Randomized Controlled Trial of Hospital-Based Hygiene and Water Treatment Intervention (CHoBI7) to Reduce Cholera


The risk for cholera infection is >100 times higher for household contacts of cholera patients during the week after the index patient seeks hospital care than it is for the general population. To initiate a standard of care for this high-risk population, we developed Cholera-Hospital-Based-Intervention-for-7-Days (CHoBI7), which promotes hand washing with soap and treatment of water. To test CHoBI7, we conducted a randomized controlled trial among 219 intervention household contacts of 82 cholera patients and 220 control contacts of 83 cholera patients in Dhaka, Bangladesh, during 2013–2014. Intervention contacts had significantly fewer symptomatic Vibrio cholerae infections than did control contacts and 47% fewer overall V. cholerae infections. Intervention households had no stored drinking water with V. cholerae and 14 times higher odds of hand washing with soap at key events during structured observation on surveillance days 5, 6, or 7. CHoBI7 presents a promising approach for controlling cholera among highly susceptible household contacts of cholera patients.

Severe cholera without adequate rehydration kills up to half of affected persons (1). The World Health Organization estimates that 3–5 million cholera cases occur worldwide each year (2). Studies have identified multiple risk factors for Vibrio cholerae infection, such as drinking street-vended water, placing one’s hands into stored household water, lack of drinking water treatment, eating food prepared by a recently ill food handler, and not washing hands with soap before eating food (3–8). These findings suggest that cholera is transmitted through contaminated water and poor hygiene practices. Therefore, interventions targeting improved water treatment and storage practices and hand washing with soap have the potential to substantially reduce cholera transmission (8).

Previous studies in Bangladesh have demonstrated that household contacts of cholera patients are at >100 times higher risk for cholera infections during the 1-week period after the index patient seeks hospital care (5,7,9–11). Although the average rate of cholera in National Institute of Health–sponsored surveillance areas of Bangladesh is 1.6 cases/1,000 persons, a study in urban Dhaka, Bangladesh, found that 210 household contacts/1,000 index patients were infected with V. cholerae during a 21-day surveillance period (>90% of these infections occurred during the first week after the index patient sought care) (7,12). This high rate of cholera among household contacts probably results from a shared contaminated environmental source, such as water or food in the household, or secondary transmission from infected household members because of poor hygiene (6,9,13).

In Bangladesh, the current standard of care for cholera patients at hospital discharge is to provide oral rehydration solution (ORS) packets. No standard of care exists for household contacts of these patients despite their very high risk for cholera (5,7). The time that patients and their accompanying family members spend at a health facility during a severe diarrheal episode provides an opportunity for health providers to communicate information about water sanitation and hygiene (WASH) behavior change when perceived severity of diarrheal disease and perceived benefits of water treatment and hand washing with soap are likely to be highest (14). However, only a few studies have evaluated the effects of health facility–based WASH interventions, and none have evaluated the effects of these interventions in reducing enteric infections among household contacts of hospitalized diarrhea patients (15–22).

To initiate a standard of care for household contacts of cholera patients during the 1-week high-risk period after the index patient seeks care, we evaluated the efficacy


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of a hospital-based intervention promoting hand washing with soap and treatment of water called Cholera-Hospital-Based-Intervention-for-7-Days (CHoBI7) in Dhaka, Bangladesh, during June 2013–November 2014. We hypothesized that, in comparison with the standard message given to cholera patients at hospital discharge on ORS, CHoBI7 would significantly reduce cholera infections and increase hand washing with soap and treatment of water among highly susceptible household contacts of cholera patients.

Methods
All study participants (household contacts and cholera index patients) provided informed consent; consent comprised adult participants (≥18 years of age) signing an informed consent and/or parental consent form and children 12–17 years of age signing an assent form. If a study participant could not read, the consent form was read to him or her, and the participant then was asked to document his or her consent with an X in the presence of a witness. All study procedures were approved by the research Ethical Review Committee of icddr,b, Dhaka, and the Institutional Review Board of The Johns Hopkins Bloomberg School of Public Health (Baltimore, MD, USA).

We evaluated the efficacy of CHoBI7 by conducting a cluster randomized controlled trial in Dhaka during June 2013–November 2014. Suspected cholera patients seeking care at the icddr,b Dhaka Hospital were defined as persons with acute watery diarrhea (≥3 loose stools during a 24-h period) and moderate to severe dehydration using the World Health Organization definition. These patients were screened for V. cholerae in their feces by using the Crystal VC Rapid Dipstick test (Span Diagnostics, Surat, India) (23,24). All dipstick-positive findings were confirmed by bacterial culture. All patients suspected to have cholera who resided within a police thana (ward) of Dhaka and were admitted to icddr,b Dhaka Hospital were screened for eligibility for the CHoBI7 trial. A cholera case was defined as a fecal bacterial culture result positive for V. cholerae in a suspected cholera patient. Cholera patients were excluded from the study if a household contact already was enrolled (currently or previously) or if they had received cholera vaccine, to avoid confounding from an ongoing cholera vaccine trial.

Household contacts were defined as persons sharing the same cooking pot as the index patient for the previous 3 days. To be eligible for the study, household contacts had to plan to reside in the household of the index patient for the following week and could not have received cholera vaccine. Eligible household contacts in the hospital attending their ill family member at the time of cholera patient enrollment were invited to participate, and the household was visited to recruit household contacts within 36 hours after patient enrollment. Typically, cholera patients stayed at Dhaka Hospital for 24–48 hours before returning home. A cluster was defined as the index cholera patient and his or her household contacts.

The design of CHoBI7 was informed by factors from the Integrated Behavioral Model for Water, Sanitation and Hygiene interventions and constructs from the Health Belief Model (25,26). CHoBI7 was tailored to residents living in slum areas of Dhaka during 3 months of piloting and previous formative research (27). CHoBI7 includes 1) a pictorial (“Chobi” in Bangla) module on how cholera can spread through the environment (e.g., contamination of household drinking water sources and stored water), how persons can spread cholera to each other by contaminating food and water in their home, and instructions on proper hand washing with soap and treatment of water (Figure 1); and 2) a cholera prevention package containing a 3-month supply of chlorine tablets (Aquatabs sodium dichloroisocyanurate; Medentech, Wexford, Ireland, UK) for water treatment, soapy water bottles (a low-cost alternative to bar soap made using detergent), a hand washing station, a sealed water vessel with cover to ensure safe water storage, and cue-to-action cards with instructions about promoted behaviors (Figure 2). A trained health promoter at Dhaka Hospital delivered this pictorial module and cholera prevention package to cholera patients and their accompanying family members during a consultation session in the hospital. These messages were reinforced through daily household visits by the health promoter for the 1-week intervention period. The cost per household for CHoBI7 was US $45.50 (online Technical Appendix Table 1, http://wwwnc.cdc.gov/EID/article/22/2/15-1175-Techapp1.pdf); the cost included intervention hardware, transport cost, and the promoter’s salary.

Study recruitment at Dhaka Hospital occurred Saturday–Thursday each week during the study period. Each
week, half of the surveillance days were randomly selected to be intervention days, and half were randomly assigned to be control days by using a random number generator. The principal investigator (C.M.G.) assigned randomization; this scheme limited the likelihood of seasonal variations in study arm assignment and selection bias. The control arm received the standard message given at health facilities in Bangladesh about the use of ORS to treat diarrhea, and the intervention arm received this standard message and CHoBI7. To minimize bias, we used 2 separate teams for intervention and evaluation activities.

Households were visited on days 1 (baseline), 3, 5, 7, and 9 (visits 1–5) after the cholera patient sought care at Dhaka Hospital for clinical surveillance and to assess intervention uptake indicators. For clinical surveillance, household contacts were asked whether they had diarrhea (≥3 loose stools during a 24-hour period) or vomiting in the previous 48 hours, and a rectal swab sample was collected from willing household contacts at each household visit to test for *V. cholerae* in feces by bacterial culture. Because of limitations in our study personnel capacity, rectal swab samples were available only from household contacts enrolled during June 2013–June 2014.

To assess indicators of intervention fidelity, we collected a water sample from the household’s water source and drinking water stored in the home at each household visit to test for *V. cholerae* by bacterial culture and for the presence of free chlorine, as a proxy measure of water treatment, by using a digital colorimeter (Hach, Loveland, CO, USA). The US Centers for Disease Control and Prevention–recommended cutoff for free chlorine of ≥0.2 mg/L in household stored drinking water was used.

Spot checks were conducted at each household visit in all study households to observe whether soap was present near (within 10 steps of) the latrine and cooking areas as a proxy measure of hand washing with soap. To observe hand washing with soap practices, a 5-hour structured observation substudy was conducted once in all households recruited during October 2013–November 2014 (59 intervention and 56 control households) on surveillance day 5, 6, or 7. Hand washing with soap was recorded at the following key events promoted in CHoBI7: 1) after using the toilet, 2) after cleaning a child’s anus, 3) before eating, and 4) before preparing food.

Rectal swab samples were collected on Cary-Blair media, and water samples were collected in 500-mL bottles and transported to the Enteric Microbiology Laboratory at icddr,b. Fecal specimens and water samples were analyzed for *V. cholerae* and serotyped according to previously
published methods (30,31). The laboratory was blinded to
the study arm of specimens received.

Our primary outcomes were 1) the incidence of V. chol-
erae--infected household contacts, defined as a culture result
positive for V. cholerae, and 2) the incidence of symptomatic
V. cholerae infection, defined as diarrhea or vomiting in the
past 48 hours in a V. cholerae--infected household contact.
Our secondary outcomes were the percentages of 1) hand
washing with soap at key events during 5-hour structured
observation, 2) households with soap at the latrine and cook-
ing areas, 3) households with stored drinking water with de-
tectable V. cholerae, and 4) households with free chlorine
concentrations ≥0.2 mg/L in stored drinking water. We ex-
cluded the baseline household visit from analyses of the in-
tervention efficacy because the intervention had not yet been
provided to household members. The cost per cholera case
and case averted was calculated by using the assumptions in
online Technical Appendix Table 2.

We used Optimal Design software (University of
Michigan, Ann Arbor, MI, USA) for the sample size cal-
culation to determine the number of cholera cases (clusters
of household contacts) needed to reject the null hypothesis
that the incidence of cholera did not significantly differ by
study arm at a 95% CI and 80% power (32). We assumed
that cholera infection would occur in 20% of household
contacts in the control arm and that the intervention would
reduce this rate to 10% with an average cluster size of 3
household contacts (7). On the basis of these assumptions,
we estimated needing 156 index cholera patients and 468
household contacts (78 cholera patients and 234 household
contacts in each study arm).

To compare baseline household- and individual-level
characteristics by study arm, we conducted a χ² test for
categorical variables, a 2-sample t test for continuous vari-
ables, and a Fisher exact test when ≤5 values were in a
category. Logistic regression models were performed to
estimate the odds of developing cholera and to compare
intervention uptake indicators during visits 2–5 by study
arm using generalized estimating equations to account for
clustering within households and approximate the 95% CI.
These analyses were performed by using SAS version 9.3
(SAS Institute Inc., Cary, NC, USA). For study variables
where 1 study arm had no events, we used Fisher exact tests
at the household level to determine a significant difference
between study arms. To calculate exact 95% CIs in this in-
stance, we used an algorithm to invert test statistics in R (R
Core Team, Vienna, Austria) (33).

Results
Of the 655 suspected cholera cases screened by Crystal VC
Rapid Dipstick for the presence of V. cholerae in feces, 255
(39%) were dipstick positive; 400 (61%) results were nega-
tive or indeterminate (Figure 3). Of persons with dipstick-
positive samples, 71 declined to participate, and 16 were
negative for V. cholerae by bacterial culture.

We invited all eligible household members in study
households during the baseline surveillance visit to par-
ticipate in the trial. Of household members in cholera pa-
tient households, 27% (229/853) were unavailable for the
baseline interview and therefore were not enrolled in the
trial. The proportion of household members available for
the baseline interview did not differ significantly by study
arm (26% intervention arm vs. 29% control arm, p = 0.45).
Of the 453 household contacts screened for eligibility, 5
declined to participate, and 6 were not home during the
clinical surveillance period.
Two intervention and 1 control household refused to participate after the baseline visit. Therefore, 84 cholera patients and 225 household contacts were allocated to the intervention arm and 84 cholera patients and 220 household contacts to the control arm. Baseline index patient, household contact, and household characteristics did not differ significantly by study arm (Table 1). During the study period, 27% of control households had at least 1 water source (e.g., water pump) sample that tested positive for V. cholerae compared with 33% of intervention households (p = 0.4).

Culture results for V. cholerae were available for 320 (73%) household contacts. Enrolled household contacts with or without rectal swab culture results available did not differ significantly in clinical or demographic characteristics (online Technical Appendix Table 3).

A total of 148 (93%) control and 140 (88%) intervention household contacts were negative for V. cholerae at the baseline visit (p = 0.30) (Table 2). Intervention contacts had a 47% lower incidence of V. cholerae infection (symptomatic and asymptomatic) than control contacts (odds ratio [OR] 0.50, 95% CI 0.21–1.18) during the intervention period. Furthermore, intervention contacts had no symptomatic V. cholerae infections, compared with 5% of control contacts (OR 0.00, 95% CI 0–0.623). On the basis of these findings, we determined the cost per cholera case (symptomatic V. cholerae infection) averted would be US $227.50 ($227.50–$598.68) (online Technical Appendix Table 2). We calculated the range for the cost estimate using the 95% CI for the OR of a symptomatic V. cholerae infection.

The odds of hand washing with soap at key events during the structured observation period were 14 times higher in the intervention arm than in the control arm (OR 14.68, 95% CI 8.32–25.90) (Table 3), and the odds of hand washing with soap after toileting were 12 times higher in the intervention arm than in the control arm (OR 12.14, 95% CI 5.68–25.93). A significantly higher proportion of household visits in the intervention arm than in the control arm had soap present at the cooking area (99.7% vs. 95% CI 5.68–25.93). A significantly higher proportion of household visits in the intervention arm than in the control arm had soap present at the cooking area (99.7% vs. 83% (46), n = 220) and latrine area (98% vs. 13%, p<0.0001) during the intervention period (Table 3). V. cholerae was

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**Table 1. Demographic and environmental characteristics of households of patients with cholera, by study arm, Dhaka, Bangladesh, June 2013–November 2014**

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Control arm</th>
<th>Intervention arm</th>
<th>p value†</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. households</td>
<td>83</td>
<td>82</td>
<td></td>
</tr>
<tr>
<td>No. enrolled household contacts, median ± SD (min–max)</td>
<td>2 ± 0.9 (2–6, n = 220)</td>
<td>2 ± 0.8 (2–5, n = 219)</td>
<td>0.9</td>
</tr>
<tr>
<td>Index patient</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Female sex, no. (%)</td>
<td>56 (67)</td>
<td>52 (63)</td>
<td>0.5</td>
</tr>
<tr>
<td>Age, y, median ± SD (min–max)</td>
<td>25 ± 17.6 (0.67–95)</td>
<td>25 ± 15 (1–65)</td>
<td>0.3</td>
</tr>
<tr>
<td>&lt;5, no. (%)</td>
<td>5 (6)</td>
<td>8 (10)</td>
<td>0.6</td>
</tr>
<tr>
<td>5–14, no. (%)</td>
<td>16 (19)</td>
<td>17 (21)</td>
<td></td>
</tr>
<tr>
<td>&gt;14, no. (%)</td>
<td>62 (75)</td>
<td>57 (70)</td>
<td></td>
</tr>
<tr>
<td>Household contact‡</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Female sex, no. (%)</td>
<td>135 (61)</td>
<td>126 (58)</td>
<td>0.3</td>
</tr>
<tr>
<td>Age, y, median ± SD (min–max)‡</td>
<td>13 ± 15 (0.75–67, n = 220)</td>
<td>13 ± 16 (0.58–75, n = 219)</td>
<td>0.3</td>
</tr>
<tr>
<td>&lt;5 years, no. (%)</td>
<td>36 (16)</td>
<td>45 (21)</td>
<td>–</td>
</tr>
<tr>
<td>5–14, no. (%)</td>
<td>84 (38)</td>
<td>68 (31)</td>
<td>0.3</td>
</tr>
<tr>
<td>&gt;14, no. (%)</td>
<td>103 (46)</td>
<td>106 (48)</td>
<td>0.4</td>
</tr>
<tr>
<td>Television ownership, no. (%)</td>
<td>42 (51)</td>
<td>45 (55)</td>
<td>0.5</td>
</tr>
<tr>
<td>Electricity, no. (%)</td>
<td>82 (99)</td>
<td>82 (100)</td>
<td>0.3</td>
</tr>
<tr>
<td>Refrigerator ownership, no. (%)</td>
<td>12 (14)</td>
<td>9 (11)</td>
<td>0.5</td>
</tr>
<tr>
<td>A household member can read and write, no. (%)</td>
<td>67 (81)</td>
<td>72 (88)</td>
<td>0.2</td>
</tr>
<tr>
<td>Educational level of person responsible for primary drinking water collection, no. (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No formal education</td>
<td>40 (48)</td>
<td>40 (49)</td>
<td>0.3</td>
</tr>
<tr>
<td>Primary school</td>
<td>31 (37)</td>
<td>24 (29)</td>
<td></td>
</tr>
<tr>
<td>Secondary school</td>
<td>11 (13)</td>
<td>17 (21)</td>
<td></td>
</tr>
<tr>
<td>Higher secondary school</td>
<td>0</td>
<td>0</td>
<td></td>
</tr>
<tr>
<td>Bachelor’s degree</td>
<td>1 (1)</td>
<td>1 (1)</td>
<td></td>
</tr>
<tr>
<td>Master’s degree</td>
<td>0</td>
<td>0</td>
<td></td>
</tr>
<tr>
<td>Water source type, no. (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Groundwater</td>
<td>45 (54)</td>
<td>46 (46)</td>
<td>0.3</td>
</tr>
<tr>
<td>Piped water supply</td>
<td>380 (46)</td>
<td>34 (41)</td>
<td></td>
</tr>
<tr>
<td>Baseline presence, no. (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Any type of soap in latrine area of household</td>
<td>13 (16)</td>
<td>9 (11)</td>
<td>0.3</td>
</tr>
<tr>
<td>Any type of soap in cooking area of household</td>
<td>10 (12)</td>
<td>8 (10)</td>
<td>0.6</td>
</tr>
<tr>
<td>Vibrio cholerae in stored drinking water</td>
<td>5 (6)</td>
<td>9 (11)</td>
<td>0.2</td>
</tr>
<tr>
<td>V. cholerae in source water</td>
<td>10 (12)</td>
<td>12 (15)</td>
<td>0.6</td>
</tr>
<tr>
<td>Presence of V. cholerae in source water during study period</td>
<td>22 (27)</td>
<td>27 (33)</td>
<td>0.4</td>
</tr>
</tbody>
</table>

*Unless otherwise specified, the denominator for the control arm is 83 and for the intervention arm 82.
†p values were calculated by using generalized estimating equations to account for the clustering of the data at the household level.
‡p values were calculated by using generalized estimating equations to account for the clustering of the data at the household level.
present in no stored drinking water samples in households in the intervention arm and in 6% of samples in the control arm during the intervention period (OR 0.95% CI 0–1.08). The proportion of households with free chlorine concentrations ≥0.2 mg/L was significantly higher in the intervention arm than in the control arm (94% vs. <1%, p<0.0001).

Discussion

CHoBI7 significantly reduced symptomatic *V. cholerae* infections and reduced overall *V. cholerae* infections by nearly half during the intervention period. Consistent with these findings, the odds of hand washing with soap at key events during the structured observation were 14 times higher in the intervention arm than in the control arm, and nearly all intervention households had free chlorine concentrations in stored drinking water in the Centers for Disease Control and Prevention–recommended range. In addition, no stored drinking water samples in intervention households had detectable *V. cholerae*. These findings demonstrate that CHoBI7 was highly effective in reducing symptomatic cholera and increasing hand washing with soap and treatment of water during the 1-week high-risk period for household contacts of cholera patients.

We attribute the success of the CHoBI7 intervention to several key factors. First, this intervention was delivered during a time of severe illness in these households, when perceived severity of diarrheal disease and perceived benefits of hand washing with soap and treatment of water were likely to be high. Previous studies have found that during outbreaks of severe disease, such as cholera, households have higher perceived severity of diarrheal disease and greater perceived benefits of water treatment (14,34,35). Consistent with this observation, in Dhaka in 2013, use of a community-level point-of-use chlorine dispenser peaked after cholera-associated deaths in a slum area (L. Unicomb, pers. comm.). Second, we provided hardware that was pretested in a pilot study and facilitated the promoted behaviors (hand washing with soap and treatment of water) (27). Third, we trained health promoters to reinforce the promoted behaviors by using the CHoBI7 pictorial module, which probably led to a favorable environment for habit formation (25).

CHoBI7 significantly reduced symptomatic *V. cholerae* infection but not overall infection. We suspect the reason is our small sample size and the intervention reducing the infecting inoculum size within households to a level below which symptomatic infection could occur. Consistent with this hypothesis, a previous challenge trial found that symptomatic infection could occur at an inoculum size of 10^5 CFUs of *V. cholerae* and that illness severity was based on the size of the infecting inoculum (36).

Major advantages of the CHoBI7 intervention are its focus on high-risk persons during the 1-week period when

<table>
<thead>
<tr>
<th>Household visit with source water with detectable <em>V. cholerae</em>, visits 2–5</th>
<th>15/83 (18)</th>
<th>22/82 (27)</th>
<th>1.66 (0.79–3.49)</th>
<th>0.18</th>
</tr>
</thead>
</table>

Table 3. Odds ratios for hand washing with soap and water treatment and indicators of water quality in an intervention study of *Vibrio cholerae*, Dhaka, Bangladesh, June 2013–November 2014

*Logistic regression using generalized estimating equations.
†Soap present within 10 steps of the latrine or cooking area at household visits during the intervention period.
‡Cutoff recommended by the Centers for Disease Control and Prevention (Atlanta, GA, USA).
§To calculate exact 95% CIs, an algorithm was used to invert test statistics.
¶Fisher exact test.
they are most susceptible to cholera infections and its dissemination in a clinical setting in which cholera cases can be rapidly identified by dipstick test. Furthermore, the intervention is relatively inexpensive (US $227.50/cholera case averted) and would be likely to be more cost effective than a similar WASH intervention implemented as a community-based intervention, given the much lower prevalence of cholera in the general population (1.6 cholera cases/1,000 general population vs. 50 cholera cases/1,000 household contacts of cholera patients) (12,37). A recent study that used a cholera vaccine cost-effectiveness calculator found that a cholera vaccination program targeting geographic hotspots for cholera (cholera incidence >10 cases/1,000 year) in Bangladesh would cost US $226 per cholera case averted, similar to the cost of CHoBI7 (38).

To our knowledge, only 1 intervention study has been published that evaluated the effectiveness of safe water storage and water treatment on cholera transmission among household contacts of cholera patients. This study, conducted in Calcutta, India, resulted in a 59% reduction in overall cholera infections in the chlorine water treatment arm and a 76% reduction in the narrow-necked water pitcher arm during the 5-day intervention period (39). An earlier intervention study in Dhaka found that promotion of hand washing with soap among household contacts of shigellosis patients resulted in an 85% reduction in symptomatic Shigella infections during the 10-day intervention period (40). These findings are consistent with those from our trial and suggest that WASH interventions directed toward the high-risk period for household contacts of hospitalized diarrhea patients might be a promising approach for reducing transmission of enteric pathogens in this susceptible population.

Our study has a few limitations. First, because CHoBI7 combined hand washing with soap and treatment of water, we cannot establish the effect of these interventions individually. Second, our sample size was small, and we were unable to obtain culture results from as many household contacts as anticipated because of limited study personnel capacity. This limitation reduced our power to detect a significant difference in primary outcome between the 2 study arms (80% vs. 69%). Third, 27% of household members of cholera patients were not present during the baseline surveillance visit and therefore were not enrolled as household contacts. These persons are likely to have been the household members who spent the most time outside the home during the study period. However, the proportion of household members who were unavailable for the baseline interview did not differ significantly by study arm. Fourth, the study could not be unblinded; however, to minimize potential bias, the evaluation and intervention teams were separate, and the laboratory was blinded to intervention assignment. Fifth, this study was an efficacy trial. Our objective was to evaluate whether hand washing with soap at key events and treatment of household stored drinking water consistently would significantly reduce V. cholerae infections. Future studies should conduct an effectiveness trial to identify whether a hospital-based intervention only (without home visits) can lead to sustained uptake of the promoted hand washing with soap and treatment of water during the 1-week high-risk period for these households.

In our study, CHoBI7 significantly reduced symptomatic V. cholerae infections among household contacts of cholera patients in urban Dhaka, Bangladesh. These findings suggest that this hospital-based intervention is a promising, cost-effective approach that could be initiated as a standard of care for household members of cholera patients. Future studies should investigate the efficacy of CHoBI7 in other settings affected by cholera globally, evaluate the effects of CHoBI7 on other enteric pathogens, and identify effective low-cost approaches to take CHoBI7 to a larger scale.

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References


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Including:
• Sequence Type 4821 Clonal Complex Serogroup B Neisseria meningitidis in China, 1978–2013
• Estimated Deaths and Illnesses Averted During Fungal Meningitis Outbreak Associated with Contaminated Steroid Injections, United States, 2012–2013

• Global Burden of Invasive Nontyphoidal Salmonella Disease, 2010
• Dose-Response Relationship between Antimicrobial Drugs and Livestock-associated MRSA in Pig Farming
• Cost-effectiveness of Chlamydia Vaccination Programs for Young Women
• Hospitalization Frequency and Charges for Neurocysticercosis, United States, 2003–2012
• Additional Drug Resistance of Multidrug-Resistant Tuberculosis in Patients in 9 Countries
• Oral Cholera Vaccination Coverage, Barriers to Vaccination, and Adverse Events following Vaccination, Haiti, 2013
• Ebola Risk Perception in Germany, 2014

http://wwwnc.cdc.gov/eid/articles/issue/21/06/table-of-contents
Randomized Controlled Trial of Hospital-Based Hygiene and Water Treatment Intervention (CHoBI7) to Reduce Cholera

Technical Appendix

Technical Appendix Table 1. Cost per household of Cholera-Hospital-Based-Intervention-for-7-Days intervention, Dhaka, Bangladesh, June 2013–November 2014†

<table>
<thead>
<tr>
<th>Item</th>
<th>Rate/unit*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Health promoter salary, US $120/mo†</td>
<td>12.00</td>
</tr>
<tr>
<td>Promoter transportation for 7 household visits</td>
<td>9.00</td>
</tr>
<tr>
<td>Hand washing station</td>
<td>12.00</td>
</tr>
<tr>
<td>12-L covered drinking water vessel</td>
<td>9.00</td>
</tr>
<tr>
<td>Detergent for soapy water for 1 wk</td>
<td>0.30</td>
</tr>
<tr>
<td>Promoter communication materials</td>
<td>3.00</td>
</tr>
<tr>
<td>Water treatment tablets for 1 wk</td>
<td>0.20</td>
</tr>
<tr>
<td>Cost for intervention/household, US $†</td>
<td>45.50</td>
</tr>
</tbody>
</table>

*These estimates assume that 1 promoter can serve 10 households/mo.
†Calculated by dividing the monthly salary per promoter by the 10 households they can serve each month.

Technical Appendix Table 2. Calculation of cost per Vibrio cholerae infection and cholera case averted for CHoBI7, Dhaka, Bangladesh, June 2013–November 2014*

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Value</th>
<th>Cost per symptomatic case of Vibrio cholerae infection averted</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cholera cases in households without CHoBI7, nonbaseline</td>
<td>0.20 Cholera cases/household</td>
<td>Without the intervention, expected cholera cases in households during visits 2–5 would be: as follows 5% (cholera case from control arm visits 2–5) × 4 (susceptible household contacts) = 0.20, assuming 4 susceptible household contacts/household during visits 2–5</td>
</tr>
<tr>
<td>Cholera cases in households with CHoBI7, nonbaseline</td>
<td>0 Cholera cases/household</td>
<td>With the intervention, expected cholera cases in household during visits 2–5 would be as follows: (100%–100% [CHoBI7 efficacy]) × 5% (cholera cases from control arm visits 2–5) × 4 (susceptible household contacts) = 0</td>
</tr>
<tr>
<td>Cholera cases averted in households with CHoBI7</td>
<td>0.20 Cholera cases/household</td>
<td>By using these assumptions the intervention would avert: 0.20 (cholera cases without CHoBI7) – 0.0 (cholera cases in households with CHoBI7) = 0.20 cholera cases per household</td>
</tr>
<tr>
<td>Cost/cholera case averted</td>
<td>US $227.50/cholera case averted</td>
<td>Cost per cholera case averted would be as follows: US $45.50 (cost of the CHoBI7)/0.20 (cholera cases averted) = US $227.50. A range of US $227.50 USD–US $598.68 was added by using the 95% CI for the odds ratio of a symptomatic V. cholerae infection.</td>
</tr>
</tbody>
</table>

*CHoBI7, cholera hospital-based intervention for 7 days.

Technical Appendix Table 2. Clinical and demographic characteristics by study arm by study participants with rectal swab results available, Dhaka, Bangladesh, June 2013–November 2014

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Yes*</th>
<th>No*</th>
<th>p value†</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. enrolled household contacts</td>
<td>320 (73)</td>
<td>122 (27)</td>
<td>0.65</td>
</tr>
<tr>
<td>Intervention arm</td>
<td>160 (49)</td>
<td>60 (50)</td>
<td>0.65</td>
</tr>
<tr>
<td>Sex, F</td>
<td>193 (60)</td>
<td>68 (56)</td>
<td>0.52</td>
</tr>
<tr>
<td>Mean age of household contact ± SD, y (min-max)</td>
<td>18 ± 15.3 (0.7–75), n = 320</td>
<td>20 ± 16.4 (0.6–72), n = 122</td>
<td>0.12</td>
</tr>
<tr>
<td>Diarrhea during surveillance period</td>
<td>103 (33)</td>
<td>40 (32)</td>
<td>0.73</td>
</tr>
<tr>
<td>Vomiting during surveillance period</td>
<td>33 (9)</td>
<td>11 (10)</td>
<td>0.68</td>
</tr>
</tbody>
</table>

*All values are no. (%) unless otherwise indicated.
†p values calculated by using generalized estimating equations to account for the clustering of data at the household level.