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<u>Authors</u>: Alfonsina Candiello, M.D; Thomas Alexander, M.D; Rhena Delport, Gabor G Toth, M.D, PhD; Paul J.L Ong, M.D; Adriaan Snyders, M.D; Jorge A Belardi, M.D; Michael K.Y Lee, M.D; Helder Pereira, M.D; Awad Mohamed, MSc, M.D; Jorge Mayol, M.D; Jan J Piek, M.D, PhD; William Wijns, M.D, PhD; Andreas Baumbach, M.D, PhD; Christoph K Naber, M.D, PhD

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2	A practical methods paper from the Stent – Save a life! Initiative (www.stentsavealife.com)
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4	Alfonsina Candiello ¹ , MD; Thomas Alexander ² , MD; Rhena Delport ³ , Gabor G Toth ⁴ , MD,
5	PhD; Paul J.L Ong ⁵ , MD; Adriaan Snyders ⁶ , MD; Jorge A Belardi ¹ MD; Michael K.Y Lee ⁷ ,
6	MD; Helder Pereira ⁸ , MD; Awad Mohamed ⁹ , MSc, MD, Jorge Mayol ¹⁰ , MD, Jan J Piek ¹¹ , MD,
7	PhD; William Wijns ¹² , MD, PhD, Andreas Baumbach ¹³ , MD, PhD, Christoph K Naber ¹⁴ , MD,
8	PhD.
9	
10	1 Instituto Cardiovascular de Buenos Aires, Buenos Aires, Argentina; 2 Kovai Medical Center,
11	Coimbatore, India; 3 Department of Family Medicine, School of Medicine, University of
12	Pretoria, Pretoria, South Africa; 4 University Heart Center Graz, Division of Cardiology,
13	Department of Medicine, Medical University Graz, Graz, Austria; 5 Tan Tock Seng Hospital,
14	Singapore, Singapore; 6 Wilgers Hospital, Pretoria, Gauteng, South Africa; 7 Division of
15	Cardiology, Queen Elizabeth Hospital, Kowloon, Hong Kong; 8 Hospital Garcia de Orta,
16	Almada, Portugal; 9 Department of Medicine, University of Khartoum, Khartoum, Sudan; 10
17	Centro Cardiológico Americano, Montevideo, Uruguay; 11 Amsterdam UMC, University of
18	Amsterdam, Heart Center; Department of Clinical and Experimental Cardiology, Amsterdam
19	Cardiovascular Sciences, Amsterdam, The Netherlands; 12 The Lambe Institute for
20	Translational Medicine and Curam, National University of Ireland Galway, Ireland; 13
21	Department of Cardiology, Barts Heart Centre, Barts Health NHS Trust, London, United
22	Kingdom; 14 Klinikum Wilhelmshaven, Wilhelmshaven, Germany
23	
24	Corresponding author: Christoph K Naber
25	Department of Medicine I, Klinikum Wilhelmshaven,
26	Friedrich Paffrathstr. 100, 26389 Wilhelmshaven
27	e-mail: christoph naher@klinikum-why de

How to Set Up Regional STEMI Networks: Providing Best Possible STEMI Care

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28	Conlfict of Interest: The authors have no conflicts of interest to declare
29 30	Abstract
31	Clinical guidelines recommend the development of ST-elevation myocardial infarction
32	(STEMI) networks at community, regional and/or national level to offer ideally primary
33	coronary angioplasty, or, at least the best available STEMI care to all patients.
34	However, there is a discrepancy between this clinical recommendation and daily
35	practice, with no coordinated care for STEMI patients in many regions of the world.
36	While this can be a consequence of lacking resources, in reality, it is more frequently a
37	lack of organizational power.
38	In this paper, the Stent-Save a life! Initiative proposes a practical methodology to
39	effectively set up a STEMI network in any region of the world, with existing resources,
40	and to continuously develop the STEMI network once established.
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41	Classifications
42 44	STEMI, Adjunctive Pharmacotherapy, Miscellaneous
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53 54 A	bbreviations	
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56	ACC/AHA	American College of Cardiology/American Heart Association
57	Cath Lab	cardiac catheterization laboratory
58	EMS	emergency medical service
59	ESC	European Society of Cardiology
60	FMC	first medical contact
61	GP/GC	general practitioner/general cardiologist
62	LMIC	low and middle-income countries
63	PI	pharmacoinvasive
64	PCI	percutaneous coronary intervention
65	pPCI	pharmacoinvasive percutaneous coronary intervention primary percutaneous coronary intervention Stent-Save a Life!
66	SSL	Stent-Save a Life!
67	STEMI	ST-elevation myocardial infarction
68	TIT	total ischemic time
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1. Introduction

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Primary percutaneous coronary intervention (pPCI) is the preferred reperfusion therapy for patients presenting with ST-elevation myocardial infarction (STEMI) as recommended by clinical guidelines (1). pPCI is clearly superior to all other treatments investigated to date regarding mortality and morbidity and is in addition cost saving for national economies (2). All health care systems should aim to provide pPCI to all STEMI patients, independent of location, nationality, race, sex or personal wealth. As a first step, available resources should be organized to provide the best available care to all patients and to optimize STEMI management nationwide. To reach this goal, systems of care for STEMI management need to be developed at community, regional and/or national level (3-5). This document proposes a universal methodology to provide the best available, guideline-adherent care for STEMI patients based on Eurolnie five general assumptions (Table 1).

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2. Review of clinical guidelines

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2.1 Therapeutic options

Reperfusion of the myocardium by pPCI within 12 h of symptom onset is the cornerstone of STEMI treatment, followed by a pharmacoinvasive strategy (PI) if pPCI cannot be performed within 120 minutes of diagnosis, or, if the latter is also not available, standalone fibrinolysis. In any PI or lysis strategy, patients should be urgently transferred to a PCI center after lysis (1, 6-8).

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2.2 Choice and timing of the optimal therapy

As a first step, all healthcare systems should develop regional networks of care for STEMI patients to encounter regional disparities as good as possible (9). Until timely pPCI can be provided to all patients, the preferred reperfusion strategy for each patient will depend on

107	resources, timing, and the entry point into the network (Figure 1; Table 3 and Supplementary
108	Table 1).
109	
110	3. Characteristics, elements and roles in a STEMI Network
111	Only an effectively organized STEMI network will ensure that all STEMI patients will be
112	optimally treated within the window of opportunity. All resources and processes in a region
113	should be organized to serve this single purpose.
114	
115	3.1 Main characteristics of a STEMI network:
116	- 24/7 treatment service for all STEMI patients
117	- structured cooperation between all parties involved following standardized protocols
118	- regular structured meetings and continuous education of all parties involved
119	- continuous self-assessment and improvement of the network
120	, rO/1,
121	3.2 Main players in a STEMI network
122	Patients
123	Ideally patients should be able to recognize symptoms of myocardial infarction and understand
124	the importance of receiving urgent treatment. They should understand how to activate the
125	emergency medical service (EMS) or otherwise seek immediate medical attention.
126	(Supplementary Table 2).
127	
128	General Practitioner/General Cardiologist (GP/GC)
129	GPs/GCs play an important role as first responders to patient consultations. GPs/GCs should be
130	integrated into a STEMI network and should be able to recognize and manage patients with
131	STEMI according to standardized protocols (Supplementary Table 2)

EMS are important coordinators of the referral pathway (3, 10). Their main actions entail pre- hospital patient management and between hospital transfer. An EMS should always coordinate
hospital patient management and between hospital transfer. An EMS should always coordinate
its actions with the network and notify the receiving hospital prior to arrival to check capacities
and allow for preparation. Ideally all EMS should be centralized and activated through a single
and well-publicized dispatch telephone number (1) (Supplementary Figure 1, Supplementary
Table 2).
Non-pPCI centers and hospitals without PCI facilities
These centers receive STEMI patients through two different pathways: directly from home or
community, or, via transfer by EMS. Non-pPCI centers should diagnose a STEMI within 10 min
after the patient's arrival and perform pPCI or transfer to a pPCI center, or handle a PI strategy
(Supplementary Figure 2; Supplementary Table 2).
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Primary PCI Centers
pPCI centers receive STEMI patients through one of three pathways: directly from home or
community, via transfer by EMS, or by secondary transportation from a non-pPCI center. They
should have a mandatory 24/7 Cath Lab available within 30 min of activation. They are obliged
to a "non-refusal" admission policy (Supplementary Figure 3, Supplementary Table 2).
4. Setting up a STEMI network
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4. Setting up a STEMI network Despite national or regional challenges, the implementation of a STEMI network is always

158	Stage 1: Preparation phase
159	The first step is to set up a local task force and an action plan for developing the network. This
160	task force is also responsible for assigning roles, developing standard protocols for diagnosis
161	and treatment in cooperation with the regional stakeholders, and, later, coordinating the
162	network.
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164	Stage 2: Mapping phase
165	In this phase, the task force identifies all potential pPCI and non-pPCI centers, estimates the
166	distances and the time needed for transportation, checks the availability of EMS services and
167	contacts the centers and the EMS to confirm their willingness to participate and ability to cope
168	with the demands. All these resources should be mapped to understand the regional situation
169	and to determine the best possible layout of regional network(s).
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171	and to determine the best possible layout of regional network(s). Stage 3: Building phase
172	Following the assumption, that the role of each player in any network is always defined by the
173	presence or absence of other players, any network can be categorized following the
174	specifications in Table 2 (Central Illustration). The task force assigns the individual roles to
175	each player, nominates the coordinators of the centers, the EMS, and the GP/GC groups.
176	
177	Stage 4: Quality assessment and continuous education phase
178	Quality assessment
179	At least one basic set (Supplementary Tables 3.1 and 3.2) of quality variables should be
180	established (7). This refers to performance parameters of all network components and includes
181	e.g. presentation timing, rate of patients treated, procedural success and in-hospital mortality.

The task force should meet periodically to analyze the performance and discuss necessary adaptations. The connection of reimbursement and compliance with standards can be a relevant steering instrument (11). One question that remains unanswered is, if having too many pPCI centers in a region may be disadvantageous, since each single center could end up having not enough experience and routine.

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Continuous education for professionals

Not all professionals involved have a basic training in cardiology. It may be important to offer specific educational and training programs for paramedics, nurses, technicians and nonrvention cardiology physicians on a recurrent basis due to staff rotation.

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Population awareness campaigns

Patient awareness of indicative symptoms and knowledge of how to effectively seek medical attention is key for the success of a STEMI network program, since the longest delays are usually caused by the patients (12). Awareness programs involving social media, entertainment industry, community organizations and scientific associations may be helpful; however, their effects quickly fade, once they are discontinued (13).

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Conclusions

The implementation of regional STEMI care systems overcomes local barriers and guarantees the best available reperfusion treatment for STEMI patients. A coordinated network of all stakeholders, guided by evidence-based, standardized protocols with a clear definition of roles and responsibilities are key, and, should be accompanied by a process of continuous improvement through evaluation of quality measures.

206	Figure Legends
207	Figure 1: Recommended reperfusion strategies according to timing and point of entry to the
208	network. EMS= emergency medical service; FL= fibrinolysis; GP/GC=general
209	practitioner/general cardiologist; PCI=percutaneous coronary intervention; pPCI= primary
210	percutaneous coronary intervention; min=minutes.
211	
212	Central Illustration: Typical combination of a hub-and-spoke network with an inner zone (green
213	circle) organized as pPCI network and an outer zone following a PI strategy (blue circle). The
214	external purple zone resembles a fibrinolysis network with no PCI center in reach.
215	GP/GC=general practitioner/general cardiologist; PI=pharmacoinvasive; pPCI= primary
216	external purple zone resembles a fibrinolysis network with no PCI center in reach. GP/GC=general practitioner/general cardiologist; PI=pharmacoinvasive; pPCI= primary percutaneous coronary intervention

Table 1. Key factors for any STEMI network

Factor	Assumption		
Players	The relevant stakeholders in any STEMI network are: patient and family, GP/GC,		
EMS, non-pPCI hospital, and pPCI-hospital.			
Roles The roles of the players are defined by i) evidence based clinical guidance a			
	presence or absence of other players.		
Scenarios The existing players and their roles determine the number of possible			
	network		
Treatment options A given scenario always defines the best available therapeutic option in a regi			
	STEMI patients.		
Quality metrics	Continuous monitoring and feedback is key to improve the network		
EMS=emergency medical service; GP/GC=general practitioner/general cardiologist; pPCI=primary percutaneous			
coronary intervention; S	STEMI=ST-segment elevation myocardial infarction		
Cob	STEWI—ST-segment elevation myocardia imalcuon		

Table 2. Network types according to existing resources.

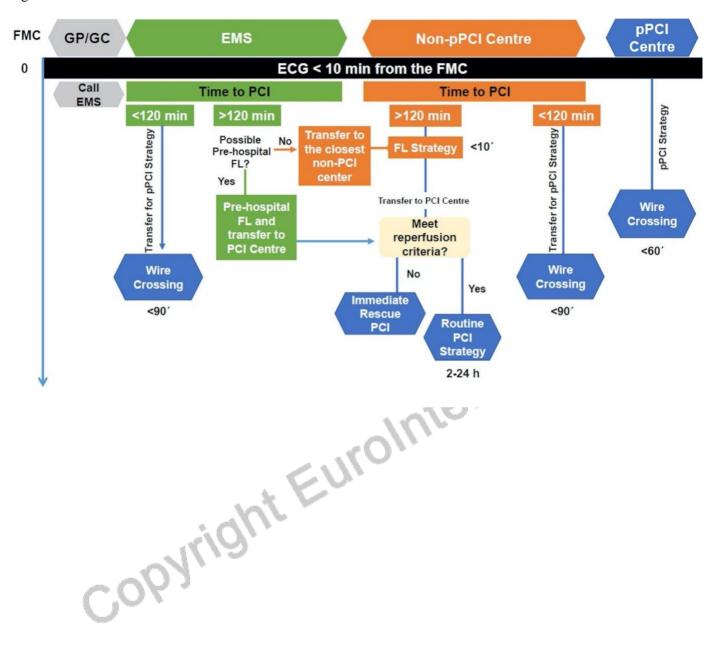
Primary PCI networks			
Ranking	Optimal long-term solution for STEMI care		
Mandatory Resources	pPCI centres which can be reached within 90 min after symptoms onset or diagnosis of		
	STEMI; EMS coordinated with the network		
Primary therapies	pPCI 24/7		
offered	If the pPCI centre is occupied or has a technical failure, a PI strategy is offered if pPCI		
	cannot be offered within guideline coherent timelines		
Processes	The EMS should bypass all other centres and transfer STEMI patients directly to the		
	closest pPCI centre		
	antilo		
Hub-and-spoke PCI ne	tworks		
Ranking	Acceptable long-term solution for STEMI care		
Mandatory Resources	pPCI centres; non-pPCI centres; EMS coordinated with the network		
Primary therapies pPCI, PI strategy			
offered	ight		
Processes	This model comprises two zones: i) the inner zone resembles a primary PCI network, the		
~ O	outer zone consists of non-pPCI centres or non-PCI hospitals which are connected to the		
00	inner zone via an EMS. They offer a PI strategy and either transfer the patient for PCI or		
	perform PCI in the same place during office hours following the recommended		
	timelines.		
Important steps to Turn non-pPCI centres into pPCI centres			
upgrade			
Pharmaco-invasive net	works		
Ranking	Transient solution for STEMI care, should be upgraded in the midterm.		
Mandatory Resources non-pPCI centres; an EMS, coordinated with the network, is highly desirable			

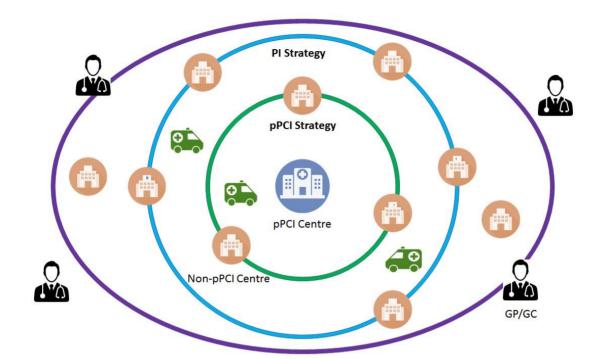
Primary therapies	PI strategy; primary PCI if patient arrives at PCI centre during office hours		
offered			
Processes	These networks offer a PI strategy 24/7 with fibrinolysis in all connected hospitals and		
	either transfer the patient for PCI or perform PCI during office hours. pPCI is offered if		
	patients arrive in a PCI hospital during office hours.		
Important steps to	Introduce an EMS, coordinated with the network		
upgrade 2. Turn non-pPCI centres into 24/7 pPCI centres			
Fibrinolysis networks			
Ranking	Transient organization which provides basic care for STEMI patients. Should be		
	upgraded as early as possible.		
Mandatory Resources	Medical centres without PCI option, able to recognize a STEMI and handle fibrinolysis;		
	an EMS, coordinated with the network is highly desirable; a remote ECG interpretation		
	service can be useful.		
Primary therapies	Standalone fibrinolysis		
offered			
Processes	These networks offer application of fibrinolysis 24/7		
Important steps to 1. Introduce an EMS, coordinated with the network			
upgrade	2. Install Cath Labs and expand their service to 24/7 pPCI		
Cath Lab=cardiac catheterization laboratory; EMS=emergency medical service; FMC=first medical contact;			
PCI=percutaneous coronary intervention; PI=pharmacoinvasive; pPCI=primary percutaneous coronary			
intervention; PI=pharmacoinvasive; STEMI=ST-elevation myocardial infarction			

Table 3. Definition of important time points and intervals in STEMI networks

Time period	Abbreviation	Definition
First medical contact	FMC	Time point when the patient is initially assessed by a physician, paramedic, nurse or trained EMS personnel who can obtain and interpret the ECG and deliver initial interventions. FMC can be in the pre-hospital setting or upon arrival at the hospital. In all scenarios, STEMI diagnosis via ECG should be obtained within 10 minutes.
Time of reperfusion	TOR	The time point of either a wire crossing the occlusion or the start of administration of lytic therapy
Total ischemic time	TIT	The time from symptom onset until reperfusion and is a strong predictor of patient outcomes. TIT comprises patient delay and system delay.
Patient delay	PD	Time interval between symptoms onset and FMC
System delay	SD	Time interval between FMC and time of reperfusion
Door-in-door-out time	DIDO	Time between patient arrival in a non-pPCI centre and the transfer to a pPCI centre
ECG=electrocardiogram; EMS=emergency medical service; pPCI=primary percutaneous coronary intervention;		
STEMI=ST-segment elevation infarction		

Figure 1





SUPPLEMENTARY MATERIAL

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