 h, without contraindication or patient refusal).

Clinical rationale: time to effective mechanical reperfusion should be reduced.

Sources of data: pre-hospital and hospital medical records, ECG, angiography.

Corresponding guidelines: timely reperfusion:

- For patients treated with fibrinolysis: <30 min FMC to needle: ESC STEMI GL, class I, level B.
- For patients admitted to centres with catheterisation laboratory facilities: <60 min door-to-arterial access for reperfusion with PCI ESC STEMI GL, class I, level B.
- For patients transferred to a non PCI-capable centre for primary PCI: should bypass the emergency department: ESC STEMI GL, class IIa, level B. <30 min door-in-door-out: ESC revascularisation GL, class IIa, level B.

Method of reporting: proportion (standard error).

2.3 Reperfusion-invasive strategy. Main QI (NSTEMI): proportion of patients with NSTEMI and no contra indication who receive coronary angiography within 72 h after admission.

Numerator: number of NSTEMI patients at high-intermediate ischaemic risk undergoing coronary angiography within 72 h after the diagnosis.

Denominator: all NSTEMI patients at high-intermediate ischaemic risk without contraindications or patient refusal. **Clinical rationale:** NSTEMI patients at high risk should be treated with early invasive strategy. Early is defined as \leq 72 h after admission. **Sources of data:** medical records, ECG, angiography.

Corresponding guidelines: invasive strategy \leq 72 h for high-intermediate risk (in patients with NSTEMI and one intermediate-risk criteria (diabetes mellitus, renal dysfunction (eGFR<30 ml/min/1.72 m²), LVEF \leq 0.40, congestive heart failure, recent PCI, prior CABG, GRACE risk score >140) or recurrent symptoms or ischaemia on non invasive testing: ESC NSTE-ACS GL class I, level A. **Method of reporting:** proportion (standard error).

2.4 Reperfusion-invasive strategy. Secondary QI (STEMI): The time between the FMC and arterial access (absolute value) for primary PCI.

Clinical rationale: improve speed and efficiency of pre-hospital care and reperfusion for STEMI patients.

Sources of data: pre-hospital and hospital medical records, ECG, angiography.

Corresponding guidelines: ESC guidelines for STEMI 2012: routine assessment of times for reperfusion: class I, level B. **Method of reporting:** median time

3.1 In hospital risk assessment. Main QI (1): the proportion of patients with NSTEMI in whom ischaemic risk assessment using the GRACE risk score is performed. GRACE score should be assessed and the numerical value of the score recorded for all patients admitted with suspected NSTEMI.

Numerator: number of NSTEMI patients who have been stratified according to the GRACE risk score.

Denominator: number of NSTEMI patients.

Clinical rationale: NSTEMI patients at high ischaemic risk should be treated with early invasive strategy.

Sources of data: medical records.

Corresponding guidelines:

- Prognostic risk assessment: the use of risk scores for estimating prognosis is recommended: ESC NSTE-ACS GL, class I, level A.
- GRACE score: recommendations class IA depending on the value of the GRACE score.

Method of reporting: proportion (standard error).

3.2 In hospital risk assessment. Main QI (2): proportion of patients admitted with STEMI or NSTEMI who have bleeding risk assessment using the CRUSADE bleeding score. The CRUSADE bleeding score should be assessed and the numerical value of the score recorded for all patients admitted with STEMI or NSTEMI.

Numerator: number of STEMI or NSTEMI patients who have been stratified according to the CRUSADE bleeding score. **Denominator:** number of STEMI or NSTEMI patients.

Clinical rationale: STEMI and NSTEMI patients at high bleeding risk should be treated with caution regarding anti-thrombotic treatment.

Sources of data: medical records.

Corresponding guidelines:

- Prognostic risk assessment: ESC NSTE-ACS GL class I, level A.
- CRUSADE bleeding score: ESC NSTE-ACS GL: class Ilb, level B.

Method of reporting: proportion (standard error).

3.3 In hospital risk assessment. Main QI (3): proportion of patients with assessment of LVEF before discharge. LVEF should be assessed and the numerical value recorded for all patients admitted with STEMI or NSTEMI. Numerator: number of AMI patients with measured LVEF.

Denominator: all AMI patients.

Clinical rationale: all AMI patients with LVEF \leq 0.40 need specific medical treatment.

Sources of data: medical records, echocardiogram, ECG.

Corresponding guidelines:

• LVEF assessment: ESC STEMI GL class I, level A.

• LVEF assessment: ESC NSTE-ACS GL class I, level C.

Method of reporting: proportion (standard error).

- 4.1 Anti-thrombotics during hospitalisation. Main QI (1): proportion of patients with 'adequate P2Y₁₂ inhibition' defined as: (number of patients discharged with prasugrel or ticagrelor or clopidogrel)/(patients eligible). Eligible is defined as follows:
- For ticagrelor: AMI patients without previous haemorrhagic stroke, high bleeding risk, fibrinolysis or oral anticoagulation.
- For prasugrel: PCI-treated AMI patients without previous haemorrhagic or ischaemic stroke, high bleeding risk (patients ≥75 years and/or <60 kg body weight are also considered as high bleeding risk), fibrinolysis or oral anticoagulation.
- For clopidogrel: no indication for prasugrel or ticagrelor and no high bleeding risk.

Numerator: number of STEMI and NSTEMI patients with 'adequate P2Y12 inhibitor' at discharge.

Denominator: STEMI and NSTEMI patients alive at discharge and without contraindications to P2Y12 inhibitors

Clinical rationale: superiority of prasugrel and ticagrelor over clopidogrel in selected patients.

Sources of data: medical records.

Corresponding guidelines: ESC STEMI, ESC NSTE-ACS and ESC revascularisation GL:

- Eligibility for ticagrelor: class I, level B.
- Eligibility for prasugrel: class I, level B.
- Eligibility for clopidogrel: class I, level A.

Method of reporting: proportion (standard error).

4.2 Anti-thrombotics during hospitalisation. Main QI (2): proportion of patients with NSTEMI treated with fondaparinux, unless candidate for immediate (≤ 2 h) invasive strategy or with eGFR<20 ml/min.

Numerator: number of NSTEMI patients with eGFR \ge 20 ml/min, not candidates for urgent invasive strategy, treated with fondaparinux. **Denominator:** all NSTEMI patients with eGFR \ge 20 ml/min, not candidates for urgent invasive strategy.

Clinical rationale: better risk/benefit profile of fondaparinux in NSTEMI patients.

Sources of data: medical records.

Corresponding guidelines: fondaparinux most favourable risk benefit profile (for NSTEMI patients not candidate for urgent angiography): ESC NSTE-ACS and Revascularisation GL class I, level B.

Method of reporting: proportion (standard error).

4.3 Anti-thrombotics during hospitalisation. Secondary QI: proportion of patients discharged on dual antiplatelet therapy, defined as: (number of patients discharged on dual antiplatelet therapy)/(number of patients with AMI without clear and documented contraindication).

Numerator: number of STEMI and NSTEMI patients, without contra indication, discharged with dual antiplatelet therapy.

Denominator: all STEMI and NSTEMI patients, without contra indications to dual antiplatelet therapy.

Clinical rationale: benefit of DAPT over single antiplatelet therapy for 12 months.

Sources of data: medical records.

Corresponding guidelines: Irrespective of the revascularisation strategy, a P2Y₁₂ inhibitor is recommended in addition to aspirin for patients with AMI: ESC STEMI GL, class I, level A; ESC NSTE-ACS GL, class I, level A.

Method of reporting: proportion (standard error).

5.1 Secondary prevention-discharge treatment. Main QI (1): proportion of patients with AMI discharged on statins, unless contra indicated, at high intensity (defined as atorvastatin ≥40 mg or rosuvastatin ≥20 mg).

Numerator: the number of patients with AMI who receive high intensity statin therapy at discharge.

Denominator: STEMI and NSTEMI patients alive at discharge and without contraindications, refusal, side effects, allergy, or history of intolerance to high-intensity statin therapy.

Clinical rationale: the use of high intensity statins is associated with reduced risk of recurrent cardiovascular events and mortality following AMI.

Sources of data: medical records.

Corresponding guidelines: statins high intensity as early as possible, unless contra indication: ESC STEMI GL, class I, level A, ESC NSTE-ACS GL, class I, level A.

Method of reporting: Proportion (standard error).

5.2 Secondary prevention-discharge treatment. Secondary QI (I): proportion of patients with AMI and clinical evidence of heart failure or a LVEF ≤0.40 who are discharged on ACEI (or ARBs if intolerant of ACEI) unless contraindicated.

Numerator: the number of patients with AMI who have heart failure or a LVEF <0.40, and who receive an ACEI/ARB before discharge. **Denominator:** all AMI patients who have heart failure or a LVEF <0.40, and who are eligible for ACEI/ARBs (no hypotension, acute renal failure, hyperkalaemia, contraindications, refusal, side effects or allergy).

Clinical rationale: the use of ACEIs/ARBs is associated with reduced mortality following AMI in patients with heart failure or left ventricular systolic dysfunction.

Sources of data: medical records.

Corresponding guidelines: ACE inhibitor in patients with LVEF < 0.40 or heart failure, hypertension or diabetes: ESC STEMI GL, class I, level A, ESC NSTE-ACS GL class I, level A

Method of reporting: proportion (standard error).

5.3 Secondary prevention-discharge treatment. Secondary QI (2): proportion of patients with AMI and clinical evidence of heart failure or an LVEF≤0.40 who are discharged on beta-blockers, unless contraindicated.

Numerator: the number of patients with AMI who have heart failure or a LVEF \leq 0.40 and receive a beta-blocker before discharge. **Denominator:** all AMI patients who have heart failure or a LVEF \leq 0.40, and are eligible for beta-blockers (no evidence of a low output state, increased risk for cardiogenic shock, PR interval >0.24 s, second- or third-degree heart block, active asthma, or reactive airways disease).

Clinical rationale: the use of beta-blockers in patients with AMI and who have heart failure or left ventricular systolic dysfunction is associated with a mortality benefit.

Sources of data: medical records.

Corresponding guidelines: beta-blocker therapy in patients with LVEF \leq 0.40, unless contraindicated: ESC STEMI GL, class I, level A, ESC NSTE-ACS GL, class I, level A

Method of reporting: proportion (standard error).

- 6.1 Patient satisfaction. Main QI: feedback regarding the patient's experience systematically collected in an organised way from all patients. It should include the following points:
- Pain control.
- Explanations provided by doctors and nurses (about the coronary disease, the benefit/risk of the discharge treatment, and medical follow-up).
- Discharge information regarding what to do in case of recurrence of symptoms and recommendation to attend a cardiac rehabilitation programme (including smoking cessation and diet counselling).

Numerator: number of STEMI and NSTEMI patients discharged alive with feedback collected.

Denominator: STEMI and NSTEMI patients discharged alive.

Clinical rationale: patient satisfaction must be considered in assessment of quality of care. Relation between patient satisfaction and adherence to guidelines, and with mortality.

Sources of data: administrative data and medical records.

Corresponding guidelines: no ESC GL to support this QI. Review paper by Anker et al. published in Eur Heart J in 2014.

- Participation in a well-structured cardiac rehabilitation programme: ESC NSTE-ACS GL, class IIa, level A.
- Smoking cessation advice/counselling: ESC STEMI GL, class I, level C; proposed as PM by ESC GL NSTE-ACS 2015, no recommendation.
- Enrolment in a secondary prevention/cardiac rehabilitation programme: : proposed as PM by ESC NSTE-ACS GL, 2015, no recommendation.

Method of reporting: proportion (standard error).

7.1 Composite QI. Main composite QI: opportunity based CQI, with the following individual indicators:

- The centre is part of a network organisation.
- Proportion of patients reperfused among eligible (STEMI with FMC<12 h after onset of pain).
- Coronary angiography in STEMI and NSTEMI patients at high ischaemic risk and without contraindications.
- Ischaemic risk assessment using the GRACE risk score in NSTEMI patients.
- Bleeding risk assessment using the CRUSADE risk score in STEMI and NSTEMI patients.
- Assessment of LVEF before discharge.
- Low dose aspirin (unless high bleeding risk or oral anticoagulation).
- Adequate P2Y₁₂ inhibition (as defined in the treatment during hospitalisation section).
- ACEI (or ARB if intolerant of ACEI) in patients with clinical evidence of heart failure or an LVEF < 0.40.
- Beta-blockers (unless clear contraindication) in patients with clinical evidence of heart failure or an LVEF < 0.40.
- High intensity statins.
- Feedback regarding the patient's experience and quality of care is systematically collected for all patients.

Numerator: all AMI patients discharged: sum of points (one point for each individual indicator, all individual indicators are weighted equally).

Denominator: all AMI patients discharged: sum of points (one point for each applicable indicator, according to patient and centre characteristics).

Clinical rationale: relation between CQI and mortality.

Sources of data: administrative data and medical records, statistical computation.

Corresponding guidelines: no recommendation.

Method of reporting: mean value (95% confidence interval).

7.2. Composite QI. Secondary composite QI: all-or-none CQI based on 3 or 5 components, according to the LVEF. *Calculated on 3 individual QIs in patients without heart failure and with LVEF>0.40.*

- Low-dose aspirin.
- P2Y₁₂ inhibitor (unless documented contraindication).
- High-intensity statins.

Calculated on 5 individual QIs in patients with heart failure or with LVEF < 0.40.

- Low-dose aspirin.
- P2Y₁₂ inhibitor (unless documented contraindication).
- High-intensity statins.
- ACEI (or ARB if intolerant to ACEI) in patients with clinical evidence of heart failure or LVEF≤0.40.

Beta-blockers (unless clear contraindication) in patients with clinical evidence of heart failure or LVEF≤0.40.

Clinical rationale: relation between CQI and mortality.

Sources of data: medical records, statistical computation.

Corresponding guidelines: no recommendation.

Method of reporting: at patient level, can be 0 or 1. At centre level, mean value (95% confidence interval).

7.3 Outcome QI. Secondary outcome QI: 30-day mortality rate adjusted for the GRACE 2.0 risk score.

Numerator: all AMI patients who died within 30 days after admission, with assessment of the GRACE risk score.

Denominator: all AMI patients with assessment of the GRACE risk score and 30-day follow-up.

Sources of data: medical records, registry data.

Clinical rationale: 30-day adjusted mortality is an established criterion for quality of care in AMI patients.

Corresponding guidelines: no recommendation.

Method of reporting: proportion (standard error).

ACEI: angiotensin-converting enzyme inhibitors; AMI: acute myocardial infarction; ARB: angiotensin receptor blocker; CQI: composite quality indicator; ESC: European Society of Cardiology; FMC: first medical contact; LVEF: left ventricular ejection fraction: NSTEMI: non-ST segment elevation myocardial infarction; PCI: percutaneous coronary intervention; QI: quality indicator; STEMI: ST segment elevation myocardial infarction.

the numerator and denominator, rationale, support from guidelines and method of reporting. Figure 1 presents a summary of the QI in each domain, with the corresponding ESC guidelines.

Support

The work is exclusively supported by the ACCA and the ESC. No commercial support was received for the development of the QIs.

QIs for STEMI/NSTEMI

Centre organisation

Dimensions of care. For patients with AMI, early diagnosis, pre-hospital medical care and quick access to revascularisation through direct admission to cardiology centres with catheterisation facilities available 24/7 have all been shown to reduce time to reperfusion, which in turn is associated with lower mortality.¹¹ To this end, organised systems of care such as structured networks are needed to determine the optimal pathways of care based on local circumstances,

centre characteristics and transfer capabilities in the area.^{12–14} Organisation of networks of care has been shown to be effective in reducing times to reperfusion, through rapid diagnosis with expeditious ECG recording and interpretation, risk assessment, safe transfer and rapid access to reperfusion strategies.^{15–19} The main QI for network organisations was based on four organisational points deemed to be the most important in clinical terms, as well as being easy to implement in practice and easy to assess. Although non-written collaborations can be efficient in practice, the ACCA Quality of Care Working Group considers that only centres with a written protocol that has been discussed and signed by both the centre and the pre-hospital system should be regarded as participating in a network for the purposes of assessing quality of care.

Clinical relevance. Organisation of care has an important impact on the implementation of recommendations for times to reperfusion.¹³ Depending on the local environment, bringing the patient to the centre in a timely manner or bringing the treatment to the patient through the administration of intravenous fibrinolytics in the pre-hospital setting have been extensively discussed. Both methods

Domain of Care	Quality Indicator	Support from ESC guidelines		
Center Organization	Main QI: The centre should be part of a Network Organization with written protocols for rapid and efficient management covering the following points:	Network: ESC GL, Class I, level B Written protocol: ESC STEMI GL Class I, level C		
8	 Single emergency phone number for the patient to be connected to a medical system for triage 	Single phone number : No ESC GL to support this QI.		
	 Pre-hospital interpretation of ECG for diagnosis and decision for immediate transfer to a center with catheterization laboratory facilities, bypassing the Emergency Department. 	Pre hospital interpretation of ECG: ESC STEMI GL, Class I level B		
	•Pre-hospital activation of the catheterization laboratory	Pre hospital easy activation of the catheterization laboratory : ESC STEMI GL, level B		
	Secondary QI (1): Routine assessment of relevant times for the reperfusion process in STEMI patients (i.e. times from "call to first medical contact", "first medical contact to door", "door to arterial access" and "door-in door-out" for centers without a catheterization laboratory on site).	Routine assessment of time to reperfusion for STEMI patients (time "call to first medical contact", first medical contact to door", door to device"): ESC STEMI GL, Class I, level C All hospital must record and monitor delay times: ESC STEMI GL, Class I, level B		
	Secondary QI (2): The center should participate in a regular registry or program for quality assessment.	The center should participate regularly in a registry for quality assessment: ESC STEMI GL, Class I, level C Development of regional or national programmes to measure performance indicators systematically and provide feedback to individual hospitals : proposed as PM by ESC GL NSTE-ACS 2015		
Reperfusion- Invasive	Main Ql (STEM1 1): Proportion of STEMI patients reperfused among eligible (onset of symptoms to diagnosis < 12h) Main Ql (STEMI 2): Proportion of patients with timely reperfusion. Timely is defined	Reperfusion STEMI patients Onset up to 12h: ESC STEMI GL, Class I, level A Timely reperfusion:		
Strategy	as: • For patients treated with fibrinolysis: < 30 mins from diagnosis (FMC) to needle • For patients treated with primary PCI and admitted to centres with catheterization laboratory facilities: < 60 mins from door to arterial access for reperfusion with PCI • For transferred patients: door-in door-out time of < 30 mins	 For patients treated with fibrinolysis: <60 min FMC to needle: ESC STEMI GL, class I, level B For patients admitted to centers with catheterization laboratory facilities: <60 min door to balloon (passage of wire) for reperfusion with PCLESC STEMI GL, class I, level B For patients transferred to a non PCL-capable centre for primary PC : should bypass the emergency department: ESC STEMI GL, Class IIa, level B For grindens is depared to generate FOR Promotion and PCLESC STEMI GL, class IIa, level B 		
	Secondary QI (STEMI): the time between the diagnosis (FMC) and arterial access time (absolute value) for primary PCI.	<30 mins door-in-door out: ESC Revascularization GL, Class IIa, level B. All hospitals must record and monitor delay times: ESC STEMI GL, Class I, level B		
	Main QI (NSTEMI): Proportion of patients with NSTEMI, and no contra-indication, who receive coronary angiography within 72 hours after admission.	Invasive strategy in moderate-high risk patients: ESC NSTE-ACS GL, Class I, level A		
In Hospital Risk Assessment	Main QI (1): Proportion of patients with NSTEMI who have ischaemic risk assessment using the GRACE risk score. GRACE risk score should be assessed and the numerical value of the score recorded for all patients admitted with suspected NSTEMI.	The use of risk scores for estimating prognosis is recommended: ESC NSTE-ACS GL, Class I, level A		
	Main QI (2): Proportion of patients admitted with STEMI and NSTEMI who have bleeding risk assessment using the CRUSADE bleeding score. The CRUSADE bleeding score should be assessed and the numerical value of the score recorded for all patients	Use of the CRUSADE scorein patients undergoing coronary angiography: ESC NSTE-ACS GL Class IIb, level B		
	admitted with STEMI and NSTEMI. <u>Main QI (3):</u> Proportion of patients with STEMI and NSTEMI who have assessment of left ventricular ejection fraction. Left ventricular ejection fraction should be assessed and the numerical value recorded for all patients admitted with STEMI and NSTEMI.	Assessment of left ventricular ejection fraction: ESC STEMI GL class I, level B ESC NSTE-ACS GL class I, level B		

Domain of Care	Quality Indicator	Support from ESC guidelines
Anti thrombotics during Hospitalization	Main QI (1): Proportion of patients with "adequate P2Y ₁₂ inhibition" defined as: number of patients discharged with prasugrel or ticagrelor or clopidogrel / patients eligible. Eligible is defined as follows: For ticagrelor: AMI patients without previous haemorrhagic stroke, high bleeding risk, fibrinolysis or oral anticoagulation For prasugrel: PCI treated AMI patients without previous haemorrhagic or ischaemic stroke, high bleeding risk (patients ≥75 years or <60 kg body weight are also considered as high bleeding risk).	Ticagrelor in absence of contra indication for all patients regardless of initial strategy (i.e. patients without previous hemorrhagic stroke, high bleeding risk, oral anticoagulation) ESC NSTE-ACS GL, Class J, level B Prasugrel: in patients without previous hemorrhagic or ischemic stroke, high bleeding risk (patients ≥75 years, <60 kg body weight are also considered as high bleeding risk), oral anticoagulation, treated with PCI. ESC NSTE-ACS GL, Class J, level B Clopidogrel for patients who cannot receive ticagrelor or prasugrel or who require oral anticoagulation: ESC NSTE-ACS GL, Class J, level B
	Main QL(2): Proportion of patients with NSTEMI treated with fondaparinux, unless candidates for immediate (≤ 2 hours) invasive strategy, or with eGRR ≥ 20 ml/min.	Fondaparinux is recommended as having the most favourable efficacy/safety profile regardless of the management strategyESC NSTE-ACS GL, Class I, level B
	Secondary QI : Proportion of patients with AMI discharged on dual antiplatelet therapy / patients with AMI without clear and documented contra-indication	Irrespective of the revascularization strategy, a P2Y ₁₂ inhibitor is recommended in addition to aspirin for patients with AMI:ESC STEMI GL, Class I, level A, ESC NSTE-ACS GL, Class I, level A.
Secondary Prevention- Discharge Treatment	$\label{eq:linear_states} \begin{tabular}{lllllllllllllllllllllllllllllllllll$	Statins high intensity as early as possible, unless contra indication: ESC STEMI GL, Class I, level A, ESC NSTE-ACS GL, Class I, level A Betablocker therapy in patients with LVEF ≤ 0.40, unless contraindicated: ESC STEMI GL, Class I, level A, ESC NSTE-ACS GL, Class I, level A ACE inhibitor in patients with LVEF ≤ 0.40 or heart failure, hypertension ordiabetes: ESC STEMI GL, class I, level A, ESC NSTE-ACS GL Class I, level A Use of aspirin, ticagrelor/prasugrel/dopidogrel, statins, betablocker and ACE inhibitor (in patients with LVEF ≤ 0.40 or heart failure), enrollment in cardiac rehabilitation at discharge : proposed as PM by ESC GL NSTE-ACS 2015, no recommendation
Patient satisfaction	Main QI: Feedback regarding the patient's experience is systematically collected for all patients. This should include the following points: •Pain control •Explanations provided by doctors and nurses (about the coronary disease, the benefit/risk of the discharge treatment, and medical follow-up) •Discharge information regarding what to do in case of a recurrence of symptoms and recommendation to attend a cardiac rehabilitation program (including smoking cession and diet counseling)	No ESC GL to support this Ql. Review paper from Anker et al published in Eur Heart J in 2014 Participation in a well-structured cardiac rehabilitation programme: ESC NSTE-ACS GL, Class IIa, level A Smoking cessation advice/counselling: ESC STEMI GL, class I, level C; proposed as PM by ESC GL NSTE-ACS 2015, no recommendation Enrollment in a secondary prevention /cardiac rehabilitation programme: : proposed as PM by ESC NSTE-ACS GL, 2015, no recommendation

Figure 1. (Continued)

Domain of Care	Quality Indicator	Support from ESC guidelines
Composite and outcome QI	Main CQI : Opportunity based CQJ, with the following individual indicators: •The center is part of a network organization •Proportion of patients reperfused among eligible (STEMI with FMC <12 hours after onset of pain)	No ESC GL to support this QI. ESC NSTE-ACS GL proposes "Performance Measures", but only individual indicators, no composite indicator.

Figure 1. Summary of the quality indicators: definitions and support from guidelines.

ACEI: angiotensin-converting enzyme inhibitors; AMI: acute myocardial infarction; ARB: angiotensin receptor blocker; CQI: composite quality indicator; ESC: European Society of Cardiology; FMC: first medical contact; LVEF: left ventricular ejection fraction: NSTEMI: non-ST segment elevation myocardial infarction; PCI: percutaneous coronary intervention; PM: performance measure; QI: quality indicator; STEMI: ST segment elevation myocardial infarction.

are not exclusive and a well-organised network organisation should provide the most appropriate treatment in the shortest delay.²⁰ Several reports relate network experiences, and all have shown an improvement in the proportion of patients reperfused in a timely manner thanks to a single call number, a physician-staffed or trained paramedic ambulance crew and direct transfer to a percutaneous coronary intervention (PCI)-capable hospital with experienced cardiologists on call. The Vienna citywide system of care involves one academic and four non-academic centres, providing a physician-staffed ambulance and a guarantee that only experienced interventionists are on duty.21 The French SAMU organisation has a physician-staffed ambulance on site and can start pre-hospital fibrinolytic therapy, antithrombotic therapy, resuscitation and transfer directly to PCI-capable centres.²² Based on similar organisations, numerous other networks have been established in large cities,²³⁻²⁵ regions²³ or nations.²⁶

Specific aspects for potential Qls. Optimal treatment delivered within a minimal time frame has been shown to reduce mortality in STEMI patients and is thus strongly recommended by guidelines.^{27–29} The main effective components of a STEMI network are pre-hospital ECG recording and interpretation,^{30–33} pre-hospital activation of the catheterisation laboratory^{34,35} with a single call number,^{7,36} adequate selection of the mode of transportation,³⁷ and direct admission to the catheterisation laboratory.¹⁹ All these strategies have been identified as predictors of short door-to-balloon times.³⁸ This type of optimal collaboration within a STEMI network has been shown to increase both the proportion of patients treated by reperfusion and the proportion of patients with timely reperfusion.²⁰

Prospective monitoring of the times to management in STEMI patients. Since the time to reperfusion is clinically important and is used to define the QI for reperfusion, the different times to management need to be recorded. Although the most clinically important overall time span is the time from symptom onset to reperfusion (that is, the total ischaemic time), several intermediate times are required, such as: time of the call, time to first medical contact, time of the first ECG, time of arrival at the PCI-capable centre (door), time of the balloon (guidewire or other device) that restores patency in the infarct-related artery. Additionally, for patients admitted to non-PCI centres, the interval between time of admission and time of discharge for transfer to a PCI-capable centre (door-in-door-out time) needs to be recorded.

The ESC guidelines recommend participation in a national or international registry or a quality programme, such as the 'Stent for Life' initiative,³⁹ to record data regarding the actual management of patients admitted with AMI. Despite a lack of firm evidence that participation in a registry has an impact on quality of care, the ACCA Quality of Care Working Group considers that regular participation in a registry which assesses quality of care is an indicator of quality at a centre level.

Definition of	the main	and second	lary QI foi	' 'centre organisation'
---------------	----------	------------	-------------	-------------------------

Name of the main QI	The centre should be part of a network organisation with written protocols for rapid and efficient management covering the following points:				
	 Single emergency phone number for the patient to be connected with a medical system for triage. Pre-hospital interpretation of ECG for diagnosis and decision for immediate transfer to a centre with catheterisation laboratory facilities. Pre-hospital activation of the catheterisation laboratory. Direct access to catheterization laboratory, bypassing the emergency department. 				
Name of the secondary QI (1)	Routine assessment of relevant times for the reperfusion process in STEMI patients (i.e. recording actual times in order to assess 'call to first medical contact', 'first medical contact to door', 'door to arterial access' and 'door-in-door-out' for centres without catheterisation laboratory on site).				
Name of the secondary QI (2)	The centre should participate in a regular registry or programme for quality assessment.				

Agreement with guidelines and existing QIs

Main Ql. The guidelines for the management of STEMI issued by the ESC explicitly recommend that each centre receiving patients with suspected AMI should be part of a network organisation. According to the ESC guidelines, the main features of the network are (a) the existence of written protocols for risk stratification and adequate transportation, (b) prehospital triage with the aim of bypassing non-PCI hospitals, (c) immediate transportation to the catheterisation laboratory for eligible STEMI patients and (d) monitoring and immediate transfer of STEMI patients admitted to non-PCI centres. In view of the guidelines recommendations, the components of the structure-network QI are strongly supported.

The ACC/AHA PM for STEMI and NSTEMI do not include any structure-network QI. The Canadian Cardiovascular Society (CCS) PM refers to 'system indicators' that contain a pre-hospital 12-lead ECG.⁴⁰

Secondary Qls. The ESC guidelines for both STEMI and NSTE-ACS recommend participation in a survey or registry, both to record times to reperfusion among STEMI patients²⁸ and to record the degree of application of guidelines for all patients with STEMI and NSTE-ACS, as is already undertaken by a number of national initiatives, such as in Sweden (through the Swedish Web-system for Enhancement and Development of Evidence-based care in Heart disease Evaluated According to Recommended Therapies (SWEDEHEART)), the UK (Myocardial Infarction National Audit Project (MINAP)) registry, Germany, Italy and Israel on a regional basis, or through intermittent programmes in other countries (French Registry on Acute ST-elevation and non ST-elevation Myocardial Infarction (FAST-MI)). The underlying justification for this recommendation is that continuous monitoring of actual patient management may enhance the quality of treatment and minimise unwarranted variation in evidence-based care.

Reperfusion/invasive strategy

Dimensions of care. Invasive strategy, myocardial revascularisation, and the speed with which it is achieved, are key

elements in the management of patients with ACS. However, the approach is different depending on the clinical presentation of the ACS, namely STEMI or NSTEMI, the myocardium at risk and the ischaemic time.

In patients with STEMI admitted during the first few hours after symptom onset, the choice of reperfusion strategy and the speed with which it is implemented have a major impact on clinical outcomes.41,42 Given this, coronary reperfusion performed within a short time frame is recommended.^{29,42,43} Both the use of reperfusion (either by fibrinolytic therapy in eligible patients or by PCI) and its timely implementation have previously been used as indicators of quality of care.8,9 Many opportunities exist to reduce the proportion of patients who do not receive reperfusion, and the delay with which this is provided. This requires active involvement from all partners involved along the management pathway, and includes high quality organisation of care within a network organisation.7,25 Accountability is an important factor in measuring quality of care, and measuring the different components of the overall time to reperfusion (such as time of call, time of first medical contact, time of arrival, transfer, arterial puncture, time at which artery patency is achieved) makes it possible to rank the quality of each in the overall pathway of care, and identify areas where there may be room for improvement.44

In patients with NSTEMI, an invasive strategy using coronary angiography with a view to myocardial revascularisation is also related to lower mortality in moderate-tohigh and in high risk patients.^{45,46} In NSTEMI, appropriate use of an invasive strategy is linked to the assessment of the patient's ischaemic and bleeding risks. In practice, according to the risk profile, invasive strategy must be immediate (<2 h), early (<24 h), <72 h or conditional. Avoiding the risks of an invasive strategy for patients at low ischaemic risk is equally as important as ensuring access to invasive strategy for patients at high ischaemic risk.

Clinical relevance. For patients with STEMI, the relation between shorter times to reperfusion and mortality has previously been established, particularly within the first 3–4 h after

onset of infarction. Beyond 12 h, the benefit of reperfusion is less well established, and although PCI is recommended within the first 24 h, measures of quality of care are limited to patients admitted less than 12 h since the onset of symptoms.^{47,48} The benefit is greater when reperfusion is performed early. The advantage of earlier reperfusion is seen more in the first 3-4 h after onset of symptoms. The estimated times to reperfusion also contribute largely to the choice of reperfusion strategy. PCI is the technique of choice except if it is estimated that it cannot be performed within 120 min after the diagnosis of STEMI has been established.^{29,42,49} A time to reperfusion therapy ≤30 min (diagnosis to injection) is recommended when fibrinolytic therapy is the strategy of choice; and when PCI is the chosen strategy, the recommended time to PCI is ≤120 min between diagnosis and opening of the infarctrelated artery. When the door-to-balloon (or to first device that opens the artery) time is considered, ESC guidelines for revascularisation propose a recommended time <60 min.

For patients with NSTEMI, the benefit of an invasive strategy is established only in patients at risk of ischaemia.45,46,50 An invasive strategy is needed to confirm the diagnosis of ACS, identify the culprit coronary lesion, establish the indication for revascularisation (by PCI or surgery) and assess the long-term risk. The timing of the invasive strategy also depends on risk assessment; a small proportion of patients require an immediate invasive strategy (within 2 h), an early invasive strategy (within 24 h) is indicated for high risk patients, while an invasive strategy can be performed within 72 h in NSTEMI patients without high risk criteria. Lastly, a selective invasive strategy can be used among patients at low ischaemic risk, according to the results of their non-invasive stress test. A decision for an early invasive strategy (i.e. performance of a coronary angiography and revascularisation if appropriate) versus a conditional invasive or medical strategy should take into account the risk-benefit ratio. Invasive treatment is recommended in the presence of ischaemic risk factors and, conversely, is not recommended in the absence of ischaemic risk factors.27,29

Specific aspects for potential Qls. For the QIs relating to reperfusion and invasive strategy, the numerator and

denominator should comprise all patients hospitalised with STEMI or NSTEMI. Only patients eligible for the invasive approach should be included, and form both the numerator and the denominator. Therefore, the numerators and denominators should not include patients who have contraindications, such as patients with STEMI who present after the first 12 h, or those who have clinical, allergic, arterial access or haemorrhagic problems, or patient-related reasons for exclusion (refusal to provide consent for angiography or PCI). Patients with clinical or patient-related reasons for exclusion are considered ineligible.

In STEMI patients, measuring time to reperfusion requires that the different times of the various stages along the management pathway be measured. Both the use of reperfusion (either by fibrinolytic therapy in eligible patients or by PCI) and its timely implementation can be used as indicators of quality of care.^{8,9} In case of fibrinolysis, the time interval to initiation of fibrinolysis is counted from first medical contact to injection, which is in line with recommendations for pre-hospital fibrinolysis. Conversely, in case of reperfusion by primary PCI, if the patient is admitted directly to a PCI-capable centre, the door-to-arterial access time is preferred over other starting time points, in order to better reflect internal hospital organisation, and also because it is easier to measure. Finally, if the patient is admitted to a centre without PCI facilities and primary PCI is chosen as the reperfusion strategy, the door-in-door-out time in the non-PCI-capable centre has been selected, as it is independent of geographical constraints that may affect transfer times.

Lastly, the median time from first medical contact (FMC) to arterial access for reperfusion of the infarct-related artery has been chosen as a secondary QI.

In NSTEMI, appropriate use of an invasive strategy is linked to the assessment of the patient's ischaemic and bleeding risks. Thus, according to the risk profile, invasive strategy must be immediate (<2 h), early (<24 h), <72 h or conditional. To avoid excessive complexity, only the decision to perform an invasive strategy within the first 72 h in NSTEMI patients without contraindication has been retained as a QI.

Name of the main QI (STEMI I)	Proportion of STEMI patients reperfused among those eligible (onset of symptoms to diagnosis <12 h)
Name of the main QI (STEMI 2)	The proportion of patients reperfused within the timeframe recommended by the ESC guidelines; these time frames vary according to whether fibrinolysis or PCI is chosen as the strategy of choice.
	 For fibrinolysis, a maximum time delay of 30 min between first medical contact and the start of injection is retained. For primary PCI, a maximum door-to-arterial access time of <60 min is retained for patients admitted to centres with PCI facilities on site. For transferred patients, an additional QI is defined, namely a 'door-in-door-out' time of less than 30 min.
Name of the secondary QI Name of the main QI (NSTEMI I)	FMC to arterial access time (absolute value) for primary PCI. Proportion of patients with NSTEMI, and no contra-indication, who receive coronary angiography within 72 h after admission.

Definition of the main and secondary QIs

Agreement with guidelines and existing Qls. <u>STEMI patients</u>. The ESC guidelines stipulate that all patients with STEMI of onset <12 h should receive reperfusion therapy as early as possible. In addition, the ESC guidelines recommend measuring the time from the onset of symptoms, FMC, diagnosis, and initiation to reperfusion, stating that an invasive strategy should be performed within 90 min in early presenters with a large area at risk.⁴⁹

Two main QIs and one secondary QI have been defined for myocardial reperfusion:

- Main QI (STEMI 1) is the percentage of patients reperfused among patients admitted with STEMI of <12 h duration. This QI is identical to the PM defined by the ACC/AHA Task Force in 2008.^{8,9}
- Main QI (STEMI 2) is the percentage of patients reperfused within the time frame recommended by the ESC guidelines; these time frames vary according to whether fibrinolysis or PCI is the strategy of choice.
 - For fibrinolysis, a maximum time delay of 30 min between FMC and the start of injection is retained, in accordance with the ESC guidelines.
 - For primary PCI, a maximum door-to-arterial access time of <60 min is retained for patients admitted to centres with PCI facilities on site.
 - For transferred patients, an additional QI is defined, namely a 'door-in-door-out' time of less than 30 min.

The 2008 ACC/AHA PM related to reperfusion in STEMI were: (a) time to fibrinolytic therapy; (b) time to primary PCI; (c) proportion of patients who received reperfusion therapy; (d) time from emergency department arrival at STEMI referral facility to emergency department discharge for transfer to PCI centre; and (e) time from emergency department arrival in PCI-centre to PCI. In the most recent ACC/AHA Task Force defining PM for reperfusion, five other PMs for reperfusion were defined. The main PM was the time to device use for PCI and four proposals for future PM relating to time to reperfusion were also defined: (a) time to reperfusion among patients transferred for PCI; (b) proportion of reperfusion-eligible patients receiving therapy; (c) diagnosis to reperfusion; and (d) time to reperfusion for patients developing STEMI in the hospital.^{8,9} The ACCA Quality of Care Working Group QIs are in agreement with these indicators from the ACC/AHA, but only three are retained in our document. Times are used with threshold values, according to the type of reperfusion and the need for transfer. The FMC to arterial access was left as an absolute value, because even though this measure depends on numerous factors, it is the most important in clinical terms.⁴³

<u>NSTEMI patients</u>: Although international guidelines concur in their recommendations of an invasive strategy among patients with high-risk features,^{27,29} this aspect of management has never before been used in measures of quality of care. To avoid excessive complexity, the decision to perform an invasive strategy within the first 72 h in NSTEMI patients without contraindication was selected as a QI, considering that all NSTEMI patients are at intermediate or high risk.

13

In-hospital risk assessment

Dimensions of care. Clinical outcomes following AMI are highly variable and are, in part, due to the wide spectrum of baseline clinical risk levels. Therefore, risk assessment is a key step in the management of patients with AMI, particularly their risk of death and recurrent MI (to determine the need for a more aggressive approach), the risk of iatrogenic complications such as bleeding (to modify the use of antithrombotic treatments) and the short- or long-term risk of HF (to enable tailoring of clinical review, investigations and guideline-indicated therapies).

Establishing the risk of in-hospital or short-term death from the ischaemic process will better inform clinicians of necessary changes to an invasive strategy, namely its timing. Indeed, for NSTEMI, the clinical benefit of an early invasive strategy has been shown to be related to the level of ischaemic risk, with greater clinical benefits seen in higher risk patients.

Bleeding is one of the main iatrogenic complications among patients with AMI and is associated with an adverse prognosis. Bleeding risk estimation can help define the best diagnostic and therapeutic strategy and should affect the type, dose and duration of anti-thrombotic treatments, as well as the choice of arterial access for invasive procedures.

Assessment of left ventricular function and the quantification of resting left ventricular ejection fraction (LVEF) before discharge from hospital are important for the selection of patients with severe left ventricular dysfunction who may benefit from additional treatments. Left ventricular systolic dysfunction is a key predictor of immediate and late episodes of HF, and ventricular arrhythmia.⁵¹ Patients with severe left ventricular dysfunction require specific secondary prevention therapies.

For STEMI, with the increasing use of primary PCI, risk assessment for ischaemia before discharge has become less important. This is because it can be assumed that the infarctrelated coronary lesion has been treated and stabilised, and the presence or absence of significant lesions in other arteries has been assessed.

Clinical relevance. Observational studies support the use of established risk scores for ischaemic and bleeding risk.

Ischaemic risk. Of the numerous ischaemic risk scores, the GRACE risk score is not widely used, but is strongly endorsed by ESC guidelines.^{27,28} A recent update of the GRACE score (GRACE v.2.0), calculated from 32,307 patients enrolled between 2001–2007 with suspected ACS, has shown, in an external population, adequate discrimination and calibration for in-hospital and three-year mortality.⁵² Categorisation of the patients into GRACE high risk, GRACE medium risk and GRACE lower risk may help physicians in their choice of whether to pursue an invasive strategy, since the invasive approach mainly benefits high risk patients.^{53,54} Studies from registries have shown that the use of coronary angiography is higher in individuals at lower risk as compared to those at higher risk, defining the 'treatment-risk paradox'. 55,56 Employing a facilitated GRACE score, like the updated one, has the potential to favour a systematic and objective evaluation of the ischaemic risk of NSTEMI patients and to increase the rate of revascularisation in high-risk patients without contraindications. This means that, based on the impact of a systematic interventional strategy in the randomised trials, there would be between 30 and 80 fewer cardiovascular deaths or MIs for every 10,000 patients with NSTEMI.45,57-60

Definition of the QIs

<u>Bleeding risk</u> can be predicted from baseline variables through a specific risk score. Bleeding risk scores have also been developed from registry or trial cohorts in the setting of AMI and PCI. The Can Rapid risk stratification of Unstable angina patients Suppress ADverse outcomes with Early implementation of the ACC/AHA guidelines (CRUSADE) bleeding risk score (http://www.crusadebleedingscore.org) was developed from a cohort of 71,277 NSTE-ACS patients (derivation cohort) and further validated.⁶¹ The CRUSADE bleeding score has been demonstrated to have a good predictive accuracy also in STEMI patients treated by primary PCI.⁶²

Assessment of infarct size and post-MI left ventricular systolic function can be performed by echocardiography, a technique available in almost all centres admitting patients with AMI.

Name of the main QI (I)	Proportion of patients with NSTEMI in whom ischaemic risk assessment using the GRACE risk score is performed. GRACE score should be assessed and the numerical value of the score recorded for all patients admitted with suspected NSTEMI.
Name of the main QI (2)	Proportion of patients admitted with STEMI or NSTEMI who have bleeding risk assessment using the CRUSADE bleeding score. CRUSADE bleeding score should be assessed and the
Name of the main QI (3)	numerical value of the score recorded for all patients admitted for NSTEMI. Proportion of patients with assessment of LVEF before discharge. LVEF should be assessed and numerical value recorded for all patients admitted with STEMI and NSTEMI.

Agreement with guidelines and existing Qls. Main QI (1): Ischaemic risk assessment. ESC NSTE-ACS guidelines recommend basing diagnosis and initial short-term ischaemic and bleeding risk stratification on a combination of clinical history, symptoms, vital signs, other physical findings, ECG and laboratory results. The quantitative assessment of ischaemic risk scores is superior to clinical assessment alone. The GRACE risk score provides the most accurate stratification of risk both on admission and at discharge and has a class IA recommendation.

In the ACC/AHA PM set, risk assessment is used to measure the risk-adjusted 30-day mortality, but does not itself represent a PM.

<u>Main QI (2)</u>: Bleeding risk assessment. The ESC NSTE-ACS guidelines stipulate that the use of the CRUSADE bleeding risk score may be considered in patients undergoing coronary angiography to quantify bleeding risk with a class IIb level B recommendation. Bleeding risk assessment is also recommended in the ESC STEMI guidelines for antiplatelet therapy (like abciximab and prasugrel), without mention of any type of score for this assessment.

In the ACC/AHA PM for STEMI/NSTEMI, no assessment of bleeding risk is stated.

Main QI (3): Evaluation of left ventricular systolic function is recommended by the ESC guidelines for STEMI (class IB) and for NSTE-ACS (class IC). Assessment of left ventricular systolic function has been a part of the ACC/ AHA PM set since 2008.

Anti-thrombotic treatment during hospitalisation

Dimensions of care. Anti-thrombotic treatment has a pivotal role in the management of AMI.⁶³ Therefore, the combination of an anticoagulant with antiplatelet therapy is mandatory in the treatment of AMI.^{27–29} This approach has been shown to reduce mortality (both cardiovascular and total), (re)infarction and stroke.^{64–69} Moreover, many patients with AMI are treated with PCI and stent implantation, and in this case dual antiplatelet therapy (DAPT) has a key role in reducing stent thrombosis, both in the short (acute/subacute) and long term. Finally, antiplatelet therapy improves longterm outcomes following AMI. Prolongation of DAPT beyond the first year is currently a matter of debate.^{70,71}

Clinical relevance. <u>Antiplatelets:</u> the benefit of aspirin for AMI has been demonstrated in four randomised studies, with the rate of ischaemic events reduced by half as compared to placebo. The benefit of the combination of aspirin with clopidogrel has also been demonstrated, leading to a high-grade recommendation for DAPT.^{27–29} More potent $P2Y_{12}$ platelet receptor inhibitors (prasugrel and ticagrelor), in addition to aspirin, are linked to improved outcome, with a reduction in the composite ischaemic outcome, including mortality from STEMI.^{72–74} The increase in bleeding is reasonable and sometimes negligible,⁷⁵ with the exception of some high-risk subgroups. Based on trial design, prasugrel use has been limited to

selected STEMI patients who receive PCI and who do not have previous stroke.⁷³ Ticagrelor is suitable for use among a broader ACS population, irrespective of history, use of PCI or pre-treatment with clopidogrel.⁷² Whatever the drug chosen for P2Y₁₂ inhibition, DAPT should be initiated as soon as the diagnosis is established, and continued for a duration of 12 months, unless bleeding complications occur.⁷⁶ The benefit of using glycoprotein GPIIb/IIIa inhibitors was established before the advent of DAPT, but since strong P2Y₁₂ platelet receptor inhibitors have become available, the use of GPIIb/IIIa inhibitors has been reserved for patients with peri-procedural thrombotic complications and patients with a high risk of thrombosis and a low risk of bleeding.

Anticoagulation: a parenteral anticoagulant is required during the acute phase, and continued until completion of revascularisation or hospital discharge, whichever occurs first. Due to multiple comparisons of a variety of anticoagulants, at different doses, with or without arterial access and associated with different combinations of antiplatelets, no specific single anticoagulant regimen can be proposed for STEMI (unfractionated heparin, the low molecular weight heparin enoxaparin, and to some extent, bivalirudin have been investigated in prospective randomised trials). Conversely, for NSTEMI, fondaparinux at a dose of 2.5 mg/day is considered by guidelines to have a favourable efficacy/safety profile, except in those proceeding directly to PCI.^{27–29,77–79}

Specific aspects for potential indicators. The complexity of selecting appropriate anti-thrombotic treatment, and the high-grade recommendations suggest that the selection of antiplatelet therapy and anticoagulants is an ideal field for the assessment of the quality of care.

Antiplatelet agents at admission: Aspirin, initiated as soon as possible in all patients without contraindication, has been used as a QI, both at admission and at discharge. Nevertheless, since the rate of use of aspirin is often higher than 95%, the interest of keeping this indicator is debatable. As regards $P2Y_{12}$ inhibitors, the choice between clopidogrel, prasugrel and ticagrelor is underpinned by clear contraindications, limitations of use, and risk assessment, and therefore might also reflect quality. Ticagrelor is recommended over clopidogrel unless the patient is at high risk of bleeding (patients with previous haemorrhagic stroke, chronic oral anticoagulation or fibrinolytic therapy, ongoing bleeding). Prasugrel is also recommended over clopidogrel in patients undergoing PCI, but also without a history of any type of cerebrovascular accident, age <75 years and body weight ≥ 60 kg.⁷³ When neither ticagrelor nor prasugrel is possible, clopidogrel is the best option. The use of GPIIb/IIIa inhibitors is decided case by case, according to criteria that are not always recorded and, therefore, is less suitable for quality assessment.

Antiplatelet agents at discharge: The prescription of DAPT at discharge, for an expected duration of one year, is well supported by guidelines, and applicable to AMI irrespective of its management. The choice of $P2Y_{12}$ inhibitor follows the same rules as at admission. In addition, DAPT prescription at discharge was suggested as a potential QI in the recent ESC 2015 NSTE-ACS guidelines,²⁷ without distinction of any single $P2Y_{12}$ inhibitor molecule. Only patients treated by chronic anticoagulants are excluded, due to a high bleeding risk, a lack of scientific evidence, and a lack of strong recommendation regarding discharge antiplatelet treatment.

Anticoagulants. The use of fondaparinux in patients with NSTEMI (except those proceeding directly to PCI or those with severe renal dysfunction) can be used as a QI. If fondaparinux is not available, enoxaparin should be preferred. In STEMI patients, the use of anticoagulant prescription to assess quality is challenging, due to the multiplicity of drugs, doses and possible combinations. Dosing errors or inappropriate combinations of different anticoagulants are associated with thrombotic or haemorrhagic complications and could be tracked for quality assessment.

Definition of the main and secondary QIs for anti-thrombotic treatments during hospitalisation

Name of the main QI (I)	 Proportion of patients (numerator and denominator detailed in appendix) with 'adequate P2Y₁₂ inhibition' defined as: number of patients discharged with prasugrel or ticagrelor or clopidogrel / patients eligible. 'Eligible' is defined as follows: For ticagrelor = AMI patients without previous haemorrhagic stroke, high bleeding risk, fibrinolysis or oral anticoagulation. For prasugrel: PCI treated AMI patients without previous haemorrhagic or ischaemic stroke, high bleeding risk (patients >75 years and/or <60 kg body weight are also considered as high bleeding risk), fibrinolysis or oral anticoagulation. For clopidogrel: no indication for prasugrel or ticagrelor and no high bleeding risk.
Name of the main QI (2)	Proportion of patients with NSTEMI treated with fondaparinux, unless candidate for immediate (≤ 2 h) invasive strategy or with eGFR <20 ml/min.
Name of the secondary QI	Proportion of patients discharged on dual antiplatelet therapy/patients with AMI, without clear and documented contraindication

Agreement with guidelines and existing Qls. For STEMI patients who receive fibrinolysis, aspirin and clopidogrel have class IB and IA indications, respectively. Enoxaparin has the best indication with fibrinolysis (class IA), followed by UFH (class IC). In STEMI patients treated with primary PCI, prasugrel and ticagrelor both have a class IB recommendation, while clopidogrel is indicated only if prasugrel and ticagrelor are not available or contraindicated (class IC). The recommendations for parenteral anticoagulants are class IC for UFH, class IIaA for bivalirudin and class IIaB for enoxaparin.

For NSTEMI patients, antiplatelet therapy with prasugrel or ticagrelor has a class IB recommendation, while clopidogrel is indicated in patients who cannot receive prasugrel or ticagrelor or who are treated with oral anticoagulants. In patients naive of $P2Y_{12}$ inhibitors undergoing PCI, cangrelor has a lower level of recommendation (IIbA).

Previous ACC/AHA PMs have not considered any PM related to anti-thrombotic therapy. However, in the 2008 update of PM for STEMI/NSTEMI, the use of clopidogrel in medically treated AMI patients only, as well as excessive initial dose of unfractionated heparin, enoxaparin, abx-icimab, eptifibatide and tirofiban, were considered as 'test measures'. In addition, the use of a standardised protocol for anticoagulants, and a tracking system for anticoagulation errors, were also considered as PMs in test.⁸

Secondary prevention – discharge treatments

Dimension of care. The secondary prevention of cardiovascular events following index AMI is critical, because patients remain at high risk of mortality and recurrent cardiovascular events long after an AMI. Long-term cohort studies have shown that mortality after hospitalisation with AMI is high.⁴⁵ Moreover, secondary prevention pharmacotherapy has been shown to reduce long-term adverse clinical outcomes in this group.⁸⁰⁻⁸² While the long-term management of patients with AMI is predominantly the responsibility of the general practitioner, secondary prevention medications will have a greater chance of being implemented if performed during the hospital stay. The caveat to this is that with declining lengths of hospital stay,⁸³ there is less of a distinction between acute and chronic therapies and it is therefore the shared responsibility of primary and secondary care physicians to ensure that all eligible patients with AMI are prescribed and maintained on guidelines-indicated medications. This domain encompasses the prescription of statins, angiotensin-converting enzyme inhibitors (ACEIs)/angiotensin receptor blockers (ARBs) and beta-blockers.

Clinical relevance. The benefit of long-term treatment with beta-blockers among patients with AMI who have HF or left ventricular systolic dysfunction is well established.⁸⁰ However, there are no contemporary randomised controlled trials testing the efficacy of beta-blockers among patients following AMI without HF. As such, current ESC guidance is to prescribe beta-blockers to haemodynamically stable patients who are eligible and who have a LVEF≤0.40. For ACEIs/ ARBs, there is also strong evidence for their use as secondary prevention therapy among eligible patients with AMI who have HF or a LVEF <0.40.81,82,84 Compelling evidence supports the use of high intensity statins for reducing recurrent cardiovascular events and mortality following AMI,85,86 and although the rate of prescription of statins at discharge from hospital is high,^{83,87,88} a substantial proportion of patients are not discharged with high intensity statins after hospitalisation for coronary heart disease.⁸⁹ This is important when high intensity statins reduce low-density lipoprotein (LDL) cholesterol by around 50% and the efficacy of statins is apparent across a range of patient groups, including both sexes, the young and the elderly, as well as those with and without diabetes. As with statins, the opportunity to reduce premature cardiovascular death through increased prescription of ACEI/ ARBs and beta-blockers is clearly apparent.83,87,88,90

The magnitude of the relationship between high intensity statin therapy and mortality is strong. In 4162 patients hospitalised with an ACS, higher intensity statins were associated with a 16% reduction in the risk of death at two years, over and above that of the reduction in risk of death associated with standard lipid therapy.85 The magnitude of the relationship between ACEIs/ARBs and mortality among patients with AMI is strong.⁸¹ For beta-blockers, the magnitude of the relationship for mortality among patients with AMI is weaker. That is, there are no contemporary randomised data testing the efficacy of beta-blockers among patients with AMI who do not have HF, and recent data from an observational study suggested no benefit among this group. In the CAPRICORN randomised trial of 1959 patients with AMI and a LVEF<0.40, the risk of all-cause mortality or non fatal AMI at a mean of 1.3 years was reduced by 29% among patients who received carvedilol.80 However, for patients with AMI and HF or a reduced LVEF, the magnitude of the evidence is strong. In a randomised trial of 2647 patients with HF, of which half had documented ischaemic heart disease, bisoprolol significantly reduced all-cause mortality by 34% at a mean of 1.3 years.91

Specific aspects for potential Qls. For the secondary prevention medication QIs, the numerator and denominator should comprise all patients hospitalised with AMI. Only patients eligible to receive the medications should be included, and form both the numerator and denominator. Therefore, the numerators and denominators should not include patients who are allergic, have contraindications, refuse, or have side effects from the medications – these patients are 'ineligible'.

Specifically, for the statins indicator, the numerator must only include patients who are prescribed a high intensity statin (such as atorvastatin 40-80 mg or rosuvastatin 20 mg)⁹² at discharge. Consequently, patients who receive a low-intermediate intensity statin after hospitalisation will be included in the denominator and not in the numerator. Patients with a history of intolerance to high-intensity statin therapy or have other characteristics that may influence safety should not be included in the numerator or denominator.

For ACEIs/ARBs, the numerator and denominator include all eligible patients with AMI who have HF or a LVEF \leq 0.40. Ineligibility criteria for the numerator and denominator include hypotension, acute renal failure and hyperkalaemia (the contraindication must be documented in the patient's file). Specifically, the numerator is the number of patients with AMI and HF or a LVEF \leq 0.40 who are prescribed an ACEI or an ARB.

Regarding beta-blockers, both STEMI and NSTE-ACS ESC guidelines are in alignment. Therefore, the numerator and denominator should comprise eligible patients with either HF or a LVEF ≤ 0.40 . Ineligibility criteria for beta-blockers include evidence of a low output state, increased risk for cardiogenic shock, PR interval >0.24 s, second- or third-degree heart block, active asthma, or reactive airways disease. Therefore, the numerator will comprise the number of patients with AMI who have HF or a LVEF ≤ 0.40 , who have no ineligibility criteria, and who are prescribed a beta-blocker, whereas the denominator is defined as the number of patients with AMI who have HF or a LVEF ≤ 0.40 , and who have no ineligibility criteria.

Definition	of	the	main	and	secondary	Qls	for	secondary
prevention	- c	lischa	arge tre	eatme	ents			

Name of the main QI	Proportion of patients with AMI discharged on statins, unless contra indicated, at high intensity (defined as atorvastatin ≥ 40 mg or rosuvastatin ≥ 20 mg).
Name of the secondary QI (I)	Proportion of patients with AMI and clinical evidence of HF or LVEF <0.40 who are prescribed, at discharge, an ACEI (or ARBs if intolerant of ACEI) unless contraindicated.
Name of the secondary QI (2)	Proportion of patients with AMI and clinical evidence of HF or LVEF≤0.40 who are prescribed, at discharge, beta-blockers, unless contraindicated.

Agreement with guidelines and existing Qls. Currently, the use of high-intensity statins initiated early after admission to hospital (the main QI) is supported by the ESC guidelines (class 1A) and ACC/AHA guidelines for STEMI (class 1B) and NSTEMI (class 1).^{27,28,93,94} The secondary QI, the use of an ACEI (or ARB if intolerant of ACEIs) unless contraindicated, before discharge in patients with clinical evidence of HF or a LVEF ≤ 0.40 is also supported by the ESC and ACC/AHA guidelines (class 1A). The other secondary QI, the use of a betablocker, unless contraindicated, before discharge in patients with clinical evidence of HF or a LVEF ≤ 0.40 is supported by the ESC guidelines (class 1A) and ACC/AHA guidelines (class 1A). The other secondary QI, the use of a betablocker, unless contraindicated, before discharge in patients with clinical evidence of HF or a LVEF ≤ 0.40 is supported by the ESC guidelines (class 1A) and ACC/AHA guidelines for STEMI (class 1A) and NSTEMI (class 1B).

Patient satisfaction

Dimensions of care. The Institute of Medicine has considered that being 'respectful of and responsive to individual patient preferences, needs and values and [ensuring] that patients guide all clinical decisions' (p.6) is a key element of the quality of care.¹ Measuring patient satisfaction, as well as patient-reported outcomes, provides information about symptoms, health-related quality of life, morbidity and satisfaction with care that are reported directly by the patient and not captured in medical records.

The ESC recognised the importance of including patientreported outcome measures (PROMs) in clinical trials and research to inform patients, clinicians, payers and policymakers.⁹⁵ PROMs provide information about health-related quality of life that can be used to assess how a treatment can improve symptoms or functional capacity, in association with other clinical endpoints such as clinical events or morbidity.⁹⁶ This aspect of the quality has been measured through the patient satisfaction QI. Patient satisfaction informs about important aspects of the quality of management, such as fast access to reliable health advice, effective treatment and information delivered by health professionals, continuity of care and smooth transitions and emotional support, empathy and respect.^{97–99} Thus, patient satisfaction is an essential and complementary approach to conventional quality of care indicators. This is the first time that PROMs have been considered in QIs for AMI.

Clinical relevance. Patient satisfaction measures a specific concept from the patient perspective that is obviously clinically relevant. Patient satisfaction can be assessed through specific questionnaires developed in the late 1970s,¹⁰⁰ which explore multiple domains such as symptoms and functional status, health perception, and also domains related to the relationship with nurses, physicians and other health professionals, to personal issues, admission, visitors, discharge, room, meals, tests and treatments.⁹⁸ In the setting of ACS, patient information is a major component of patient satisfaction, whereby the more information patients receive, the more they report being satisfied.99 In a study using data from the CRU-SADE registry and including 6467 patients with AMI, patient satisfaction was associated with guidelines adherence and with mortality rates.97

Specific aspects for potential Qls. Given the complexity of assessment of patient satisfaction, the main QI is limited to the routine and continuous assessment of patient satisfaction through a specific scale. This scale should explore at

Definition of the main and secondary Qls for patient satisfaction

Name of the	Feedback regarding patient's experiences systematically collected in an organised way		
main	from all patients. It should include the following		
QI (I)	points:		
	Pain control.		
	• Explanations provided by health professionals		
	about the coronary disease, the benefit/risk of		
	discharge treatment, and medical follow-up.		
	 Discharge information regarding what to do in case 		
	of recurrence of symptoms and recommendation		
	to attend a cardiac rehabilitation programme		
	(including smoking cession and diet counselling).		

Agreement with guidelines and existing Qls. The ESC NSTE-ACS guidelines recognised the need to consider patients' preferences for the decision regarding invasive strategy and revascularisation, particularly in older patients. Pain control is usually not an issue for NSTEMI patients. In STEMI patients, relief of pain is considered of paramount importance, both for humane reasons but also to avoid excessive adrenaline activation and anxiety. Use of morphine, if needed, has a class IC recommendation.

Patient information about the disease, need for treatment compliance and risk factor control (like smoking cessation, diet and weight control, physical activity) is recommended, as well as education about the recognition of symptoms.

No existing QI set has incorporated patient satisfaction. Conversely, a large number of hospitals routinely assess patient satisfaction. Centres for Medicare and Medicaid Services (CMS) in the USA have developed a national standardised survey instrument for measuring patient satisfaction: the Hospital Consumer Assessment of Healthcare Providers and Systems (HCAHPS).^{101–103} The UK's National Health Service Patient Survey also systematically gathers the views of the patients about the quality of care they have received (http://www.pickereurope.org/wp-content/uploads/2014/10/Inpatients_2015_spec_v12.pdf?gclid=COWclcmIk8oCFWo CwwodhRUC0Q).

CQIs and outcomes

Dimensions of care. <u>CQI</u>. In view of the growing interest in quality assessment among healthcare providers, insurances, agencies, press and the general public, CQIs have been developed. A CQI is a combination of two or more indicators into a single number to summarise multiple dimensions and to facilitate comparisons. In the field of assessment of quality of care, a CQI presents three potential advantages. Firstly, it comprehensively represents quality of care, making it

possible to reduce the information into a single summary. Secondly, the information may contain a broader range of measures, including multiple dimensions of care. Thirdly, the presentation of the CQI as a single number facilitates its use by providers for decision-making, benchmarking or financial incentives. In addition, a CQI reduces the visible size of a set of indicators, allowing comparison and categorisation of the centres, and it can be used to assess progress over time and facilitate communication and accountability. The disadvantage of the CQI is that it may be misinterpreted and invite simplistic policy conclusions. Another criticism

levelled at CQIs is that they can be like a 'black box', with loss of information; moreover, the selection of indicators and

the models used to construct the CQI can be disputed. Despite widespread enthusiasm for the development of COIs, methodological controversies have arisen over their robustness, in the absence of an established model. In 2010, the AHA/ACC Task Force on Performance Measures published a position statement for the creation and interpretation of CQIs in healthcare assessment.⁶ The ACCA Quality of Care Working Group has decided to include two COIs. A first CQI (opportunity scoring), using a greater number of individual QIs, is suitable with a view to promoting high quality standards. The second (all or none) is based on discharge treatment, and is suitable for use in survivors and in all centres, irrespective of on-site facilities. These CQI can serve as a comparator between different healthcare systems or within centres to compare quality over time, and to determine categories of centres by comparison with the average value.

Outcome QI. Although clinical outcome is the final aim of quality of care, the use of outcome measures in quality assessment is the subject of controversy, since the variation in outcomes only partially depends on the quality of care. Furthermore, reporting outcomes might have adverse effects, such as restriction of admission for more severe patients. Nevertheless, since outcome measures are the most easily interpretable and also potentially important for patients,³ the Quality of Care Working Group has decided to include an outcome QI.

Clinical relevance. Whereas the relation between a single QI and clinical outcome is difficult to demonstrate, the relation between CQI and mortality has been established through different approaches. In early studies, a trend towards lower in-hospital mortality was observed across categories of centres defined according to the quartiles of a composite indicator.¹⁰⁴ Similarly, a significant trend towards lower 30-day mortality, adjusted for the GRACE risk score, was also observed at patient level, by quartiles of a composite score.¹⁰⁵ The strength of association is higher with CQIs as compared to individual QIs.^{104,106}

The association between CQIs and mortality has been observed in-hospital, but also at 30 days^{105–108} and one year.^{106,108} The magnitude of the relation between a CQI and mortality is usually modest, but significant. In a study from the National Registry of Myocardial Infarction, 6% of the

variance in 30-day mortality after STEMI was explained by a CQI including 'only' timely reperfusion and smoking measures.⁷ Similarly, the correlation between CQI and mortality¹⁰⁷ or risk-adjusted mortality^{105,106,109} is not strong. Indeed, the magnitude of the relation between a CQI and mortality depends on the individual QIs used, and on the type of QI.

Specific aspects for potential Qls. Explicit criteria exist for the development of composite performance measures so that they can accurately reflect healthcare quality, including explicit quantification of the numerator and denominator of potential measures and explicit evaluation of the interpretability, actionability, and feasibility of the proposed measure. Among the different methods of aggregation, the 'opportunity-based' score (with or without weighting) and the 'all or none' are the most frequently used in the assessment of quality of care. These two methods provide different results and the appropriate approach should be selected according to the purpose of the assessment.^{107,108,110,111}

<u>Opportunity scoring</u>: The main CQI includes all the individual QIs selected by the group, including structure, process and patient satisfaction QIs. However, since it is possible that not all the components are available in all centres, the opportunity-based scoring method has been chosen for the calculation of this CQI, without weighting.

All or none: The secondary CQI included here uses discharge prescriptions, and is calculated using the 'all or none' method. This method best reflects the interest of the patient, since even one missing component in the score may influence outcome. This CQI makes it possible to track excellence. For each patient, the CQI is rated=1 if all components of the CQI are present and it is rated=0 if one or more components are missing. In patients with HF or LVEF \leq 0.40, the CQI is calculated from five individual QIs, and for patients without HF or with LVEF>0.40, the CQI is calculated using three individual QIs.

30-Day mortality rate, adjusted for GRACE risk score: Although there is no question as to the importance of outcome in quality of care, the use of outcome OIs is still controversial. Limitations on the use of outcomes as OI include the complex and multifactorial determinants of outcome, as well as the low proportion of variance in outcomes that can be explained by quality of care. The least controversial outcome OI in the setting of AMI is the adjusted 30-day mortality. Given the relatively low 30-day mortality rate after AMI in European countries, reliable assessment of mortality rate over a short period of time is challenging. In addition, the risk model used for adjustment can influence the results. In their latest position paper, the ACC/AHA recognised adjusted 30-day mortality as an acceptable outcome PM.³ Our writing group has selected the GRACE risk score for adjustment, since this score has been recently updated and validated in European registries and because guidelines recommend the calculation of GRACE risk score in patients with NSTEMI.52

Definition	of the mair	and secondo	ary QI for	CQIs an	nd outcomes
------------	-------------	-------------	------------	---------	-------------

Name of the main	Composite QI = main CQI: opportunity-based CQI, with the following individual indicators:			
CQI	 The centre is part of a network organisation. 			
	 Proportion of patients reperfused among eligible (STEMI with onset of symptoms to diagnosis <12 h). 			
	 Coronary angiography in STEMI and NSTEMI patients at high ischaemic risk and without contraindications. 			
	 Ischaemic risk assessment using the GRACE risk score in NSTEMI patients. 			
	 Bleeding risk assessment using the CRUSADE risk score in STEMI and NSTEMI patients. 			
	Assessment of LVEF before discharge.			
	 Low-dose aspirin (unless high bleeding risk or oral anticoagulation). 			
	Adequate P2Y12 inhibition (as defined in treatment during hospitalisation section).			
	 ACEI (or ARBs if intolerant of ACEI, unless contra-indicated) before discharge in patients with clinical evidence of HF or LVEF < 0.40. 			
	 beta-Blockers (unless clear contraindication) in patients with clinical evidence of HF or LVEF < High-intensity statins. 			
	• Feedback regarding the patient's experience and quality of care is systematically collected for all patients.			
Name of the	Secondary CQI: all-or-none CQI based on 3 or 5 components, according to the LVEF.			
secondary CQI (I)	 In patients without HF and with LVEF>0.40, CQI calculated on 3 individual QI: 			
	 Low-dose aspirin. 			
	 P2Y₁₂ inhibitor (unless documented contraindication). 			
	• High-intensity statins.			
	• In patients with HF or with LVEF \leq 0.40, CQI calculated on 5 individual QI:			
	 Low-dose aspirin. 			
	 P2Y₁₂ inhibitor (unless documented contraindication). 			
	 High-intensity statins. 			
	• ACEI (or ARB if intolerant of ACEI) in patients with clinical evidence of HF or LVEF≤0.40.			
	• Beta-blockers (unless clear contraindication) in patients with clinical evidence of HF or LVEF≤0.40			
Name of the	Outcome QI: risk-adjusted 30-day mortality rate adjusted on the GRACE 2.0 risk score.			
secondary QI (2)				

Agreement with existing QI and guidelines. Currently, no CQI is included in the ACC/AHA set of PMs,⁸ or in the Canadian PMs.¹¹² Conversely, CQIs are used for assessment of quality of care by the Joint Commission on Accreditation of Healthcare Organisations (JCAHO), the Agency for Healthcare Research and Quality (AHRQ), the CMS, the MINAP¹⁰⁸ and by the French National Authority for Health (Haute Autorité de Santé).¹¹³

Outcome QI: The use of the GRACE 2.0 risk calculator is recommended in the ESC guidelines for ischaemic risk assessment and for prognostic assessment in NSTEMI patients. The risk-adjusted 30-day mortality rate is recommended by the ACC/AHA task force, but is not included in the ACC/AHA QI set.

Discussion

The ACCA brought together the current Quality of Care Working Group with a view to defining a set of QIs for the management of AMI. This is the first such initiative undertaken within the ESC by one of its constituent associations. The absence of any official publication of QIs in STEMI and NSTEMI by the ESC to date is in contrast with the historical experience of the ACC and AHA in this area,^{8,9} and with individual national initiatives. Among the missions of the ACCA, 'improving the quality of care of patients with acute cardiovascular disease' is fundamental, thus justifying the creation of this Working Group to define QIs suitable for use in this clinical setting.

Specificities of the QIs defined by the ACCA

Compared with previous QIs defined by the ACC-AHA,^{8,9} the CCS⁴⁰ or the National Service Framework for Coronary Heart Disease,¹¹⁴ the selection of suitable QIs by the ACCA Quality of Care Working Group followed a methodology that is comparable in many respects, but different on a certain number of important points.

While the OIs developed by our group are in line with the official guidelines published by the ESC, they are not simply the reflection of high grade recommendations, but rather incorporate measures that have a grade II recommendation, or even no recommendation at all, because the Quality of Care Working Group feels that it is a critically measurable reflection of the quality of care. Indeed, therein lies the difference between guidelines and QIs.115 Conversely, some key features of management that hold a strong grade recommendation were not retained as QI, when it was judged that there was little room for improvement. A typical example of this is the prescription of aspirin at admission or discharge, which is among the QIs issued by the ACC-AHA, but registry data show that the already widespread implementation of this measure leaves little room to further improve practices on this point. Although some of the QIs do not fulfil the criteria for PM, the ACCA Quality of Care Working Group decided to keep all selected QIs, even if the set of QI might not be suitable for public reporting, benchmarking or pay-for-performance. The main reason was that the QI focus on processes of care for which failure to follow the recommendations is likely to result in suboptimal patient outcomes.

Another point of divergence between the QIs defined here and previous publications from other societies concerns the wide spectrum of topics covered by the ACCA OIs. Indeed, seven domains were selected to define one or more main QIs as well as secondary QIs. Two domains are related to the organisation at the level of the centre, as opposed to the quality of individual management. In particular, the structure-network domain emphasises the importance of the working environment in each centre, as this has been clearly shown to have an impact on clinical outcomes in STEMI management. Patient satisfaction is the second domain that deals with centre organisation rather than individual patterns of care. To date, neither the ACC-AHA QIs nor those of the CCS have defined any indicators relating to centre organisation or patient satisfaction. The ACCA Quality of Care Working Group considered that certain contributors to patient satisfaction (e.g. pain management, respect, education) can reflect the quality of care better than medical criteria.

In the five other domains, the QIs selected by the ACCA Quality of Care Working Group present some differences and specificities as compared to existing QIs from other sources. For example, in addition to a QI for reperfusion, a QI for 'early invasive strategy' has been added, which is applicable to patients with NSTEMI. Estimation of the ischaemic and bleeding risks using the GRACE and CRUSADE risk scores respectively has been added as a QI, going beyond the simple measure of LVEF recommended by US PMs. The QIs related to secondary prevention focus on the use of high-intensity statins and on the treatment of patients with a LVEF≤0.40, which are two situations where these treatments are especially beneficial. Lastly, the composite criterion proposed in this paper is computed using the 'opportunity-based' method, based on all the main OIs from all the domains, while a second 'all or none' composite QI is proposed, with a limited number of QIs.108,116

Implementation of QIs

The QIs defined here by the ACCA Quality of Care Working Group are not intended for ranking, benchmarking or payfor-performance, but merely contributing to the aim of monitoring and improving quality of care through meaningful surveillance, in line with the founding principles of quality of care first described by Donabedian in 1966.¹⁰ A second major point is the wide spectrum of the care pathway that is covered by the different domains, namely centre organisation, risk assessment, acute management and secondary prevention through to outcome. A composite indicator is proposed to provide a summary indicator for all the information from these different domains.

In this document, we propose main QIs and secondary level QIs. A total of 12 main QIs are proposed, representing criteria that we consider to be major, requiring preferential measurement. The eight secondary QIs are supplementary measures of quality that are suitable for use as a complementary approach. In the continuity of this document, the Quality of Care Working Group plans to design a registry specifically intended for performance measurement, as well as annual updates of the QIs to reflect any changes in ESC guidelines.

Assessment of QIs and perspectives

Among the main characteristics of a QI, feasibility and reliability of the assessment are important, and have an impact on the selection of the QI. The ACCA Quality of Care Working Group QI set has been defined with this view. To date, national initiatives for assessment of quality have used administrative databases, registries (specifically designed or not) or observational cohort studies. Across Europe, leading examples include the SWEDEHEART registry, which is a comprehensive and voluntary database of all patients admitted for ACS in all Swedish hospitals and including approximately 80,000 new cases per year;¹¹⁷ the MINAP in England and Wales (the participation of all institutions in England and Wales is mandatory and MINAP is used as a tool to improve quality of care);¹¹⁸ the French FAST-MI registries, which consist in one-month snapshot surveys performed every five years since 1995,119 Acute Coronary Syndrome Israeli Survey (ACSIS),120 biennial two-month registries in all Israeli ICCUs¹¹⁹ and, lastly, the German Maximal Individual Therapy in Acute Myocardial Infarction (MITRA) and Arbeitsgemeinschaft Leitender Kardiologischer Krankenhausärzte (ALKK) registries.¹²¹ The Spanish Society of Cardiology and of Thoracic and Cardiovascular Surgery has defined quality markers for clinical cardiology, cardiac imaging acute care, interventional cardiology, cardiac rehabilitation and cardiac surgery.¹²² The proposed indicators for AMI are in line with the ACCA selection, but not precisely defined and limited to direct transfer for PCI and recording FMC to balloon time for STEMI patients, risk assessment, revascularisation for high risk patients and adherence to guidelines for discharge treatment. Several other European countries have successfully implemented large-scale population-based registries to collect information about the incidence, management and outcome of ACS, such as the Acute Myocardial Infarction and Unstable Angina in Switzerland (AMIS plus) registry,¹²³ the Italian BLITZ registry,¹²⁴ the PRIAMHO I and II125 in Spain and in the Central and Eastern European countries.¹²⁶ Registries for assessment of quality have also been organised by the ESC, such as the Euro Heart Survey-ACS programme.88,90,127 Results from specific European national campaigns using QIs have also been reported.113

Conclusions

In agreement with the missions of the ACCA, the Quality of Care Working Group has planned a quality assessment programme through a dedicated registry using the main and secondary QIs developed here. Despite its limitations, the publication of this set of QIs will offer the possibility to assess the quality of management of AMI, which in turn will provide a clear picture of the management of AMI in Europe and serve to identify the domains of care where improvements are most needed.

Conflict of interest

The authors declare that there is no conflict of interest.

Funding

This research received no specific grant from any funding agency in the public, commercial, or not-for-profit sectors.

References

- Crossing the quality chasm: A new health system for the 21st century. Washington DC: Committee on Quality Health Care in America. The National Academies Press, 2001. Available at: http://www.nap.edu/catalog/10027/ crossing-the-quality-chasm-a-new-health-system-for-the (accessed 11 April 2016).
- Bonow RO, Masoudi FA, Rumsfeld JS, et al. ACC/AHA classification of care metrics: Performance measures and quality metrics: A report of the American College of Cardiology/American Heart Association Task Force on Performance Measures. *J Am Coll Cardiol* 2008; 52: 2113–2117.
- Spertus JA, Bonow RO, Chan P, et al. ACCF/AHA new insights into the methodology of performance measurement: A report of the American College of Cardiology Foundation/American Heart Association Task Force on performance measures. *Circulation* 2010; 122: 2091–2106.
- Spertus JA, Eagle KA, Krumholz HM, et al. American College of Cardiology and American Heart Association methodology for the selection and creation of performance measures for quantifying the quality of cardiovascular care. *Circulation* 2005; 111: 1703–1712.
- Krumholz HM, Keenan PS, Brush JE Jr, et al. Standards for measures used for public reporting of efficiency in health care: A scientific statement from the American Heart Association Interdisciplinary Council on Quality of Care and Outcomes Research and the American College of Cardiology Foundation. *Circulation* 2008; 118: 1885–1893.
- Peterson ED, Delong ER, Masoudi FA, et al. ACCF/ AHA 2010 Position Statement on Composite Measures for Healthcare Performance Assessment: A report of the American College of Cardiology Foundation/American Heart Association Task Force on Performance Measures (Writing Committee to develop a position statement on composite measures). *Circulation* 2010; 121: 1780–1791.
- Bradley EH, Herrin J, Elbel B, et al. Hospital quality for acute myocardial infarction: Correlation among process measures and relationship with short-term mortality. *JAMA* 2006; 296: 72–78.

- 8. Krumholz HM, Anderson JL, Bachelder BL, et al. ACC/AHA 2008 performance measures for adults with ST-elevation and non-ST-elevation myocardial infarction: A report of the American College of Cardiology/American Heart Association Task Force on Performance Measures (Writing Committee to develop performance measures for ST-elevation and non-ST-elevation myocardial infarction): Developed in collaboration with the American Academy of Family Physicians and the American College of Emergency Physicians: Endorsed by the American Association of Cardiovascular and Pulmonary Rehabilitation, Society for Cardiovascular Angiography and Interventions, and Society of Hospital Medicine. *Circulation* 2008; 118: 2596–2648.
- Masoudi FA, Bonow RO, Brindis RG, et al. ACC/AHA 2008 statement on performance measurement and reperfusion therapy: A report of the ACC/AHA Task Force on Performance Measures (Work Group to address the challenges of performance measurement and reperfusion therapy). *Circulation* 2008; 118: 2649–2661.
- Donabedian A. Evaluating the quality of medical care. Milbank Mem Fund Q 1966; 44: S166–S206.
- Armstrong PW, Westerhout CM and Welsh RC. Duration of symptoms is the key modulator of the choice of reperfusion for ST-elevation myocardial infarction. *Circulation* 2009; 119: 1293–1303.
- Fox KA and Huber K. A European perspective on improving acute systems of care in STEMI: We know what to do, but how can we do it? *Nat Clin Pract Cardiovasc Med* 2008; 5: 708–714.
- Huber K, Gersh BJ, Goldstein P, et al. The organization, function, and outcomes of ST-elevation myocardial infarction networks worldwide: Current state, unmet needs and future directions. *Eur Heart J* 2014; 35: 1526–1532.
- Rathore SS, Epstein AJ, Nallamothu BK, et al. Regionalization of ST-segment elevation acute coronary syndromes care: Putting a national policy in proper perspective. *J Am Coll Cardiol* 2006; 47: 1346–1349.
- Huber K, Goldstein P, Danchin N, et al. Network models for large cities: The European experience. *Heart* 2010; 96: 164–169.
- Nallamothu BK, Krumholz HM, Ko DT, et al. Development of systems of care for ST-elevation myocardial infarction patients: Gaps, barriers, and implications. *Circulation* 2007; 116: e68–e72.
- Rathore SS, Curtis JP, Nallamothu BK, et al. Association of door-to-balloon time and mortality in patients > or =65 years with ST-elevation myocardial infarction undergoing primary percutaneous coronary intervention. *Am J Cardiol* 2009; 104: 1198–1203.
- Nielsen PH, Terkelsen CJ, Nielsen TT, et al. System delay and timing of intervention in acute myocardial infarction (from the Danish Acute Myocardial Infarction-2 [DANAMI-2] trial). Am J Cardiol 2011; 108: 776–781.
- Steg PG, Cambou JP, Goldstein P, et al. Bypassing the emergency room reduces delays and mortality in ST elevation myocardial infarction: The USIC 2000 registry. *Heart* 2006; 92: 1378–1383.
- Danchin N. Systems of care for ST-segment elevation myocardial infarction: Impact of different models on clinical outcomes. *JACC Cardiovasc Interv* 2009; 2: 901–908.

- Kalla K, Christ G, Karnik R, et al. Implementation of guidelines improves the standard of care: The Viennese registry on reperfusion strategies in ST-elevation myocardial infarction (Vienna STEMI registry). *Circulation* 2006; 113: 2398–2405.
- 22. Danchin N, Coste P, Ferrieres J, et al. Comparison of thrombolysis followed by broad use of percutaneous coronary intervention with primary percutaneous coronary intervention for ST-segment-elevation acute myocardial infarction: Data from the French registry on acute ST-elevation myocardial infarction (FAST-MI). *Circulation* 2008; 118: 268–276.
- Henry TD, Sharkey SW, Burke MN, et al. A regional system to provide timely access to percutaneous coronary intervention for ST-elevation myocardial infarction. *Circulation* 2007; 116: 721–728.
- 24. Henry TD, Unger BT, Sharkey SW, et al. Design of a standardized system for transfer of patients with ST-elevation myocardial infarction for percutaneous coronary intervention. *Am Heart J* 2005; 150: 373–384.
- 25. Ting HH, Krumholz HM, Bradley EH, et al. Implementation and integration of prehospital ECGs into systems of care for acute coronary syndrome: A scientific statement from the American Heart Association Interdisciplinary Council on Quality of Care and Outcomes Research, Emergency Cardiovascular Care Committee, Council on Cardiovascular Nursing, and Council on Clinical Cardiology. *Circulation* 2008; 118: 1066–1079.
- 26. Bohmer E, Hoffmann P, Abdelnoor M, et al. Efficacy and safety of immediate angioplasty versus ischemia-guided management after thrombolysis in acute myocardial infarction in areas with very long transfer distances results of the NORDISTEMI (NORwegian study on DIstrict treatment of ST-elevation myocardial infarction). J Am Coll Cardiol 2010; 55: 102–110.
- Roffi M, Patrono C, Collet JP, et al. 2015 ESC Guidelines for the management of acute coronary syndromes in patients presenting without persistent ST-segment elevation: Task Force for the Management of Acute Coronary Syndromes in Patients Presenting without Persistent ST-Segment Elevation of the European Society of Cardiology (ESC). *Eur Heart J* 2016; 37: 267–315.
- Steg PG, James SK, Atar D, et al. ESC Guidelines for the management of acute myocardial infarction in patients presenting with ST-segment elevation: The Task Force on the management of ST-segment elevation acute myocardial infarction of the European Society of Cardiology (ESC). *Eur Heart J* 2012; 33: 2569–2619.
- 29. Windecker S, Kolh P, Alfonso F, et al. 2014 ESC/EACTS Guidelines on myocardial revascularization: The Task Force on Myocardial Revascularization of the European Society of Cardiology (ESC) and the European Association for Cardio-Thoracic Surgery (EACTS) Developed with the special contribution of the European Association of Percutaneous Cardiovascular Interventions (EAPCI). *Eur Heart J* 2014; 35: 2541–2619.
- Canto JG, Rogers WJ, Bowlby LJ, et al. The prehospital electrocardiogram in acute myocardial infarction: Is its full potential being realized? National Registry of Myocardial Infarction 2 Investigators. J Am Coll Cardiol 1997; 29: 498–505.
- 31. Brown JP, Mahmud E, Dunford JV, et al. Effect of prehospital 12-lead electrocardiogram on activation of the

cardiac catheterization laboratory and door-to-balloon time in ST-segment elevation acute myocardial infarction. *Am J Cardiol* 2008; 101: 158–161.

- Dhruva VN, Abdelhadi SI, Anis A, et al. ST-Segment Analysis Using Wireless Technology in Acute Myocardial Infarction (STAT-MI) trial. *J Am Coll Cardiol* 2007; 50: 509–513.
- Quinn T, Johnsen S, Gale CP, et al. Effects of prehospital 12-lead ECG on processes of care and mortality in acute coronary syndrome: A linked cohort study from the Myocardial Ischaemia National Audit Project. *Heart* 2014; 100: 944–950.
- Camp-Rogers T, Dante S, Kontos MC, et al. The impact of prehospital activation of the cardiac catheterization team on time to treatment for patients presenting with ST-segmentelevation myocardial infarction. *Am J Emerg Med* 2011; 29: 1117–1124.
- Kontos MC, Kurz MC, Roberts CS, et al. Emergency physician-initiated cath lab activation reduces door to balloon times in ST-segment elevation myocardial infarction patients. *Am J Emerg Med* 2011; 29: 868–874.
- 36. Goyal NK, Giglio J, Lorberbaum M, et al. A rapid-response alphanumeric paging design decreases door-to-balloon times in patients undergoing primary percutaneous coronary intervention for ST elevation acute myocardial infarction. *Crit Pathw Cardiol* 2004; 3: 150–153.
- Clemmensen P, Schoos MM, Lindholm MG, et al. Prehospital diagnosis and transfer of patients with acute myocardial infarction–a decade long experience from one of Europe's largest STEMI networks. *J Electrocardiol* 2013; 46: 546–552.
- Peterson MC, Syndergaard T, Bowler J, et al. A systematic review of factors predicting door to balloon time in ST-segment elevation myocardial infarction treated with percutaneous intervention. *Int J Cardiol* 2012; 157: 8–23.
- KaifoszovaZ, KalaP, AlexanderT, et al. Stent for Life Initiative: Leading example in building STEMI systems of care in emerging countries. *EuroIntervention* 2014; 10: T87–T95.
- Tu JV, Khalid L, Donovan LR, et al. Indicators of quality of care for patients with acute myocardial infarction. *CMAJ* 2008; 179: 909–915.
- Boersma E, Primary Coronary Angioplasty vs. Thrombolysis G. Does time matter? A pooled analysis of randomized clinical trials comparing primary percutaneous coronary intervention and in-hospital fibrinolysis in acute myocardial infarction patients. *Eur Heart J* 2006; 27: 779–788.
- 42. Steg PG, Bonnefoy E, Chabaud S, et al. Impact of time to treatment on mortality after prehospital fibrinolysis or primary angioplasty: Data from the CAPTIM randomized clinical trial. *Circulation* 2003; 108: 2851–2856.
- Steg PG, James SK, Atar D, et al. ESC Guidelines for the management of acute myocardial infarction in patients presenting with ST-segment elevation. *Eur Heart J* 2012; 33: 2569–2619.
- Chassin MR, Loeb JM, Schmaltz SP, et al. Accountability measures–using measurement to promote quality improvement. *N Engl J Med* 2010; 363: 683–688.
- 45. Fox KA, Carruthers KF, Dunbar DR, et al. Underestimated and under-recognized: The late consequences of acute

coronary syndrome (GRACE UK-Belgian Study). *Eur Heart J* 2010; 31: 2755–2764.

- Katritsis DG, Siontis GC, Kastrati A, et al. Optimal timing of coronary angiography and potential intervention in non-ST-elevation acute coronary syndromes. *Eur Heart J* 2011; 32: 32–40.
- Hochman JS, Lamas GA, Buller CE, et al. Coronary intervention for persistent occlusion after myocardial infarction. *N Engl J Med* 2006; 355: 2395–2407.
- Schomig A, Mehilli J, Antoniucci D, et al. Mechanical reperfusion in patients with acute myocardial infarction presenting more than 12 h from symptom onset: A randomized controlled trial. *JAMA* 2005; 293: 2865–2872.
- 49. Pinto DS, Frederick PD, Chakrabarti AK, et al. Benefit of transferring ST-segment-elevation myocardial infarction patients for percutaneous coronary intervention compared with administration of onsite fibrinolytic declines as delays increase. *Circulation* 2011; 124: 2512–2521.
- O'Donoghue M, Boden WE, Braunwald E, et al. Early invasive vs conservative treatment strategies in women and men with unstable angina and non-ST-segment elevation myocardial infarction: A meta-analysis. *JAMA* 2008; 300: 71–80.
- 51. Dickstein K, Cohen-Solal A, Filippatos G, et al. ESC guidelines for the diagnosis and treatment of acute and chronic heart failure 2008: The Task Force for the diagnosis and treatment of acute and chronic heart failure 2008 of the European Society of Cardiology. Developed in collaboration with the Heart Failure Association of the ESC (HFA) and endorsed by the European Society of Intensive Care Medicine (ESICM). *Eur J Heart Fail* 2008; 10: 933–989.
- 52. Fox KA, Fitzgerald G, Puymirat E, et al. Should patients with acute coronary disease be stratified for management according to their risk? Derivation, external validation and outcomes using the updated GRACE risk score. *BMJ Open* 2014; 4: e004425.
- Bavry AA, Kumbhani DJ, Rassi AN, et al. Benefit of early invasive therapy in acute coronary syndromes: A metaanalysis of contemporary randomized clinical trials. *J Am Coll Cardiol* 2006; 48: 1319–1325.
- Hoenig MR, Aroney CN and Scott IA. Early invasive versus conservative strategies for unstable angina and non-ST elevation myocardial infarction in the stent era. *Cochrane Database Syst Rev* 2010: CD004815.
- 55. Cohen MG, Filby SJ, Roe MT, et al. The paradoxical use of cardiac catheterization in patients with non-ST-elevation acute coronary syndromes: Lessons from the Can Rapid Stratification of Unstable Angina Patients Suppress Adverse Outcomes With Early Implementation of the ACC/AHA Guidelines (CRUSADE) Quality Improvement Initiative. *Am Heart J* 2009; 158: 263–270.
- Fox KA, Anderson FAJr, Dabbous OH, et al. Intervention in acute coronary syndromes: Do patients undergo intervention on the basis of their risk characteristics? The Global Registry of Acute Coronary Events (GRACE). *Heart* 2007; 93: 177–182.
- Lagerqvist B, Husted S, Kontny F, et al. 5-Year outcomes in the FRISC-II randomised trial of an invasive versus a non-invasive strategy in non-ST-elevation acute coronary syndrome: A follow-up study. *Lancet* 2006; 368: 998–1004.
- Fox KA, Poole-Wilson P, Clayton TC, et al. 5-Year outcome of an interventional strategy in non-ST-elevation

acute coronary syndrome: The British Heart Foundation RITA 3 randomised trial. *Lancet* 2005; 366: 914–920.

- 59. Damman P, Hirsch A, Windhausen F, et al. 5-Year clinical outcomes in the ICTUS (Invasive versus Conservative Treatment in Unstable coronary Syndromes) trial a randomized comparison of an early invasive versus selective invasive management in patients with non-ST-segment elevation acute coronary syndrome. *J Am Coll Cardiol* 2010; 55: 858–864.
- Damman P, van Geloven N, Tijssen JG, et al. Timing of intervention in patients undergoing invasive management for acute coronary syndromes. *J Am Coll Cardiol* 2010; 56: 998, author reply: 999.
- Subherwal S, Bach RG, Chen AY, et al. Baseline risk of major bleeding in non-ST-segment-elevation myocardial infarction: The CRUSADE (Can Rapid risk stratification of Unstable angina patients Suppress ADverse outcomes with Early implementation of the ACC/AHA Guidelines) Bleeding Score. *Circulation* 2009; 119: 1873–1882.
- Flores-Rios X, Couto-Mallon D, Rodriguez-Garrido J, et al. Comparison of the performance of the CRUSADE, ACUITY-HORIZONS, and ACTION bleeding risk scores in STEMI undergoing primary PCI: Insights from a cohort of 1391 patients. *Eur Heart J Acute Cardiovasc Care* 2013; 2: 19–26.
- 63. Toutouzas K, Benetos G, Karanasos A, et al. Vulnerable plaque imaging: Updates on new pathobiological mechanisms. *Eur Heart J* 2015; 36: 3147–3154.
- Randomised trial of intravenous streptokinase, oral aspirin, both, or neither among 17,187 cases of suspected acute myocardial infarction: ISIS-2. ISIS-2 (Second International Study of Infarct Survival) Collaborative Group. *Lancet* 1988; 2: 349–360.
- 65. Antithrombotic Trialists' Collaboration. Collaborative meta-analysis of randomised trials of antiplatelet therapy for prevention of death, myocardial infarction, and stroke in high risk patients. *Br Med J* 2002; 324: 71–86.
- Steinhubl SR, Berger PB, Mann JT 3rd, et al. Early and sustained dual oral antiplatelet therapy following percutaneous coronary intervention: A randomized controlled trial. *JAMA* 2002; 288: 2411–2420.
- Yusuf S, Zhao F, Mehta SR, et al. Effects of clopidogrel in addition to aspirin in patients with acute coronary syndromes without ST-segment elevation. *N Engl J Med* 2001; 345: 494–502.
- Chen ZM, Jiang LX, Chen YP, et al. Addition of clopidogrel to aspirin in 45,852 patients with acute myocardial infarction: Randomised placebo-controlled trial. *Lancet* 2005; 366: 1607–1621.
- 69. Yusuf S, Mehta SR, Zhao F, et al. Early and late effects of clopidogrel in patients with acute coronary syndromes. *Circulation* 2003; 107: 966–972.
- Bonaca MP, Braunwald E and Sabatine MS. Long-term use of ticagrelor in patients with prior myocardial infarction. N Engl J Med 2015; 373: 1274–1275.
- Mauri L, Kereiakes DJ, Yeh RW, et al. Twelve or 30 months of dual antiplatelet therapy after drug-eluting stents. *N Engl J Med* 2014; 371: 2155–2166.
- Wallentin L, Becker RC, Budaj A, et al. Ticagrelor versus clopidogrel in patients with acute coronary syndromes. N Engl J Med 2009; 361: 1045–1057.

- 73. Wiviott SD, Braunwald E, McCabe CH, et al. Prasugrel versus clopidogrel in patients with acute coronary syndromes. *N Engl J Med* 2007; 357: 2001–2015.
- 74. Montalescot G, Wiviott SD, Braunwald E, et al. Prasugrel compared with clopidogrel in patients undergoing percutaneous coronary intervention for ST-elevation myocardial infarction (TRITON-TIMI 38): Double-blind, randomised controlled trial. *Lancet* 2009; 373: 723–731.
- Wiviott SD, Desai N, Murphy SA, et al. Efficacy and safety of intensive antiplatelet therapy with prasugrel from TRITON-TIMI 38 in a core clinical cohort defined by worldwide regulatory agencies. *Am J Cardiol* 2011; 108: 905–911.
- Navarese EP, Andreotti F, Schulze V, et al. Optimal duration of dual antiplatelet therapy after percutaneous coronary intervention with drug eluting stents: Meta-analysis of randomised controlled trials. *Br Med J* 2015; 350: h1618.
- Yusuf S, Mehta SR, Chrolavicius S, et al. Comparison of fondaparinux and enoxaparin in acute coronary syndromes. *N Engl J Med* 2006; 354: 1464–1476.
- Steg PG, Jolly SS, Mehta SR, et al. Low-dose vs standarddose unfractionated heparin for percutaneous coronary intervention in acute coronary syndromes treated with fondaparinux: The FUTURA/OASIS-8 randomized trial. *JAMA* 2010; 304: 1339–1349.
- 79. Navarese EP, Andreotti F, Kolodziejczak M, et al. Comparative efficacy and safety of anticoagulant strategies for acute coronary syndromes. Comprehensive network meta-analysis of 42 randomised trials involving 117,353 patients. *Thromb Haemost* 2015; 114: 933–944.
- Dargie HJ. Effect of carvedilol on outcome after myocardial infarction in patients with left-ventricular dysfunction: The CAPRICORN randomised trial. *Lancet* 2001; 357: 1385–1390.
- Pfeffer MA, Braunwald E, Moye LA, et al. Effect of captopril on mortality and morbidity in patients with left ventricular dysfunction after myocardial infarction. Results of the survival and ventricular enlargement trial. The SAVE Investigators. N Engl J Med 1992; 327: 669–677.
- Pfeffer MA, McMurray JJ, Velazquez EJ, et al. Valsartan, captopril, or both in myocardial infarction complicated by heart failure, left ventricular dysfunction, or both. *N Engl J Med* 2003; 349: 1893–1906.
- Gale CP, Cattle BA, Woolston A, et al. Resolving inequalities in care? Reduced mortality in the elderly after acute coronary syndromes. The Myocardial Ischaemia National Audit Project 2003–2010. *Eur Heart J* 2012; 33: 630–639.
- Granger CB, McMurray JJ, Yusuf S, et al. Effects of candesartan in patients with chronic heart failure and reduced left-ventricular systolic function intolerant to angiotensinconverting-enzyme inhibitors: The CHARM-Alternative trial. *Lancet* 2003; 362: 772–776.
- Cannon CP, Braunwald E, McCabe CH, et al. Intensive versus moderate lipid lowering with statins after acute coronary syndromes. *N Engl J Med* 2004; 350: 1495–1504.
- Cholesterol Treatment Trialists C, Baigent C, Blackwell L, et al. Efficacy and safety of more intensive lowering of LDL cholesterol: A meta-analysis of data from 170,000 participants in 26 randomised trials. *Lancet* 2010; 376: 1670–1681.

- 87. Hasdai D, Behar S, Wallentin L, et al. A prospective survey of the characteristics, treatments and outcomes of patients with acute coronary syndromes in Europe and the Mediterranean basin: The Euro Heart Survey of Acute Coronary Syndromes (Euro Heart Survey ACS). *Eur Heart J* 2002; 23: 1190–1201.
- Simms AD, Weston CF, West RM, et al. Mortality and missed opportunities along the pathway of care for ST-elevation myocardial infarction: A national cohort study. *Eur Heart J Acute Cardiovasc Care* 2015; 4: 241–253.
- Rosenson RS, Kent ST, Brown TM, et al. Underutilization of high-intensity statin therapy after hospitalization for coronary heart disease. *J Am Coll Cardiol* 2015; 65: 270–277.
- Puymirat E, Battler A, Birkhead J, et al. Euro Heart Survey 2009 Snapshot: Regional variations in presentation and management of patients with AMI in 47 countries. *Eur Heart J Acute Cardiovasc Care* 2013; 2: 359–370.
- 91. The Cardiac Insufficiency Bisoprolol Study II (CIBIS-II): A randomised trial. *Lancet* 1999; 353: 9–13.
- 92. Stone NJ, Robinson JG, Lichtenstein AH, et al. 2013 ACC/ AHA guideline on the treatment of blood cholesterol to reduce atherosclerotic cardiovascular risk in adults: A report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines. J Am Coll Cardiol 2014; 63: 2889–2934.
- 93. Amsterdam EA, Wenger NK, Brindis RG, et al. 2014 AHA/ ACC guideline for the management of patients with non-ST-elevation acute coronary syndromes: Executive summary: A report of the American College of Cardiology/ American Heart Association Task Force on Practice Guidelines. *Circulation* 2014; 130: 2354–2394.
- 94. O'Gara PT, Kushner FG, Ascheim DD, et al. 2013 ACCF/ AHA guideline for the management of ST-elevation myocardial infarction: Executive summary: A report of the American College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines: Developed in collaboration with the American College of Emergency Physicians and Society for Cardiovascular Angiography and Interventions. *Catheter Cardiovasc Interv* 2013; 82: E1–E27.
- Anker SD, Agewall S, Borggrefe M, et al. The importance of patient-reported outcomes: A call for their comprehensive integration in cardiovascular clinical trials. *Eur Heart* J 2014; 35: 2001–2009.
- Ahmed S, Berzon RA, Revicki DA, et al. The use of patientreported outcomes (PRO) within comparative effectiveness research: Implications for clinical practice and health care policy. *Med Care* 2012; 50: 1060–1070.
- Glickman SW, Boulding W, Manary M, et al. Patient satisfaction and its relationship with clinical quality and inpatient mortality in acute myocardial infarction. *Circ Cardiovasc Qual Outcomes* 2010; 3: 188–195.
- Lee DS, Tu JV, Chong A, et al. Patient satisfaction and its relationship with quality and outcomes of care after acute myocardial infarction. *Circulation* 2008; 118: 1938–1945.
- Oterhals K, Hanestad BR, Eide GE, et al. The relationship between in-hospital information and patient satisfaction after acute myocardial infarction. *Eur J Cardiovasc Nurs* 2006; 5: 303–310.

- Van Campen C, Sixma H, Friele RD, et al. Quality of care and patient satisfaction: A review of measuring instruments. *Med Care Res Rev* 1995; 52: 109–133.
- Goldstein E, Farquhar M, Crofton C, et al. Measuring hospital care from the patients' perspective: An overview of the CAHPS hospital survey development process. *Health Serv Res* 2005; 40: 1977–1995.
- Crofton C, Darby C, Farquhar M, et al. The CAHPS hospital survey: Development, testing, and use. *Jt Comm J Qual Patient Saf* 2005; 31: 655–659, 601.
- Darby C, Hays RD and Kletke P. Development and evaluation of the CAHPS hospital survey. *Health Serv Res* 2005; 40: 1973–1976.
- Granger CB, Steg PG, Peterson E, et al. Medication performance measures and mortality following acute coronary syndromes. *Am J Med* 2005; 118: 858–865.
- Schiele F, Meneveau N, Seronde MF, et al. Compliance with guidelines and 1-year mortality in patients with acute myocardial infarction: A prospective study. *Eur Heart J* 2005; 26: 873–880.
- Werner RM and Bradlow ET. Relationship between Medicare's hospital compare performance measures and mortality rates. *JAMA* 2006; 296: 2694–2702.
- 107. Eapen ZJ, Fonarow GC, Dai D, et al. Comparison of composite measure methodologies for rewarding quality of care: An analysis from the American Heart Association's Get With The Guidelines program. *Circ Cardiovasc Qual Outcomes* 2011; 4: 610–618.
- 108. Simms AD, Batin PD, Weston CF, et al. An evaluation of composite indicators of hospital acute myocardial infarction care: A study of 136,392 patients from the Myocardial Ischaemia National Audit Project. *Int J Cardiol* 2013; 170: 81–87.
- 109. Kontos MC, Rennyson SL, Chen AY, et al. The association of myocardial infarction process of care measures and inhospital mortality: A report from the NCDR(R). *Am Heart* J 2014; 168: 766–775.
- 110. Couralet M, Guerin S, Le Vaillant M, et al. Constructing a composite quality score for the care of acute myocardial infarction patients at discharge: Impact on hospital ranking. *Med Care* 2011; 49: 569–576.
- Nolan T and Berwick DM. All-or-none measurement raises the bar on performance. JAMA 2006; 295: 1168–1170.
- Tu JV, Khalid L, Donovan LR, et al. Indicators of quality of care for patients with acute myocardial infarction. *CMAJ* 2008; 179: 909–915.
- 113. Schiele F, Capuano F, Loirat P, et al. Hospital case volume and appropriate prescriptions at hospital discharge after acute myocardial infarction: A nationwide assessment. *Circ Cardiovasc Qual Outcomes* 2013; 6: 50–57.
- Weston CF. Performance indicators in acute myocardial infarction: A proposal for the future assessment of good quality care. *Heart* 2008; 94: 1397–1401.
- 115. Walter LC, Davidowitz NP, Heineken PA, et al. Pitfalls of converting practice guidelines into quality measures: Lessons learned from a VA performance measure. *JAMA* 2004; 291: 2466–2470.
- 116. Simms AD, Baxter PD, Cattle BA, et al. An assessment of composite measures of hospital performance and associated mortality for patients with acute myocardial infarction. Analysis of individual hospital performance and outcome

for the National Institute for Cardiovascular Outcomes Research (NICOR). *Eur Heart J Acute Cardiovasc Care* 2013; 2: 9–18.

- 117. Jernberg T, Attebring MF, Hambraeus K, et al. The Swedish Web-system for enhancement and development of evidence-based care in heart disease evaluated according to recommended therapies (SWEDEHEART). *Heart* 2010; 96: 1617–1621.
- Herrett E, Smeeth L, Walker L, et al. The Myocardial Ischaemia National Audit Project (MINAP). *Heart* 2010; 96: 1264–1267.
- Hanssen M, Cottin Y, Khalife K, et al. French Registry on Acute ST-elevation and non ST-elevation Myocardial Infarction 2010. FAST-MI 2010. *Heart* 2012; 98: 699–705.
- Behar S, Battler A, Porath A, et al. A prospective national survey of management and clinical outcome of acute myocardial infarction in Israel, 2000. *Isr Med Assoc J* 2003; 5: 249–254.
- 121. Koeth O, Zahn R, Gitt AK, et al. Clinical benefit of early reperfusion therapy in patients with ST-elevation myocardial infarction usually excluded from randomized clinical trials (results from the Maximal Individual Therapy in Acute Myocardial Infarction Plus [MITRA Plus] registry). *Am J Cardiol* 2009; 104: 1074–1077.
- 122. Lopez-Sendon JL, Gonzalez-Juanatey JR, Pinto F, et al. Quality markers in cardiology: Measures of outcomes and

clinical practice–a perspective of the Spanish Society of Cardiology and of Thoracic and Cardiovascular Surgery. *Eur Heart J* 2016; 37: 12–23.

- 123. Radovanovic D and Erne P. AMIS Plus: Swiss registry of acute coronary syndrome. *Heart* 2010; 96: 917–921.
- 124. Olivari Z, Steffenino G, Savonitto S, et al. The management of acute myocardial infarction in the cardiological intensive care units in Italy: The 'BLITZ 4 Qualita' campaign for performance measurement and quality improvement. *Eur Heart J Acute Cardiovasc Care* 2012; 1: 143–152.
- 125. Cabades A, Echanove I, Cebrian J, et al. The characteristics, management and prognosis of the acute myocardial infarct patient in the Valencian Community in 1995: The results of the PRIMVAC Registry (The Registry Project of Acute Myocardial Infarct in Valencia, Alicante and Castellon). As representatives of the PRIMVAC investigators]. *Rev Esp Cardiol* 1999; 52: 123–133.
- 126. Smith FG, Brogan RA, Alabas O, et al. Comparative care and outcomes for acute coronary syndromes in Central and Eastern European Transitional countries: A review of the literature. *Eur Heart J Acute Cardiovasc Care* 2015; 4: 537–554.
- 127. Schiele F, Hochadel M, Tubaro M, et al. Reperfusion strategy in Europe: Temporal trends in performance measures for reperfusion therapy in ST-elevation myocardial infarction. *Eur Heart J* 2010; 31: 2614–2624.