

Private Bag X828, PRETORIA, 0001, Room 401, Civitas Building, Pretoria

Reference: EDP022019/01

#### RECOMMENDATION FOR ALTERNATIVE THERAPY FOR STREPTOKINASE

The Primary Health Care and Adult Hospital Level Standard Treatment Guidelines (STGs) and Essential Medicines List (EML) currently recommends the use of streptokinase as thrombolytic therapy for the treatment of ST elevation myocardial infarction (STEMI) in adult patients. The supplier of streptokinase, however, is unable to meet supply demands due to unforeseen operational delays<sup>1</sup>. The supplier anticipates that the supply of streptokinase will resume from March 2019.

While streptokinase is unavailable, it is recommended that alteplase be used as thrombolytic therapy (when indicated) in patients with STEMI.

See Primary Health Care STGs and EML, 2018: Section 4.4 Myocardial infarction, acute (AMI)/ST elevation myocardial infarction (STEMI) and Adult Hospital Level STGs and EML, 2015: Section 3.2.1 ST elevation myocardial infarction (STEMI) for the treatment of STEMI, including indications and contraindications for thrombolysis.

## Alteplase dose and directions for use:

# A: PRIMARY HEALTH CARE LEVEL -with urgent referral to secondary level

- Alteplase, IV infusion<sup>2 3:</sup> (Doctor initiated)
  - Do not exceed 100 mg<sup>4</sup>.
  - o If within 6 hours of symptom onset:

Weight	Bolus	Next 30 minutes	Next 60 minutes
>67 kg	15 mg	50 mg	35 mg
≤67 kg	15 mg	0.75 mg/kg	0.5 mg/kg

Indications and contraindications are similar to those for streptokinase, as indicated in the table below (except that prior use of strentokinase is not a contraindication).

Indications	Contra-indications	
<ul> <li>For acute myocardial infarction with ST elevation or left bundle branch block:         <ul> <li>if history of onset &lt; 6 hours (Beyond 6 hours treat as NSTEMI (see below)</li> <li>if on-going ischaemic pain</li> </ul> </li> </ul>	<ul> <li>» Absolute: <ul> <li>streptokinase used within the last year,</li> <li>previous allergy,</li> <li>CVA within the last 3 months,</li> <li>history of recent major trauma,</li> <li>bleeding within the last month,</li> <li>aneurysms,</li> <li>brain or spinal surgery or head injury within the preceding month, or</li> <li>active bleeding or known bleeding disorder.</li> </ul> </li> <li>» Relative: <ul> <li>refractory hypertension,</li> <li>warfarin therapy,</li> <li>recent retinal laser treatment,</li> <li>subclavian central venous catheter,</li> <li>pregnancy,</li> <li>TIA in the preceding 6 months,</li> <li>traumatic resuscitation.</li> </ul> </li> </ul>	

<sup>&</sup>lt;sup>1</sup> Mirren (Pty) Ltd, Actor Speciality: communication on file at NDoH

<sup>-</sup> Single exit price of streptokinase 1.5 million units ≈R 4000.00, 21 December 2018.

2 Dundar Y, Hill R, Dickson R, Walley T. Comparative efficacy of thrombolytics in acute myocardial infarction: Asystematic review. QJM - Monthly Journal of the Association of Physicians. 2003;96(2):103-13.https://www.ncbi.nlm.nih.gov/pubmed/12589008

Thrombolytics (Therapeutic class): National Department of Health: Affordable Medicines, EDP-Adult Hospital level. Medicine Review: Thrombolytics, therapeutic class for STEMI, July

<sup>2015.</sup> http://www.health.gov.za/ 4 Quotation price of alteplase 100mg: R 11 830.88, 29 January 2019

Monitor the following, continuously and also during transfer:

- » pulse
- » blood pressure
- » respiration depth and rate (count for a full minute)
- » ECG

Note: Defibrillator should be readily available at all times including during transport.

## B: ADULT HOSPITAL LEVEL - management by doctor/medical officer

Alteplase, IV infusion, as per recommendations in the PHC STGs and EML, above.

### Adjunctive treatment

- Low molecular weight heparin<sup>5</sup> <sup>6</sup>, e.g.:
- Enoxaparin.
  - Loading dose: IV, 30 mg as a bolus, followed by SC, 1 mg/kg as a single dose (total cumulative dose not to exceed 100mg).
  - o Maintenance dose: SC, 1mg/kg 12 hourly.

In the elderly (> 75 years of age), omit IV loading dose and reduce SC dose:

o Loading dose: SC, 0.75 mg/kg as a single dose.

**Note:** These recommendations are provided as an interim measure and the National Department of Health will advise when the supply of streptokinase is stabilised and acquisition can be processed.

Provinces and Health Care Facilities are requested to distribute and communicate this information in consultation with the Pharmaceutical and Therapeutics Committees.

Kindly share with all healthcare professionals. Comments may be submitted via e-mail or post:

Stock queries:

Contract Management Unit Private Bag X828

**PRETORIA** 

0001

Tel: 012 3959539

E-mail: Buhle.Mbongo@health.gov.za

Clinical gueries:

**Essential Drugs Programme** 

Private Bag X828

**PRETORIA** 

0001

Tel: 012 3958287

E-mail: SAEDP@health.gov.za

Kind regards

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**DIRECTOR: AFFORDABLE MEDICINES** 

DATE: 1/2/2019

<sup>&</sup>lt;sup>5</sup> Eikelboom JW, Quinlan DJ, Mehta SR, Turpie AG, Menown IB, Yusuf S. Unfractionated and low-molecular-weight heparin as adjuncts to thrombolysis in aspirin-treated patients with ST-elevation acute myocardial infarction: a meta-analysis of the randomized trials. Circulation. 2005 Dec 20;112(25):3855-67. https://www.ncbi.nlm.nih.gov/pubmed/16344381

<sup>6</sup> De Luca G, Marino P. Adjunctive benefits from low-molecular-weight heparins as compared to unfractionated heparin among patients with ST-segment elevation myocardial infarction treated with thrombolysis. A meta-analysis of the randomized trials. Am Heart J. 2007 Dec;154(6):1085.e1-6. https://www.ncbi.nlm.nih.gov/pubmed/18035079