SECOND EDITION

## **CLINICAL DECISION-MAKING**

# TOOLKIT

**ACUTE CARDIOVASCULAR CARE ASSOCIATION** 





The ACCA Clinical Decision-Making Toolkit is produced by the Acute Cardiovascular Care Association. Developed and distributed through an educational grant from AstraZeneca and Novartis Pharma AG. AstraZeneca and Novartis Pharma AG were not involved in the development of this publication and in no way influenced its contents.





# The Acute Cardiovascular Care Association Clinical Decision-Making TOOLKIT

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Preface

The best care of patients with acute cardiovascular syndromes relies not only on specialists but also on systems of care that involve many non-cardiologists. Several of these syndromes require immediate diagnosis and decisions on treatment, some of them life-saving. Critical decisions must often be made quickly by professionals with different backgrounds and levels of expertise with limited resources. This poses a significant clinical challenge.

Against this background, the ACCA Clinical Decision-Making Toolkit was created as a comprehensive resource encompassing all aspects of acute cardiovascular care but structured as an easy-to-use instrument in environments where initial acute cardiovascular care is typically initiated. Comprehensive tables, clear diagrams and algorithms, based on the ESC clinical practice guidelines as well as in clinical experience should provide diagnostic and therapeutic guidance at a glance.

The Second Edition of the ACCA Toolkit has been updated with the 2014 and 2015 ESC Guidelines, and enriched with a new chapter with up-to-date coverage of drugs most frequently used in acute cardiovascular care. However, it does not replace textbooks and other sources of information that need to be consulted to reach an optimal management of these patients.

The ACCA Toolkit is available through different platforms:

Printed booklet, available at congresses where ESC-ACCA is represented

Web-based pdf file downloadable at www.escardio.org/ACCA

Mobile application for smartphones/tablets available in both Apple & Googleplay stores

Héctor Bueno, M.D., PhD., FESC, FAHA Editor in Chief

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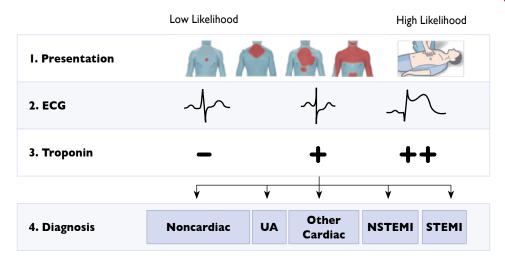
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### **CHAPTER I: KEY SYMPTOMS**

I.I CHEST PAIN	p.2
M. Lettino, F. Schiele	
<b>1.2 DYSPNEA</b>	p.9
1.3 <b>SYNCOPE</b>	p.16

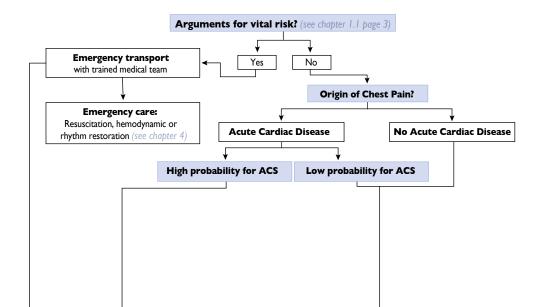
#### Initial assessment of patients with CHEST PAIN

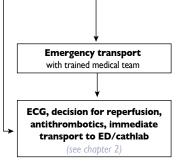
p.2

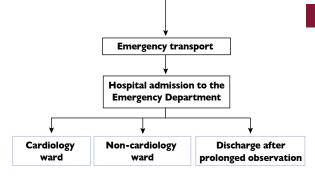


## Factors to be considered in the evaluation after the first call for CHEST PAIN

First call for chest pain	Higher risk / probability	Lower risk / probability
Arguments for vital risk	<ul> <li>Cardiorespiratory arrest, syncope / loss of consciousness, neurological defect</li> <li>Dyspnea</li> <li>Nausea – vomiting</li> <li>Arrhythmias – tachycardia</li> </ul>	Normal consciousness Normal breathing (see chapter 1.1 page 9) Normal heart rhythm
Context, CV risk	Age > 40 years, previous CV disease (MI, stroke, PE), modifiable CV risk factors (smoker, HTN, hypercholesterolemia, diabetes), chronic CV treatment	<ul> <li>Age &lt; 40 years,</li> <li>No previous CV disease</li> <li>No CV risk factors</li> <li>No chronic treatment</li> </ul>
Chest Pain	Medial / lateral thoracic pain, intense, with dyspnea	Depends on position/ palpation/ movements     Variable intensity, short duration (<1 min)     Hyperthermia
Cardiac Ischemic Pain	Retro-sternal, constriction, jaw/cervical/arm/back irradiation, spontaneous, prolonged > 20 min + dyspnea, sweating, lightheadedness, nausea	Lateral, abdominal irradiation     No neuro-vegetative symptoms



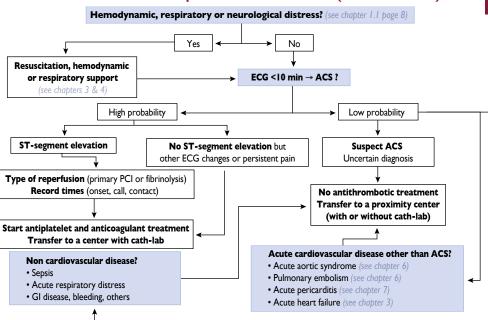




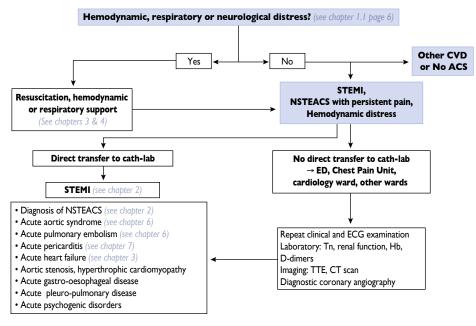
# Factors to be considered in the evalutation during the first medical contact for CHEST PAIN

First medical contact	Higher risk / probability	Lower risk / probability
Hemodynamic, respiratory, neurological distress	<ul> <li>Cardiopulmonary arrest, hypotension, tachycardia, shock</li> <li>Dyspnea, hypoxemia, lung rales (Killip class &gt;2)</li> <li>ECG: ST segment deviation</li> </ul>	<ul> <li>Normal consciousness, no motion defects</li> <li>Normal HR and BP</li> <li>Normal breathing and SpO<sub>2</sub>, no loss of pulse</li> </ul>
Probability for ACS	<ul> <li>Context, typical symptoms consistent with myocardial ischemia</li> <li>ECG changes</li> <li>Bedside Tn</li> </ul>	<ul> <li>No CV risk, atypical symptoms, normal ECG</li> <li>Negative bedside Tn only if onset of pain &gt;6 hours (see chapter 2.1 page 24)</li> </ul>
STEMI NSTEACS Uncertain diagnosis (see chapter 2.1 page 24)	<ul> <li>ECG criteria for STEMI (see chapter 2.3 page 35)</li> <li>ST depression or normal ECG</li> <li>Normal ECG → Repeat 12-lead ECG recording</li> </ul>	Other ST-segment abnormalities not related to STEMI (see chapter 2.3)
Type of reperfusion  Time assessment	Primary PCI or thrombolysis? Primary PCI if delay <120 (preferably <90) min or <60 min if onset of pain <120 min Consider age, anterior wall location Times: Onset of pain, call, first medical contact, ECG, door, balloon inflation or needle (lytic drug) administration	No reperfusion if delay >12 h, no symptoms, no ST-segment elevation

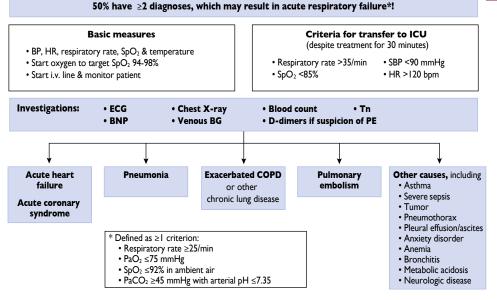
First medical contact in patients with CHEST PAIN (home-ambulance)



Management of patients with CHEST PAIN (emergency room)



#### **DYSPNEA:** Diferential diagnosis



Reference: Ray P et al. Acute respiratory failure in the elderly: etiology, emergency diagnosis and prognosis. Critical Care (2006), 10 (3):R82.

#### DYSPNEA: Acute heart failure (see chapter 3.1)

#### **BASIC WORK-UP**

 Immediate 12-lead ECG, cardiac monitor, BP, respiratory rate, pulse oximetry

Clinical findings

Most commonly: lower extremity edema, jugular venous distension, rales; work up for underlying cardiac disease and triggers

· Laboratory findings

Complete blood count, chemistries, cardiac enzymes, BNP, TSH, ABG as needed

· Chest X-ray (lung ultrasound)

Echocardiogram

During admission (earlier if decompensated aortic stenosis or endocarditis are suspected)

Coronary angiography
 Emergent in patients with ACS; delayed in patients with suspected coronary artery disease

Positioning Keep head of bed elevated above level of legs

 $\bullet$  Oxygen Up to 12 L/min via non-rebreather, titrate oxygen saturation to 95%

• Nitroglycerin I-2 SL tablets or 2-3 patches 10 mg (1st choice). In pulmonary edema with severe shortness of breath:

NTG drip 0.05% (100 mg in 200 ml)

- Start with 25 µg/min = 3 ml/h, check BP after 5 and 10 min

- Increase dose per SHO/attending recommendations by 25  $\mu$ g/min at a time as long as SBP >90 mmHg

- Additional BP check 5 and 10 min after each increase in dosing

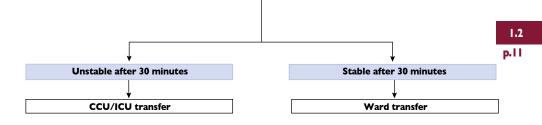
- Check BP every 20 min once a steady drip rate is reached

• Furosemide 40-120 mg i.v. (adjust based on kidney function and clinical findings; monitor creatinine)

• Morphine 2 mg i.v. (preceeded by 10 mg i.v. metoclopramide PRN)

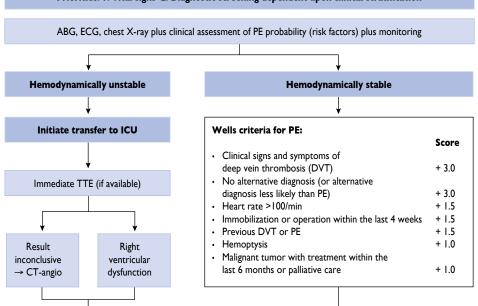
• Consider digoxin 0.5 (-1.0) mg i.v. in patients with atrial fibrillation

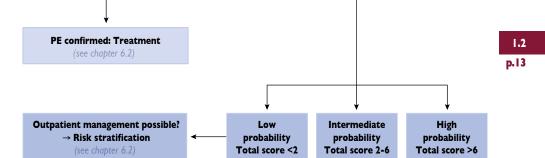
Anticoagulation Therapeutic dosing in ACS and atrial fibrillation: Enoxaparin 1 mg/kg body weight as 1st dose



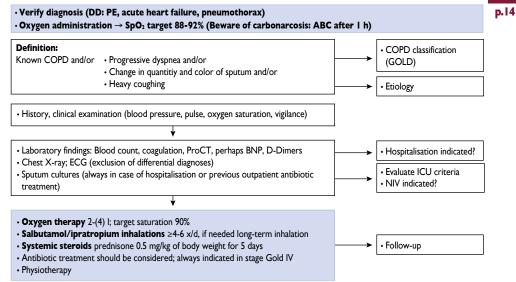
#### DYSPNEA: Acute pulmonary embolism (see chapter 6.2)

#### Priorities: I. Vital signs 2. Diagnostic screening dependent upon clinical stratification



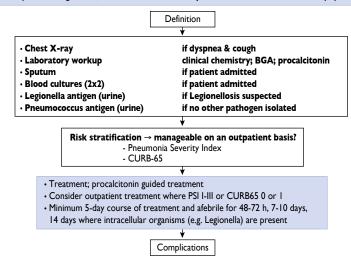


#### **DYSPNEA: COPD exacerbation**



#### **DYSPNEA: Community-acquired pneumonia**

#### Objective: diagnostics, risk stratification & empirical immediate treatment <2(-4) hrs.



Copyrights: Mandell LA et al. Infectious Diseases Society of America/American Thoracic Society consensus guidelines on the management of community-acquired pneumonia in adults. Clin Infect Dis. (2007);44 Suppl 2:527-72. - Halm EA and Teirstein AS. Management of Community-Acquired Pneumonia New Engl J Med (2002); 347:2039-2045 - Woodhead M et al. Guidelines for the management of adult lower respiratory tract infections ERJ December 1, (2005); 26 (6) 1138-1180.

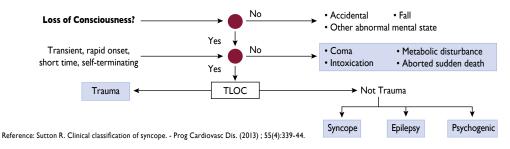
# **SYNCOPE:** Assessment of patients with transient loss of conscioussness (TLOC)

Syncope is a transient loss of consciousness due to global cerebral hypoperfusion (usually, itself due to a period of low blood pressure) characterised by rapid onset, short duration, spontaneous and complete recovery.

The differentiation between syncope and non-syncopal conditions with real or apparent LOC can be achieved in most cases with a **detailed clinical history** but sometimes can be extremely difficult. The following questions should be answered:

- Was LOC complete?
- Was LOC transient with rapid onset and short duration?
- Did the patient recover spontaneously, completely and without sequelae?
- Did the patient lose postural tone?

If the answers to these questions are positive, the episode has a high likelihood of being syncope. If the answer to one or more of these questions is negative, exclude other forms of LOC before proceeding with syncope evaluation.



# SYNCOPE: Diagnostic criteria (I) Diagnostic criteria with initial evaluation

**Vasovagal syncope** is diagnosed if syncope is precipitated by emotional distress or orthostatic stress and is associated with typical prodrome.

Situational syncope is diagnosed if syncope occurs during or immediately after specific triggers.

**Orthostatic syncope** is diagnosed when it occurs after standing up and there is documentation of orthostatic hypotension.

Arrhythmia related syncope is diagnosed by ECG when there is:

- Persistent sinus bradycardia <40 bpm in awake or repetitive sinoatrial block or sinus pauses >3 s
- Mobitz II 2nd or 3rd degree AV block
- Alternating left and right BBB
- VT or rapid paroxysmal SVT
- Non-sustained episodes of polymorphic VT and long or short QT interval
- Pacemaker or ICD malfunction with cardiac pauses

Cardiac ischemia related syncope is diagnosed when syncope presents with ECG evidence of acute ischemia with or without myocardial infarction.

Cardiovascular syncope is diagnosed when syncope presents in patients with prolapsing atrial myxoma, severe aortic stenosis, pulmonary hypertension, pulmonary embolus or acute aortic dissection.

Reference: Moya A et al. Eur Heart J(2009) 30, 2631-2671 (I).

#### SYNCOPE: Evaluation and risk stratification of patients with suspected syncope

Once syncope is considered to be the likely diagnosis, risk stratification is required to determine further management.

Patients with suspected syncope presenting to ED or clinic "Uncertain" or unexplained syncope Certain diagnosis of syncope Initiate therapy Inpatient SMU, outpatient SMU or Risk stratification personal physician as appropriate High risk Intermediate risk Low risk Observation Unit Home Home if stable. Outpatient SMU Admit to hospital referral if evidence of high risk Outpatient SMU Hospital admission for diagnosis, treatment

and follow-up as appropriate

Copyright: Sutton R, Brignole M, Benditt DG. Key challenges in the current management of syncope. Nat Rev Cardiol. (2012);(10):590-8.

Inpatient SMU

# SYNCOPE: Diagnostic criteria (2) Diagnostic criteria with provocation maneuvers

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Carotid sinus massage	Orthostatic Hypotension
<ul> <li>Indications</li> <li>CSM is indicated in patients &gt;40 years with syncope of unknown aetiology after initial evaluation;</li> <li>CSM should be avoided in patients with previous MI, TIA or stroke within the past 3 months and in patients with carotid bruits (except if carotid Doppler studies excluded significant stenosis)</li> </ul>	Recommendations: Active standing Indications  Manual intermittent determination with sphygmomanometer of BP supine and, when OH is suspected, during active standing for 3 min is indicated as initial evaluation;  Continuous beat-to-beat non-invasive pressure measurement may be helpful in cases of doubt
CSM is diagnostic if syncope is reproduced in presence of asystole longer than 3 s and/or a fall in systolic BP >50 mmHg	Diagnostic criteria The test is diagnostic when there is a symptomatic fall in systolic BP from baseline value ≥20 mmHg or diastolic BP ≥10 mmHg or a decrease in systolic BP to <90 mmHg; The test should be considered diagnostic when there is an asymptomatic fall in systolic BP from baseline value ≥20 mmHg or diastolic BP >10 mmHg or a decrease in systolic BP to <90 mmHg

Reference: Moya A et al. Eur Heart J(2009) 30, 2631-2671 (2).

#### Treatment according to type of SYNCOPE (I)

p.20

	·
Treatment of reflex syncope	Treatment of orthostatic hypotension
<ul> <li>Explanation of the diagnosis, provision of reassurance and explanation of risk of recurrence are in all patients</li> <li>Isometric PCM are indicated in patients with prodrome</li> <li>Cardiac pacing should be considered in patients with dominant cardioinhibitory CSS</li> <li>Cardiac pacing should be considered in patients with frequent recurrent reflex syncope, age &gt; 40 years and documented spontaneous cardioinhibitory response during monitoring</li> <li>Midodrine may be indicated in patients with VVS refractory to lifestyle measures</li> <li>Tilt training may be useful for education of patients but long-term benefit depends on compliance</li> <li>Cardiac pacing may be indicated in patients with tilt-induced cardioinhibitory response with recurrent frequent unpredictable syncope and age &gt; 40 after alternative therapy has failed</li> <li>Triggers or situations inducing syncope must be avoided as much as possible</li> <li>Hypotensive drugs must be modified or discontinued</li> <li>Cardiac pacing is not indicated in the absence of a documented cardioinhibitory reflex</li> <li>Beta-adrenergic blocking drugs are not indicated</li> <li>Fluid consumption and salt in the diet should be increased</li> </ul>	Adequate hydration and salt intake must be maintained     Midodrine should be administered as adjunctive therapy if needed     Fludrocortisone should be administered as adjunctive therapy if needed     PCM may be indicated     Abdominal binders and/or support stockings to reduce venous pooling may be indicated     Head-up tilt sleeping (>10°) to increase fluid volume may be indicated     Triggers or situations inducing syncope must be avoided as much as possible     Hypotensive drugs administered for concomitant conditions must be discontinued or reduced

#### Treatment according to type of SYNCOPE (2)

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#### Treatment of arrhythmic syncope

#### **Cardiac Pacing**

- Pacing is indicated in patients with sinus node disease in whom syncope is demonstrated to be due to sinus arrest (symptom-ECG correlation) without a correctable cause
- Pacing is indicated in sinus node disease patients with syncope and abnormal CSNRT
- Pacing is indicated in sinus node disease patients with syncope and asymptomatic pauses > 3 sec. (with possible exceptions of young trained persons, during sleep and in medicated patients)
- Pacing is indicated in patients with syncope and 2nd degree Mobitz II, advanced or complete AV block
- · Pacing is indicated in patients with syncope, BBB and positive EPS
- Pacing should be considered in patients with unexplained syncope and BBB
- Pacing may be indicated in patients with unexplained syncope and sinus node disease with persistent sinus bradycardia itself asymptomatic
- Pacing is not indicated in patients with unexplained syncope without evidence of any conduction disturbance

#### Catheter ablation

- Catheter ablation is indicated in patients with symptom/ arrhythmia ECG correlation in both SVT and VT in the absence of structural heart disease (with exception of atrial fibrillation)
- Catheter ablation may be indicated in patients with syncope due to the onset of rapid atrial fibrillation

#### **Antiarrhythmic drug therapy**

- Antiarrhythmic drug therapy, including rate control drugs, is indicated in patients with syncope due to onset of rapid atrial fibrillation
- Drug therapy should be considered in patients with symptom/ arrhythmia ECG correlation in both SVT and VT when catheter ablation cannot be undertaken or has failed

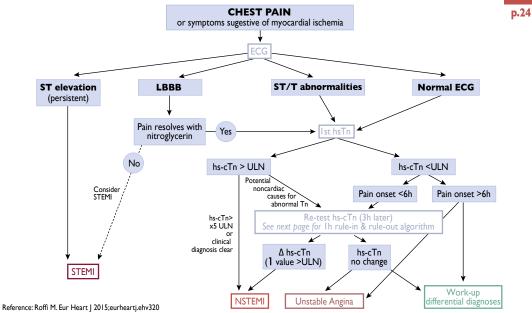
#### Implantable Cardioverter Defibrillator (ICD)

- ICD is indicated in patients with documented VT and structural heart disease
- ICD is indicated when sustained monomorphicVT is induced at EPS in patients with previous myocardial infarction
- ICD should be considered in patients with documented VT and inherited cardiomyopathies or channelopathies

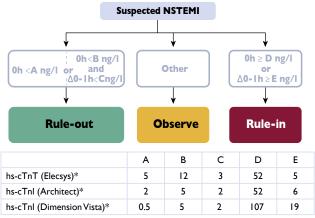
# CHAPTER 2: ACUTE CORONARY SYNDROMES

2.1 GENERAL CONCEPTS	p.2
2.2 NON ST-SEGMENT ELEVATION ACS	р.2
2.3 ST-SEGMENT ELEVATION MI (STEMI)	p.3

#### **ACUTE CORONARY SYNDROMES: Diagnosis (I)**



## ACUTE CORONARY SYNDROMES: Diagnosis (2) 0-1 H Rule-in & rule out test for NSTEMI



\*Cut-off levels are assay-specific.

- NSTEMI can be ruled-out at presentation, if hs-cTn concentration is very low
- NSTEMI can be ruled out by the combination of low baseline levels and the lack of a relevant increase within 1 h
- NSTEMI is highly likely if initial hs-cTn concentration is at least moderately elevated or hs-cTn concentrations show
  a clear rise within the first hour

#### **ACUTE CORONARY SYNDROMES: Differential diagnosis (I)**

Causes of chest pain Not related to ACS	Causes of troponin elevation  Not related to ACS
Primary cardiovascular  Acute pericarditis, pericardial effusion  Acute myocarditis  Severe hypertensive crisis  Stress cardiomyopathy (Tako-Tsubo syndrome)  Hypertrophic cardiomyopathy, aortic stenosis  Severe acute heart failure  Acute aortic syndrome (dissection, hematoma)  Pulmonary embolism, pulmonary infarction  Cardiac contusion	Primary cardiovascular  Acute myo(peri)carditis  Severe hypertensive crisis  Pulmonary edema or severe congestive heart failure  Stress cardiomyopathy (Tako-Tsubo syndrome)  Post- tachy- or bradyarrhythmias  Cardiac contusion or cardiac procedures (ablation, cardioversion, or endomyocardial biopsy)  Aortic dissection, aortic valve disease or hypertrophic cardiomyopathy  Pulmonary embolism, severe pulmonary hypertension
Primary non-cardiovascular  Oesophageal spasm, oesophagitis, Gastro Esophageal Reflux (GER)  Peptic ulcer disease, cholecystitis, pancreatitis Pneumonia, bronchitis, asthma attack Pleuritis, pleural effusion, pneumothorax Pulmonary embolism, severe pulmonary hypertension Thoracic trauma Costochondritis, rib fracture Cervical / thoracic vertebral or discal damage Herpes Zoster	Primary non-cardiovascular  Renal dysfunction (acute or chronic) Critical illness (sepsis, repiratory failure) Acute neurological damage (i.e. stroke, subarachnoid hemorrhage) Severe burns (affecting >30% of body surface area) Rhabdomyolysis Drug toxicity (chemotherapy with adriamycin, 5-fluorouracil, herceptin, snake venoms) Inflammatory or degenerative muscle diseases Hypothyroidism Infiltrative diseases (amyloidosis, hemochromatosis, sarcoidosis) Scleroderma

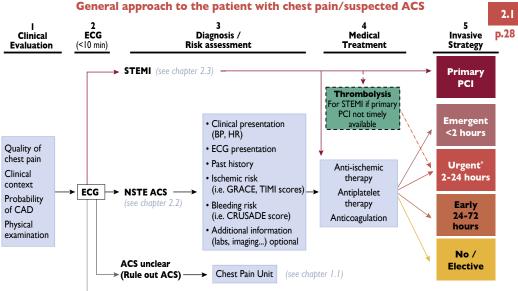
**Negative T waves** 

# ACUTE CORONARY SYNDROMES: Differential diagnosis (2) Causes of repolarisation abnormalities in the ECG not related to ACS

**ST-**segment elevation

Fixed  • LV aneurysm  • LBBB, WPW, hypertrophic cardiomyopath  • Pacemaker stimulation  • Early repolarisation (elevated J-point)  Dynamic  • Acute (myo)pericarditis  • Pulmonary embolism  • Electrolyte disturbances (hyperkalemia)  • Acute brain damage (stroke, subarachnoide)  • Tako Tsubo syndrome	,	Normal variants, i.e. women (right precordial leads), children, teenagers Fevolutive changes post myocardial infarction Chronic ischemic heart disease Acute (myo)pericarditis, cardiomyopathies BBB, LYH, WPW Post-tachycardia or pacemaker stimulation Metabolic or ionic disturbances
ST-segment dep	ression	Prominent T waves
Fixed  • Abnormal QRS (LBBB, WPW, pacemaker • LVH, hypertrophic cardiomyopathy • Chronic ischemic heart disease  Dynamic • Acute (myo)pericarditis • Acute pulmonary hypertension • Electrolyte disturbances (hyperkalemia) • Intermitent LBBB, WPW, pacing • Post-tachycardia / cardioversion	Severe hypertensive crisis Drug effects (digoxin) Shock, pancreatitis Hyperventilation Tako Tsubo syndrome	Normal variants, i.e. early repolarisation Metabolic or ionic disturbances (i.e. hyperkalemia) Acute neurological damage (stroke, subarachnoid haemorrhage)





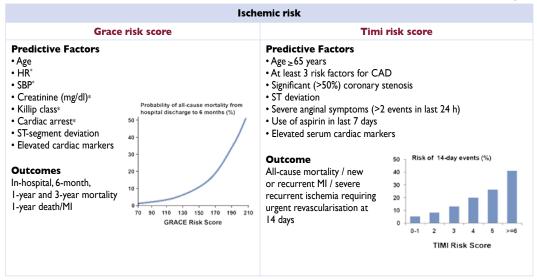
Rule out noncardiac causes

No ACS -

<sup>\* 3-12</sup> hours after thrombolysis.

#### NON ST-SEGMENT ELEVATION ACS: Risk stratification (I)

p.29



<sup>\*</sup> At admission.

#### NON ST-SEGMENT ELEVATION ACS: Risk stratification (2)

# Bleeding risk Crusade risk score Predictive Factors • Sex • HR\* • SBP\* • Creatinine (mg/dl)\* • Baseline hematocrit\* • GFR: Cockcroft-Gault\* • Diabetes • Prior vascular disease • Signs of congestive heart failure\*

**Outcome** 

In-hospital major bleeding

Copyrights: Eagle KA et al. A validated prediction model for all forms of acute coronary syndrome: estimating the risk of 6-month post-discharge death in an international registry. JAMA. (2004) ;291(22):2727-33.

CRUSADE Bleeding Score

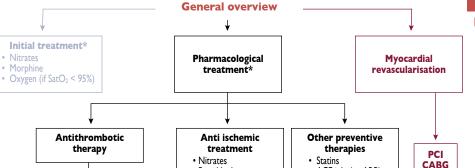
Antman EM, et al. The TIMI risk score for unstable angina/non-ST elevation MI: A method for prognostication and therapeutic decision making. JAMA. (2000);284(7):835-42.

Subherwal S, 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(14):1873-82.

<sup>\*</sup> At admission.



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Beta-blockers

Calcium antagonists

**NON ST-SEGMENT ELEVATION ACS: Treatment (I)** 

#### Anticoagulation

One of the following:

- Fondaparinux
- Enoxaparin
- UFH

#### Bivalirudin

#### **Antiplatelets**

Aspirin + one of:

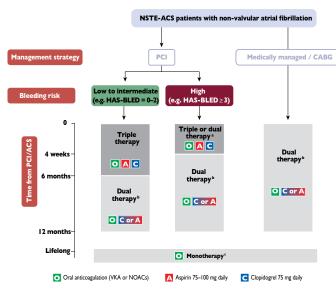
- Ticagrelor
- Prasugrel
- Clopidogrel
- Optionally:
  - GP IIb/IIIa inhibitors
  - Cangrelor

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

· ACE inh. (or ARB) Aldosterone inhibitors.

### NON ST-SEGMENT ELEVATION ACS: Treatment (2) Antithrombotic strategies in patients with NSTE-ACS and non-valvular atrial fibrillation

p.32



CHA2DS2-VASc = Cardiac failure, Hypertension, Age ≥ 75 [2 points], Diabetes, Stroke [2 points] – Vascular disease, Age 65–74, Sex category.

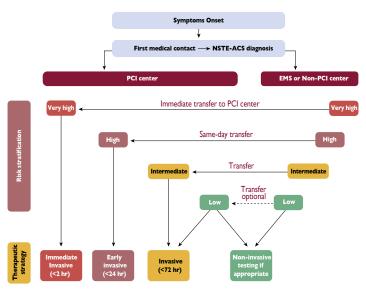
- <sup>a</sup> Dual therapy with oral anticoagulation and clopidogrel may be considered in selected patients (low ischaemic risk).
- Aspirin as an alternative to clopidogrel may be considered in patients on dual therapy (i.e., oral anticoagulation plus single antiplatelet); triple therapy may be considered up to 12 months in patients at very high risk for ischaemic events.
- c Dual therapy with oral anticoagulation and one antiplatelet agent (aspirin or clopidogrel) beyond one year may be considered in patients at very high risk of coronary events.
- In patients undergoing coronary stenting, dual antiplatelet therapy may be an alternative to triple or a combination of anticoagulants and single antiplatelet therapy if the CHA2DS2-VASc score is I (males) or 2 (females).

Reference: Eur Heart | 2015;eurheartj.ehv320- Figure 5.

## NON ST-SEGMENT ELEVATION ACS: Treatment (3) Risk criteria mandating invasive strategy in NSTE-ACS

Very-high-risk criteria	Haemodynamic instability or cardiogenic shock     Recurrent or ongoing chest pain refractory to medical treatment     Life-threatening arrhythmias or cardiac arrest     Mechanical complications of MI     Acute heart failure     Recurrent dynamic ST-T wave changes, particularly with intermittent ST-elevation
High-risk criteria	Rise or fall in cardiac troponin compatible with MI Dynamic ST- or T-wave changes (symptomatic or silent) GRACE score > 140
Intermediate- risk criteria	Diabetes mellitus Renal insufficienty (eGFR <60 mL/min/1.73 m²)  LVEF <40% or congestive heart failure Early post-infarction angina Prior PCI Prior CABG GRACE risk score >109 and <140
Low-risk criteria	Any characteristics not mentioned above

## NON ST-SEGMENT ELEVATION ACS: Treatment (4) Timing and strategy for invasive management



#### **STEMI: Electrocardiographic diagnosis**

STEMI is diagnosed according to the presence of the following acute ischemic ECG changes:

#### In the absence of LVH and LBBB:

- New ST elevation at the J point in 2 contiguous leads with ≥ 0.2 mV in men or ≥ 0.15 mV in women in leads V<sub>2</sub>-V<sub>3</sub> and/or ≥ 0.1 mV in other leads
  - $\rightarrow$  Contiguous leads mean lead groups such as anterior leads (V<sub>1</sub>-V<sub>6</sub>), inferior leads (II, III, aVF) or lateral/apical leads (I, aVL).

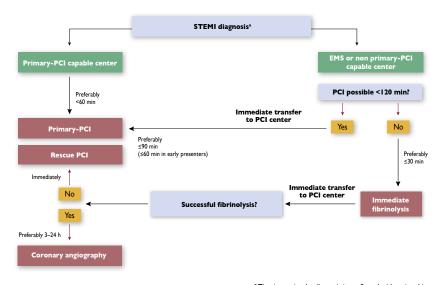
#### In the presence LBBB or ST depression:

- New LBBB, and symptoms suggestive of ACS
- ST depression in leads  $V_1 V_3$  indicate inferobasal myocardial ischemia (especially when the terminal T-wave is positive)

#### In suspected posterior (circumflex artery- related) or right ventricle-related infarction:

- ST elevation in V<sub>7</sub> (at the left posterior axillary line), V<sub>8</sub> (at the left midscapular line), and V<sub>9</sub> (at the left paraspinal border), using a cut-point > 0.05 mV
  - ightarrow Capture an overlooked left dominant circumflex using posterior leads in the fifth interspace
- ST elevation in right precordial leads ( $V_3R$  and  $V_4R$ ), using a cut-off point > 0.05 mV, and > 0.1 mV in men <30 years
  - → Capture suspected right ventricular infarction using right precordial leads

### STEMI: Treatment (I) General overview of initial management



<sup>&</sup>lt;sup>a</sup> The time point the diagnosis is confirmed with patient history and ECG ideally within 10 min from First Medical Contact (FMC). All delays are related to FMC.

### STEMI: Treatment (2) Primary PCI - First 24 hours and days 2-7

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

Pre hospital	PCI	 	CCU/I	CCU
Acetylicsalisylic Acid Heparin	300 mg 70 IU/kg	Bivalirudir or GPI:	n Eptifibatide Tirofiban	
or Prasugrel 60	mg mg mg	Follow loc	Abxicimab cal in-lab instr	ruction / dosing
		or c	oprolol arvedilol isoprolol	25 mg x 2 3,25 mg x 2 2,5 mg x 2
		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	rvastatin Kosuvastatin	80 mg x I 40 mg x I

Medication							
Titration Day 2-7							
75 mg x I							
90 mg x 2							
10/5 mg x 1							
75 mg x I							
200mg x I							
25 mg x 2							
5 mg x 2							
or Ca-antagonist (see chapter 2.2)							
Start ACE-i or ARB in DM, LVSD, CHF, or to control BP Aldosterone RB Start or continue anti-diabetic medication							

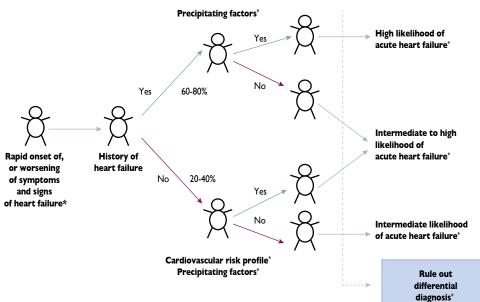
Reference: Steg G et al. Eur Heart J. (2012);33:2569-619  $^{(7)}$ .

Pre-hospital management of patients with chest pain and/or dyspnoea of cardiac origin. A position paper of the Acute Cardiovascular Care Association (ACCA) of the ESC - European Heart Journal: Acute Cardiovascular Care August 27, 2015 2048872615604119.

### **CHAPTER 3: ACUTE HEART FAILURE**

3.1 HEART FAILURE AND PULMONARY OEDEMA	. р.40
I.C.C. van der Horst, G. Filippatos	
3.2 CARDIOGENIC SHOCK	. р. <b>5</b> 2
P. Vranckx, U. Zeymer	

#### **ACUTE HEART FAILURE: Diagnosis and causes (I)**



<sup>\* (</sup>See page 41).

#### **ACUTE HEART FAILURE: Diagnosis and causes (2)**

- 1. Symptoms: Dyspnea (on effort or at rest)/breathlessness, fatigue, orthopnea, cough, weight gain/ankle swelling
- 2. Signs: Tachypnea, tachycardia, low or normal blood pressure, raised jugular venous pressure, 3<sup>rd</sup>/4<sup>th</sup> heart sound, rales, oedema, intolerance of the supine position
- 3. Cardiovascular risk profile: Older age, HTN, diabetes, smoking, dyslipidemia, family history, history of CVD
- 4º Precipitating factors: Myocardial ischemia, rhythm disturbances, medication (NSAID, negative inotropic agents), infection, noncompliance
- 5• Differential diagnosis: Exacerbated pulmonary disease, pneumonia, pulmonary embolism, pneumothorax, acute respiratory distress syndrome, (severe) anaemia, hyperventilation (acidosis), sepsis/septic shock, redistributive/hypovolemic shock
- 6• Likelihood: Depending on the site off presentation the underlying cause of acute heart failure is likely to differ. Cardiologists see more often worsening heart failure and physicians at the Emergency Department more often see patients with preserved systolic left ventricular function

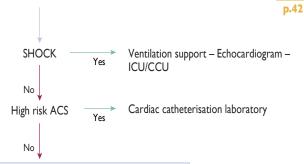
#### MAIN CAUSES OF ACUTE HEART FAILURE

- Coronary artery disease
- Hypertension
- Cardiomyopathy (familial, acquired)
- Valvular heart disease
- Peri-/endocardial disease

- Congenital heart disease
- Arrhythmia (tachy-, brady-)
- Conduction disorder (blocks)
- Volume overload (renal, iatrogenic)
- Tumor

- Pleural effusion
- Anxiety disorder
- Neurologic disease

Reference: McMurray JJ et al, Eur Heart J (2012);33(14):1787-847 (19).

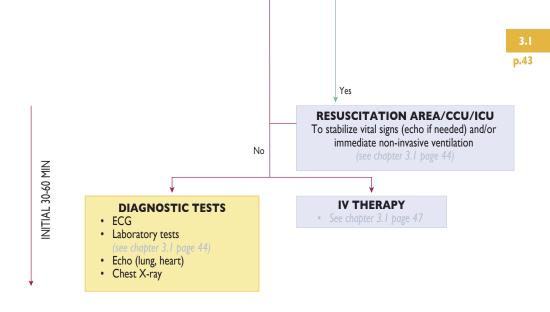


#### **SEVERITY SCORE** (excluding shock)

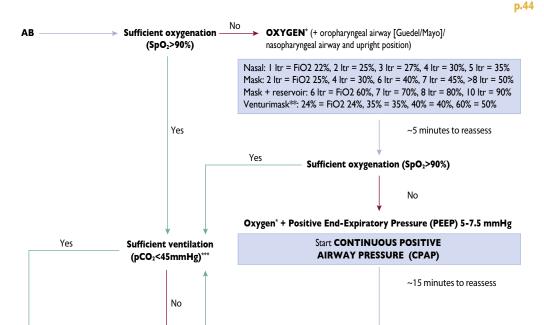
# Respiratory distress RR > 25/min, SpO<sub>2</sub><90% on O<sub>2</sub>, or increased work of breathing

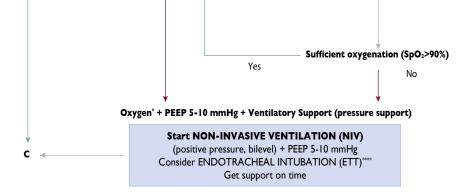
### Haemodynamic instability

Low or high blood pressure, Severy arrhythmia, HR <40 or >130/min



### ACUTE HEART FAILURE: Initial diagnosis and treatment Airway (A) & Breathing (B)





<sup>\*</sup> Goal SpO, 94-98%.

<sup>\*\*</sup> Use the predefined liters of oxygen. When using higher flows the  ${\rm FiO_2}$  will drop.

<sup>\*\*\*</sup> For a patient with COPD, a pCO, of 45-50 mmHg may be optimal. Aim for a normal pH.

<sup>\*\*\*\*</sup> Consider if the above fails or when patient is fatigued.

#### **ACUTE HEART FAILURE: Initial diagnosis (CDE)**

C - CIRCULATION\* HR (bradycardia [<60/min], normal [60-100/min], tachycardia [>100/min]), rhythm (regular, irregular), SBP (very low [<90t mmHg], low, normal [110-140 mmHg], high [>140 mmHg]), and elevated jugular pressure should be checked

p.46

#### INSTRUMENTATION & INVESTIGATIONS:

Consider intravenous (central) & arterial line (BP monitoring)

#### Laboratory measures

- Cardiac markers (troponin, (BNP/NT-proBNP, MR-proANP)
- Complete blood count, electrolytes, creatinine, urea, glucose, inflammation, TSH

#### Standard 12-lead FCG

- Venous blood gases, D-dimer (suspicion of acute pulmonary embolism)
- · Rhythm, rate, conduction times?
- Signs of ischemia/myocardial infarction? Hypertrophy?

#### **Echocardiography**

- Ventricular function (systolic and diastolic)?
- Presence of valve dysfunction (severe stenosis/insufficiency)?
- Pericardial effusion/tamponade?

#### **ACTIONS:**

Rule in/out diagnosis of acute heart failure as diagnosis for symptoms and signs

Establish cause of disease

Determine severity of disease

Start treatment as soon as possible. i.e. both heart failure and the factors identified as triggers

#### D - DISABILITY DUE TO NEUROLOGICAL DETERIORATION

Normal consiousness/altered mental status? Measurement of mental state with AVPU (alert, visual, pain or unresponsive) Glasgow Coma Scale: FMV score <8 → Consider FTT

Anxiety, restlessness? -> Consider morphine 2.0-5 mg i.y. bolus (diluted in normal saline), preceded by metoclopramide 10 mg i.y. PRN

#### **E-EXPOSURE & EXAMINATION**

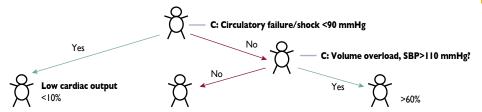
Temperature/fever: central and peripheral

Weight

Skin/extremities: circulation (e.g. capilary refill), color

Urinary output (<0.5ml/kg/hr) → Insert indwelling catheter; the benefits should outweigh the risks of infection and long-term complications

#### **ACUTE HEART FAILURE: Initial treatment (C) IV therapy**



#### I Inotropic drugs

- Dobutamine 2.5 μg/kg/min
- Milrinone bolus 25 μg/kg in I 0-20 min, continuous 0.375 μg/kg/min
- 2 Vasopressor i.v.
  - Norepinephrine 0.2 µg/kg/min
- 3 Diuretics i.v.
- Furosemide 20-40 mg bolus, continuous 100 mg/6 h
- 4 Consider hypertonic saline + diuretic
- 5 Consider mechanical circulatory support

#### I Diuretics i.v.

 Furosemide 20-40 mg bolus, continuous 100 mg/6 h\*

#### 2 Inotropic drugs

- Dobutamine continuous 2.5 µg/kg/min
- Milrinone bolus 25 µg/kg in 10-20 min, continuous 0.375 µg/kg/min
- Levosimendan bolus 12 µg/kg in 10 min, continuous 0.1 µg/kg/min
- 3 Consider to start ACE-I/ARB, beta-blocker, MRA.

\*See Chapter 8: use of drugs in acute Cardiovascular Care.

(See table page 50-51)

#### I Vasodilators

- Nitroglycerine spray 400 μg sublingual, repeat ~5-10 min
- Nitroglycerine i.v. continuously
- ~10 µg/min, increase ~5 µg/min • Nitroprusside 0.3 µg/kg/min
- increase to 5 microg/kg/min

#### 2 Diuretics i.v.

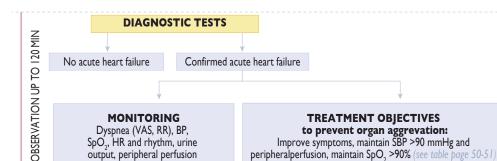
- Furosemide 20-40 mg bolus, continuous 100 mg/6 h
- 3 Consider to start ACE-I/ARB, beta-blocker, MRA.

\*See Chapter 8: use of drugs in acute Cardiovascular Care.

(See table page 50-51)

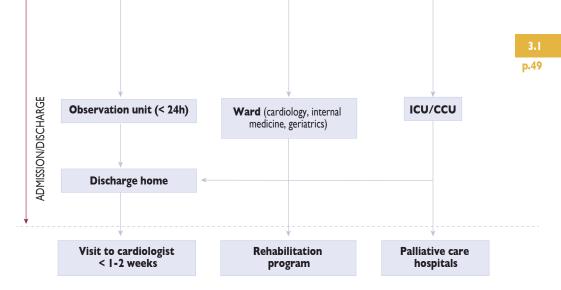
<sup>\*</sup> Use higher dose in patients on chonic diuretic treatment for HF (i.e. 2.5 times normal dose).

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#### REASSESSMENT

Clinical, biological and phychosocial parameters by trained nurses



#### **ACUTE HEART FAILURE: Treatment (C) and preventive measures**

#### Management of oral therapy in AHF in the first 48 hours

p.su

	Normotension/ Hypertension	Hypotension		Low Heart rate		Potassium		Renal impairment	
		<100 >85 mmHg	<85 mmHg	<60 ≥50 bpm	<50 bpm	≤3.5 mg/dL	>5.5 mg/dL	Cr < 2.5, eGFR > 30	Cr > 2.5, eGFR < 30
ACE-I/ARB	Review/increase	Reduce/ stop	Stop	No change	No change	Review/ increase	Stop	Review	Stop
Beta-blocker	No change	Reduce/ stop	Stop	Reduce	Stop	No change	No change	No change	No change
MRA	No change	No change	Stop	No change	No change	Review/ increase	Stop	Reduce	Stop
Diuretics	Increase	Reduce	Stop	No change	No change	Review/ No change	Review/ increase	No change	Review

CCB, calcium channel blockers (mg/dL); Cr, creatinine blood level (mg/dL); eGFR, estimated glomerular filtration rate ml/min/1.73 m2; MRA, mineralocorticoid receptor antagonist; (\*) amiodarone. - Depicted from Mebazaa A et al. Eur | Heart Fail. (2015):17(6):544-58.

#### **ACUTE HEART FAILURE: Treatment (C) and preventive measures (Cont.)**

#### Management of oral therapy in AHF in the first 48 hours

p.5 l

	Normotension/ Hypertension	Hypotension		Low Heart rate		Potassium		Renal impairment	
		<100 >85 mmHg	<85 mmHg	<60 ≥50 bpm	<50 bpm	≤3.5 mg/dL	>5.5 mg/dL	Cr < 2.5, eGFR > 30	Cr > 2.5, eGFR < 30
Other vasodilators (Nitrates)	Increase	Reduce/ stop	Stop	No change	No change	No change	No change	No change	No change
Other heart rate slowing drugs (amiodarone, CCB, Ivabradine)	Review	Reduce/ stop	Stop	Reduce/ stop	Stop	Review/ stop (*)	No change	No change	No change

Thrombosis prophylaxis should be started in patients not anticoagulated (enoxaparin 1 mg/kg as first dose)

Maintain an adequate nutritional status with a nutritional support of 20-25 kcal/kg/day within the first 48 hours

CCB, calcium channel blockers (mg/dL); Cr, creatinine blood level (mg/dL); eGFR, estimated glomerular filtration rate ml/min/1.73 m2; MRA, mineralocorticoid receptor antagonist; (\*) amiodarone. - Depicted from Mebazaa A et al. Eur | Heart Fail. (2015);17(6):544-58.

Clinical condition defined as the inability of the heart to deliver an adequate amount of blood to the tissues to meet resting metabolic demands as a result of impairment of its pumping function.

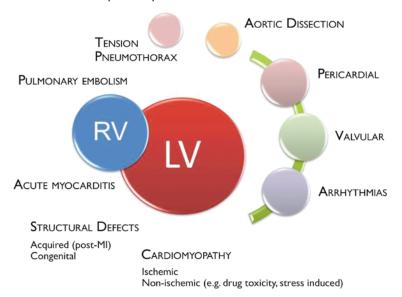
#### Hemodynamic criteria to define cardiogenic shock

- $\bullet$  Systolic blood pressure <80 to 90 mmHg or mean arterial pressure 30 mmHg lower than baseline
- · Severe reduction in cardiac index:
  - <1.8 L/min/m<sup>2</sup> without support or
  - <2.0 to 2.2 L/min/m<sup>2</sup> with support
- Adequate or elevated filling pressure:

Left ventricular end-diastolic pressure >18 mmHg or Right ventricular end-diastolic pressure >10 to 15 mmHg

#### **CARDIOGENIC SHOCK: Causes**

LV pump failure is the primary insult in most forms of CS, but other parts of the circulatory system contribute to shock with inadequate compensation or additional defects.



#### **CARDIOGENIC SHOCK: Initial triage and management**

This protocol should be initiated as soon as cardiogenic shock/end organ hypoperfusion is recognised and should not be delayed pending intensive care admission.

p.54

EPARTMENT	0 min	
EMERGENCY D	5 min	
<u></u>		E CARE UNIT
		CARDIAC INTENSIVI

#### **EARLY TRIAGE & MONITORING**

**INITIAL RESUSCITATION** 

catheterization with a catheter capable of measuring central venous oxygen

Standard transthoracic echocardiogram

mechanical complications following MI

· Early coronary angiography in specialized

and/or symptoms of ongoing myocardial

ischemia (e.g. ST segment elevation

myocardial infarction).

myocardial intervention center when signs

function and for the detection of potential

to assess left (and right) ventricular

· Arterial and a central venous

Start high flow O<sub>2</sub> Establish i.v. access

saturation

- Age: 65-74, ≥75
- Heart rate >100 beats per minute
- Systolic blood pressure <100 mmHg
- Proportional pulse pressure ≤25 mmhg (Cl <2.2l/min/m²)</li>
- Orthopnea (PCWP >22 mmHg)
- Tachypnea (>20/min), >30/min (!)
- Killip class II-IV
- Clinical symptoms of tissue hypoperfusion/hypoxia:
- cool extremities, decreased urine output (urine output <40 ml/h)
- decreased capillary refill or mottling alteration in mental status
- CORRECT: hypoglycemia & hypocalcemia,
- TREAT: sustained arrhythmias: brady- or tachy-
- Isotonic saline-fluid challenge of 20 to 30 ml per kilogram of body weight over a 30-minute period to achieve a central venous pressure of 8 to 12 mmHg or until perfusion improves (with a maximum of 500 ml)
- CONSIDER NIVmechanical ventilation for comfort (fatigue, distress) or as needed:
  - To correct acidosis To correct hypoxemia
- INOTROPIC SUPPORT (dobutamine and/or vasopressor support)

#### TREATMENT GOALS

- a mean arterial pressure of 60 mmHg or above.
- a mean pulmonary artery wedge pressure of 18 mmHg or below,
- a central venous pressure of 8 to 12 mmHg,
- a urinary ouput of 0,5 ml or more per hour per kilogram of body weight
- an arterial pH of 7.3 to 7.5
- a central venous saturation (ScvO<sub>2</sub>)  $\geq$ 70% (provided SpO<sub>2</sub>  $\geq$ 93% and Hb level  $\geq$ 9 g/dl)

In persistent drug-resistant cardiogenic shock, consider mechanical circulatory support

#### **CARDIOGENIC SHOCK: Treatment and ventilator procedures**

#### For more informations on individual drug doses and indications:

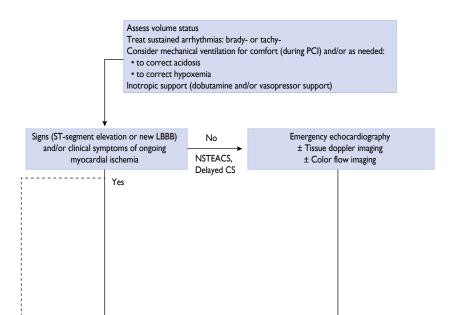
\*See Chapter 8: Use of drugs in Acute Cardiovascular Care.

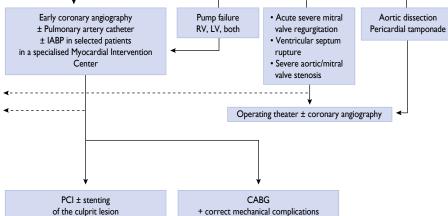
Ventilator mode	Pressure assist/control
Tidal Volume goal	Reduce tidal volume to 6-8 ml/kg lean body weight
Plateau Pressure goal	≤ 30 cm H <sub>2</sub> O
Anticipated PEEP levels	5-10 cm H <sub>2</sub> O
Ventilator rate and pH goal	12-20, adjusted to achieve a pH ≥ 7.30 if possible
Inspiration: Expiration time	1:1 to 1:2
Oxygenation goal:	
• PaO <sub>2</sub>	50-80 mmHg
• SpO <sub>2</sub>	> 90%

#### Predicted body weight calculation:

- Male: 50 + 0.91 (height in cm 152.4)
- Female: 45.5 + 0.91 (height in cm 152.4)

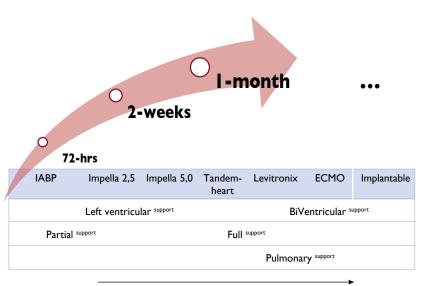
Some patients with CS will require increased PEEP to attain functional residual capacity and maintain oxygenation, and peak pressures above 30 cm H2O to attain effective tidal volumes of 6-8ml/kg with adequate CO2 removal.





Short-term mechanical circulatory support

### CARDIOGENIC SHOCK: Mechanical circulatory support, basic characteristics



Level of support

	Туре	Support		Access
Intra-aortic balloon pump	Balloon counterpulsation	Pulsatile flow	<0.5 L	Arterial: 7.5 French
Impelia Recover LP 2.5 CP LP 5.0	Axial flow	Continuous flow	<2.5 L <4,0 L <5.0 L	Arterial: 12 French Arterial: 14 French Arterial: 21 French
Tandemheart  Cardiohelp	Centrifugal flow	Continuous flow	<5.0 L	Venous: 21 French Arterial: 15-17 French Venous: 15-29 French Arterial: 15-29 French

Different systems for mechanical circulatory support are available to the medical community. The available devices differ in terms of the insertion procedure, mechanical properties, and mode of action. A minimal flow rate of 70 ml/kg/min, representing a cardiac index of at least  $2.5 \, \text{L/m}^2$ , is generally required to provide adequate organ perfusion. This flow is the sum of the mechanical circulatory support output and the remaining function of the heart.

The SAVE-score may be a tool to predict survival for patients receiving ECMO for refractory cardiogenic shock ( ).

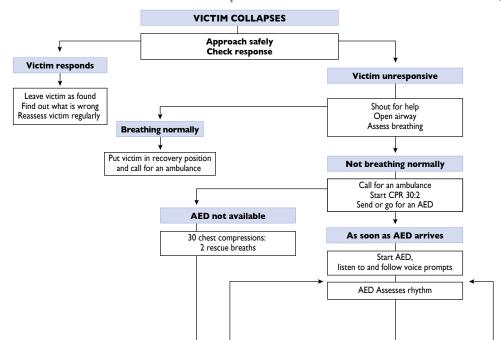
## CHAPTER 4: CARDIAC ARREST AND CARDIOPULMONARY RESUSCITATION

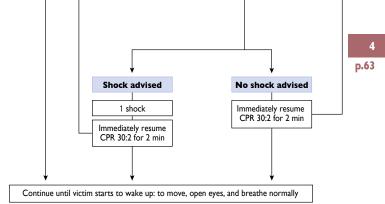


Monsieurs KG, et al. European Resuscitation Council Guidelines for Resuscitation 2015. Section 1. Executive Summary. Resuscitation 2015; 95C:1-80, DOI:10.1016/j.resuscitation.2015.07.038

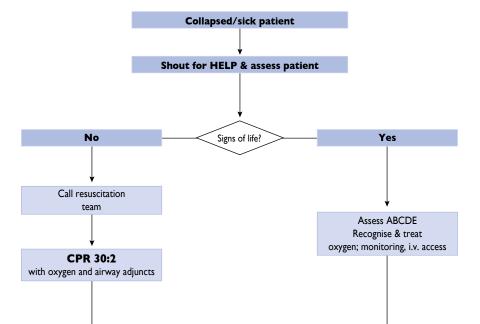
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### OUT OF HOSPITAL CARDIAC ARREST: Assessment of a collapsed victim and initial treatment

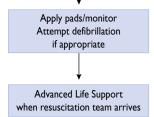


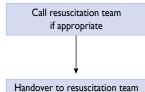


### IN-HOSPITAL CARDIAC ARREST: Assessment of a collapsed victim and initial treatment

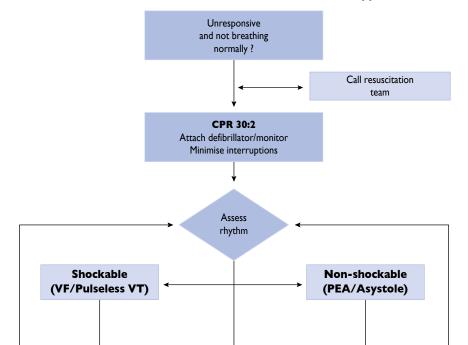


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#### **IN-HOSPITAL CARDIAC ARREST: Advanced life support**



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CPR for 2 min

Minimise interruptions

#### **DURING CPR**

- Ensure high-quality chest compressions
- Minimise interruptions to compressions
- Give Oxygen
- Use waveform capnography
- Continuous chest compressions when advanced airway in place
- Vascular access (intravenous, intraosseous)
- Give adrenaline every 3-5 min
- · Give amiodarone after 3 shocks
- Correct reversible causes

Return of spontaneous circulation

# IMMEDIATE POST CARDIAC ARREST TREATMENT

- Use ABCDE approach
- Aim for SaO<sub>2</sub> 94-98%
- Aim for normal PaCO<sub>2</sub>
- 12-lead ECG
- Treat precipitating cause
- Temperature control / Therapeutic hypothermia

#### **CONSIDER**

- Ultrasound imaging
- Mechanical chest compressions to facilitate transfer/treatment
- Coronary angiography and PCI
- Extracorporeal CPR

Immediately resume:

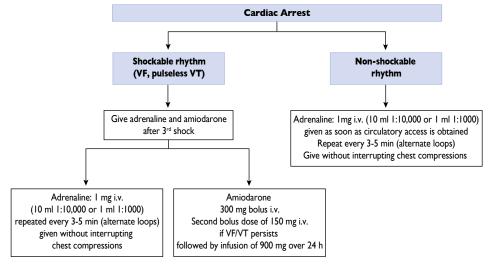
**CPR for 2 min**Minimise interruptions

#### **REVERSIBLE CAUSES**

- Hypoxia
- Hypovolaemia
- Hypo-/hyperkalaemia/metabolic
- Hypothermia
- Thrombosis
- Tamponade cardiac
- Toxins
- Tension pneumothorax

#### IN-HOSPITAL CARDIAC ARREST: Drug therapy during advanced life support

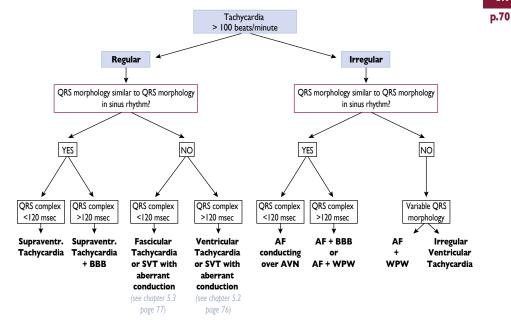




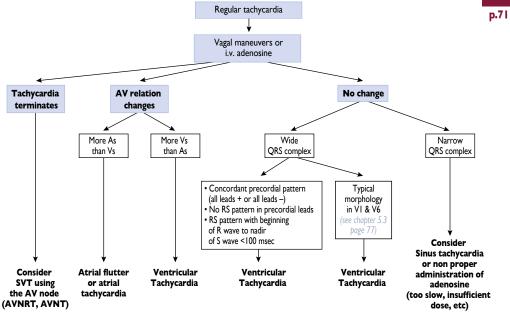
## **CHAPTER 5: RHYTHM DISTURBANCES**

5.1 SUPRAVENTRICULAR TACHYO AND ATRIAL FIBRILLATION	
J. Brugada	·
5.2 VENTRICULAR TACHYCARDIA M. Santini, C. Lavalle, S. Lanzara	<b>S</b> p.7
5.3 BRADYARRHYTHMIAS	p.7

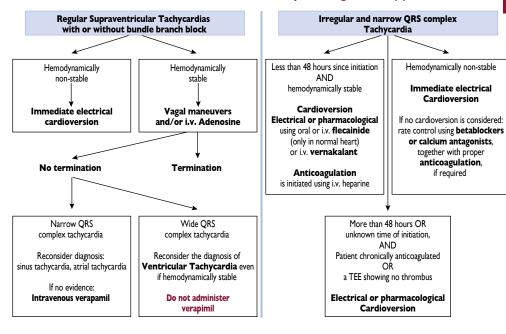
#### **TACHYARRHYTHMIAS: Diagnostic criteria**



#### **TACHYARRHYTHMIAS: Diagnostic maneuvers**

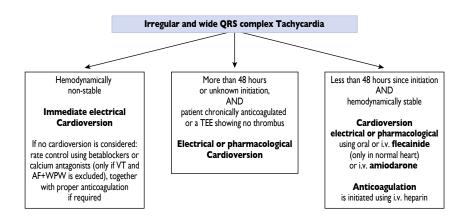


#### **TACHYARRHYTHMIAS: Therapeutic algorithms (I)**

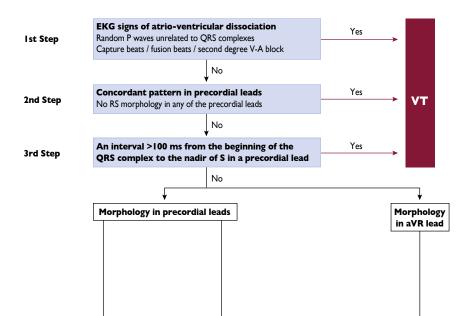


#### **TACHYARRHYTHMIAS:** Therapeutic algorithms (2)

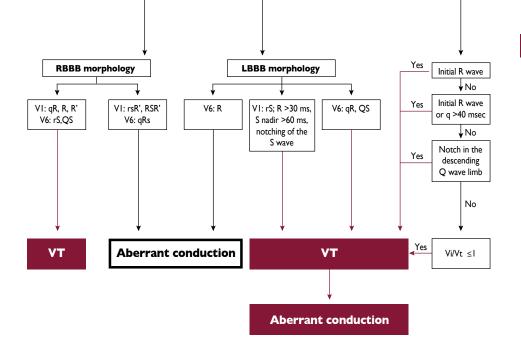
p.73



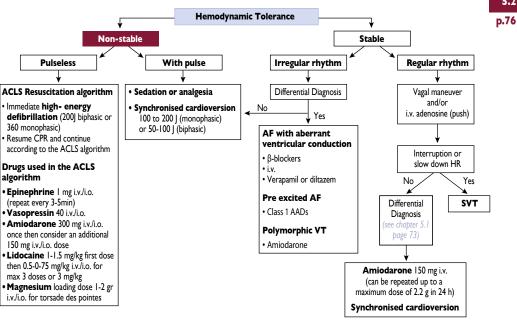
# VENTRICULAR TACHYSCARDIAS: Diferential diagnosis of wide QRS tachyscardias



p.75



#### Management of wide QRS TACHYSCARDIAS



#### **BRADYARRHYTHMIAS: Definitions and diagnosis**



#### Sinus node dysfunction



- Sinus bradycardia. It is a rhythm that originates from the sinus node and has a rate of under 60 beats per minute
- Sinoatrial exit block. The depolarisations that occur in the sinus node cannot leave the node towards the atria
- Sinus arrest. Sinus pause or arrest is defined as the transient absence of sinus P waves on the ECG



#### Atrioventricular (AV) blocks



- First degree AV block. Atrioventricular impulse transmission is delayed, resulting in a PR interval longer than 200 msec
- Second degree AV block. Mobitz type I (Wenckebach block): Progressive PR interval prolongation, which precedes a nonconducted P wave
- Second degree AV block. Mobitz type II: PR interval remains unchanged prior to a P wave that suddenly fails to conduct to the ventricles
- Third degree (complete) AV block.
   No atrial impulses reach the ventricle

#### **BRADYARRHYTHMIAS: Treatment (I)**

- Rule out and treat any underlying causes of bradyarrhythmia
- Treat symptomatic patients only

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

#### **Temporary transvenous pacing**

#### Be Careful!

- Complications are common!
- Shall not be used routinely
- Use only as a last resource when chronotropic drugs are insufficient
- Every effort should be made to implant a permanent pacemaker as soon as possible,
   if the indications are established

#### **Indications** limited to:

- High-degree AV block without escape rhythm
- Life threatening bradyarrhythmias, such as those that occur during interventional procedures, in acute settings such as acute myocardial infarction, drug toxicity.

# BRADYARRHYTHMIAS: Treatment (2) Pacemaker therapies in sinus node dysfunction

#### Permanent pacemaker is indicated in the following settings:

- Documented symptomatic bradycardia, including frequent sinus pauses that produce symptoms
- Symptomatic chronotropic incompetence
- Symptomatic sinus bradycardia that results from required drug therapy for medical conditions

#### Permanent pacemaker is <u>not</u> recommended in the following settings:

- Asymptomatic patients
- Patients for whom the symptoms suggestive of bradycardia have been clearly documented to occur in the absence of bradycardia
- Symptomatic bradycardia due to nonessential drug therapy

# BRADYARRHYTHMIAS: Treatment (3) Pacemaker therapies in atrioventricular blocks

p.80

# Permanent pacemaker therapy is indicated in the following settings regardless of associated symptoms:

- Third-degree AV block
- Advanced second-degree AV block
- Symptomatic Mobitz I or Mobitz II second-degree AV block
- Mobitz II second-degree AV block with a wide QRS or chronic bifascicular block
- Exercise-induced second- or third-degree AV block
- Neuromuscular diseases with third- or second-degree AV block
- Third- or second-degree (Mobitz I or II) AV block after catheter ablation or valve surgery when block is not expected to resolve

#### Permanent pacemaker is <u>not</u> recommended in the following settings:

- Asymptomatic patients
- Patients for whom the symptoms suggestive of bradycardia have been clearly documented to occur in the absence of bradycardia
- Symptomatic bradycardia due to nonessential drug therapy

# **CHAPTER 6: ACUTE VASCULAR SYNDROMES**

6.1	ACUTE AORTIC SYNDROMES	<b>8</b> .8
A. E	Evangelista	
6.2	ACUTE PULMONARY EMBOLISM	) 5.9'
	Torbicki	

# ACUTE AORTIC SYNDROMES: Concept and classification (I) Types of presentation

p.82



on ce all.

Separation of the aorta media with presence of extraluminal blood within the layers of the aortic wall. The intimal flap divides the aorta into two lumina, the true and the false



Intramural hematoma (IMH)
Aortic wall hematoma with no entry tear
and no two-lumen flow



Penetrating aortic ulcer (PAU)
Atherosclerotic lesion penetrates
the internal elastic lamina of the aorta wall



Aortic aneurysm rupture (contained or not contained)

# ACUTE AORTIC SYNDROMES: Concept and classification (2) Anatomic classification and time course

#### **DeBakey's Classification**

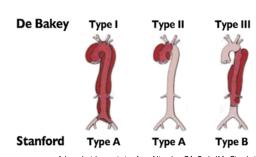
- Type I and type II dissections both originate in the ascending aorta
   In type I, the dissection extends distally to the descending aorta
   In type II, it is confined to the ascending aorta
- Type III dissections originate in the descending aorta

#### Stanford Classification

- Type A includes all dissections involving the ascending aorta regardless of entry site location
- Type B dissections include all those distal to the brachiocephalic trunk, sparing the ascending aorta

#### Time course

- Acute: < I4 days</li>
- Subacute: 15-90 days
- Chronic: > 90 days



Adapted with permission from Nienaber CA, Eagle KA, Circulation 2003;108(6):772-778. All rights reserved.

# ACUTE AORTIC SYNDROME: Clinical suspicion and differential diagnosis

# SYMPTOMS AND SIGNS SUGGESTIVE OF AAS

- Abrupt and severe chest/back pain with maximum intensity at onset
- · Pulse/pressure deficit
  - Peripheral or visceral ischemia
  - Neurological deficit
- Widened mediastinum on chest X -ray
- · Risk factors for dissection
- Other
  - Acute aortic regurgitation
  - Pericardial effusion
  - Hemomediastinum/hemothorax

#### **DIFFERENTIAL DIAGNOSIS**

- Acute coronary syndrome (with/without ST-segment elevation)
- · Aortic regurgitation without dissection
- · Aortic aneurysms without dissection
- Musculoskeletal pain
- Pericarditis
- Pleuritis
- · Mediastinal tumours
- Pulmonary embolism
- Cholecystitis
- · Atherosclerosis or cholesterol embolism

# General approach to the patient with suspected ACUTE AORTIC SYNDROME

#### Consider acute aortic dissection in all patients presenting with:

- · Chest, back or abdominal pain
- Syncope
- Symptoms consistent with perfusion deficit (central nervous system, visceral myocardial or limb ischemia)

#### Pre-test risk assessment for acute aortic dissection

#### **High-risk conditions**

- · Marfan's syndrome
- Connective tissue disease
- · Family history of aortic disease
- Aortic valve disease
- Thoracic aortic aneurysm

#### High-risk pain features

### Chest, back or abdominal pain described as:

Abrupt at onset, severe in intensity, and ripping/sharp or stabbing quality

#### High-risk exam features

- Perfusion deficit:
  - Pulse deficit
  - SBP differential
- Focal neurological deficit
- Aortic regurgitation murmur
- Hypotension or shock

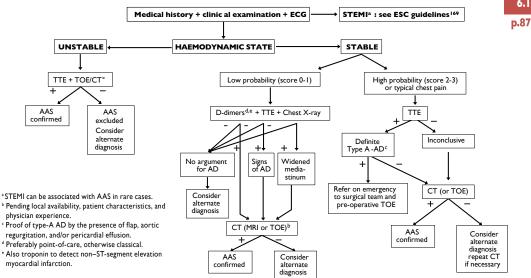
# Laboratory tests required for patients with ACUTE AORTIC dissection

p.86

Laboratory tests	To detect signs of:
Red blood cell count	Blood loss, bleeding, anaemia
White blood cell count	Infection, inflammation (SIRS*)
C-reactive protein	Inflammatory response
ProCalcitonin	Differential diagnosis between SIRS* and sepsis
Creatine kinase	Reperfusion injury, rhabdomyolysis
TroponinlorT	Myocardial ischaemia, myocardial infarction
D-dimer	Aortic dissection, pulmonary embolism, thrombosis
Creatinine	Renal failure (existing or developing)
Aspartate transaminase/ alanine aminotransferase	Liver ischaemia, liver disease
Lactate	Bowel ischaemia, metabolic disorder
Glucose	Diabetes mellitus
Blood gases	Metabolic disorder, oxygenation

\*SIRS = systemic inflammatory response syndrome.

#### **ACUTE CHEST PAIN**



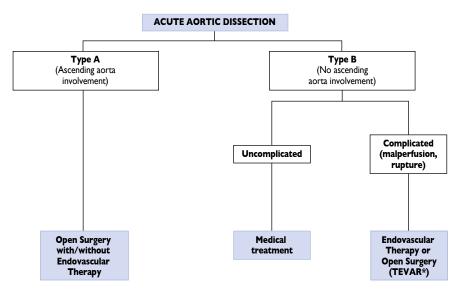
Flowchart for decision-making based on pre-test sensitivity of acute aortic syndrome. Reference: Eur Heart I 2014:eurhearti.ehu281

#### **Details required from imaging in ACUTE AORTIC dissection**

Aortic dissection	Visualization of intimal flap Extent of the disease according to the aortic anatomic segmentation Identification of the false and true lumens (if present) Localization of entry and re-entry tears (if present) Identification of antegrade and/or retrograde aortic dissection Identification grading, and mechanism of aortic valve regurgitation Involvement of side branches Detection of malperfusion (low flow or no flow) Detection of organ ischaemia (brain, myocardium, bowels, kidneys, etc.) Detection of pericardial effusion and its severity Detection and extent of pleural effusion Detection of peri-aortic bleeding Signs of mediastinal bleeding
Intramural haematoma	Localization and extent of aortic wall thickening     Co-existence of atheromatous disease (calcium shift)     Presence of small intimal tears
Penetrating aortic ulcer	Localization of the lesion (length and depth) Co-existence of intramural haematoma Involvement of the peri-aortic tissue and bleeding Thickness of the residual wall
In all cases	Co-existence of other aortic lesions: aneurysms, plaques, signs of inflammatory disease, etc.

#### **ACUTE AORTIC SYNDROMES MANAGEMENT: General approach**

p.89



<sup>\*</sup>TEVAR Thoracic Endovascular Aortic Repair.

#### **ACUTE AORTIC SYNDROMES: Initial management**

p.90

- 1 Detailed medical history and complete physical examination (when possible)
- 2 Standard 12-lead ECG: Rule-out ACS, documentation of myocardial ischemia
- 3 Intravenous line, blood sample (CK, Tn, myoglobin, white blood count, D-dimer, hematocrit, LDH)
- 4 Monitoring: HR and BP
- **5** Pain relief (morphine sulphate) (see chapter 3)
- **6 Noninvasive imaging** (see previous page)
- 7 Transfer to ICU

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

#### p.91

#### TYPE A ACUTE AORTIC DISSECTION

#### **URGENT SURGERY (<24h)**

Graft replacement of ascending aorta +/- arch
with/without aortic valve or aortic root
replacement/repair (depending on aortic regurgitation and
aortic root involvement)

#### **Emergency Surgery**

- Haemodynamic instability (hypotension/shock)
- Tamponade
- Severe acute aortic regurgitation
- · Impending rupture
- Flap in aortic root
- Malperfusion syndrome

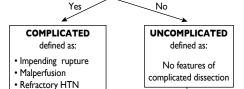
#### Elective/individualised Surgery

- Non-complicated intramural hematoma
- Comorbidities
- Age >80 years

#### TYPE B ACUTE AORTIC DISSECTION

#### **Definitive diagnosis**

by clinical presentation and imaging



#### MEDICAL MANAGEMENT and TEVAR

Shock

• SBP < 90 mmHg)

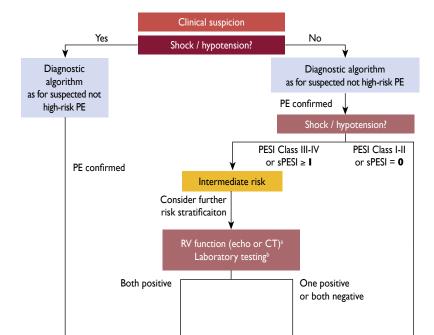
MEDICAL
MANAGEMENT
and
OPEN SURGERY

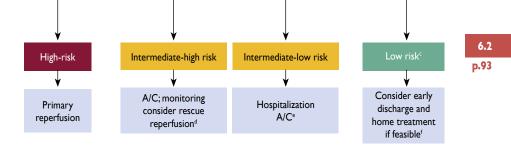
REPAIR if TEVAR contraindicated

# MEDICAL MANAGEMENT and imaging surveillance protocol

- On admission
- At 7 days
- At discharge
- Every 6 months thereafter

#### Risk-adjusted management strategies in ACUTE PULMONARY EMBOLISM





- alf echocardiography has already been performed during diagnostic work-up for PE and detected RV dysfunction, or if the CT already performed for diagnostic work-up has shown RV enlargement (RV/LV (left ventricular) ratio ≥0.9, a cardiac troponin test should be performed except for cases in which primary reperfusion is not a therapeutic option (e.g. due to severe comorbidity or limited life expectancy of the patient).
- b Markers of myocardial injury (e.g. elevated cardiac troponin I or T concentrations in plasma), or of heart failure as a result of (right) ventricular dysfunction (elevated natriuretic peptide concentrations in plasma). If a laboratory test for a cardiac biomarker has already been performed during initial diagnostic work-up (e.g. in the chest pain unit) and was positive, then an echocardiogram should be considered to assess RV function, or RV size should be (re)assessed on CT.
- Patients in the PESI Class I-II, or with sPESI of 0, and elevated cardiac biomarkers or signs of RV dysfunction on imaging tests, are also to be classified into the intermediate-low risk category. This might apply to situations in which imaging or biomarker results become available before calculation of the clinical severity index. These patients are probably no candidates for home treatment.
- <sup>d</sup> Thrombolysis, if (and as soon as) clinical signs of haemodynamic decompensation appear; surgical pulmonary embolectomy or percutaneous catheter-directed treatment may be considered as alternative options to systemic thrombolysis, particularly if the bleeding risk is high.
- e Monitoring should be considered for patients with confirmed PE and a positive troponin test, even if there is no evidence of RV dysfunction on echocardiography or CT.
- <sup>†</sup> The simplified version of the PESI has not been validated in prospective home treatment trials; inclusion criteria other than the PESI were used in two single-armed (non-randomized) management studies.

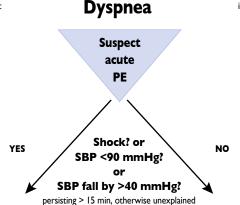
Reference: Eur Heart | 2014;35:3033-3073.

#### **ACUTE PULMONARY EMBOLISM: Diagnosis**

p.94



- · Chest pain (angina)
- Syncope
- Tachycardia
- ECG changes
- NT-proBNP ↑
- Troponin ↑



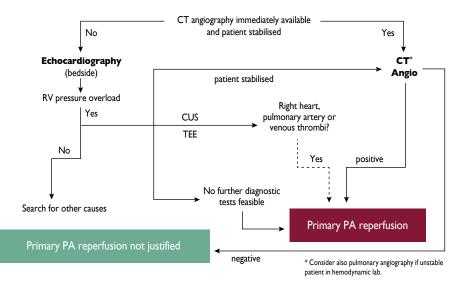
#### RESPIRATORY Symptoms/Signs including but not limited to:

- Chest pain (pleural)
- Pleural effusion
- Tachypnea
- Hemoptysis
- Hypoxemia
- Atelectasis

Management algorithm for UNSTABLE patients

Management algorithm for initially STABLE patients

#### Management algorithm for unstable patients with suspected ACUTE PULMONARY EMBOLISM

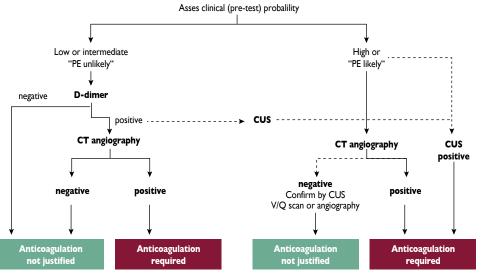


# ACUTE PE: Management strategy for initially unstable patients with confirmed high risk pulmonary embolism

Shock or hypotension	YES		
Contraindications for thrombolysis	No Relative Absolute		
Primary PA reperfusion strategy	Thrombolysis	Low-dose/ transcatheter thrombolysis/ clot fragmetation	Surgical or Percutaneous catheter embolectomy (availability/experience)
Supportive treatment	i.v. UFH, STABILISE SYSTEMIC BLOOD PRESSURE, CORRECT HYPOXEMIA		

p.97

#### Management algorithm for initially stable patients with suspected ACUTE PULMONARY EMBOLISM



#### Suggested management strategy for initially stable patients with (non-high risk) confirmed

STRATECY	Monitoring (ICU)*	Hospitalisation**	Early
Suggested initial anticoagulation	UFH i.v /LMWH s.c.	LMWH/Fonda/ apixaban/ rivaroxaban	apixaban/rivaroxaban
Clinical risk assessment score (PESI)	Positive (class III-V)	Positive (class III-V)	Negative (class I-II)
Markers for RV overload	Positive	Positive	Negative
Markers for myocardial injury	Positive	Positive	Negative

rescue thrombolysis

#### **Pharmacological treatment:**

**STRATEGY** 

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

<sup>\*</sup> When all markers are positive.

<sup>(</sup>telemonitoring) \*\* When at least one marker is positive.

discharge\*\*\* \*\*\* When all markers are negative.

#### **PULMONARY EMBOLISM: Pharmacological treatment**

#### Key drugs for initial treatment of patients with confirmed PE

For more information on individual drug doses and indications, See chapter 8: Use of drugs in Acute Cardiovascular Care.

Unstable	Alteplase (rtPA) (intravenous)	100 mg/2 h or 0.6 mg/kg/15 min (max 50 mg)
	Urokinase (intravenous)	3 million IU over 2 h
	Streptokinase (intravenous)	1.5 million IU over 2 h
	Unfractionated heparin (intravenous)	80 IU/kg bolus + 18 IU/kg/h
	Enoxaparine (subcutaneous)	I.0 mg/kg BID or I.5 mg/kg QD
	Tinzaparin (subcutaneous)	175 U/kg QD
Stable	Fondaparinux (subcutaneous)	7.5 mg (50-100 Kg of body weight) 5 mg for patients <50 kg, 10 mg for patients >100 kg
	Rivaroxaban (oral)	15 mg BID (for 3 weeks, then 20 mg QD)
	Apixaban (oral)	10mg bid (for 7 days, than 5mg bid)

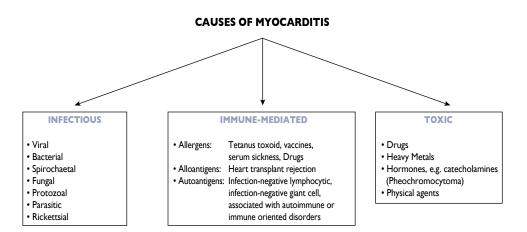
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# CHAPTER 7: ACUTE MYOCARDIAL / PERICARDIAL SYNDROMES

7.1 ACUTE MYOCARDITIS	р.102
A. Keren, A. Caforio	
7.2 ACUTE PERICARDITIS	
AND CARDIAC TAMPONADE	р.107
C Vrints S Price	

p.102

MYOCARDITIS (WHO /ISFC): Inflammatory disease of the myocardium diagnosed by established histological, immunological and immunohistochemical criteria.



# ACUTE MYOCARDITIS: Diagnostic criteria (I) Diagnostic criteria for clinically suspected myocarditis

p.103

# Clinical presentations with or without ancillary findings

- Acute chest pain (pericarditic or pseudo-ischemic )
- New-onset (days up to 3 months) or worsening dyspnea or fatigue, with or without left/right heart failure signs
- Palpitation, unexplained arrhythmia symptoms, syncope, aborted sudden cardiac death
- Unexplained cardiogenic shock and/or pulmonary oedema

## Ancillary findings which support the clinical suspicion of myocarditis

- Fever ≥38.0°C within the preceding 30 days
- A respiratory or gastrointestinal infection
- Previous clinically suspected or biopsy proven myocarditis
- Peri-partum period
- · Personal and/or family history of allergic asthma
- · Other types of allergy
- Extra-cardiac autoimmune disease
- · Toxic agents
- · Family history of dilated cardiomyopathy, myocarditis

#### Diagnostic criteria

- I. ECG/Holter/stress test features: Newly abnormal ECG and/or Holter and/or stress testing, any of the following:
- I to III degree atrioventricular block, or bundle branch block, ST/T wave changes (ST elevation or non ST elevation, T wave inversion),
- Sinus arrest, ventricular tachycardia or fibrillation and asystole, atrial fibrillation, frequent premature beats, supraventricular tachycardia
- Reduced R wave height, intraventricular conduction delay (widened ORS complex), abnormal Q waves, low voltage
- II. Myocardiocytolysis markers: Elevated TnT/Tnl

#### III. Functional/structural abnormalities on echocardiography

 New, otherwise unexplained LV and/or RV structure and function abnormality (including incidental finding in apparently asymptomatic subjects): regional wall motion or global systolic or diastolic function abnormality, with or without ventricular dilatation, with or without increased wall thickness, with or without pericardial effusion, with or without endocavitary thrombi

#### IV. Tissue characterisation by CMR:

Edema and/or LGE of classical myocarditic pattern

Reference: Caforio ALP et al. Eur Heart J. (2013) Jul 3 (15).

#### **ACUTE MYOCARDITIS: Diagnostic criteria (2)**

## Acute myocarditis should be clinically suspected in the presence of:

One or more of the clinical presentations shown in the Diagnostic Criteria\*
with or without Ancillary Features\*
AND

One or more Diagnostic Criteria from different categories (I to IV)\*

OR

when the patient is asymptomatic, two or more diagnostic criteria from different categories (I to IV)\*

#### in the absence of:

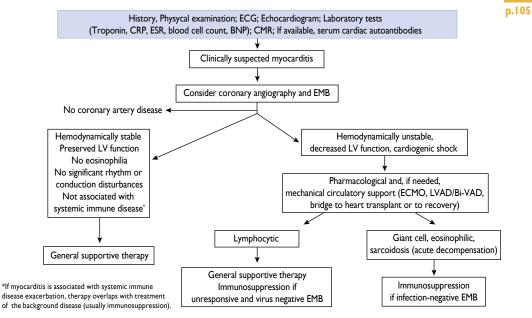
- I) angiographically detectable coronary artery disease
- 2) known pre-existing cardiovascular disease or extra-cardiac causes that could explain the syndrome (e.g. valve disease, congenital heart disease, hyperthyroidism, etc.)

#### Suspicion is higher with higher number of fulfilled criteria\*

Endomyocardial biopsy is necessary to: 1) confirm the diagnosis of clinically suspected myocarditis,

3) identify the type and aetiology of inflammation, and 2) provide the basis for safe immunosuppression (in virus negative cases).

#### **ACUTE MYOCARDITIS: Diagnostic and management protocol**

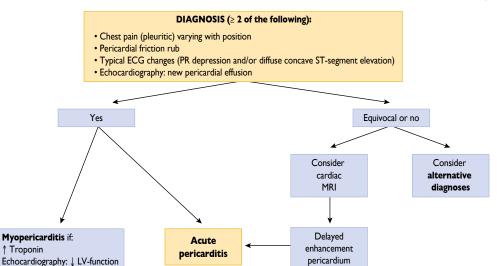


# Management of patients with life-threatening ACUTE MYOCARDITIS

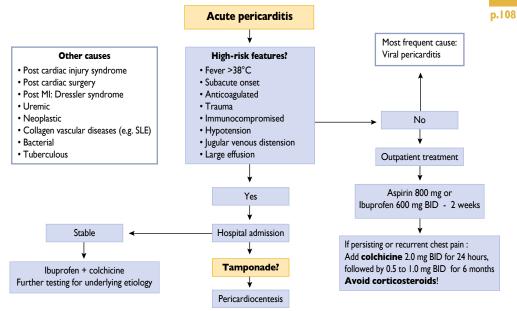
- Patients with a life-threatening presentation should be sent to specialised units with capability for hemodynamic monitoring, cardiac catheterisation and expertise in endomyocardial biopsy.
- In patients with hemodynamic instability a mechanical cardio-pulmonary assist device
  may be needed as a bridge to recovery or to heart transplantation.
- Heart transplant should be deferred in the acute phase, because recovery may occur, but can be considered
  for hemodynamically unstable myocarditis patients, including those with giant cell myocarditis,
  if optimal pharmacological support and mechanical assistance cannot stabilise the patient
- ICD implantation for complex arrhythmias should be deferred until resolution of the acute episode, with possible
  use of a lifevest during the recovery period.

#### **ACUTE PERICARDITIS: Diagnosis**

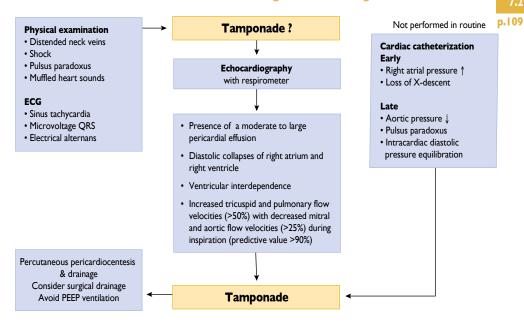
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#### **ACUTE PERICARDITIS: Management**



#### **CARDIAC TAMPONADE: Diagnosis and management**



# CHAPTER 8: DRUGS USED IN ACUTE CARDIOVASCULAR CARE

Ana de Lorenzo

#### **Oral antiplatelets**

p.112

Drug	Indications	Dose	Dose adjustments	Comments
Aspirin	Primary (not universally approved) and secondary cardiovascular disease prevention	LD (if ACS): 150-300 mg oral MD: 75-100 mg oral QD	-	Major contraindications: GI bleeding-active peptic ulcer
relor	ACS (all patients at moderate-to-high risk of ischaemic events, e.g. elevated cardiac troponins)	LD: 180 mg oral MD: 90 mg oral BID	-	Major contraindications: previous intracerebral hemorrhage
Ticagrelor	Secondary prevention I-3 years post-MI	MD: 60 mg oral BID	-	Major contraindications: previous intracerebral hemorrhage
Prasugrel	ACS with planned PCI	LD: 60 mg oral MD: 10 mg oral QD	MD: 5 mg QD weight < 60 kg	Contraindication: previous stroke/TIA Prasugrel is generally not recommended in elderly, and if positive benefit/risk 5 mg is recommended
Clopidogrel	ACS + PCI or medical management (patients who cannot receive ticagrelor or prasugrel) and in ACS patients at high bleeding risk (e.g. patients who require oral anticoagulation)	LD: 300-600 mg oral MD: 75 mg oral QD	-	-

#### **Oral antiplatelets (Cont.)**

p.113

Drug	Indications	Dose	Dose adjustments	Comments
Clopidogrel	STEMI + fibrinolysis < 75 years	LD: 300 mg oral MD: 75 mg oral QD	-	
	STEMI + fibrinolysis ≥ 75 years	LD: 75mg oral. MD: 75 mg oral QD	-	Prasugrel and ticagrelor have not been studied as adjuncts to fibrinolysis and oral anticoagulants
	Secondary prevention >12 months post coronary stenting	MD: 75 mg oral QD	-	
Vorapaxar	Co-administered with aspirin and, where appropriate, clopidogrel, in patients with a history of MI or peripheral artery disease	2.08 mg oral QD	-	Initiated at least 2 weeks after a MI and preferably within the first 12 months Major contraindications: active pathologic bleeding or increased risk of bleeding, history of stroke / TIA or intracranial bleeding, severe hepatic dysfunction

#### **Intravenous Antiplatelets**

p. l l 4

Drug	Indications	Dose	Dose adjustments	Comments
Abciximab	Adjunct to PCI for bailout situations or thrombotic complications	LD: 0.25 mg/Kg i.v. MD: 0.125 ug/Kg/min i.v. (max: 10 µg/min) for 12h	-	Contraindications: Active internal bleeding - History of CVA within 2 years - Bleeding diathesis - Preexisting thrombocytopenia - Recent (within 2 months) intracranial or intraspinal surgery or trauma - Recent (within 2 months) major surgery - Intracranial neoplasm, arteriovenous malformation, or aneurysm - Severe uncontrolled hypertension - Presumed or documented history of vasculitis - Severe hepatic failure or severe renal failure requiring haemodialysis - Hypertensive retinopathy
Eptifibatide	ACS treated medically or with PCI	LD: 180 µg/Kg i.v. (at a 10 min interval) If STEMI and PCI: add a second 180 mcg/kg i.v. bolus at 10 min MD: 2 µg/Kg/min i.v. infusion	Reduce infusion dose to I µg/kg/ min if CrCl 30-50ml/min	Contraindications: Bleeding diathesis or bleeding within the previous 30 days - Severe uncontrolled hypertension - Major surgery within the preceding 6 weeks - Stroke within 30 days or any history of hemorrhagic stroke - Coadministration of another parenteral GP Illb/Illa inhibitor - Dependency on renal dialysis - Known hypersensitivity to any component of the product

#### **Intravenous Antiplatelets (Cont.)**

p. l l 5

Drug	Indications	Dose	Dose adjustments	Comments
Tirofiban	ACS treated medically or with PCI	LD: 25 µg/Kg i.v. over 5 min MD: 0.15 µg/Kg/min i.v. Infusion to 18 hour	CrCl < 30ml/min: decrease 50% bolus and infusion dose	Contraindications: Severe hypersensitivity reaction to tirofiban A history of thrombocytopenia following prior exposure Active internal bleeding or a history of bleeding diathesis, major surgical procedure or severe physical trauma within the previous month
Cangrelor	All patients undergoing PCI (elective + ACS) immediate onset + rapid offset (platelet recovery in 60 min)	IV Bolus of 30 µg/Kg + IV infusion of 4 µg/kg/min For at least 2 hours from start of PCI	-	Major contraindications: significant active bleeding or stroke Transition to oral P2Y12 inhibitors variable according to type of agent

#### **Oral Anticoagulants**

p.116

Drug	Indications	Dose	Dose adjustments	Comments
Warfarin Acenocoumarol	Treatment and prophylaxis of thrombosis	INR goal of 2-3 (INR: 2.5-3.5 for mechanical mitral valve prostheses or double valve replacement)	Assessing individual risks for thromboembolism and bleeding	-
	Prevention of stroke and systemic embolism in NVAF	150 mg oral BID	110 mg BID (if age ≥ 80, increased bleeding risk	Contraindicated if CrCl < 30ml/min
Dabigatran	Treatment of DVT and PE in patients who have been treated with a parenteral anticoagulant for 5-10 days and prevention of recurrent DVT and PE in patients who have been previously treated	150 mg oral BID	or concomitant use of verapamil)	or severe hepatic impairment Active pathological bleeding Idarucizumab: specific antidote (not yet available)

#### **Oral Anticoagulants (Cont.)**

p. l l 7

Drug	Indications	Dose	Dose adjustments	Comments	
_	Prevention of stroke and systemic embolism in NVAF	20 mg oral QD	CrCl < 50ml/min: 15 mg QD	Contraindicated if CrCl < 15ml/min	
Rivaroxaban	Treatment of DVT and PE and prevention of recurrent DVT and PE	15 mg oral BID for the first 3 weeks followed by 20 mg QD	Reduce the maintenance dose to 15 mg QD if bleeding risk outweighs the risk for recurrent DVT and PE (not formally approved)	or hepatic disease associated with coagulopathy and clinically relevant bleeding risk	
<b>E</b>	Prevention of atherothrombotic events after an ACS	2.5 mg oral BID	-	Diccomy risk	
Apixaban	Prevention of stroke and systemic embolism in NVAF	5 mg oral BID	2.5 mg oral BID  1) when at least 2 of the following characteristics: age ≥ 80,  Cr > 1.5 mg/dl or weight < 60Kg  2) when CrCl 15-29 mL/min	Contraindicated if CrCl < 15ml/min or severe hepatic impairment	
	Treatment of DVT and PE	10 mg oral BID for the first 7 days followed by 5 mg oral BID	-		
	Prevention of recurrent DVT and PE	2.5 mg oral BID	-		

#### **Intravenous/Subcutaneous Anticoagulants**

p. I I 8

Drug	Indications	Dose	Dose adjustments	Comments
	NSTE-ACS	LD: 4000 IU i.v. MD: 1000 IU/h i.v.	Target aPTT: 50-70s or 1.5 to 2.0 times that of control to be monitored at 3, 6, 12 and 24h	Monitoring for heparin-induced thrombocytopenia (HIT) Dose-independent reaction
P.	STEMI	Primary PCI: 70-100 IU/Kg i.v. when no GP-IIb/IIIa inhibitor is planned. 50-60 IU/Kg i.v. bolus with GP-IIb/IIIa inhibitors - Fibrinolysis/No reperfusion: 60 IU/kg i.v. bolus (max: 4000 IU) followed by an i.v. infusion of 12 IU/kg (max: 1000 IU/h) for 24-48h	Target aPTT: 50-70s or 1.5 to 2.0 times that of control to be monitored at 3, 6, 12 and 24h	
	Treatment of DVT and PE	80 IU/Kg i.v. bolus followed by 18 IU/Kg/h	According to aPTT, thromboembolic and bleeding risk	
×	NSTE-ACS 2.5 mg QD s.c.		-	Severe hepatic impairment: caution
Fondaparinux	STEMI	Fibrinolysis/No reperfusion: 2.5 mg i.v. bolus followed by 2.5 mg QD s.c. up to 8 days or hospital discharge	-	advised Contraindicated if CrCl < 20ml/min
Fonda	Treatment of DVT and PE	5 mg QD s.c. (< 50 kg); 7.5 mg QD s.c. (50-100 kg); 10 mg QD s.c.(> 100 kg)	If > 100Kg and CrCl 30-50ml/min: 10 mg followed by 7.5 mg/24h s.c.	Contraindicated for DVT/PE treatment
	Prevention of VTE	2.5 mg QD s.c.	CrCl 20-50ml/min: I.5 mg QD s.c.	if CrCl < 30ml/min

#### Intravenous/Subcutaneous Anticoagulants (Cont.)

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Drug	Indications	Dose	Dose adjustments	Comments
	PCI for NSTE-ACS	0.75 mg/kg i.v. bolus followed immediatelly by 1.75 mg/kg/h infusion which may be continued for up to 4h post PCI as clinically warranted and further continued at a reduced infusion dose of 0.25 mg/kg/h for 4-12h as clinically necessary	Patients undergoing PCI with CrCl 30-50ml/min should receive a lower infusion rate of 1.4 mg/kg/h. No change for the bolus dose.	Contraindicated if CrCl < 30ml/min
Bivalirudin	PCI for STEMI	0.75 mg/kg i.v. bolus followed immediatelly by 1.75 mg/kg/h infusion which should be continued for up to 4h after the procedure After cessation of the 1.75 mg/kg/h infusion, a reduced infusion dose of 0.25 mg/kg/h may be continued for 4-12h		
	PCI for elective cases	0.75 mg/kg i.v. bolus followed immediatelly by 1.75 mg/kg/h infusion which may be continued for up to 4h post PCI as clinically waranted		

#### Intravenous/Subcutaneous Anticoagulants (Cont.)

p. I 20

Drug	Indications	Dose	Dose adjustments	Comments
	NSTE-ACS	30 mg i.v.+ I mg/kg s.c. BID	If > 75 years: no LD and MD 0.75 mg/Kg BID s.c. CrCl < 30ml/min: no LD and MD I mg/Kg QD s.c. If > 75 years and CrCl < 30ml/min: no LD and 0.75 mg/Kg QD s.c.	Monitoring for HIT - Anti Xa monitoring during treatment with LMWH might be helpful in pregnancy, extreme body weights and renal impairment.
Enoxaparin	STEMI	Primary PCI: 0.5 mg/Kg i.v. bolus Fibrinolysis/No reperfusion: a) Age < 75y: 30 mg i.v. bolus followed by I mg/Kg BID s.c. until hospital discharge for a max of 8 days - The first two doses should not exceed 100 mg b) Age > 75y: no bolus; 0.75 mg/Kg BID s.c The first two doses should not exceed 75 mg	In patients with CrCl < 30 ml/min: regardless of age, the s.c. doses are given once daily	·
	Treatment of DVT and PE	I mg/Kg s.c. BID or I.5 mg/Kg s.c. QD	CrCl < 30ml/min: I mg/Kg/24h s.c.	
	Prevention of VTE	40 mg s.c. QD	CrCl < 30ml/min: 20 mg s.c. QD	

#### Intravenous/Subcutaneous Anticoagulants (Cont.)

p. | 2 |

Drug	Indications	Dose	Dose adjustments	Comments
Tinzaparin	Prevention of VTE	3500 IU s.c. QD (moderate risk) 4500 IU s.c. QD (high risk)	-	Monitoring for HIT - Anti Xa monitoring during treatment with LMWH
Tinza	Treatment of DVT and PE	175 IU/Kg s.c. QD	-	might be helpful in pregnancy, extreme body weights and renal impairment - Dalteparin:
Dalteparin	Prevention of VTE	2500 IU s.c. QD (moderate risk) 5000 IU s.c. QD (high risk)	-	In cancer patients, dose of 200 IU/kg (max:18000 IU)/24h for I month, followed by 150 IU/
	Treatment of DVT and PE	200 IU/Kg QD or 100 IU/Kg BID s.c.	Anti Xa monitoring if renal impairment	kg/24h for 5 months - After this period, vitamin K antag or a LMWH should be continued indefinitely or until the cancer is considered cured
Argatroban	Anticoagulant in patients with HIT	Initial i.v. infusion dose: 2 µg/kg/min (not to exceed 10 µg/kg/min) Patients undergoing PCI: 350 µg/kg i.v. followed by 25 µg/kg/min i.v.	Renal and hepatic impairment: caution advised	Monitored using aPTT goal: 1.5 to 3.0 times the initial baseline value PCI:ACT goal: 300-450s

#### **Fibrinolytics**

p. l 22

Drug	Indications	Dose	Dose adjustments	Comments
(SK)	STEMI	1.5 million units over 30-60min i.v.	-	Absolute contraindications to fibrinolytics:
Streptokinase	Treatment of PE	250000 IU as a LD over 30min, followed by 100000 IU/h over 12-24h	-	Previous intracranial haemorrhage or stroke of unknown origin at any time Ischaemic stroke in the preceding 6 months Central nervous system damage or neoplasms or atrioventricular malformation Recent major trauma/surgery/head injury (within the preceding 3 weeks)
ase (tPA)	STEMI	15 mg i.v. bolus: 0.75 mg/kg over 30 min (up to 50 mg) then 0.5 mg/kg over 60 min i.v. (up to 35 mg)	-	Gastrointestinal bleeding within the past month Known bleeding disorder (excluding menses) Aortic dissection Non-compressible punctures in the past 24h (e.g. liver biopsy, lumbar puncture)
Alteplase	Treatment of PE	Total dose of 100 mg: 10 mg i.v. bolus followed by 90 mg i.v. for 2h	If weight < 65 Kg: max dose < 1.5 mg/kg	

#### **Fibrinolytics (Cont.)**

p. I 23

Drug	Indications	Dose	Dose adjustments	Comments
Reteplase (rt-PA)	STEMI	10 units + 10 units i.v. bolus given 30 min apart	Renal and hepatic impairment: caution advised	Absolute contraindications to fibrinolytics:  Previous intracranial haemorrhage or stroke of unknown origin at any time Ischaemic stroke in the preceding 6 months Central nervous system damage or neoplasms
Tenecteplase (TNK-tPA)	STEMI	Over 10 seconds; Single i.v. bolus: 30 mg if < 60 kg 35 mg if 60 to < 70 kg 40 mg if 70 to < 80 kg 45 mg if 80 to < 90 kg 50 mg if ≥ 90 kg	-	or atrioventricular malformation Recent major trauma/surgery/head injury (within the preceding 3 weeks) Gastrointestinal bleeding within the past month Known bleeding disorder (excluding menses) Aortic dissection Non-compressible punctures in the past 24h (e.g. liver biopsy, lumbar puncture)

#### **Antiischemic drugs**

p. I 24

Drug	Indications	Dose	Dose adjustments	Comments
Beta-block	ers: Preferred over ca	alcium channel blockers - Contraindicated if coronary spass	m, severe bradycardia, AV bl	ock, severe bronchospasm
9	NSTE-ACS	LD: 25-100 mg oral MD: 25-100 mg QD	Elderly: start at a lower dose	Only if normal LVEF
Atenolol	STEMI	25-100 mg QD, titrate as tolerated up to 100 mg QD only if no LVSD or CHF	CrCl: 15-35ml/min: max dose 50 mg/day; CrCl < 15ml/min: max dose 25 mg/day	
loliba	NSTE-ACS	LD: 3.125-25 mg oral MD: 3.125-25 mg BID	Caution in elderly and hepatic impairment	Preferred if LVSD/HF
Carvedilol	STEMI	3.125-6.25 mg BID, titrated as tolerated up to 50 mg BID		
Bisoprolol	NSTE-ACS	LD: 1.25-10 mg oral MD: 1.25-10 mg QD	Caution in renal or hepatic impairment	Preferred if LVSD/HF
	STEMI	1.25-5 mg QD, titrate as tolerated up to 10 mg QD		

p. I 25

Drug	Indications	Dose	Dose adjustments	Comments
Beta-block	ers: Preferred over ca	lcium channel blockers - Contraindicated if coronary spass	m, severe bradycardia, AV blo	ock, severe bronchospasm
Metoprolol	NSTE-ACS	LD: 25-100 mg oral MD: 25-100 mg BID	Caution in hepatic impairment	Preferred if LVSD/HF
Met	STEMI	5-25 mg BID, titrate as tolerated up to 200 mg QD		
Calcium an	tagonists: Consider	if beta-blockers are contraindicated. First option in vasospa	astic angina	
Verapamil	ACS	LD: 80-120 mg oral MD: 80-240 mg TID-QD	Caution in elderly, renal or hepatic impairment	Contraindicated if bradycardia, HF, LVSD
Diltiazem	ACS	LD: 60-120 mg oral MD: 60-300 mg TID-QD	Caution in elderly and hepatic impairment	Contraindicated if bradycardia, HF, LVSD

p. I 26

Drug		Indications	Dose	Dose adjustments	Comments
Cal	cium ant	tagonists: Consider	if beta-blockers are contraindicated. First option in vasosp	astic angina	
	Amlodipine	ACS	LD: 5-10 mg oral, MD: 5-10 mg QD	Caution in hepatic impairment	Contraindicated if hypotension
Nit	rates				
Nitroglycerin	<u>.</u> .	ACS	If intolerant or unresponsive to nitroglycerin s.1.5 µg/min - Increase by 5 mcg/min q3-5min up to 20 µg/min - If 20 mcg/min is inadequate, increase by 10 to 20 µg/min every 3 to 5min - Max dose: 400 µg/min	-	Contraindicated if severe hypotension and co-administration with phosphodiesterase inhibitors The most common adverse effects
	spray	Angina	I-2 puff s.l. every 5min as needed, up to 3 puff in I5min	-	use glass bottles for nitroglycerin
Ž	sublingual tablet	Angina	0.3 to 0.6 mg s.l. or in the buccal pouch every 5min as needed, up to 3 doses in 15min	-	- I.V. AUTHIIISU AUOH

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Drug	Indications	Dose	Dose adjustments	Comments
Isosorbide mononitrate	Angina	5-10 mg BID with the two doses given 7h apart (8am and 3pm) to decrease tolerance development - then titrate to 10 mg BID in first 2-3 days  Extended release tablet: Initial: 30-60 mg given in the morning as a single dose  Titrate upward as needed, giving at least 3 days between increases  Max daily single dose: 240mg	-	Contraindicated if severe hypotension and co-administration with phosphodiesterase inhibitors The most common adverse effects are headache and dizziness
Isosorbide dinitrate	Angina	Initial dose: 5 to 20 mg orally 2 or 3 times/day MD: 10 to 40 mg orally 2 or 3 times a day Extended release: 40 to 160 mg/day orally	-	
Nitroglycerin transdermal patch	Angina	0.2 to 0.4 mg/h patch applied topically once a day for 12 to 14h per day; titrate as needed and tolerated up to 0.8 mg/h	-	

p. I 28

Drug	Indications	Dose	Dose adjustments	Comments
Other antii	shemic drugs			
Ivabradine	Stable angina	5-7.5 mg oral BID	Caution in elderly and CrCl < I5ml/min	Contraindicated if severe hepatic impairment
Ranolazine	Stable Initial dose: 375 mg oral BID  After 2-4 weeks, the dose should be titrated to 500 mg BID and, according to the patient's response, further titrated to a recommended max dose of 750 mg BID		Use with caution in renal and hepatic impairment, CHF, elderly, low weight	Contraindicated if CrCl < 30ml/ min, concomitant administration of potent CYP3A4 inhibitors, moderate or severe hepatic impairment
Trimetazidine	Stable angina Modified-release: 35 mg oral BID		Caution in elderly and 30 < CrCl < 60ml/min	Contraindicated in parkinson disease, parkinsonian symptoms, tremors, restlessleg syndrome, movement disorders, severe renal impairment

#### **Hypolipidemic drugs**

p. I 29

Drug	Indications		Dose		Dose adjustments	Comments	
	Statins: Secondary prevention of cardiovascular disease: start with high doses and down titrate if side effects [arget LDL-C levels < 70 mg/dl initiated early after admission						
Atorvastatin		LDL-C r	eduction		-	Contraindicated in	
Rosuvastatin	<30%	30-40%	40-50%	>50%	CrCl < 30ml/min: start 5 mg QD,	patients with active liver disease or with	
	Simva 10 mg	Simva 20-40 mg	Simva 40 mg	Ator 80 mg	max: 10 mg QD	unexplained elevation of liver function	
Pitavastatin	Lova 20 mg	Ator 10 mg	Ator 20-40 mg	Simva/ezet 40/10 mg	CrCl 30-59ml/min: start I mg QD, max 2 mg/day; CrCl 10-29ml/min: not defined	enzyme levels	
Simvastatin	Prava 20-40 mg	Prava 40 mg	Rosu 10-20 mg	Rosu 40 mg	Severe renal impairment: start 5 mg QPM		
Fluvastatin	Fluva 40 mg	Fluva 80 mg	Pita 4 mg		Caution in severe renal impairment		
Pravastatin	Pita I mg	Rosu 5 mg	Simva/ezet 20/10 mg		Significant renal impairment: start 10 mg QD		
Lovastatin		Pita 2 mg			CrCl < 30ml/min: caution if dose > 20 mg QD		

## **Hypolipidemic drugs (Cont.)**

p. I 30

Drug	Indications	Dose	Dose adjustments	Comments
Others				
Ezetimibe	Hyperlipidemia	10 mg oral QD	Avoid use if moderate-severe hepatic impairment	-
Fenofibrate	Hyperlipidemia	48-160 mg oral QD May adjust dose q4-8 weeks	CrCl 50-90ml/min: start 48-54 mg QD	Contraindicated if CrCl < 50ml/min or hepatic impairment
Gemfibrozil	Hyperlipidemia	900-1200 mg/day oral		Contraindicated if severe renal impairment or hepatic dysfunction Statins may increase muscle toxicity: avoid concomitant use
Evolocumab	PCSK9 inhibitor (not yet available). Most common side effects: nasopharyngitis, upper respiratory tract infection, headache and back pain			

#### **Heart failure & hypertension**

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Drug	Indications	Dose	Dose adjustments	Comments				
ACEI	ACEI							
pri	HF	Start: 6.25 mg oral TID Target dose: 50 mg TID	CrCl > 50 ml/min: 75-100% of the normal dose CrCl 10-50ml/min: 25-50% CrCl < 10ml/min: 12.5%	Check renal function, electrolytes, drug interactions				
Captopril	HTN	Start: 12.5 mg oral BID Target dose: 25-50 mg TID Max 450 mg/day		Major contraindications: History of angioedema, known				
Enalapril	HE,HTN	Start: 2.5 mg oral BID Target dose: 10-20 mg BID	CrCl 10-30ml/min: start 5 mg/day CrCl 10-30ml/min: start 2.5 mg/day	bilateral renal artery stenosis, pregnancy (risk)				
Lisinopril	HF	Start: 2.5-5.0 mg oral QD Target dose: 20-35 mg QD	CrCl 31-80ml/min:start 5-10 mg/day CrCl 10-30ml/min:start 2.5-5 mg/day CrCl < 10ml/min:start 2.5 mg/day					
	HTN	10-20 mg oral QD Max: 80 mg QD						

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Drug	Indications	Dose	Dose adjustments	Comments
ř.	HF	Start: 2.5 mg oral QD Max: 5mg QD	CrCl > 60ml/min: start 5 mg/day CrCl 31-60ml/min: start 2.5 mg/day	Check renal function, electrolytes,
Perindopril	HTN	Start: 2.5-5 mg QD Target dose: 10 mg QD	CrCl 15-30ml/min: start 2.5 mg alternate days CrCl < 15ml/min: start 2.5 mg/day on the day of dialysis	drug interactions  Major contraindications: History of angioedema, known
Ramipril	HF, HTN	Start: 2.5 mg oral QD Target dose: 5 mg BID	CrCl < 40ml/min: start 1.25 mg QD, max 5 mg/day Caution in elderly and hepatic impairment	bilateral renal artery stenosis, pregnancy (risk)
Trandolapril	HF	Start: 0.5 mg oral QD Target dose: 4 mg QD	CrCl < 30ml/min or severe hepatic impairment start 0.5 mg	
F	HTN	2-4 mg oral QD	CrCl < 30ml/min or severe hepatic impairment: start 0.5 mg	

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Drug	Indications	Dose	Dose adjustments	Comments
ARB				
Candesartan	НЕНТИ	Start: 4-8 mg oral QD Target dose: 32 mg QD	If renal or hepatic impairment: start 4 mg/day	If ACEI is not tolerated. Check renal function, electrolytes, drug interactions  Major contraindications: History of angioedema, known bilateral renal artery stenosis,
Losartan	HF	Start: 50 mg oral QD Target dose: 150 mg QD	CrCl < 20ml/min: 25 mg QD Caution if hepatic impairment	pregnancy (risk)
Losa	HTN	50-100 mg oral QD	CrCl < 20ml/min: 25 mg QD Caution if hepatic impairment	
ırtan	HF	Start: 40 mg oral BID Target dose: 160 mg BID	If mild-moderate hepatic impairment: max dose 80 mg/day	
Valsartan	HTN	80-160 mg QD	If mild-moderate hepatic impairment: max dose 80 mg/day	

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Dr	ug	Indications	Dose	Dose adjustments	Comments
Bet	ta-bl	ockers: Check 12- lea	id ECG		
	Atenolol	HTN	Start: 25 mg oral QD Usual dose: 50-100 mg QD	CrCl 10-50ml/min: decrease dose 50% CrCl < 10ml/min: decrease dose 75%	Major contraindications: asthma, 2nd or 3rd degree AV block
tive	lolo	HF	Start: 1.25 mg oral QD Target dose: 10 mg QD	CrCl < 20ml/min: max dose 10 mg QD Hepatic impairment: avoid use	
Cardioselective	Bisoprolol	HTN	Start: 2.5-5 mg oral QD Usual dose: 5-10 mg QD Max dose: 20 mg QD		
S	Metoprolol	HF	Start: 12.5-25 mg oral QD Target dose: 200 mg QD	Hepatic impairment: start with low doses and titrate gradually	
	Meto	HTN	100-400 mg QD Max dose: 400 mg QD		

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Dr	ug	Indications	Dose	Dose adjustments	Comments
Bet	ta-bl	ockers: Check 12- lea	ad ECG		
elective	Nebivolol	HF	Start: I.25 mg oral QD Target dose: I0 mg QD	Renal impairment or elderly: start dose 2.5 mg QD, titrate to 5 mg QD Hepatic impairment: contraindicated	Major contraindications: asthma, 2nd or 3rd degree AV block
Cardioselective	Nebi	HTN	Start: 2.5 mg oral QD Usual dose: 5 mg QD		
selective	dilol	HF	Start: 3.125 mg oral BID Target dose: 25-50 mg BID	Caution in elderly Contraindicated if hepatic impairment	
Non-cardioselective	Carvedilol	HTN	Start: 12.5 mg oral QD Usual dose: 25 mg QD and max dose: 25 mg BID or 50 mg QD		

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Drug	Indications	Dose	Dose adjustments	Comments					
Other v	Other vasodilators								
Amlodipine	HTN	Start: 5 mg oral QD, increase after 1-2 weeks Max: 10 mg/day	Elderly or secondary agent: start 2.5 mg QD Hepatic impairment: start 2.5 mg QD	Contraindicated if cardiogenic shock, 2nd or 3rd degree AV block, severe hypotension					
Nifedipine	HTN	Extended-release form: Start 20 mg oral BID or TID Max: 60 mg BID	Renal and hepatic impairment: caution advised						
Clevidipine	HTN	Initiate the IV infusion at 4 ml/h (2 mg/h); the dose may be doubled every 90 seconds Uptitration until desired BP range is achieved Half life of I-2min	The desired therapeutic response for most patients occurs at doses of 8-12 ml/h (4-6 mg/h) The max recommended dose is 64 ml/h (32 mg/h)	Hypersensitivity to soy, peanut, or egg products Critical Aortic stenosis, mitral stenosis, HOCM					

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Drug	Indications	Dose	Dose adjustments	Comments			
Other vasodilators							
Verapamil	HTN	Immediate-release form: Dose: 80-120 mg oral TID; Start: 80 mg TID; Max: 480 mg/day	Start 40 mg oral TID in elderly or small stature patients	Contraindicated if bradycardia, HF, LVSD			
Loop diuretics							
Furosemide	HF	20-40 mg i.v. bolus, continuous 100 mg/6h (adjust based on kidney function and clinical findings; monitor creatinine)	Anuria: contraindicated Cirrhosis/ascites: caution advised	-			
	HTN	10-40 mg oral BID					
Torsemide	HF	10-20 mg oral or i.v. QD	Hepatic impairment: initial dose should be reduced by 50% and dosage adjustments made cautiously	-			
	HTN	5 mg oral or i.v. QD Max 10 mg QD					

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Drug	Indications	Dose	Dose adjustments	Comments				
Thiazid	Thiazides							
Chlorthalidone	HF	50-100 mg oral QD MD: 25-50 mg QD	Elderly: max dose 25 mg/day CrCl < 25ml/min: avoid use	-				
	HTN	Start 12.5-25 mg oral QD; Max: 50 mg/day	Elderly: max dose 25 mg/day CrCl < 25ml/min: avoid use	-				
Hydrochlorothiazide	HF	25-200 mg oral/day	CrCl < 25 ml/min: avoid use Hepatic impairment: caution advised	-				
	HTN	Start 12.5-25 mg oral QD MD: may increase to 50 mg oral as a single or 2 divided doses	CrCl < 25 ml/min:avoid use Hepatic impairment: caution advised	-				

## **Heart failure & hypertension (Cont.)**

p. 139

Drug	Indications	Dose	Dose adjustments	Comments	
Thiazid	es				
Indapamide	HTN	Start I.25 mg PO QAM x4weeks, then increase dose if no response Max: 5 mg/day	CrCl < 25 ml/min: avoid use Hepatic impairment: caution advised	-	
Aldoste	rone-antagonists				
tone	HF	Start 25 mg oral QD Target dose: 25-50 mg QD	CrCl < 10ml/min, anuria or acute renal impairment: contraindicated Severe hepatic impairment and elderly:	Check renal function, electrolytes, drug interactions	
Spironolactone	HTN	50-100 mg/day oral	caution advised	Produces gynecomastia	

## **Heart failure & hypertension (Cont.)**

p. 140

Drug	Indications	Dose	Dose adjustments	Comments	
Aldoste	rone-antagonists				
Eplerenone	HF	Start 25 mg oral QD Target dose: 50 mg QD	Elderly: caution advised CrCl < 50ml/min: contraindicated	Check renal function, electrolytes, drug interactions Major contraindications: strong CYP3A4 inhibitors	
Eplere	HTN	50 mg oral QD-BID Max: I 00 mg/day			
Others					
Ivabradine	HF	5-7.5 mg oral BID	Caution in elderly and CrCl < 15ml/min	Contraindicated if severe hepatic impairment	

## **Inotropics & vasopressors**

p. 141

Drug	Indications	Dose	Dose adjustments	Comments
Levosimendan	HF/cardiogenic shock	LD: 6 to 12 µg/kg i.v. over 10 min (given only if immediate effect is needed) followed by 0.05 to 0.2 µg/kg/min as a continuous infusion for 24h	Avoid use if CrCl < 30ml/min or severe hepatic impairment	Calcium sensitizer and ATP-dependent potassium channel opener
Milrinone	HF/cardiogenic shock	50 μg/kg i.v. in 10-20 min, continuous 0.375-0.75 μg/kg/min	Renal: Same bolus. Adjust infusion: CrCl 50ml/min: start 0.43 µg/kg/min CrCl 40ml/min: start 0.38 µg/kg/min CrCl 30ml/min: start 0.33 µg/kg/min CrCl 20ml/min: start 0.28 µg/kg/min CrCl 10ml/min: start 0.23 µg/kg/min CrCl 5ml/min: start 0.20 µg/kg/min	Phosphodiesterase inhibitor  Caution if atrial flutter  Hypotensive drug
line/ renol	Cardiogenic shock	0.5-5 μg/min (0.25-2.5 ml of a 1:250,000 dilution) i.v. infusion	-	ß1, ß2 agonist. Contraindicated in patients with tachyarrhythmia,
Isoprenaline/ Isoproterenol	Bradyarrhythmias	Bolus: 20-40 µg i.v. Infusion: 0.5 µg/min of 2 mg/100 ml normal saline		tachycardia or heart block caused by digitalis intoxication, ventricular arrhythmias which require inotropic therapy, angina pectoris, recent ACS, hyperthyroidism

## **Inotropics & vasopressors (Cont.)**

p. I 42

Drug	Indications	Dose	Dose adjustments	Comments
Dobutamine	Cardiogenic shock	2-20 μg/kg/min i.v.	-	$B1, \alpha1/B2$ agonist. Increases contractility with little effect on heart rate and blood pressure. Reduces pulmonary and systemic VR, PCP
Dopamine	Cardiogenic shock	Dopaminergic effect: 2-5 $\mu$ g/Kg/min i.v. $\beta$ effect : 5-15 $\mu$ g/Kg/min i.v. $\alpha$ effect : 15-40 $\mu$ g/Kg/min i.v.	-	$B, \alpha$ , dopaminergic agonist Increases BP, PAP, heart rate, cardiac output and pulmonary and systemic VR More arrhythmogenic than dobutamine and noradrenaline
Noradrenaline	Cardiogenic shock	0.05-0.2 µg/kg/min i.v. titrate to effect	-	α I, βI agonist Increases BP and PAP Little arrhythmogenic

## **Antiarrhythmics**

p. 143

Drug	Indications	Dose	Dose adjustments	Comments
Group	1			
Procainamide i.v.	AF (termination); stable VT (with a pulse)	15-18 mg/kg i.v. over 60 min, followed by infusion of 1-4 mg/min	Reduce LD to 12 mg/kg in severe renal impairment Reduce MD by one-third in moderate renal impairment and by two-thirds in severe renal impairment Caution in elderly and asthma	Hypotension (negative inotropic agent) Lupus-like syndrome Contraindicated if myasthenia gravis,AV block, severe renal impairment
Lidocaine i.v.	Pulseless VT/VF	I-1.5 mg/kg i.v./i.o. bolus (can give additional 0.5-0.75 mg/kg i.v./i.o. push every 5-10 min if persistent VT/VF, max cumulative dose = 3 mg/kg), followed by infusion of I-4 mg/min	1-2 mg/min infusion if liver disease or HF	Contraindicated if advanced AV block, bradycardia, hypersensitivity to local anesthetics
	Stable VT (with a pulse)	I-1.5 mg/kg i.v. bolus (can give additional 0.5-0.75 mg/kg i.v. push every 5-10min if persistentVT, max cumulative dose = 3 mg/kg), followed by infusion of I-4 mg/min	1-2 mg/min infusion if liver disease or HF	Caution in HF, renal impairment and elderly May cause seizures, psychosis. Stop if QRS widens > 50%

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Drug	Indications	Dose	Dose adjustments	Comments
Group	1			
Flecainide i.v.	SVT, ventricular arrhythmias	2 mg/kg (max 150 mg) i.v. over 30min This may be followed by an infusion at a rate of 1.5 mg/kg/h for 1 h, reduced to 0.1-0.25 mg/kg/h for up to 24h, max cumulative dose = $600$ mg	Severe renal impairment: caution advised	Contraindicated if cardiogenic shock, recent MI, 2nd or 3rd degree AV block
Propafenone i.v.	PSVT, ventricular arrhythmias	LD: 0.5-2 mg/kg i.v. direct over a minimum of 3-5min MD: 0.5-2.5 mg/kg i.v. direct q8h (max 560 mg/day) or continuous infusion up to 23 mg/h	May need to reduce dose in renal or hepatic failure	Contraindicated if unstable HF, cardiogenic shock, AV block, bradycardia, myasthenia gravis severe hypotension, bronchospastic disorders, Brugada syndrome

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Drug	Indications	Dose	Dose adjustments	Comments
Group	П			
Atenolol i.v.	Arrhythmias	2.5 mg i.v. over 2.5 min every 5 min (max 10 mg)	Caution in elderly and/or severe renal impairment	Contraindicated if cardiogenic shock, bradycardia and greater than first-degree block, unstable HF
Metoprolol i.v.	Arrhythmias	2.5-5mg i.v. over 5 min, may repeat every 5 min (max 15mg)	Caution if severe hepatic impairment	Contraindicated if cardiogenic shock, bradycardia and greater than first-degree block, unstable HF
Propranolol i.v.	Arrhythmias	Initially given as slow i.v. boluses of 1 mg, repeated at 2 min intervals (max: 10 mg in conscious patients and 5 mg if under anesthesia)	-	Contraindicated if cardiogenic shock, bradycardia and greater than first-degree block, asthma, unstable HF

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Drug	Indications	Dose	Dose adjustments	Comments
Group	III			
Amiodarone i.v.	AF (termination)	5 mg/Kg i.v. over 30 min, followed by infusion of 1 mg/min for 6h, then 0.5 mg/min	-	Reduce infusion rate if bradycardia, AV block, hypotension
	Stable VT (with a pulse)	150 mg i.v. over 10 min followed by infusion of 1 mg/min for 6h, then 0.5 mg/min	-	Bolus should be avoided if hypotension or severe LV dysfunction Highly vesicant agent
	Pulseless VT/VF	300 mg bolus i.v. (can give additional 150 mg i.v. bolus if VF/VT persists) followed by infusion of 900 mg over 24h	-	
Dronedarone	Paroxysmal or persistent AF prevention	400 mg oral BID	-	Contraindicated if severe renal or liver dysfunction, LVSD, symptomatic HF, permanent AF, bradycardia (multiple contraindications)

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Drug	Indications	Dose	Dose adjustments	Comments
Group	IV			
Diltiazem i.v.	PSVT;AF (rate control)	0.25 mg/kg i.v. over 2 min (may repeat with 0.35 mg/kg i.v. over 2 min), followed by infusion of 5-15 mg/h	Hepatic impairment: caution advised	-
Verapamil i.v.	PSVT;AF (rate control)	2.5-5 mg i.v. over 2 min (may repeat up to max cumulative dose of 20 mg); can follow with infusion of 2.5-10 mg/h	-	Contraindicated if AF+WPW, tachycardias QRS (except RVOT-VT), fascicular VT, bronchospasm, age>70 Antidote: - LVD: Calcium gluconate, dobutamine - Bradycardia/AV block: Atropine, Isoproterenol
Adenosine i.v.	Rapid conversion to a normal sinus rhythm of PSVT including those associated with accessory by-pass tracts (WPW syndrome)	Rapid i.v. boluses separated by 2 min: 6 mg $\rightarrow$ 6 mg $\rightarrow$ 12 mg	-	Contraindicated if sick sinus syndrome, second or third degree Atrio-Ventricular (AV) block (except in patients with a functioning artificial pacemaker), chronic obstructive lung disease with evidence of bronchospasm (e.g. asthma bronchiale), long QT syndrome, severe hypotension; decompensated states of heart failure - Adenosine can cause AF

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Drug	Indications	Dose	Dose adjustments	Comments
Other	s			
Magnesium sulfate	VT-Torsades de Pointes	Bolus: 1-2 g i.v/i.o. over 5 min Perfusion: 5-20 mg/min i.v.	Caution if severe renal failure	Contraindicated if myasthenia gravis
Vernakalant	Acute atrial fibrillation	3 mg/kg i.v. over 10 min. If AF persists, a second 10-min-infusion of 2 mg/kg, 15 min later may be administered	-	Contraindicated if ACS within the last 30 days, severe aortic stenosis, SBP < 100mmHg, HF class NYHA III/IV, severe bradycardia, sinus node dysfunction or 2nd or 3rd degree heart block

APTT = Activated partial thromboplastin time

AB = Airway and breathing ABG = Arterial blood gas

AADs = Antiarrhythmic drugs

AAS = Acute aortic syndrome

ACEI = Angiotensin converting enzyme inhibitor

ACLS = Advanced cardiovascular life support

ACS = Acute coronary syndrome

ACT = Activated clotting time

AD = Aortic Dissection

AED = Automated external defibrillator

AF = Atrial fibrillation

Ao = Aortic

aPRR = Activated partial thromboplastin time

ARB = Angiotensin receptor blockers

AS = Aortic stenosis AV = Atrioventricular

AVN = Atrioventricular node

AVNRT = Atrioventricular nodal re-entrant tachycardia

AVNT = Atrioventricular nodal tachycardia

BID = Twice a day

BBB = Bundle branch block

BLS = Basic life support

BNP = Brain natriuretic peptide

BP = Blood pressure

CABG = Coronary artery bypass grafting

CAD = Coronary artery disease

Cath Lab = Catheterisation laboratory

CCU = Coronary care unit CHF = Congestive heart failure

CMR = Cardiovascular magnetic resonance

COPD = Chronic obstructive pulmonary disease CPAP = Continuous positive airway pressure

CPR = Cardiopulmonary resuscitation

CrCI = Creatinine clearance

CS = Cardiogenic shock

CSM = Carotid sinus massage

CSNRT = Corrected sinus node recovery time

CSS = Carotid sinus syndrome CT = Computed tomography

CT-angio = Computed tomography angiography

CUS = Compression venous ultrasound

CV = Cardiovascular CXR = Chest X-ray

DD = Dyastolic dysfunction

DM = Diabetes mellitus

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DVT = Deep vein thrombosis ECG = Electrocardiogram ED = Emergency department

ED = Emergency department

EG = Electrograms

EMB = Endomyocardial biopsy

EMS = Emergency medical services

EPS = Electrophysiological study

ERC = European Resuscitation Council

ESR = Erythrocyte sedimentation rate

ETT = Exercice treadmill testing

FMC = First medical contact

GER = Gastroesophageal reflux

GFR = Glomerular flow rate

GI = Gastrointestinal

GP = Glycoprotein

HF = Heart failure

HTN = Hypertension

HR = Heart rate

hsTn = High-sensitive troponin

IABP = Intra-aortic balloon pump

ICC = Intensive cardiac care

ICCU = Intensive cardiac care unit

ICD = Implantable cardioverter defibrillator

IHD = Ischemic heart disease

IMH = Intramural hematoma

ISFC = International Society and Federation of

Cardiology

i.o. = Intraosseous

IV = Invasive ventilation

i.v. = Intravenous

KD = Kidney disease

LBBB = Left bundle branch block

LD = Loading dose

LGE = Late gadolinium enhancement

LMWH = Low-molecular weight heparin

LOC = Loss of consciousness

LV = Left ventricular

LVD = Left ventricular dysfunction

LVEF = Left ventricular ejection fraction

LVH = Left ventricular hypertrophy

LVSD = Left ventricular systolic dysfunction

MCS = Mechanical circulatory support

MD = Maintenance dose

MDCT = Computed tomography with >4 elements

MI = Myocardial infarction

MRI = Magnetic resonance imaging

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Mvo = Microvascular obstruction NIV = Non-invasive ventilation NOAC = New oral anticoagulants

NSAID = Non-steroidal anti-inflammatory drugs

NSTEACS = Non-ST-elevation ACS

NSTEMI = Non ST-segment elevation myocardial infarction

NTG = Nitroglycerin

NT-proBNP = N-terminal pro brain natriuretic peptide

NVAF = Non-valvular atrial fibrillation NYHA = New York Heart Association OH = Orthostatic hypotension

PAP = Pulmonary arterial pressure

PAU = Penetrating aortic ulcer

PCI = Percutaneous coronary intervention

PCM = Physical counter-measures PCP = Pulmonary capillary pressure

PE = Pulmonary embolism

PEA = Pulmonary endarterectomy

PEEP = Positive end expiratory pressure

PR = Pulmonary regurgitation

ProCT = Procalcitonin
PRN = Pro re nata

PSVT = Paroxysmal supraventricular tachycardia

QD = Once a day

QPM = Every evening

rtPA = Recombinant tissue plasminogen

activator

RV = Right ventricular

RVOT-VT = Right ventricular outflow tract ventricular

tachycardia

SBP = Systemic blood pressure

s.c = Subcutaneous

SLE = Systemic lupus erythematosus SMU = Syncope management units

STE-ACS = ST-segment elevation acute coronary syndrome

STEMI = ST-segment elevation myocardial infarction

SVT = Supraventricular tachycardia

 $SpO_2 = Oxygen saturation$ 

TEE = Transesophageal echocardiography

TEVAR = Thoracic endovascular aortic aneurysm repair

TIA = Transient ischemic attack

TID = Three times a day

TLOC = Transient loss of consciousness

Tn = Troponin

TOE = Transoesopageal echocardiography

TSH = Thyroid-stimulating hormone

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TTE = Transthoracic echocardiography

UFH = Unfractionated heparin

 ${\sf ULN} = {\sf Upper\ limit\ of\ normal}$ 

VF = Ventricular fibrillation

VR = Vascular resistance

VT = Ventricular tachycardia

VTE = Venous thromboembolism

VVS = Vasovagal syncope

WHO = World Health Organization

WPW = Wolff-Parkinson-White

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# Notes

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# Notes

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## References and copyright acknowledgments Acknowledgements

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# Acute Cardiovascular Care Association Clinical Decision-Making TOOLKIT



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