

WHO activities towards harmonization of diagnosis and clinical management of *Echinococcus granulosus* infection/Cystic Echinococcosis

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Background



Review

Expert consensus for the diagnosis and treatment of cystic and alveolar echinococcosis in humans[☆]

Enrico Brunetti ^{a,*},¹ Peter Kern ^b, Dominique Angèle Vuitton ^c, Writing Panel for the WHO-IWGE²

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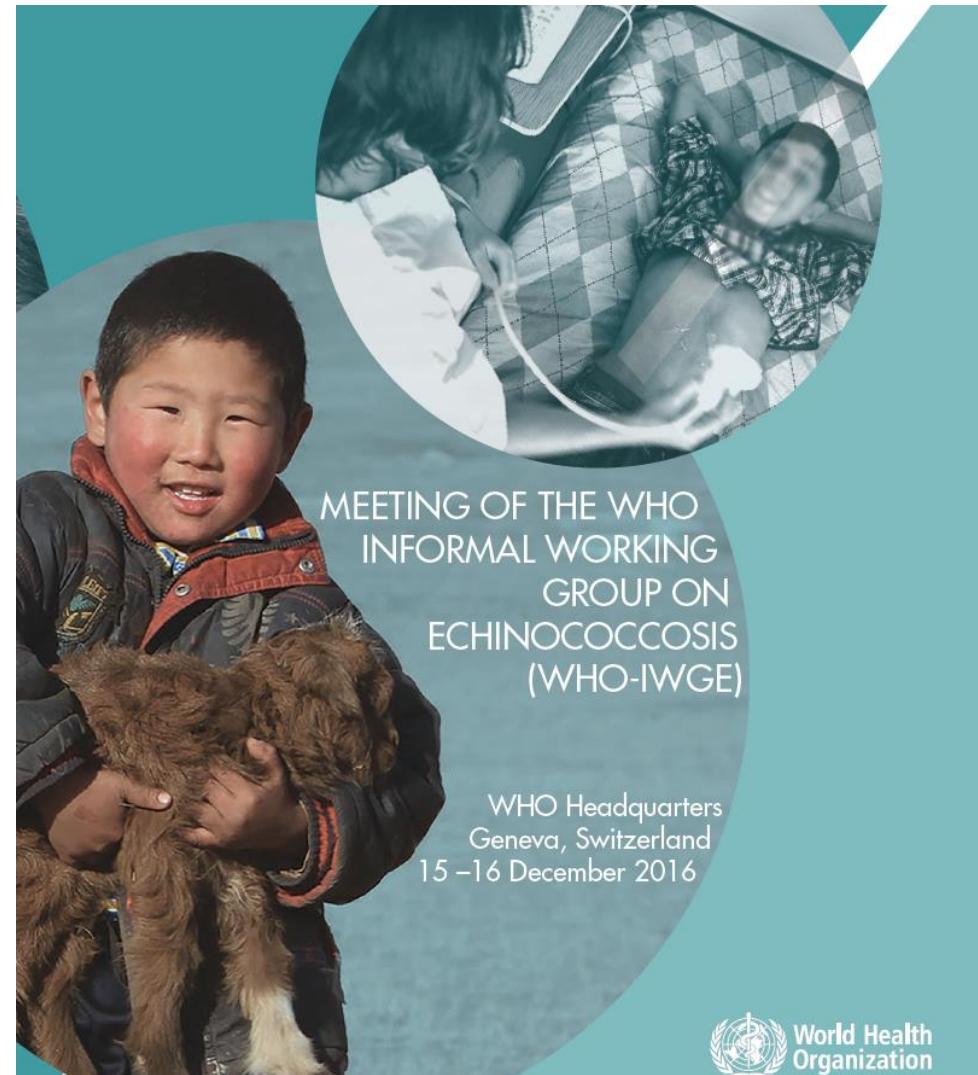
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<https://doi.org/10.1016/j.actatropica.2009.11.001>

A series of meetings of the WHO Informal Working Group on Echinococcosis have taken place:

- Geneva, Switzerland, 15-16 December 2016
- Bern, Switzerland, 9-11 October 2019
- Lima, Peru, 29 November 2019



Guideline Development Group



Health Topics Countries Newsroom Emergencies

Home / Newsroom / Article / Call for public consultation – for experts to join the Guideline Development Group (GDG) on treatment

Call for public consultation – for experts to join the Guideline Development Group (GDG) on treatment of patients with cystic echinococcosis (CE)

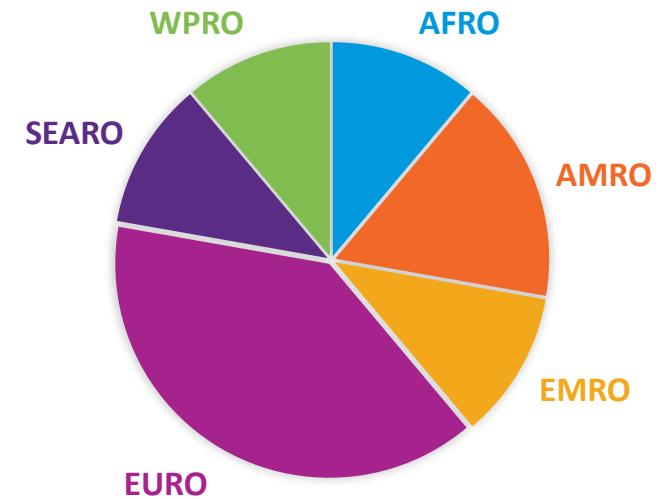
Deadline: 22 March 2022

7 March 2022 | Call for consultation

bit.ly/36JVNMY



17 Experts



Systematic Review Team

Paul Garner, Rebecca Kuehn (Liverpool School of Tropical Medicine, UK)

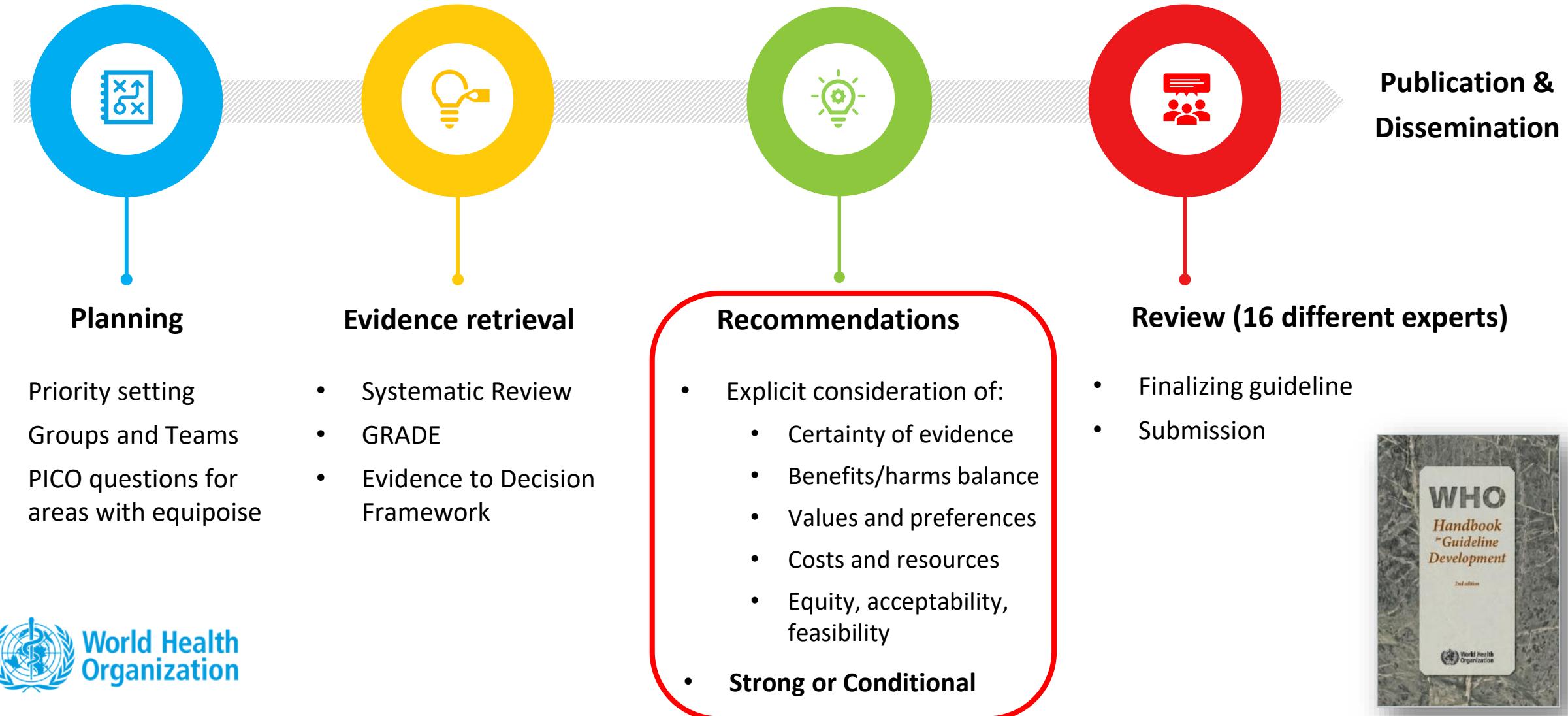
Methodologist

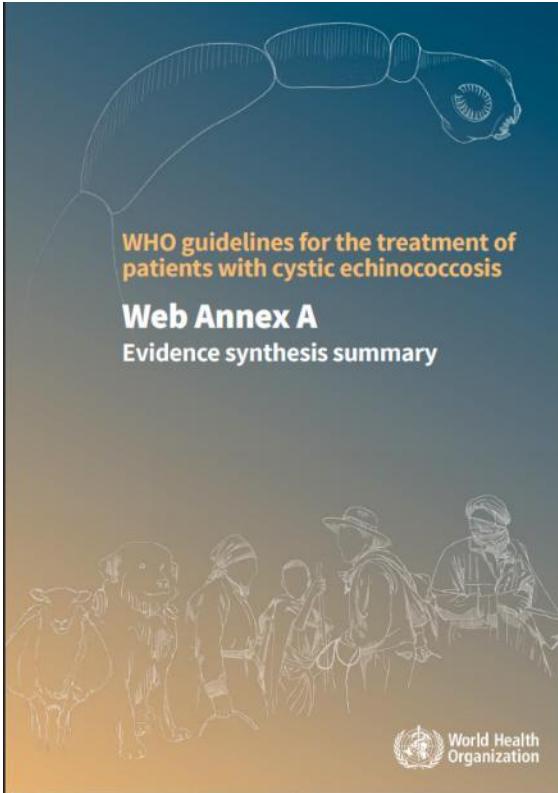
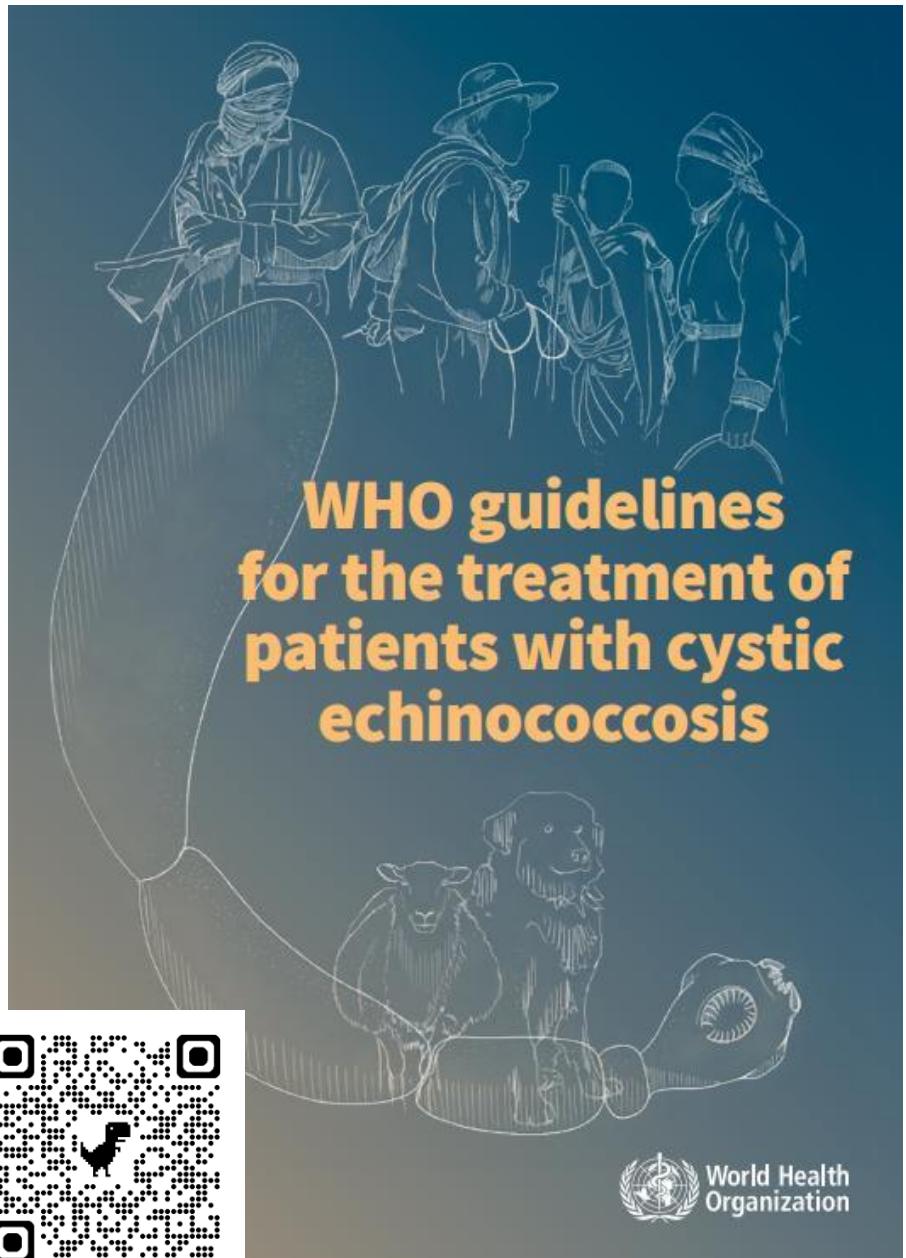
Priscilla Rupali (Christian Medical College, Vellore, India)

Chair

Timothy Pennel (University of Cape Town and Groote Schuur Hospital, South Africa) FOLLOWED BY Peter Chioldini (Hospital for Tropical Diseases and London School of Hygiene & Tropical Medicine, UK)

Guideline Development Overview



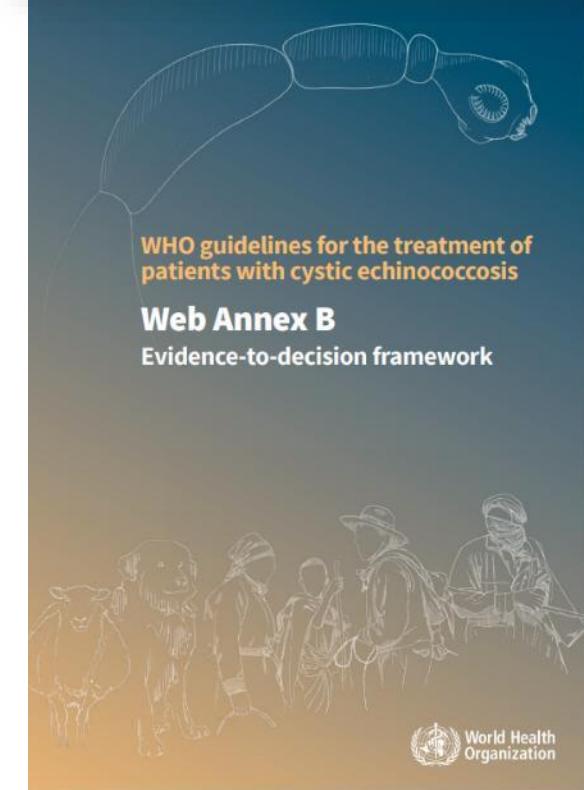


16 June 2025

Cochrane Library
Cochrane Database of Systematic Reviews

Treatment of uncomplicated hepatic cystic echinococcosis (hydatid disease) (Review)

Kuehn R, Uchiumi LJ, Tamarozzi F



Writing Team

Meritxell Donadeu (WHO Consultant, Australia); Bernadette Abela and Katie Corridan (WHO/NTD); Priscilla Rupali, Hanna Alexander and Jisha Sara John (Christian Medical College, Vellore, India); Rebecca Kuehn and Paul Garner (Liverpool School of Tropical Medicine, UK).

Scope of the Guideline -1

- Enable clinicians in their **respective healthcare environments** to manage CE patients **appropriately** and at the **highest standards of care** possible
- Reduce and avoid over- and mistreatment of patients
- Providing recommendations on the indications of the four main management modalities:
 - (1) anti-parasitic drug treatment,
 - (2) percutaneous methods,
 - (3) surgery,
 - (4) “watch & wait”
- depending on the stage and localization of the cysts
- For UNCOMPLICATED LIVER CE and small uncomplicated lung CE
- Where areas of equipoise exist

1.3 Objectives and scope of these guidelines

The purpose of these guidelines is to provide guidance on the choice of treatment so that patients (adults and children) with CE cysts can be offered and receive appropriate and equitable treatment. The aim is to ensure that patients receive the most appropriate and affordable management in the context of infrastructure and expertise sufficient to ensure its safety, and without unnecessary invasive procedures or treatment, to avoid iatrogenic complications by using invasive interventions.

For complicated CE liver cysts, surgery is usually the treatment of choice, based on best medical practice as conveyed by the WHO-IWGE (9). These guidelines are focused on the different choices for uncomplicated liver cysts. Pulmonary CE is usually managed by surgical intervention, but these guidelines evaluate the option of using ALB alone to treat small pulmonary cysts.

For uncomplicated inactive cysts, there are no recommendations in these guidelines since current best medical practice is follow-up with imaging (ultrasonography, CT or MRI), also known as the “watch and wait” approach (9, 14). Surgery should be avoided as far as possible unless the inactive cyst is causing complications (e.g. cyst causing portal hypertension).

Table 2. Health care system tiers for managing different treatment options for CE, according to available expertise and resources

Tier	Health care worker technical expertise required	Surgical infrastructure required	Radiological capacity required	Laboratory required	Intervention(s) possible
Tier 1	Medical doctor	Not available	Referral access to ultrasonography	Access to facilities for complete blood cell count, liver function tests.	Albendazole
Tier 2	General surgeon Anaesthesiologist Nursing care	Operating theatre Inpatient facility with monitoring	Ultrasound on site	Laboratory tests as needed for anaesthesia	Tier 1 and Surgery (non-radical only)
Tier 3 (includes expertise and facilities available in tier 2)	Surgeon with laparoscopic skills and surgeon, radiologist or physician with a relevant speciality trained in PAIR and S-CAT	General surgery and laparoscopic surgery facilities Inpatient facility with monitoring and access to ICU	CT scan Fluoroscopy	Laboratory tests as needed for anaesthesia	Tier 2 and Surgery (radical and non-radical) Laparoscopic Surgery PAIR Standard catheterization
Tier 4 (includes expertise and facilities available in tier 3)	General and laparoscopic surgeons Interventional Radiologists Thoracic Surgeon	Interventional Radiology Facilities and Procedure Room	MRI and MRCP	Routine clinical pathology, biochemistry and microbiology	Tier 3 and Modified Catheterization Technique Thoracic (lung) surgery

Scope of the Guideline -2

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Scope of the Guideline -3

- Enable clinicians in their **respective healthcare environments** to manage CE patients **appropriately** and at the **highest standards of care** possible
- **Reduce and avoid over- and mistreatment of patients**
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- For **UNCOMPLICATED LIVER CE** and small uncomplicated lung CE
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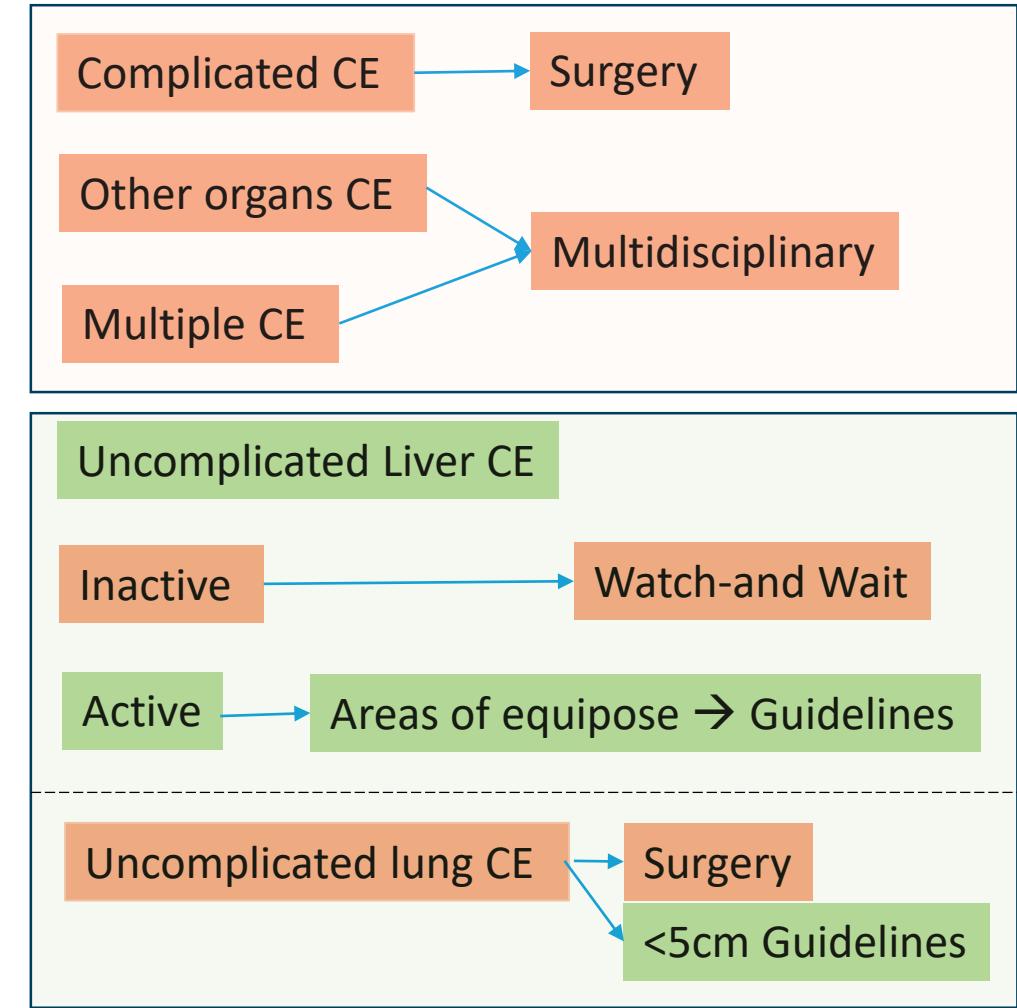
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Scope of the Guideline - 3 cont.

- Enable clinicians in their **respective healthcare environments** to manage CE patients **appropriately** and at the **highest standards of care** possible
- **Reduce and avoid over- and mistreatment of patients**
- Providing recommendations on the indications of the four main management modalities:
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- depending on the stage and localization of the cysts
- For **UNCOMPLICATED LIVER CE** and small uncomplicated lung CE
- **Where areas of equipoise exist**



Uncomplicated inactive cysts are not covered in these guidelines. Current practice is to “watch and wait”, that is, to follow-up with imaging (ultrasonography or MRI). Surgery should be avoided as far as possible unless the inactive cyst is causing complications (e.g. cyst causing portal hypertension).

Scope of the Guideline - 4

- Enable clinicians in their respective healthcare environments to manage CE patients appropriately and at the highest standards of care possible
- Reduce and avoid over- and mistreatment of patients
- Providing recommendations on the indications of the four main management modalities:
 - (1) anti-parasitic drug treatment,
 - (2) percutaneous methods,
 - (3) surgery,
 - (4) “watch & wait”
- depending on the stage and localization of the cysts
- For UNCOMPLICATED LIVER CE and small uncomplicated lung CE
- Where areas of equipoise exist

Ultrasound image	Particulars of pathognomonic sign	Stage	Viability*
Cystic echinococcal cysts		CE1	Viable
		CE2	Viable
		CE3a	Viable or non-viable
		CE3b	Viable
		CE4	Low viability or non-viable
		CE5	Non-viable

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Active cysts, likely to contain viable protoscoleces:	
CE1.	Active, unilocular, liquid content
CE2.	Active, multivesicular, liquid content
CE3a.	Transitional unilocular, liquid content with detached parasitic layers
CE3b.	Active multivesicular cyst, with partially solid content with daughter cysts
Inactive stages:	
CE4.	Solid content
CE5.	Solid content with eggshell calcified wall

1.1.2 Diagnosis

The diagnosis of CE is based on imaging techniques, primarily ultrasound or magnetic resonance imaging (MRI), while computed tomography (CT) is less reliable (6), complemented by serology when imaging is not conclusive. Contrast-enhancement imaging allows excluding CE diagnosis in a case where the cyst takes contrast. No antigen detection tests are commercially available. Antibody detecting serological tests complement imaging findings, yet their limitations warrant careful consideration. In cases of sero-negativity, confirming a presumptive diagnosis might involve demonstrating the presence of protoscoleces and/or hooks by microscopic examination of the cyst fluid, histology, polymerase chain reaction (PCR) of cyst material (7) or observation of changes in the cyst ultrasound appearances on treatment, such as detachment of parasite layers in an unilocular cyst (suspected CE1) after percutaneous puncture or administration of ALB. Currently, there are no WHO guidelines for the diagnostic of CE, and this has been identified as a key research priority (section 6).

6. Research priorities

Research priorities

- 1 Prospective comparative trials to update recommendations.
- 2 Health services research on provision of access to services in endemic areas.
- 3 Survey on patient preferences for the treatment and management of CE.
- 4 For all PICO questions the duration of the ALB regimen needs to be assessed in RCT. In addition, for those where a procedure is recommended, the potential role of combination with praziquantel also needs to be addressed with a proper RCT.
- 5 Include additional questions to these guidelines such as the management of treatment failures after PAIR.
- 6 Develop WHO CE diagnostic guidelines.
- 7 Improved diagnostic tools for specific use cases.

PICOs



Research (PICO) questions

P1 For treating uncomplicated hepatic cyst types CE1 or CE3a < 5 cm, is PAIR combined with ALB as effective and safe as ALB alone?

P2 For treating uncomplicated hepatic cyst types CE1 or CE3a 5–10 cm, is PAIR combined with ALB as effective and safe as ALB alone?

P3 For treating uncomplicated hepatic cyst types CE1 or CE3a 5–10 cm, is surgery combined with ALB as effective and safe as PAIR combined with ALB?

P4 For treating uncomplicated hepatic cyst types CE1 or CE3a > 10 cm, is standard catheterization combined with ALB as effective and safe as PAIR combined with ALB?

P5 For treating uncomplicated hepatic cyst types CE1 or CE3a > 10 cm, is standard catheterization combined with ALB as effective and safe compared to surgery combined with ALB?

P6 For treating uncomplicated hepatic cyst types CE2 or CE3b ≤ 5 cm, is surgery combined with ALB as effective and safe as ALB alone?

P7 For treating uncomplicated hepatic cyst types CE2 or CE3b 5–10 cm, is surgery combined with ALB as effective and safe as ALB alone?

P8 For treating uncomplicated hepatic cyst types CE2 or CE3b of any size, is laparoscopic surgery combined with ALB as effective and safe as open surgery combined with ALB?

P9 For treating uncomplicated hepatic cyst types CE2 or CE3b of any size, is modified catheterization technique (Mo-CAT) combined with ALB as effective and safe as surgery combined with ALB?

P10 Is praziquantel combined with ALB as effective and safe as ALB alone for treating active cysts (cyst types CE1, CE2 or CE3a, CE3b) when given pre- and post- percutaneous or surgical interventions?

P11 For treating uncomplicated lung CE cysts of ≤ 5 cm, is ALB as effective and safe as surgery combined with ALB?

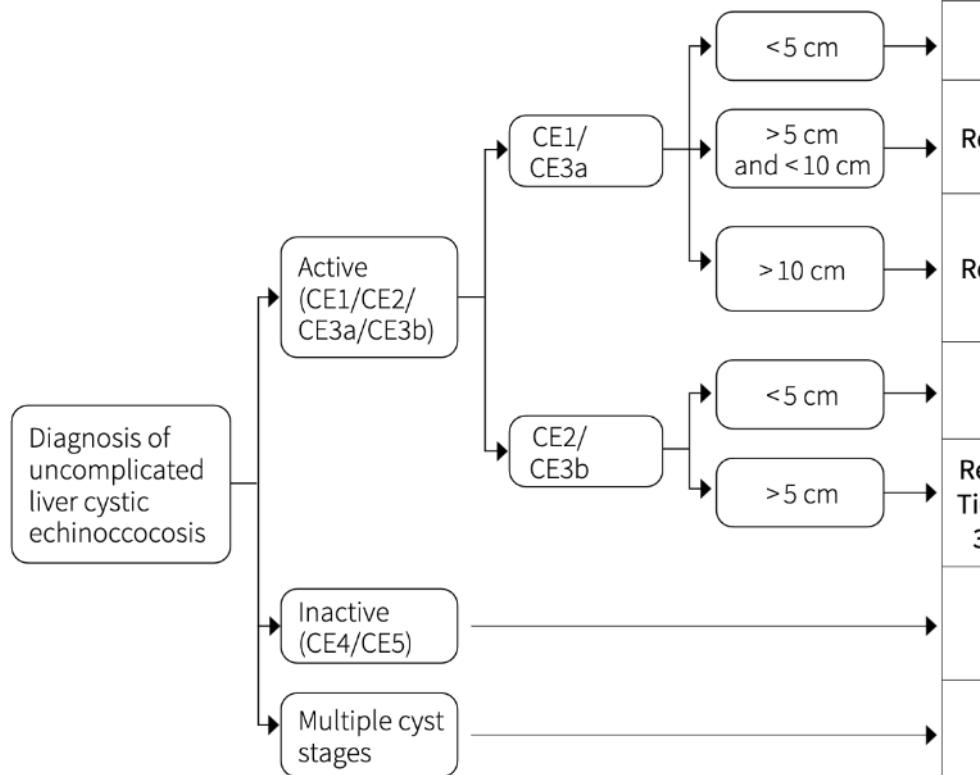
Recommendations

Key:

ALB: Albendazole

PAIR: Puncture, Aspiration, Injection, Re-aspiration

S-CAT: Standard Catheterization



First-line treatment recommendations			
Tier 1	Tier 2	Tier 3	Tier 4
ALB			
		Refer to Tier 3 or 4	PAIR and ALB
		Refer to Tier 3 or 4	Percutaneous treatment and ALB (PAIR preferred over S-CAT)
ALB			
Refer to Tier 2 or 3 or 4		Surgery and ALB (open or laparoscopy)	
Monitoring with imaging «watch and wait»			
Management recommendations individualized			

Uncomplicated lung CE cysts
≤ 5 cm

Recommendation 7:

In patients with uncomplicated active lung CE cysts < 5 cm, surgery is suggested. ALB should not be given before surgery. When spillage is suspected or has occurred, ALB after surgery is suggested. Lung surgery requires tier 4 settings.

Conditional recommendation based on expert consensus.

Use of praziquantel combined with ALB post-percutaneous/ surgical procedures for hepatic cyst types CE1, CE2, CE3a, CE3b

Recommendation 6:

In CE patients undergoing percutaneous or surgical interventions, when spillage is suspected or has occurred, the combination praziquantel and ALB is suggested.

Conditional recommendation based on expert consensus.

Background

ALB is most often used in the treatment of CE, alone or in addition to invasive interventions. Recently, attention has been given to the addition of praziquantel pre- and post-intervention, combined with ALB.

Summary of the evidence

This recommendation is based on PICO question 10. No trials were identified. The GDG formulated the recommendation based on pharmacological data, expert consensus, risk benefit assessment and clinician experience.

Praziquantel has been reported to have a protoscolecidal effect but is not parasitocidal for the cysts (14). Pharmacological data indicate that the combination of praziquantel and ALB enhances efficacy by increasing ALB sulfoxide levels, the pharmacologically active metabolite, resulting in markedly increased protoscolecidal activity, enhancing the efficacy of treatment and reducing the risk of recurrence or complications associated with spillage. Biological plausibility has been reported by Cobo et al. (29).

Certainty of the evidence

There is no evidence available to support the use of praziquantel combined with ALB when performing invasive interventions. The recommendation was formulated by the GDG using expert consensus within the evidence-to-decision framework.

Additional factors considered

- Benefits and harms**
The GDG acknowledges potential benefits, such as enhanced therapeutic activity, broader applicability and risk mitigation, associated with the praziquantel and ALB combination. The concerns raised include limited experience, cost issues and uncertainties regarding specific outcomes. There is a lack of data regarding undesirable effects.
- Health equity, acceptability, resource implications, and feasibility**
The high cost of praziquantel in some countries could potentially create access barriers, highlighting a concern for health equity, acceptability and feasibility. Efforts should be made to make praziquantel more affordable and accessible, especially in low- and middle-income regions where CE is endemic.

Implementation considerations

1. In case of suspected or ascertained cyst fluid spillage, ALB should be given at a dose of 10–15 mg/kg/day in two divided doses (up to 400 mg twice a day) for a minimum of 3 months, usually, 6–12 months after the intervention, as considered appropriate by the clinician.
2. Praziquantel should be given at a dose of 40–50 mg/kg/day divided into two daily doses for 2 weeks after the intervention. Because praziquantel does not have an effect on the cyst (as compared to ALB), 2 weeks are suggested. However, the period can be increased to a maximum of 4 weeks if considered appropriate by the clinician.
3. ALB and praziquantel can be given simultaneously during a fat-rich meal to increase their bioavailability.
4. Some clinicians use praziquantel in combination with ALB for 2 weeks prior to procedure (29). More evidence is needed to make this practice a recommendation.

Target Product Profiles (TPP) for diagnostics for *E. granulosus*/Cystic echinococcosis in animals and humans for PUBLIC HEALTH PURPOSES



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SOON!

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TPP structure

World Health Organization

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Home / Newsroom / Article / Call for nomination of experts – WHO Diagnostic Technical Advisory Group (DTAG) for Neglected Tropical Diseases. One Health sub-group: neglected zoonotic diseases

Call for nomination of experts – WHO Diagnostic Technical Advisory Group (DTAG) for Neglected Tropical Diseases. One Health sub-group: neglected zoonotic diseases

Deadline: 5 April 2024

DTAG One Health subgroup - Members

Core group (12 members)

AFRO: P. Gichuki, R. Miambo
AMRO: R. Wallace, V. Periago, A. Strailey
EMRO: M. Fasih Harandi
EURO: C. Freuling, F. Tamarozzi
SEARO: G. Singh
WPRO: M. Lighthowers, MB. Qian, SH. Kim

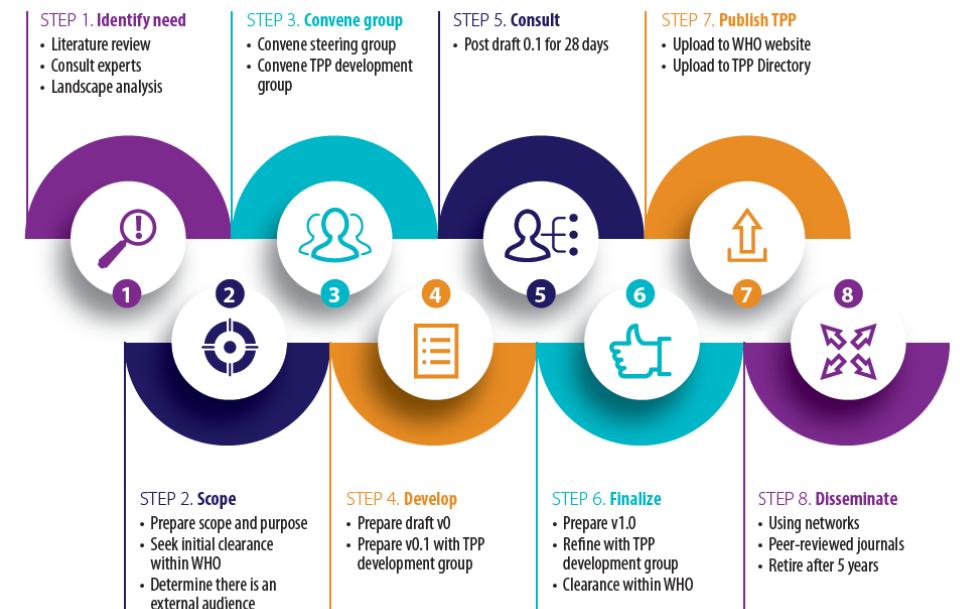
Subject matter expert

Taeniasis/cysticercosis: A. Fleury, S. Gabriel, R. Jambou, V. Khieu, B. Ngowi
Echinococcosis: G. Minbaeva, L. Uchiumi, T. Manciulli
FBT: S. Mas Coma, B. Sripa, ST. Hong, M. Adriko, MD Bargues, V. Khieu
Rabies: A. Latz

World Health Organization

The role of the subgroups is:

1. Understand the current diagnostics landscape and set diagnostic priorities
2. To prepare new or review existing TPPs for **teniasis/cysticercosis, cystic echinococcosis, FBT and rabies**
3. To describe the programmatic use case to scientists and product developers



1- Product use summary	2- Design	3- Performance	4- Product configuration	5- Product cost and channels
1.1- Use case	2.1- Portability	3.1- Species differentiation	4.1- Shipping conditions	5.1- Target pricing per test
1.2- Target population	2.2- Instrument power/requirement	3.2- Diagnostic/ Clinical sensitivity	4.2- Storage conditions	5.2- Capital cost
1.3- Lowest infrastructure level	2.3- Water requirement	3.3- Diagnostic/ Clinical specificity	4.3- Service and support	5.3- Product lead times
1.4- Lowest user level	2.4- Maintenance and calibration	3.4- Time to results	4.4- Labelling and instructions for use	5.4- Target launch countries
1.5- Training requirements	2.5- Sample type/collection	3.5- Result stability		5.5- Product registration
	2.6- Sample preparation – transfer to device	3.6- Throughput		5.6- Procurement
	2.7- Sample volume	3.7- Target shelf life/ stability	3.8- Ease of use	5.7- Test pack size
			3.9- Ease of results interpretation	
			3.10- Operating temperature	
			3.11- Equivalence of matrices*	
			3.12 Reproducibility and robustness	