





		EMLc	Codes ATC: J01CA04
Indication	Bacterial meningitis	Code ICD11: 8E71.0Z	
INN	Amoxicillin		
Type de médicament	Chemical agent		
Groupes d'antibiotiques	 ACCESS		
Type de liste	Liste de base		
Formulations	Oral > Liquid: 125 mg per 5 mL (as trihydrate) powder for oral liquid ; 250 mg per 5 mL (as trihydrate) powder for oral liquid (EMLc) Oral > Solid: 250 mg (as trihydrate) ; 500 mg (as trihydrate) Parenteral > General injections > unspecified: 250 mg in vial (as sodium) powder for injection ; 500 mg in vial (as sodium) powder for injection ; 1 g in vial (as sodium) powder for injection		
Historique des statuts LME	Ajouté pour la première fois en 2017 (TRS 1006)		
Sexe	Tous		
Âge	Aussi recommandé pour les enfants		
Équivalence thérapeutique	La recommandation concerne ce médicament spécifique		
Renseignements sur le brevet	Patents have expired in most jurisdictions Lire la suite sur les brevets. 		
Wikipédia	Amoxicillin 		
DrugBank	Amoxicillin 		

Recommandation du comité d'experts

The Expert Committee endorsed the inclusion on the EML and EMLc of ceftriaxone or cefotaxime as first-choice option for use in suspected acute bacterial meningitis and of chloramphenicol, benzylpenicillin, ampicillin or amoxicillin as second-choice therapy, recognizing that the last three beta-lactams may be added as first-choice options in some countries for suspected acute bacterial meningitis in particular when *Listeria* is suspected. The Committee recommended the addition of meropenem to the EMLc for use in neonates as a second-choice option to treat suspected acute bacterial meningitis where resistant Gram-negative organisms are the common causative agents.

Contexte

Acute bacterial meningitis is a medical emergency requiring prompt administration of antibiotics that penetrate well into inflamed meninges. Because of the severity of this infection, evidence from randomized controlled trials (RCTs) is limited; recommendations for antimicrobials are driven largely by susceptibility patterns of the most common pathogens together with experimental work in animal models.

Résumé des preuves

In a 2015 systematic review, chloramphenicol was compared with two penicillins, two cephalosporins and one tetracycline (5 RCTs; 1753 patients) (1). Chloramphenicol was associated with higher mortality than the other antibiotics (risk ratio (RR) 1.27; 95% confidence interval (CI) 1.00–1.60). In contrast, a 2007 Cochrane review (19 RCTs; 1496 patients) that compared third-generation

cephalosporins with treatment with penicillin or ampicillin-chloramphenicol found no differences in mortality (risk difference (RD) 0%; 95% CI 3% to 2%), risk of deafness (RD -4%; 95% CI -9% to 1%), or risk of treatment failure (RD -1%; 95% CI -4% to 2%) (2). There was a reduced risk of CSF culture positivity after 10–48 hours (RD -6%; 95% CI -11% to 0%) and an increased risk of diarrhoea (RD 8%; 95% CI 3% to 13%) for third-generation cephalosporins compared with penicillin/ampicillin-chloramphenicol (2). A 2009 systematic review compared short-course (4–7 days) and long-course (7–14 days) antibiotics in children (5 RCTs; 426 patients) and found no difference in clinical success (odds ratio (OR) 1.24; 95% CI 0.73–2.11), long-term neurological complications (OR 0.60; 95% CI 0.29–1.27) or long-term hearing impairment (OR 0.59; 95% CI 0.28–1.23) (3).

Recommandations

The National Institute for Health and Care Excellence (NICE) guidelines recommend ceftriaxone for patients aged 3 months and older, while younger infants should be treated with IV cefotaxime plus amoxicillin or ampicillin (4). It is also recommended that vancomycin be added for patients who have received prolonged or multiple exposures to antibiotics within the previous 3 months and for those who have recently travelled outside the United Kingdom. IDSA (Infectious Diseases Society of America) guidelines recommend ampicillin and cefotaxime or an aminoglycoside for children less than 1 month of age, vancomycin and ceftriaxone or cefotaxime for children older than 23 months and adults up to 50 years of age, and addition of ampicillin for patients over 50 years for coverage of *Listeria monocytogenes* (5). Vancomycin plus cefepime, ceftazidime or meropenem is recommended for patients with penetrating trauma, who are post-neurosurgery or have a cerebrospinal shunt in place.

Justification de la sélection des antibiotiques

Systematic review evidence suggests that chloramphenicol is associated with higher mortality than other antibiotics; it was therefore not proposed as a core antibiotic. Ampicillin, ceftriaxone and cefotaxime are proposed for multiple indications and are categorized as core, while aminoglycosides and vancomycin have more specific indications (e.g. by age or indication) and are therefore categorized as targeted, as are ceftazidime and meropenem.

Considérations du comité

For common community-acquired infections, the main focus has been on empirical treatment choices that are broadly applicable in most countries. Generally, alternatives for use in case of allergy were not considered. The Committee considered the various antibiotics proposed in the application under the guiding principle of parsimony and selected first- and second-choice antibiotics for this indication for inclusion on the EML and/or EMLc. Ceftazidime, amikacin, gentamicin and vancomycin were excluded, because the Committee considered that these antibiotics have limited or no indications in community-acquired acute bacterial meningitis. The Committee recommended the inclusion of chloramphenicol as a second-choice option, particularly for epidemic bacterial meningitis. Recommended first- and second-choice antibiotics are reported above. The first-choice antibiotics are those generally recommended on the basis of available evidence and are usually narrow-spectrum agents.

Recommandations de la LME : Bacterial meningitis

Premier choix

cefotaxime

ceftriaxone

Second choix

amoxicillin

ampicillin

benzylpenicillin

chloramphenicol

1. Eliakim-Raz N, Lador A, Leibovici-Weissman Y, Elbaz M, Paul M, Leibovici L. Efficacy and safety of chloramphenicol: joining the review
2. Prasad K, Kumar A, Gupta PK, Singhal T. Third generation cephalosporins versus conventional antibiotics for treating acute bacterial meningitis
3. Karageorgopoulos DE, Valkimadi PE, Kapaskelis A, Rafailidis PI, Falagas ME. Short versus long duration of antibiotic therapy for bacterial meningitis
4. Visintin C, Mugglestone MA, Fields EJ, Jacklin P, Murphy MS, Pollard AJ et al. Management of bacterial meningitis and meningococcal meningitis
5. Tunkel AR, Hartman BJ, Kaplan SL, Kaufman BA, Roos KL, Scheld WM et al. Practice guidelines for the management of bacterial meningitis

