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Wagstaff, D.^a, Amausi, J.^b, Arfin, S.^{c d e}, Aryal, D.^f, Nor, M.B.M.^g, Bonney, J.^h, Dondorp, A.^{e i j}, Dongelmans, D.^{k l}, Dullawe, L.^m, Fazla, F.^m, Ghose, A.ⁿ, Hanciles, E.^{o p}, Haniffa, R.^{e m q}, Hashmi, M.^r, Smith, A.H.^{s t}, Kumar, B.^u, Minh, Y.L.^v, Moonesinghe, R.^a, Pisani, L.^e, Sendagire, C.^{w x}, Hasan, M.S.^g, Ghalib, M.S.^y, Frimpong, M.S.^z, Ranzani, O.^{aa ab}, Sultan, M.^{ac}, Thomson, D.^{ad}, Tripathy, S.^{q ae}, Thwaites, L.^v, Uddin, R.A.M.E.ⁿ, Mazlan, M.Z.^{af}, Waweru-Siika, W.^{ag}, Beane, A.^{e m q}

Evidence based QUALity Improvement for Prescribing Stewardship in ICU (EQUIPS-ICU): protocol for type III hybrid implementation-effectiveness study

(2025) *Implementation Science*, 20 (1), art. no. 12, . Cited 1 time.

DOI: 10.1186/s13012-024-01413-4

^a Centre for Preoperative Medicine, University College London, London, United Kingdom

^b Department of Global Health, Kwame Nkrumah University of Science and Technology, Kumasi, Ghana

^c The George Institute for Global Health, New Delhi, India

^d Amsterdam Public Health (APH), Amsterdam University Medical Centre(UMC), Amsterdam, Netherlands

^e Mahidol Oxford Tropical Medicine Research Unit, Bangkok, Thailand

^f Department of Critical Care, Nepal Intensive Care Research Foundation, Kathmandu, Nepal

^g Department of Intensive Care Anaesthesiology, International Islamic University Malaysia, Kuala Lumpur, Malaysia

^h Virology Department, Noguchi Memorial Institute for Medical Research, Accra, Ghana

ⁱ Amsterdam Institute for Global Health and Development, Amsterdam, Netherlands

^j Nuffield Department of Medicine, University of Oxford, Oxford, United Kingdom

^k National Intensive Care Evaluation (NICE) Foundation, Amsterdam, Netherlands

^l Department of Intensive Care Medicine, UMC, University of Amsterdam, Amsterdam, Netherlands

^m Nat-Intensive Care Surveillance, Mahidol Oxford Tropical Medicine Research Unit, Colombo, Sri Lanka

ⁿ Department of Medicine, Chittagong Medical College Hospital, Chattogram, Bangladesh

^o Connaught Hospital, Freetown, Sierra Leone

^p Ministry of Health and Sanitation, Freetown, Sierra Leone

^q Pandemic Sciences Hub and Institute for Regeneration and Repair, University of Edinburgh, Edinburgh, United Kingdom

^r Department of Critical Care Medicine, Ziauddin University, Karachi, Pakistan

^s Faculty of Health Sciences, Busitema University, Mbale, Uganda

^t William Harvey Research Institute, Queen Mary University of London, London, United Kingdom

^u Department of Critical Care Medicine, Apollo Hospitals Educational and Research Foundation, Chennai, India

^v Clinical Research Unit, Oxford University, University of Oxford, Ho Chi Minh City, Viet Nam

^w D'Or Institute for Research and Education, Sao Paulo, Brazil

^x Uganda Heart Institute, University of Makerere, Makerere, Uganda

^y General Surgery, Wazir Akbar Khan Hospital, Kabul, Afghanistan

^z Department of Anaesthesiology and Intensive Care, Komfo Anokye Teaching Hospital, Kumasi, Ghana

^{aa} Barcelona Institute for Global Health, ISGlobal, Barcelona, Spain

^{ab} Pulmonary Division, Heart Institute, Faculty of Medicine, Hospital das Clínicas da Faculdade de Medicina da Universidade de São Paulo, São Paulo, Brazil

^{ac} Department of Emergency Medicine and Critical Care, St. Paul's Hospital Millennium Medical College, Addis Ababa, Ethiopia

^{ad} Department of Anaesthesia and Perioperative Medicine, University of Cape Town, Cape Town, South Africa

^{ae} All India Institute of Medical Sciences, New Delhi, India

^{af} Hospital Universiti Sains Malaysia, Kota Bharu, Malaysia

^{ag} Department of Anaesthesia, The Aga Khan University, Nairobi, Kenya

Abstract

Background: Approximately half of all antimicrobial prescriptions in intensive care units (ICUs) may be inappropriate, including those prescribed when not needed, in unnecessary combinations or for longer durations than needed. Inappropriate prescribing is costly, exposes patients to unnecessary side-effects and drives population-level antimicrobial resistance, the prevalence and consequences of which are greatest in low- and middle-income countries. However, the implementation of interventions to improve the appropriateness of antimicrobial prescribing has been variable and requires further study. **Methods:** We propose a type III hybrid implementation/effectiveness interventional cohort trial in 35 ICUs in up to 11 low- and middle- income countries. The study intervention is a structured review of antimicrobial prescriptions as

recommended by the World Health Organisation. Strategies to support stakeholder-led implementation include development of local protocols, registry-enabled audit and feedback, and education. Evaluation of implementation, and the determinants of its success, is informed by the RE-AIM framework and the Consolidated Framework for Implementation Research respectively. The primary outcome is a composite measure of fidelity, reach and adoption. Secondary outcomes describe the effectiveness of the intervention on improving antimicrobial prescribing. Qualitative interviews will assess relevant implementation acceptability, adaptations and maintenance. A baseline survey will investigate ICU-level antimicrobial stewardship structures and processes. Discussion: This study addresses global policy priorities by supporting implementation research of antimicrobial stewardship, and strengthening associated healthcare professional competencies. It does this in a setting where improvement is sorely needed: low- and middle- income country ICUs. The study will also describe the influence of pre-existing antimicrobial stewardship structures and processes on implementation and improve understanding about the efficacy of strategies to overcome barriers to implementation in these settings. Trial registration: This study protocol has been registered with ClinicalTrials.gov (ref NCT06666738) on 31 Oct 2004. <https://clinicaltrials.gov/study/NCT06666738?term=NCT06666738&rank=1>. © The Author(s) 2025.

Author Keywords

Antimicrobial Resistance; Antimicrobial Stewardship; Audit & Feedback; Critical Care; Implementation; Intensive Care; Low- and Middle- Income Countries; Quality Improvement

Index Keywords

antifungal agent; antimicrobial stewardship, human, implementation science, intensive care unit, organization and management, prescribing error, prevention and control, total quality management; Anti-Bacterial Agents, Antimicrobial Stewardship, Humans, Implementation Science, Inappropriate Prescribing, Intensive Care Units, Quality Improvement

Chemicals/CAS

Anti-Bacterial Agents

References

- Kaki
Impact of antimicrobial stewardship in critical care: A systematic review
 (2011) *J Antimicrob Chemother*, 66 (6), pp. 1223-1230.
 1:CAS:528:DC%2BC3MXmt1ajurs%3D, 21460369
- (2024) *Antimicrobial resistance : accelerating national and global responses*, 3, pp. 3-8.
- Murray
Global burden of bacterial antimicrobial resistance in 2019: a systematic analysis
 (2022) *Lancet*, 399, pp. 629-655.
 10325, 1:CAS:528:DC%2BB38XpvFGrtb0%3D
- (2016) *Tackling drug-resistant infections globally: Final report and recommendations*,
<https://>
- Chiotos
Antibiotic stewardship in the intensive care unit: Challenges and opportunities
 (2019) *Infect Control Hosp Epidemiol*, 40 (6), pp. 693-698.
 31046851
- Wunderink
Antibiotic Stewardship in the Intensive Care Unit. An Official American Thoracic Society Workshop Report in Collaboration with the AACN, CHEST, CDC, and SCCM
 (2020) *Ann Am Thorac Soc*, 17 (5), pp. 531-540.
 32356696, 7193806
- **Antimicrobial Stewardship in the Intensive Care Unit**
 (2022) *Antimicrob. Steward. Non-Traditional Settings a Pract. Guid.*, 3, pp. 161-183.
- Thomas
A Multicenter Evaluation of Prolonged Empiric Antibiotic Therapy in Adult ICUs in the United States
 (2015) *Crit Care Med*, 43 (12), pp. 2527-2534.
 1:CAS:528:DC%2BC2MXhvVykbtzE, 26457751
- Quirós
Antimicrobial stewardship programs in adult intensive care units in Latin America: Implementation, assessments, and impact on outcomes

(2022) *Infect Control Hosp Epidemiol*, 43 (2), pp. 181-190.
33829982

- Marquet
Incidence and outcome of inappropriate in-hospital empiric antibiotics for severe infection: a systematic review and meta-analysis
(2015) *Crit Care*, 19 (1), p. 63.
25888181, 4358713
- (2019) *Antimicrobial stewardship programmes in health-care facilities in low- and middle-income countries: a WHO practical toolkit*,
<https://>
- **Opportunities to Improve Antibiotic Appropriateness in U.S. ICUs: A Multicenter Evaluation**
(2020) *Crit. Care Med*, 48 (7).
- **Hospital-acquired colonization and infections in a Vietnamese intensive care unit**
(2018) *PLoS One*, 13 (9).
no., p., <https://>
- McBride
Catastrophic health care expenditure due to septic shock and dengue shock in Vietnam
(2019) *Trans R Soc Trop Med Hyg*, 113 (10), pp. 649-651.
31340045, 6792161
- Pierce
Global Antimicrobial Stewardship with a Focus on Low- and Middle-Income Countries
(2020) *Int J Infect Dis*, 96, pp. 621-629.
32505875, 7271868
- Hashimoto
Impact of ward pharmacist-led antimicrobial stewardship in intensive care units
(2023) *J Chemother*, 35 (3), pp. 188-197.
35748502
- Wu
Barriers and facilitators of implementing interventions to improve appropriate antibiotic use in low- and middle-income countries: a systematic review based on the Consolidated Framework for Implementation Research
(2022) *Implement Sci*, 17 (1), p. 30.
35550169, 9096759
- Singh
Investigating infection management and antimicrobial stewardship in surgery: a qualitative study from India and South Africa
(2021) *Clin Microbiol Infect*, 27 (10), pp. 1455-1464.
33422658
- **The impact of training informal health care providers in India: A randomized controlled trial**
Science (80-.), 354 (6308).
Oct. 2016
- **A scoping review and behavioural analysis of factors underlying overuse of antimicrobials**
(2023) *Jac-Antimicrobial Resist*, 5 (3), p. dlad043.
Jun

- **Clinical Microbiology in Low Resource Settings**
(2020) *Frontiers in Medicine*, 7, p. 258.
Switzerland
- **Interventions to improve antibiotic prescribing practices for hospital inpatients**
(2017) *Cochrane Database Syst. Rev*, 2 (2).
Feb
- Pollack
Antibiotic Stewardship Programs in U.S. Acute Care Hospitals: Findings From the 2014 National Healthcare Safety Network Annual Hospital Survey
(2016) *Clin Infect Dis*, 63 (4), pp. 443-449.
27199462
- Howard
An international cross-sectional survey of antimicrobial stewardship programmes in hospitals
(2015) *J Antimicrob Chemother*, 70 (4), pp. 1245-1255.
1:CAS:528:DC%2BC2MXhsFahtL3J, 25527272
- O'Riordan
Quality indicators for hospital antimicrobial stewardship programmes: a systematic review
(2021) *J Antimicrob Chemother*, 76 (6), pp. 1406-1419.
1:CAS:528:DC%2BB3MXhslyjtb3O, 33787876
- Kotwani
Prescriber and dispenser perceptions about antibiotic use in acute uncomplicated childhood diarrhea and upper respiratory tract infection in New Delhi: Qualitative study
(2017) *Indian J Pharmacol*, 49 (6), pp. 419-431.
29674796, 5892023
- Chan
SPIRIT 2013 explanation and elaboration: guidance for protocols of clinical trials
(2013) *BMJ*, 346, pp. 1-42.
- **Effectiveness-implementation Hybrid Designs: Combining Elements of Clinical Effectiveness and Implementation Research to Enhance Public Health Impact**
(2012) *Med. Care*, 50 (3).
no
- Branch-Elliman
Promoting de-implementation of inappropriate antimicrobial use in cardiac device procedures by expanding audit and feedback: protocol for hybrid III type effectiveness/implementation quasi-experimental study
(2022) *Implement Sci*, 17 (1), pp. 1-12.
- Becker
Implementing contingency management for stimulant use in opioid treatment programs: protocol of a type III hybrid effectiveness-stepped-wedge trial
(2023) *Implement Sci*, 18 (1), pp. 1-15.
- Yardley
The Person-Based Approach to Intervention Development: Application to Digital Health-Related Behavior Change Interventions
(2015) *J Med Internet Res*, 17 (1).
25639757, 4327440
- Glasgow
Evaluating the public health impact of health promotion interventions: the RE-AIM

framework

(1999) *Am J Public Health*, 89 (9), pp. 1322-1327.

1:STN:280:DyaK1MvgtlOqtw%3D%3D, 10474547, 1508772

- Glowacki
Antibiotic Combinations with Redundant Antimicrobial Spectra: Clinical Epidemiology and Pilot Intervention of Computer-Assisted Surveillance
(2003) *Clin Infect Dis*, 37 (1), pp. 59-64.
1:CAS:528:DC%2BD3sXmsVWmur0%3D, 12830409
- Kim
Redundant combinations of antianaerobic antimicrobials: impact of pharmacist-based prospective audit and feedback and prescription characteristics
(2020) *Eur J Clin Microbiol Infect Dis*, 39 (1), pp. 75-83.
1:CAS:528:DC%2BC1MXhsleju7nO, 31482420
- Schultz
Economic impact of redundant antimicrobial therapy in US hospitals
(2014) *Infect Control Hosp Epidemiol*, 35 (10), pp. 1229-1235.
25203175, 6487658
- Llewelyn
Antibiotic review kit for hospitals (ARK-Hospital): a stepped-wedge cluster-randomised controlled trial
(2023) *Lancet Infect Dis*, 23 (2), pp. 207-221.
36206793
- Tamma
Rethinking How Antibiotics Are Prescribed: Incorporating the 4 Moments of Antibiotic Decision Making Into Clinical Practice
(2019) *JAMA*, 321 (2), pp. 139-140.
30589917
- **Impact of pharmacist-led antibiotic stewardship program in a PICU of low/middle-income country**
(2018) ,” *BMJ open Qual*, 7 (1).
no., p., <https://>
- **Interventions for improving critical care in Low- and Middle-Income Countries: A systematic review.,”**
Intensive Care Med.,
vol. In press
- **Effectiveness and efficiency of guideline dissemination and implementation strategies**
(2004) *Health Technol. Assess*, 8 (6), pp. iii-iv.
Feb
- **Performance evaluation of a multinational data platform for critical care in Asia**
(2022) *Wellcome Open Res*, 6 (251).
- Beane
Establishing a critical care network in Asia to improve care for critically ill patients in low- and middle-income countries
(2020) *Crit Care*, 24 (1), p. 608.
- Pulcini
Design of a ‘day 3 bundle’ to improve the reassessment of inpatient empirical antibiotic prescriptions
(2008) *J Antimicrob Chemother*, 61 (6), pp. 1384-1388.
1:CAS:528:DC%2BD1cXmtVajtro%3D, 18367462

- **Audit and feedbackâ€™ : Effects on professional practice and healthcare outcomes (Review)**
(2012) *Cochrane Database Syst. Rev*, 6 (6).
- Salluh
National ICU Registries as Enablers of Clinical Research and Quality Improvement
(2024) *Crit Care Med*, 52 (1), pp. 125-135.
37698452
- **Organisation, staffing and resources of critical care units in Kenya**
(2023) *PLoS One*, 18 (7).
no., p., <https://>
- Satterfield
The role of education in antimicrobial stewardship
(2020) *J Hosp Infect*, 105 (2), pp. 130-141.
1:STN:280:DC%2BB38zgs1Sjsg%3D%3D, 32243953
- **Feasibility and acceptability of implementing a practice guideline for the use of high flow nasal cannula in critically ill patients who have hypoxemia: A multi-centre study in Nepal**
(2023) *Wellcome Open Res*, 8 (196).
- **AIIMS ICU Rehabilitation (AIR): Development and description of intervention for home rehabilitation of chronically ill tracheostomized patients**
(2024) *Wellcome Open Res*, 8 (285).
- **A learning health systems approach to improving the quality of care for patients in South Asia**
(2019) *Glob. Health Action*, 12 (1).
- Damschroder
Conceptualizing outcomes for use with the Consolidated Framework for Implementation Research (CFIR): the CFIR Outcomes Addendum
(2022) *Implement Sci*, 17 (1), p. 7.
35065675, 8783408
- Harris
Research electronic data capture (REDCap)–a metadata-driven methodology and workflow process for providing translational research informatics support
(2009) *J Biomed Inform*, 42 (2), pp. 377-381.
18929686
- Harris
The REDCap consortium: Building an international community of software platform partners
(2019) *J Biomed Inform*, 95.
31078660, 7254481
- Ahuja
An evaluation of the implementation of interventions to reduce postoperative infections and optimise antibiotic use across the surgical pathway in India: a mixed-methods exploratory study protocol
(2022) *Pilot Feasibility Stud*, 8 (1), p. 237.
36335367, 9636821
- **Performance evaluation of a multinational data platform for critical care in Asia**
(2021) *Wellcome Open Res*, 6.
- *The TRUST Code – A Global Code of Conduct for Equitable Research Partnerships,*

- *University of Oxford, "OxTREC."*,
- (2017) *Effective strategies for scaling up evidence- based practices in primary care*: A systematic review, pp. 1-13.
- **Implementation Matters : A Review of Research on the Influence of Implementation on Program Outcomes and the Factors Affecting Implementation**
(2008) *pp*, pp. 327-350.
–, <https://>
- An
What Really Works in Intervention? Using Fidelity Measures to Support Optimal Outcomes
(2020) *Phys Ther*, 100 (5), pp. 757-765.
31944249
- (2023) *Evaluating the Implementation Fidelity to a Successful Nurse-Led Model (INTERCARE) Which Reduced Nursing Home Unplanned Hospitalisations*, 8, pp. 1-17.
- **Establishing Treatment Fidelity in Evidence-Based Parent Training Programs for Externalizing Disorders in Children and Adolescents**
(2014) *pp*, pp. 230-247.
–, <https://>
- **Implementation fidelity, student outcomes, and cost - effectiveness of train - the - trainer strategies for Masters - level therapists in urban schools**: Results from a cluster randomized trial
(2024) *Implement. Sci.*, pp. 1-19.
- *Consolidated Framework for Implementation Research*,
- Sheron
Healthcare provider and patient perspectives on access to and management of atrial fibrillation in the Northern Province, Sri Lanka: a rapid evaluation of barriers and facilitators to care
(2022) *BMC Health Serv Res*, 22 (1), pp. 1-13.

Correspondence Address

Wagstaff D.; Centre for Preoperative Medicine, United Kingdom; email: d.wagstaff@ucl.ac.uk

Publisher: BioMed Central Ltd

ISSN: 17485908

PubMed ID: 40001051

Language of Original Document: English

Abbreviated Source Title: *Implement. Sci.*

2-s2.0-85219608666

Document Type: Article

Publication Stage: Final

Source: Scopus

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