Contents lists available at ScienceDirect



# African Journal of Emergency Medicine

journal homepage: www.elsevier.com/locate/afjem



# ORIGINAL ARTICLE

# Essential medicines for emergency care in Africa<sup> $\star$ </sup>,

M.C. Broccoli<sup>a</sup>, J.L. Pigoga<sup>b</sup>, M. Nyirenda<sup>c</sup>, L.A. Wallis<sup>d</sup>, E.J. Calvello Hynes<sup>e,\*</sup>

<sup>a</sup> Department of Emergency Medicine, Boston Medical Center, Boston, MA, USA

<sup>b</sup> Rollins School of Public Health, Emory University, Atlanta, GA, USA

<sup>c</sup> Queen Elizabeth Central Hospital, Ministry of Health and University of Malawi College of Medicine, Blantyre, Malawi

<sup>d</sup> Division of Emergency Medicine, University of Cape Town, Cape Town, South Africa

<sup>e</sup> Department of Emergency Medicine, University of Colorado School of Medicine, CO, USA

ARTICLE INFO	A B S T R A C T	
<i>Keywords:</i> Essential medicines Emergency care Africa Health system	Objectives: Essential medicines lists (EMLs) are efficient means to ensure access to safe and effective medications. The WHO has led this initiative, generating a biannual EML since 1977. Nearly all countries have implemented national EMLs based on the WHO EML. Although EMLs have given careful consideration to many public health priorities, they have yet to comprehensively address the importance of medicines for treating acute illness and injury.         Methods: We undertook a multi-step consensus process to establish an EML for emergency care in Africa. After a review of existing literature and international EMLs, we generated a candidate list for emergency care. This list was reviewed by expert clinicians who ranked the medicines for overall inclusion and strength of recommendation. These medications and recommendations were then evaluated by an expert group. Medications that reached consensus in both the online survey and expert review were included in a draft emergency care EML, which underwent a final in-person consensus process.         Results: The final emergency care EML included 213 medicines, 25 of which are not in the 2017 WHO EML but were deemed essential for clinical practice by regional emergency providers. The final EML has associated recommendations of desirable or essential, and is subdivided by facility level. Thirty-nine medicines were recommended for basic facilities (e.g. tertiary centres).         Conclusion: The 25 novel medications not currently on the WHO EML should be considered by planners when making rational 76 for easier implementation, and will be a useful tool for practical expansion of emergency care delivery in Africa.	

# African relevance

What is already known about this subject:

- Essential medicines lists are efficient means to ensure access to safe and effective medications.
- Although essential medicines lists have given careful consideration to many public health priorities, they have yet to comprehensively address the importance of medicines for treating acute illness and injury.
- Low-resource settings with underdeveloped emergency care systems are in urgent need of emergency care-specific essential medicines

lists to enhance facility and system planning.

What this study adds:

- This manuscript proposes an emergency care essential medicines lists that includes 213 medicines, 25 of which are not in the 2017 WHO essential medicines lists.
- These medications were selected and agreed upon by expert clinicians practicing emergency care in Africa using a multi-phase consensus process.
- The final essential medicines lists has associated recommendations of desirable or essential, and is subdivided by facility level (basic,

\* Corresponding author.

https://doi.org/10.1016/j.afjem.2018.05.002

Available online 29 June 2018

2211-419X/ 2018 African Federation for Emergency Medicine. Publishing services provided by Elsevier. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/BY-NC-ND/4.0/).

<sup>\*</sup> This article has been co-published in the Emergency Medicine Journal; published Online First: 07 April 2018. Doi: http://doi.org/10.1136/emermed-2017-207396. Peer review under taken by the Emergency Medicine Journal.

E-mail address: emilie.calvellohynes@ucdenver.edu (E.J. Calvello Hynes).

intermediate, advanced).

### Introduction

Since the introduction of the World Health Organization's (WHO) Model List of Essential Medicines in 1977, essential medicines lists (EMLs) have become a core component of primary healthcare [1]. EMLs have nearly doubled access to essential medicines and are often cited as one of the most cost-effective interventions in modern healthcare [2,3]. By effectively limiting the number of medicines available, EMLs encourage prescription of higher-quality medicines [4], guide proper use and dosage of these medicines, and facilitate clinical education and training [1]. EMLs can also address public health concerns such as antibiotic resistance by promoting responsible use of these medicines [5].

The WHO EML is evidence-based, with medicines being prioritised based on public health relevance, safety, cost-effectiveness, and efficacy to ensure that they can be available in adequate doses at all times [6,7]. Ninety-five percent of countries have national EMLs guided by the WHO EML [4,7]; national EMLs are then used to inform treatment guidelines and allow for mandates surrounding procurement, provision, and training. Aid groups also rely on the WHO EML to create widely-applicable purchase lists and field guidelines [8,9].

From a population perspective, EMLs represent cost-effective interventions directed toward preventing and treating diseases [1]. However, the current WHO list has not been oriented toward a crucial form of secondary prevention – the appropriate treatment of patients with emergent clinical presentations. Emergency care is recognised as an effective means of reducing morbidity and mortality, especially in low-resource settings where the majority of the global burden of injury and illness resides [10–12]. Emergency care remains an underdeveloped aspect of healthcare in much of the world [10,13,14], and the essential medicines necessary for a general approach to treating acute illness and injury across multiple diseases have not been formally specified.

Some countries include medicines lists in their emergency care treatment guidelines, but these lists are not vetted for cost-effectiveness, public health relevance, and evidence of efficacy and safety [7]. The WHO EML itself receives minimal input from emergency care specialists [11]; the resultant list prioritizes vaccines and medications for the chronic treatment of infectious and non-communicable diseases but has notable gaps in vital medications required for emergency care [7]. In high-income countries the current WHO inclusion list is not problematic; systems are able to tailor the WHO EML to their already-functioning emergency units. But low-resource settings where emergency care systems are far less developed lack the capacity to complete such a process, yet are in urgent need of emergency care-specific medication lists to enhance facility and system planning [10,14,15].

More than half of Africans lack access to essential medicines [2] and many are required to pay for medicines prior to receiving care [16]. This is particularly problematic and often grossly unethical in emergency situations when medicines are needed immediately, regardless of ability to pay. A simplified emergency care EML would ensure the safety, efficacy, and availability of life-saving medicines [17]. It would serve to guide an essential component of emergency service delivery, leading to a more reliable drug supply and improved prescribing patterns, likely with improved patient outcomes and cost-effectiveness [4,8].

Over the coming decades, many African countries are expected to develop their emergency care systems. Currently, only one example of a country-level essential emergency medication list exists in Africa [18]. This paper describes a multi-step consensus process which we undertook to establish an EML for emergency care in Africa. We believe that such a list will be a useful tool for practical expansion of emergency care delivery in Africa, and that increasing the availability of essential emergency medicines will ultimately improve outcomes from acute African Journal of Emergency Medicine 8 (2018) 110-117

injury and illness.

## Literature review

A comprehensive literature review of both peer-reviewed publications and grey literature was conducted in late 2015 to identify EMLs specific to emergency care. A list of key search terms surrounding emergency medications and EMLs was devised. PubMed, MEDLINE, and Google Scholar were first searched to identify EMLs specific to emergency care; however, this search only returned one such list, published by the Ethiopian Food, Medicine and Health Care Administration and Control Authority in 2014 [18].

The search was then broadened to look for lists of essential medicines designed for low-resource settings. Although many national- and regional-level EMLs are available online, they are not published in indexed journals, so a detailed search of the grey literature was necessary [19]. Through the use of online search engines and key search terms we were able to identify numerous EMLs on academic, organisational, and government sites, including the 2013 WHO Essential Medicines List, the 2013 Médecins Sans Frontiers Essential Drugs list, and National Medicines Lists from 44 (81%) African countries. There was mention of EMLs for several other African countries, but they were not specific to emergency care and were unable to be obtained online.

The 2013 WHO EML and the 2013 Essential Drugs list from Médecins Sans Frontiers were compared to the retrieved National Medicines lists. We abstracted all medications known to be used in emergency care settings. Drugs used only to treat chronic conditions, such as chemotherapeutics and anti-retroviral medications, were not included. Drugs were also excluded if they were not listed on a majority (> 50%) of identified African EMLs, as this strongly indicated their lack of relevance and/or availability in the African setting.

This initial process generated a candidate list of 250 potential medicines. Drugs were grouped by categories deemed relevant to emergency care in order to facilitate later presentation for consensus gathering. We undertook a year-long, multi-step consensus process to refine the candidate list into a final EML for emergency care in 2015–16 (Fig. 1). The study design did not require our team to seek IRB approval.

## **Online survey**

The candidate list of medicines was used to generate a survey using an online tool (SurveyMonkey\*). In order to capture opinions from the breadth of emergency care systems across Africa, a wide variety of healthcare providers who deliver emergency care in a range of contexts across the continent were sampled via convenience. Survey participants were identified through AFEM's membership pool: members who had been practising emergency care (whether as a specialist or any other level of healthcare provider) for at least one year at any type of facility were targeted. Of these, 109 had current contact details on file and were able to be reached via email to request participation.

The online survey contained 250 total medications, divided into 33 categories. Due to the large francophone population in Africa, it was made available to participants in both English and French. Participants were asked to rank each medicine as "essential", "desirable", "not necessary", or "not applicable" [20,21]. Medicines were also ranked for three levels of facilities (basic, intermediate, advanced) to allow the EML to build upon the previously published African Federation for Emergency Medicine's (AFEM) sub-Saharan African emergency care facility designations [21]. This approach is synergistic with other initiatives for African emergency care development that base recommendations on facility capacity [22].

Respondents were asked to propose other medications and provide commentary. Medication routes were not specified at the time of the survey; however, respondents were informed about the facility level capacity of drug administration (e.g., basic level facilities can



Fig. 1. Multi-step emergency care EML refinement process.

administer PO and IM medications, but not IV).

The survey was completed by 83 (76%) providers. The participant group comprised emergency medicine specialists (43%) and registrars (18%), non-emergency medicine specialist physicians (10%), emergency nurses (11%), and midlevel providers (8%) with collective

experience in 25 countries (22 of which are in Africa). When the data were analysed, it was discovered that the number of respondents completing each section decreased over the course of the survey: in the initial sections, 100% of respondents answered the questions, but by the last section only 64% of respondents replied.

#### Table 1

Essential emergency medicines for basic facilities. Resources available in one category are not repeated in another. The majority of essential medications for trauma presentations are included in categories respiratory failure, shock, altered mental status and severe pain. For a complete listing of drugs by categories, please see Appendix 2.

RESPIRATORY FAILURE	SHOCK	ALTERED MENTAL STATUS	SEVERE PAIN
Ipratropium bromide	Adrenaline	Diazepam	Acetylsalicylic acid
Oxygen	Oral rehydration salts	Oral glucose <sup>*</sup>	Diclofenac
Prednisolone			Ibuprofen
Salbutamol			Lidocaine (local anaesthesia)
			Paracetamol
ANTIDOTES & VACCINATIONS	ANTIMICROBIALS	OTHER	OPHTHALMOLOGIC DROPS/OINTMENT
Rabies vaccine	Albendazole/Mebendazole	Chlorhexidine	Aciclovir
Tetanus vaccine	Amoxicillin	Diphenhydramine	Erythromycin
	Amoxicillin + clavulanic acid	Metformin	Ofloxacin
	Azithromycin	Metoclopramide	Tetracaine
	Benzathine benzylpenicillin	Povidone iodine	Tetracycline
	Ceftriaxone	Zinc sulfate	TOPICAL
	Ciprofloxacin		Hydrocortisone
	Clindamycin		Miconazole
	Doxycycline		Silver sulfadiazine
	Metronidazole		
	Nystatin		
	Praziquantel		
	Regionally recommended oral antimalarial		
	Sulfamethoxazole + trimethoprim		

According to the African Federation for Emergency Medicine's emergency facility designation, a basic level emergency care facility minimally offers oral and IM medications. This differs from standard Basic Emergency Obstetric and Newborn Care capacity facilities, which are considered to be able to give IV medications. Thus, no medicines are recommended in the category of "Maternal Health" at basic facilities [21,25].

\* Not listed on 2017 WHO Model List of Essential Medicines. Recommendation strength: desirable; essential.

In order to reduce the potential for weak results for medications near the end of the survey secondary to respondent attrition, we undertook a second expert review prior to the formal consensus meeting. Due to this attrition, no consensus level was defined to exclude medications, and all were considered in the expert review.

# Expert review

The expert review included 307 medications: 2 of the 250 original medications were excluded due to duplication and 59 additional medications were suggested. Survey data were analysed to generate descriptive statistics about responses. For simplicity, the medications from the online survey were regrouped from the initial 33 categories into 11 clinical presentation-based categories. For further context, the survey results were presented along with information on whether or not the medications were currently included on the WHO EML and the newly-released sidHARTe program Emergency Services Resource Assessment Tool (unpublished).

The survey results were presented for online review by expert physicians practicing emergency care in Africa. Recommendations were made for type of medication, location for inclusion (basic, intermediate and advanced) and whether the medication should be essential or desirable. The expert review group was comprised of a smaller cohort of participants who had not participated in the online survey. Fifteen physicians were included: three practicing in each AFEM region (North, South, East, and West) and three international subject matter experts. To be included, physicians were required to be qualified emergency medicine specialists, with experience practising directly in emergency units. Potential experts were identified via AFEM's membership pool, and contacted via email to request participation. The first three respondents to confirm availability to complete the full review process from each region were included.

A threshold of 70% agreement was predetermined for inclusion and the expert review recommended a total of 225 medications: 15 essential and 37 desirable medications at the basic level, 76 essential and 115 desirable medications at the intermediate level, and 154 essential and 71 desirable medications at the advanced level. These results were combined with the feedback from the online survey and used to create a further-refined emergency care EML to be presented at the consensus meeting.

# **Consensus** meeting

Of the 307 medicines considered in the expert review, 225 reached consensus and were used as the basis on which to draft a tiered emergency care EML for basic, intermediate, and advanced facility levels. A medication was proposed for a given facility level if it received greater than 70% agreement in the expert review, either as essential or desirable.

A consensus meeting was convened at the 2016 African Conference on Emergency Medicine in Cairo, Egypt to review and provide further refinement of these preliminary EMLs. The consensus panel was composed of 21 clinicians currently practicing emergency care in Africa. These clinicians were selected based on agreement and availability to attend the meeting, longitudinal involvement in emergency care systems development and capacity building, and expertise in the African context. Participants could not have participated in previous reviews of the list. Again, participants were identified via AFEM's membership and contacted via email to request participation.

At the meeting, the draft list for each facility level was presented, and each medication was discussed in depth. Simple majority consensus opinion (> 50% agreement) was sought. Participants also had the opportunity to make comments on paper copies of the lists; these comments were collated for consideration in the final EMLs.

The final emergency care EML contained 213 medicines, grouped into 11 categories reflective of potential emergency presentations; these categories included the six AFEM sentinel conditions (difficulty in breathing, shock, altered mental status, severe pain, trauma, and maternal health) as well as five other categories identified as critical to emergency care. The final EML was further subdivided by facility level (Tables 1–3) to align with other emergency care development tools used in the region [21]. Of the original 250 medications derived from the literature review and proposed in the online survey, 36 medications were excluded as they did not reach consensus (Appendix 1). Most

#### Table 2

Essential emergency medicines for intermediate facilities. Resources available in one category are not repeated in another. The majority of essential medications for trauma presentations are included in categories respiratory failure, shock, altered mental status and severe pain. For a complete listing of drugs by categories, please see Appendix 2.

RESPIRATORY FAILURE	SHOCK	ALTERED MENTAL STATUS	SEVERE PAIN	MATERNAL HEALTH
Adrenaline Dexamethasone Hydrocortisone Ipratropium bromide Oxygen Prednisolone Propofol Salbutamol Suxamethonium Vecuronium	Fresh frozen plasma Oral rehydration salts Phytomenadione (vitamin K) Platelet concentrates Red blood cells (packed) Sodium chloride 0.9% Sodium lactate (Ringer's lactate) Tranexamic acid Whole blood	Chlorpromazine Clozapine Diazepam Glucagon Glucose (10%, 50%) Haloperidol Insulin injection Intermediate-acting insulin Lorazepam Mannitol Midazolam Oral glucose Phenobarbital Phenotoin Thiamine	Bupivacaine Codeine Diclofenac Droperidol Fentanyl Ibuprofen Ketamine Lidocaine (local anaesthesia) Lidocaine + epinephrine Morphine Paracetamol Pethidine Tramadol	<b>Ergometrine</b> <b>Magnesium sulfate</b> Mifepristone Misoprostol <b>Oxytocin</b> Prostaglandin E
ANTIDOTES & VACCINATIONS	ANTIMICROBIALS	Valproic acid CARDIAC	OTHER	OPHTHALMOLOGIC DROPS/ OINTMENT
Acetylcysteine <sup>1</sup>	Aciclovir	Acetylsalicylic acid	5% glucose, 0.45% sodium chloride	Aciclovir
Activated charcoal	Albendazole/Mebendazole	Adenosine <sup>*</sup>	5% glucose, 0.9% normal saline	Atropine
Anti-rabies immunoglobulin	Amoxicillin	Amiodarone	Acetazolamide	Chloramphenicol*
Anti-tetanus immunoglobulin	Amoxicillin + clavulanic acid	Amlodipine	Aluminium hydroxide*	Erythromycin
Anti-venom immunoglobulin	Amphotericin B	Bisoprolol	Chlorhexidine	Fluorescein
Anti-D immunoglobulin	Ampicillin	Captopril <sup>*</sup>	Diphenhydramine <sup>*</sup>	Gentamicin
Atropine	Artemether	Digoxin	Iodine	Ofloxacin
Calcium gluconate	Artesunate	Diltiazem	Levothyroxine	Prednisolone
Diphtheria antitoxin	Azithromycin	Dopamine	Loratadine	Tetracaine
Fomepizole <sup>2</sup> Methylthionium chloride (methylene blue) <sup>4°</sup>	Benzathine benzylpenicillin Benzylpenicillin	Enalapril Enoxaparin	Magnesium hydroxide <sup>®</sup> Metformin	Tetracycline Timolol
Naloxone	Cefazolin	Furosemide	Metoclopramide	Tropicamide
Pyridoxine	Ceftriaxone/Cefotaxime	Glyceryl trinitrate	Omeprazole	TOPICAL
Rabies vaccine	Chloramphenicol	Heparin sodium	Ondansetron	Betamethasone
Sodium nitrite + sodium thiosulfate <sup>3</sup>	Ciprofloxacin	Hydralazine	Potassium iodide	Clotrimazole
Tetanus vaccine	Clarithromycin	Hydrochlorothiazide	Promethazine	Ethanol
	Cloxacillin/Dicloxacillin	Isosorbide dinitrate	Propythiouracil	Hydrocortisone
	Diethylcarbamazine	Labetalol	Pyridoxine	Miconazole
	Doxycycline Fluconazole	Lidocaine Methyldopa	<b>Ranitidine</b> Sodium chloride 0.45% <sup>*</sup>	Mupirocin Potassium permanganate (solution)
	Ivermectin	Potassium chloride (powder)	Zinc sulfate	Povidone iodine
	Metronidazole	Potassium chloride (solution)		Selenium sulfide
	Nystatin	Sodium nitroprusside		Silver sulfadiazine
	Praziquantel	Spironolactone		Terbinafine
	Quinine	Verapamil		
	Regionally recommended oral antimalarial			
	Sulfamethoxazole + trimethoprim			

\* Not listed on 2017 WHO Model List of Essential Medicines Recommendation strength: desirable; essential.

<sup>1</sup> Treatment for paracetamol overdose.

<sup>2</sup> Treatment for methanol or ethylene glycol poisoning.

<sup>3</sup> Treatment for cyanide poisoning.

<sup>4</sup> Treatment for cyanide poisoning and methemoglobinemia.

medicines recommended in the final EML (88%) are included in the current WHO EML. The expert panel acknowledged the robust internationally accepted guidance on regionally appropriate oral anti-malarials and thus deferred specific guidance on this class.

## Limitations of process

Using an online literature review strategy to find emergency care

EMLs will only identify those emergency care EMLs that have been published online. As emergency care is a rapidly developing field, there may be existing emergency care EMLs for low-resource settings that have not yet been published online, and therefore would have been missed by our literature review.

Although we attempted to reach a broad range of emergency care providers across Africa, there was limited input from those in Francophone nations. The online survey was offered in both English

# Table 3

Essential emergency medicines for advanced facilities. Resources available in one category are not repeated in another. The majority of essential medications for trauma presentations are included in categories respiratory failure, shock, altered mental status and severe pain. For a complete listing of drugs by categories, please see Appendix 2.

RESPIRATORY FAILURE	SHOCK	ALTERED MENTAL STATUS	SEVERE PAIN	MATERNAL HEALTH
drenaline	Desmopressin	3% Sodium chloride*	Bupivacaine	Ergometrine
Atracurium	Dobutamine <sup>*</sup>	Chlorpromazine	Codeine	Magnesium sulfate
Dexamethasone	Factor IX concentrate	Clozapine	Diclofenac	Mifepristone
tomidate	Fludrocortisone	Diazepam	Droperidol	Misoprostol
Iydrocortisone	Fresh frozen plasma	Ethosuximide	Fentanyl	Oxytocin
pratropium bromide	Oral rehydration salts	Fluphenazine	Ibuprofen	Prostaglandin E
<b>Methylprednisolone</b>	Phytomenadione (vitamin K)	Glucagon	Ketamine	Surfactant
Dxygen	Platelet concentrates	Glucose (5%, 10%, 50%)	Lidocaine + epinephrine	
Prednisolone	Red blood cells (packed)	Haloperidol	Lidocaine (local anaesthesia)	
Propofol	Sodium chloride 0.9%	Insulin injection	Morphine	
albutamol	Sodium lactate (Ringer's lactate)	Intermediate-acting	Nitrous oxide	
Suxamethonium	Tranexamic acid	<b>insulin</b> Levetiracetam <sup>®</sup>	Paracetamol	
"hiopental <sup>*</sup>	Whole blood		Pethidine <sup>*</sup>	
/ecuronium	Whole blood	Lorazepam Mannitol	Tramadol <sup>*</sup>	
ecuromum		Midazolam	ITallauoi	
		Oral glucose		
		Phenobarbital		
		Phenytoin		
		Thiamine Valencia poid		
		Valproic acid		
ANTIDOTES &	ANTIMICROBIALS	CARDIAC	OTHER	OPHTHALMOLOGIC DROP
VACCINATIONS				OINTMENT
Acetylcysteine <sup>1</sup>	Aciclovir	Acetylsalicylic acid	5% glucose, 0.45% sodium chloride	Aciclovir
Activated charcoal	Albendazole/Mebendazole	Adenosine	5% glucose, 0.9% normal saline	Atropine
Anti-D immunoglobulin	Amoxicillin	Amiloride	Acetazolamide	Azithromycin
Anti-rabies immunoglobulin	Amoxicillin + clavulanic acid	Amiodarone	Albumin <sup>*</sup>	Chloramphenicol <sup>*</sup>
Anti-tetanus	Amphotericin B	Amlodipine	Aluminium hydroxide*	Erythromycin
immunoglobulin Anti-venom	Ampicillin	Bisoprolol	Diphenhydramine	Fluorescein
immunoglobulin	-			
Atropine	Artemether	Captopril	Iodine	Gentamicin
Calcium chloride <sup>2*</sup>	Artesunate	Clopidogrel	Levothyroxine	Hydrocortisone
Calcium gluconate	Azithromycin	Digoxin	Loratadine	Latanoprost
Dantrolene	Benzylpenicillin	Diltiazem	Magnesium hydroxide	Ofloxacin
Deferoxamine <sup>3</sup>	Benzathine benzylpenicillin	Dopamine	Metformin	Pilocarpine
Dimercaprol <sup>4</sup>	Cefalexin	Enalapril	Metoclopramide	Prednisolone
Diphtheria antitoxin	Cefazolin	Enoxaparin	Omeprazole	Tetracaine
<sup>5</sup> omepizole <sup>5</sup>	Ceftriaxone/Cefotaxime/Cefixime/ Ceftazidime	Furosemide	Ondansetron	Tetracycline
Iydroxocobalamin <sup>6</sup>	Chloramphenicol	Glyceryl trinitrate	Potassium iodide	Timolol
ntralipid <sup>®</sup>	Ciprofloxacin	Heparin sodium	Promethazine	Tropicamide
Aethylthionium chloride	Clarithromycin	Hydralazine	Propythiouracil	Topicaliliae
(methylene blue) <sup>9*</sup>			Topythiouruch	
laloxone	Clindamycin	Hydrochlorothiazide	Ranitidine	TOPICAL
ralidoxine <sup>7*</sup>	Cloxacillin/Dicloxacillin	Isosorbide dinitrate	Sodium chloride 0.45% <sup>*</sup>	Betamethasone
yridoxine	Diethylcarbamazine	Labetalol	Sulfasalazine	Chlorhexidine
abies vaccine	Doxycycline	Lidocaine	Zinc sulfate	Clotrimazole
odium calcium edetate <sup>8</sup>	Fluconazole	Losartan		Ethanol
odium nitrite + sodium thiosulfate <sup>6</sup>	Flucytosine	Methyldopa		Hydrocortisone
uccimer <sup>8</sup>	Gentamicin	Nifedipine		Miconazole
etanus vaccine	Ivermectin	Potassium chloride		Mupirocin
		(powder)		
	Levofloxacin	Potassium chloride		Potassium permanganate
		(solution)		(solution)
	Metronidazole	Procainamide		Povidone iodine
	Nitrofurantoin	Sodium nitroprusside		Selenium sulfide
	Nystatin	Spironolactone		Silver sulfadiazine
	Pentamidine	Streptokinase		Terbinafine
	Praziguantel	Verapamil		
	Quinine	Tenecteplase (tPA,		
	Periopally recommended and	metalyse) <sup>10*</sup>		
	Regionally recommended oral antimalarial			
	Sulfamethoxazole + trimethoprim			
	Sunametrioxazore + trimetrioprim			(continued on next p

#### Table 3 (continued)

RESPIRATORY FAILURE	SHOCK	ALTERED MENTAL STATUS	SEVERE PAIN	MATERNAL HEALTH
	Vancomycin			

\* Not listed on 2017 WHO Model List of Essential Medicines Recommendation strength: desirable; essential.

<sup>1</sup> Treatment for paracetamol overdose.

 $^{2}\,$  Treatment for hypocalcaemia and magnesium sulfate overdose.

<sup>3</sup> Treatment for iron toxicity.

<sup>4</sup> Treatment for arsenic, mercury, lead poisoning.

<sup>5</sup> Treatment for methanol or ethylene glycol poisoning.

<sup>6</sup> Treatment for cyanide poisoning.

<sup>7</sup> Treatment for organophosphate poisoning.

<sup>8</sup> Treatment for lead poisoning.

<sup>9</sup> Treatment for cyanide poisoning and methemoglobinemia.

<sup>10</sup> Treatment for massive pulmonary embolism.

and French, but the expert review and consensus meeting were conducted only in English. In future iterations of the emergency care EML we will work with translators to ensure better engagement of Frenchspeaking emergency care representatives.

Pharmacists bring diverse and indispensable expertise to EMLs. Our process is limited by the fact that we did not engage enough pharmacists throughout the study. There are few, if any, pharmacists in Africa that specialise in emergency care. As such, we could not identify a robust pool of pharmacists that could be considered experts in this field. The consultation of a pharmacist with significant experience providing emergency care in low-resource settings was sought later in the consensus process. Future iterations of the list should include perspectives from a larger group of pharmacists.

A potential limitation of any consensus group discussion is that not all views may end up being included, as some participants conform with the majority opinion. This may have occurred in the consensus meeting; however, we tried to avoid this by allowing participants to also make written comments for later consideration.

# Discussion

The process to generate this emergency care EML was iterative, engaging a diverse group of experienced clinicians practicing at all levels in a range of African settings. In some categories, individuals practicing in tertiary care facilities recommended more specialised drugs (e.g. imipenem), but the multi-stage consensus process appropriately excluded these resource-intensive, likely unrealistic medicines from the EML.

This is the first non-country-specific emergency care EML for lowresource settings. The EML addresses two challenges to implementing emergency care: it identifies context-specific medicines essential to emergency care, and it identifies the appropriate locations for distribution of those medications based on fundamental emergency capacities of facilities [21].

In order to ensure that emergency care EMLs are appropriate for a range of facility levels within countries and regions, we divided the comprehensive list first by facility level and then by strength of recommendation. The three facility levels were defined according to minimum expected emergency capacity via consensus across African emergency care experts in 2013 [21]. Medication recommendations for each level were matched to expected performance of emergency service delivery.

Certain medications are notably absent from the emergency care EML. These were excluded specifically because of lack of data concerning their efficacy in resource limited settings (e.g., tissue plasminogen activator for cerebral vascular accident [23]) or lack of evidence to show significant improvement in mortality, morbidity, or patientcentred outcomes in the first 24 h of care (e.g., anti-tuberculosis medications). Instead, experts recommended additional medications deemed essential for treating emergent conditions, yet were absent from the WHO EML. Some examples include etomidate, a hemodynamically neutral induction agent for rapid sequence intubation; labetalol and diltiazem, both used for immediate rate control in supraventricular tachycardias and for blood pressure reduction; dobutamine for cardiogenic shock, and diphenhydramine for allergic reactions.

Medications were stratified as being either essential or desirable; this terminology is modelled off of prior WHO checklists [24]. Essential and desirable were used in place of WHO's 'core' and 'complementary' recommendation levels, respectively, because these terms have previously been used to define capacity in the African setting [21]. *Essential* medicines should be available at all facilities of a given level at all times, while the availability and use of *desirable* medicines might vary based on availability of clinical expertise and resource constraints at a facility. Of note, processes should be in place at health care facilities to ensure timely access by health care providers to essential medications; for example, drugs required for emergent airway management should not exclusively be stocked in the operating theatre, far from the emergency unit.

Prior to completing this process, it was anecdotally held that most medicines sufficient to treat emergencies in Africa were contained in the WHO EML. In fact, 88% of medicines included in the final emergency care EML *are* listed on the WHO EML. Since national EMLs are typically based off of the WHO list, this implies that most emergency medicines would also already be included at the national level, and that dosage and administration route guidance provided in the WHO EML will be well known. However, of the 213 medicines ultimately recommended for inclusion, 25 were not included on the 2017 WHO EML. This highlights a potential gap, and suggests that the WHO EML is missing medications important for emergency care in Africa. It is our hope that these resource-stratified lists will allow for easier implementation in countries, while maintaining the rigorous recommendations necessary to improve availability of essential emergency medications.

#### Authors' contributions

MCB, LAW, and EJCH conceived and designed the study. All authors contributed substantially to the EML refinement process. MCB, JLP, LAW, and EJCH drafted the manuscript. All the authors contributed to the manuscript's revision.

# Acknowledgements

The authors and the AFEM Scientific Committee would like to thank the 83 participants of the online survey. We would also like to thank the 15 physicians who completed expert review:

AFEM East: S Kassamali, S Kilindimo, A Alenyo Ngabirano. AFEM South: M Botha, K Diango, K Sikuvi. AFEM West: J Jeong, E Ami Momade, C Kwaku Oppong, AFEM North: H Benmasaud, S Helmy, H Zaki.

International: H Harrison, A Levine, C Taubman.

*Consensus panel:* J Becker, M Bizanso, C Chang, K Diango, J Dixon, S Dunlop, A Hankin-Wei, S Helmy, S Kassamali, J Lin, G Mbanjumucyo, J Mfinanga, N Mould-Millman, M Mutendi Muller, H Sawe, H Zaki.

The following countries (number of participants) were represented throughout the process- some participated in more than one phase: Botswana (3), Cameroon (2), DR Congo (2), Egypt (3), Ethiopia (2), Ghana (6), Kenya (1), Libya (1), Madagascar (5), Malawi (1), Namibia (2), Netherlands (1), Nigeria (2), Rwanda (5), Saudi Arabia (1), Sierra Leone (1), Somalia (3), South Africa (15), Sudan (6), Tanzania (11), U.A.E (1), Uganda (2), U.K. (1), U.S.A. (4), Zambia (2).

The authors are grateful for the pharmacological expertise of Dr Lance Ray and the extensive contributions of Dr Teri Reynolds.

### **Conflicts of interest**

The authors declare no conflict of interest.

#### Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at https://doi.org/10.1016/j.afjem.2018.05.002.

#### References

- Executive Summary: The Selection and Use of Essential Medicines Geneva; 2017.
   Equitable access to essential medicines a framework for collective action. WHO
- Policy Perspectives on Medicines Geneva; 2004.
  [3] Bansal D, Purohit VK. Accessibility and the use of essential medicines in health care: Current progress and challenges in India. J Pharmacol Pharmacother 2013;4(1):13-8. http://dx.doi.org/10.4103/0976-500X.107642.
- [4] The Selection of Essential Medicines WHO Policy Perspective on Medicines Geneva; 2002
- Gulland A. WHO targets antimicrobial resistance in new essential medicines list. BMJ 2017;357:j2809. http://dx.doi.org/10.1136/bmj.j2809.
- [6] Laing R, Waning B, Gray A, et al. 25 years of the WHO essential medicines lists: progress and challenges. Lancet 2003;361(9370):1723–9. http://dx.doi.org/10. 1016/S0140-6736(03)13375-2.

- [7] Understanding the Role and Use of Essential Medicines Lists; 2015.[8] UNHCR's Essential Medicines and Medical Supplies: Policy and Guidance 2011.
- Geneva; 2011. [9] Pilon S. Essential Drugs: 2016 Edition Medicins Sans Frontieres; 2016.
- [10] Reynolds TA, Mfinanga JA, Sawe HR, et al. Emergency care capacity in Africa: a clinical and educational initiative in Tanzania. J Public Health Policy 2012;33(Suppl 1):S126–37. http://dx.doi.org/10.1057/jphp.2012.41.
- [11] Thind A, Hsia R, Mabweijano JR, et al. Essential Surgery. Disease Control Priorities in Developing Countries. 3 ed; 2015:246.
- [12] Wang HNM, Allen C, Barber R, Bhutta Z, Carter A, Casey D, et al. Global, regional, and national life expectancy, all-cause mortality, and cause-specific mortality for 249 causes of death, 1980–2015: a systematic analysis for the Global Burden of Disease Study 2015. Lancet 2016;388(10053):1459–544. http://dx.doi.org/10. 1016/S0140-6736(16)31012-1.
- [13] Hamedani AG, Wears RL. Anyone, anything, anytime...All the time. Ann Emerg Med 2009;53(6):724–6. http://dx.doi.org/10.1016/j.annemergmed.2008.12.002.
- [14] Nowacki AK, Landes M, Azazh A, et al. A review of published literature on emergency medicine training programs in low- and middle-income countries. Int J Emerg Med 2013;6(1):26. http://dx.doi.org/10.1186/1865-1380-6-26.
- [15] Tumuslime P, Gonani A, Walker O, et al. Health systems in sub-Saharan Africa: What is their status and role in meeting the health Millennium Development Goals? Afr Health Monitor 2012;14.
- [16] Leive A, Xu K. Coping with out-of-pocket health payments: empirical evidence from 15 African countries. Bull World Health Org 2008;86(11):849–56.
- [17] Hutchings J, Neroutsos K, Donnelly K. Making the list: the role of essential medicines lists in reproductive health. Int Perspect Sex Reprod Health 2010;36(4):205–8. http://dx.doi.org/10.1363/3620510.
- [18] Emergency Medicines List 1ed. Addis Ababa; 2014.
- [19] Saleh AA, Ratajeski MA, Bertolet M. Grey literature searching for health sciences systematic reviews: a prospective study of time spent and resources utilized. Evid Based Libr Inf Pract 2014;9(3):29–50.
- [20] Germa F, Bayleyegn T, Kebede T, et al. Emergency medicine development in Ethiopia: Challenges, progress and possibilities. Afr J Emerg Med 2013;3:3–9.
- [21] Calvello EJ, Tenner AG, Broccoli MC, et al. Operationalising emergency care delivery in sub-Saharan Africa: consensus-based recommendations for healthcare facilities. Emerg Med J 2016;33(8):573–80. http://dx.doi.org/10.1136/emermed-2015-204994.
- [22] Wallis LA, Reynolds TR. AFEM handbook of acute and emergency care. Oxford University Press; 2014.
- [23] Aaron L, Berkowitz AL, Mittal MK, McLane HC, et al. Worldwide reported use of IV tissue plasminogen activator for acute ischemic stroke. Int J Stroke 2014;9:349–55.
- [24] World Health Organization and International Association for the Surgery of Trauma and Surgical Intensive Care (IATSIC). Essential trauma care project: checklists for surveys of trauma care capabilities. Geneva: World Health Organization; 2004. http://www.who.int/emergencycare/trauma/essential-care/estc\_checklist.pdf?ua = 1.
- [25] Bailey P, Lobis S, Maine D, et al. Monitoring emergency obstetric care: a handbook. Geneva: World Health Organization; 2009.