



Case-area targeted interventions (CATI) for cholera outbreaks: protocol for a prospective observational evaluation

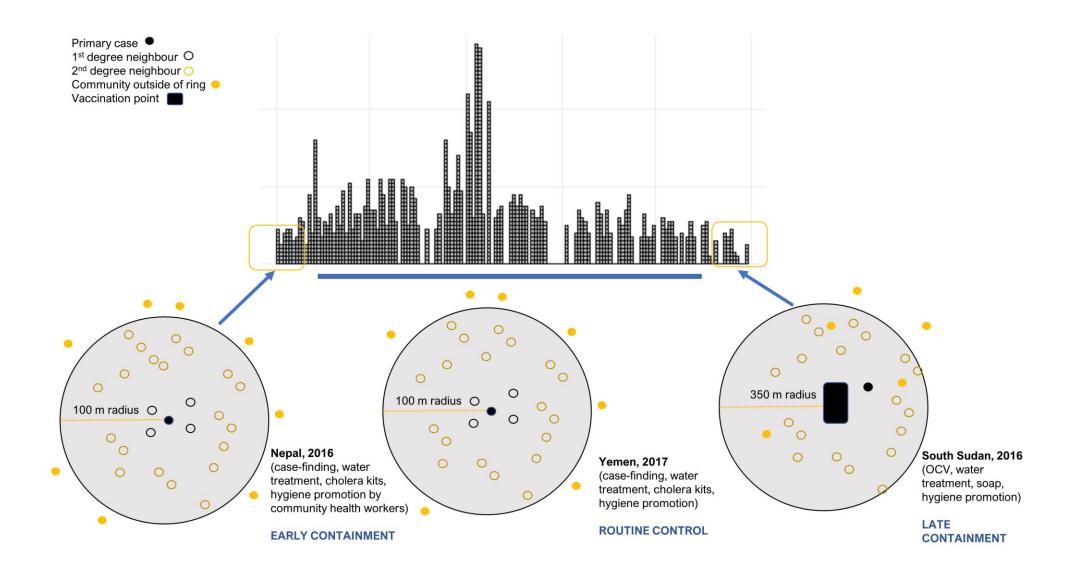
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CATI – Definition







CATI – Current evidence and gaps

Haiti (Centre Dept, 2015-7): retrospective data analysis

Reduction in size and duration of small-scale outbreaks when CATI was used (WASH and antibiotic chemoprophylaxis) (Michel et al, eLife, 2020)

• Bangladesh (Dhaka, 2013): randomized trial of culture-confirmed cases

Reduction in intra-household transmission when their household contacts were targeted with hygiene promotion and WASH (versus standard messaging) (George et al, EID, 2016)

- WASH Rapid Response Teams in Haiti, Yemen, Nigeria, South Sudan (Ramos et al, UNICEF, 2019)
- Scoping review of current evidence (Ratnayake et al, Lancet ID, 2020)
- Gaps:
 - Most studies are retrospective
 - No CATI evaluations including OCV
 - Based on suspected cholera





Including OCV in CATI

- Increased effectiveness
- Longer duration of protection
- Single dose
 - Short term protection demonstrated
- But:
 - ramp-up period of protection
 - OCV needs to be combined with other, faster acting interventions
- Little experience (South Sudan, very recently in Cameroon) and limited evidence so far
- Can be complementary to mass OCV campaigns





Using CATI with OCV in outbreak response

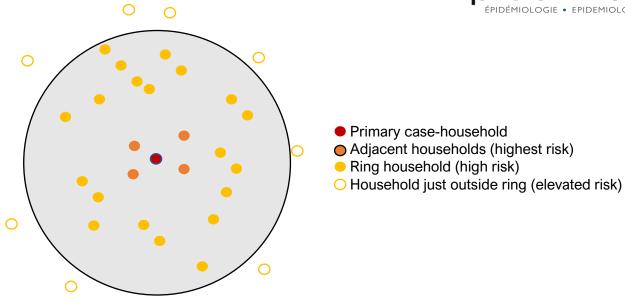
- MSF is interested in the concept of CATI with OCV as an additional tool for cholera outbreak response
- Combining single-dose OCV with household WaSH, possibly chemoprophylaxis
- Different MSF Operational Centres involved in developing strategies
- Discussions with MoH of DRC, Zimbabwe and Cameroon about first implementations
- Evaluation using research protocol developed in collaboration with Epicentre & LSHTM





Example

Target CATI <u>only</u> to enriched RDTpositive cases which are most likely to be true cases



	Primary case household	Adjacent households	Ring households
Single-dose oral cholera vaccine (1-dose, oral)			
Water treatment, safe storage, soap			
Doxycycline or azithromycin (1-dose, oral)			
Intensive hygiene promotion			
Community hygiene promotion			





Evaluation – research questions

Aim: to evaluate the effectiveness of CATI in the rapid containment of case-clusters, at the start of the outbreak

Primary objective: to evaluate the effectiveness of CATI in the reduction of incidence of enriched RDT-positive cholera within targeted rings

Secondary objectives:

- 1. Population-based coverage
- 2. Spatiotemporal transmission patterns of the outbreak
- 3. Effectiveness in the reduction of household transmission
- 4. Antimicrobial resistance related to chemoprophylaxis (if used)
- 5. Resources and costs required





Evaluation – principles

- Observational study of CATI performed by MSF
 - CATI can include OCV, WaSH, chemoprophylaxis
 - Evaluation can adapt to ring-size of intervention
- Use a study design appropriate for an outbreak (e.g., no explicit control group)
- Use real-time data collection to evaluate a CATI strategy prospectively
 - CTC linelist data & location of primary cases
- Compare immediately-implemented CATIs with naturally delayed CATIs on incidence of enriched RDT-positive cholera
- Understand the pathway to impact:
 - Coverage of CATI (each individual component)
 - Impact on household transmission and ring transmission





Conclusions

- CATI is an interesting new tool for cholera response
- Including OCV could increase effectiveness
- Complementary to preventive mass OCV campaigns and long-term WaSH improvements
- MSF planning on using CATI with OCV
- Discussions ongoing with several MoH about first implementations
- Systematic and prospective evaluation to provide more evidence
- Generic study protocol for evaluation of CATI interventions developed, currently under ethical review





Thank you!

This is a large and interdisciplinary project with many contributions:

- MSF: Nicolas Peyraud, Iza Ciglenecki, Catherine Bachy, Andrew Azman, Isabella Panunzi, Claire Dorion, Rob D'Hondt, Caroline Henry-Ostian
- Epicentre: María Lightowler, Etienne Gignoux, Francisco Luquero
- LSHTM: Ruwan Ratnayake, Francesco Checchi, John Edmunds
- MoH of DRC
- MoH of Zimbabwe
- UNICEF DRC