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Vaz Nery, Susana; Pickering, Amy J; Abate, Ebba; Asmare, Abraham; Barrett, Laura; Benjamin-Chung, Jade; Bundy, Donald AP; Clasen, Thomas; Clements, Archie CA; Colford, John M; +13 more... Ercumen, Ayse; Crowley, Siobhan; Cumming, Oliver; Freeman, Matthew C; Haque, Rashidul; Mengistu, Birhan; Oswald, William E; Pullan, Rachel L; Oliveira, Rita G; Einterz Owen, Katey; Walson, Judd L; Youya, Ashrafedin; Brooker, Simon J; (2019) The role of water, sanitation and hygiene interventions in reducing soil-transmitted helminths: interpreting the evidence and identifying next steps. Parasites & Vectors, 12 (1). DOI: https://doi.org/10.1186/s13071-019-3532-6

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REVIEW

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The role of water, sanitation and hygiene interventions in reducing soil-transmitted helminths: interpreting the evidence and identifying next steps

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Abstract

The transmission soil transmitted helminths (STH) occurs *via* ingestion of or contact with infective stages present in soil contaminated with human faeces. It follows therefore that efforts to reduce faecal contamination of the environment should help to reduce risk of parasite exposure and improvements in water, sanitation and hygiene (WASH) are seen as essential for the long-term, sustainable control of STH. However, the link between WASH and STH is not always supported by the available evidence from randomised controlled trials, which report mixed effects of WASH intervention on infection risk. This review critically summarises the available trial evidence and offers an interpretation of the observed heterogeneity in findings. The review also discusses the implications of findings for control programmes and highlights three main issues which merit further consideration: intervention design, exposure assessment, and intervention fidelity assessment.

Keywords: Soil-transmitted helminths, Water, Sanitation, Hygiene, WASH

Background

Soil-transmitted helminths (STH) are a group of intestinal nematodes that include *Ascaris lumbricoides, Trichuris trichiura,* and the hookworm species, *Necator americanus, Ancylostoma duodenale* and *An. ceylanicum.* These species are some of the most common infections among humans, affecting over 1.5 billion individuals globally [1]. Infection occurs through accidental ingestion of eggs of *A. lumbricoides* and *T. trichiura* (and occasionally *A. duodenale*) or larval penetration of the skin by hookworm larvae present in contaminated soil [2]. In recent decades, the burden of STH has declined

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markedly: the 2016 Global Burden of Disease study estimated there was a 43–78% (depending on STH species) reduction in disability adjusted life years caused by STH between 1990 and 2016 [3]. These reductions likely reflect the direct impact of a scale-up in school- or community-based deworming programmes [4] as well as increased access to self-treatment.

It is also likely that economic development and increased access to improved water, sanitation and hygiene (WASH) infrastructure and services have contributed to the reduction in the STH disease burden, by reducing exposure to STH infective stages in the environment. Yet today too many people still lack access to basic WASH services, including 4.5 billion people without access to safely managed sanitation, 844 million without access to a basic water service, and 892 million people

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still practicing open defecation [5]. Addressing such inequalities has the potential to further reduce the burden of STH and potentially interrupt transmission.

Recent years have seen increased coordination and collaboration between the WASH and neglected tropical diseases sectors [6, 7]. However, key policy questions remain on the role of WASH in STH control, including: (i) whether improved access to WASH is an essential adjunct to deworming in order to control and eliminate STH; (ii) what are the appropriate WASH interventions and behaviours to achieve these goals; and (iii) how to best deliver those improvements. In this viewpoint, we report on an expert meeting convened by the Bill & Melinda Gates Foundation and the Children's Investment Fund Foundation in London, January 29, 2018. The available evidence on the impact of WASH interventions on STH infection is reviewed and the main discussion from the meeting is presented.

The current evidence

The potential for WASH to reduce STH transmission is supported by observational studies that report lower risk of STH infection associated with improved WASH access and practices. A 2014 systematic review and meta-analysis found that improved WASH, including piped water access, access to sanitation facilities, wearing shoes, and handwashing with soap, were associated with a 33–70% lower odds of STH infection [8]. A separate systematic review on the impact of sanitation found that sanitation was associated with 27% lower odds of *A. lumbricoides* infection, 20% lower odds of *T. trichiura* and 35% lower odds of hookworm infection [9].

Whilst observational studies can be useful and relatively quick and low-cost to conduct, they are vulnerable to systematic error or bias, including confounding. The strongest source of evidence to evaluate WASH interventions are cluster randomised controlled trials (RCTs), where interventions are randomised by groups of individuals, such as schools or communities. Cluster RCTs have their own limitations however, as they often focus on internal validity to assess a specific intervention within a specific context, sometimes at the detriment of external validity (i.e. generalisability). Table 1 summarises the designs and results of published cluster RCTs reporting the impact of school- and community-based WASH interventions on STH infections. Trials were identified by meeting participants who conducted many of the trials and had extensive knowledge of the field. The selection was supplemented by PubMed searches using key WASH and STH medical subject headings. The included trials have a variety of designs and outcomes, resulting in differing degrees of rigor, including the potential for contamination between arms, and this can hinder comparability across studies. However, some general results do emerge from the trial findings.

School-based hand hygiene and sanitation interventions can reduce STH reinfection among school children in some settings, but the impact varies by species. For example, hygiene promotion reduced prevalence and intensity of *A. lumbricoides* and *T. trichiura* in a trial in China [10] and intensity of *A. lumbricoides* infections, but not other species, in Peru [11]. WASH was also found to have an impact on *A. lumbricoides* alone in a trial of school-based WASH interventions in Kenya [12]. By contrast, a comprehensive school WASH intervention in Laos PDR did not reduce STH among school-aged children or their parents or under 5 siblings (Chard AN & Freeman MC, unpublished data).

Of the available community trials, most have focused on evaluating sanitation interventions aiming to reduce open defaecation. Three trials in India reported mixed effects from sanitation interventions on STH infection. Trials in Madhya Pradesh and Odisha in India reported no protective effects of latrine construction campaigns on STH infections [13, 14]. In both cases, however, open defaecation was still widespread in intervention communities due to low coverage and limited use of latrines. In a more recent study using a non-randomised, matchedcohort study design to assess the effect of a combined household water connection and latrine programme in Odisha, investigators reported a 44% reduction in overall STH infections (mainly hookworm, the most prevalent species) (Reese H et al., unpublished data). In that case, latrine coverage and coverage and use of household water supply connections were relatively high (>85%) and the intervention had been in place for at least three years before the study was undertaken, whereas follow-up in other studies has been shorter.

Other community randomised trials in different settings have evaluated the impact of single and combined WASH interventions on STH as secondary outcomes. Two recent, large factorial RCTs in rural Kenya and Bangladesh, known collectively as WASH Benefits, used similar trial designs to evaluate the effect of WASH interventions, alone and in combination, on STH infections in children in each setting. In Kenya, the trial achieved moderate-to-high levels of WASH coverage and found a lower prevalence of A. lumbricoides in the arm that received chlorine water treatment and in the combined WASH arm [15, 16]. In Bangladesh, sustained high uptake of the WASH interventions was achieved, and the trial found reductions of T. trichiura prevalence in the sanitation arm and hookworm in the chlorine water treatment arm and the integrated WASH intervention arm [17]. It is important to note that STH infections were not the primary outcomes for which the trials were

Setting	Design	Follow-up duration	Intervention and control	STH outcomes	Impact on STH
School-based trials Peru, Belén [11]	Pair-matched cluster-rand- omized controlled trial in 18 schools	4 months	Intervention: deworming fol- lowed by hygiene education which was supported by fortnightly support visits <i>Control:</i> deworming alone	STH infection; knowledge about STH among grade 5 school children (~12 years of age)	Intensity of A. <i>lumbricoides</i> 58% lower in the intervention arm but no impact on intensity of <i>T. trichura</i> or hookworm; no impact on prevalence of any STH
China, Hunan Province [10]	Single-blind, unmatched, cluster-randomized trial in 38 urban schools	1 school year	Intervention: Behavior change intervention, including a cartoon video <i>Control</i> , display of a health- education poster	A. Iumbricoides and T. trichiura infection, knowledge about STH; hand-washing behavior (hookworm not present)	50% reduction in STH infection
Kenya, Nyanza Province [12]	Cluster-randomized trial in 40 schools	10 months	Intervention: deworming fol- lowed by hygiene promotion, water treatment technology and supplies, latrine con- struction and hand washing and drinking water storage containers <i>Control</i> : deworming alone	STH reinfection among school children	44% reduction on A. <i>lumbricoides</i> reinfection, but no impact on other STH species
Community-based trials					
Northern Ethiopia [24]	Individual 2 × 2 factorial clustered randomized in 216 households	6 months	Intervention: promotion of handwashing with (provided) soap and nail clipping	STH reinfection among chil- dren aged 6–15 years	Lower rates of STH reinfection among children receiving handwashing intervention (14 vs 29%) and nail clipping (17 vs 26%)
India, Odisha [13]	Cluster-randomized trial in 100 rural villages	30 months	Intervention: latrine promotion and construction, as part of a CLTS campaign	STH infection among children aged <5 years	No impact on STH infection
India, Madhya Pradesh [14]	Cluster-randomized trial in 80 rural villages	21 months	Intervention: subsidies for and promotion of household latrines, as part of a CLTS campaign	STH infection among children aged <2 years	No impact on STH infection
Kenya, Western Province [16]	Cluster-randomized trial in 1226 villages grouped into 702 clusters	24 months	Intervention: 6 interventions including water, sanitation, handwashing, nutrition, com- bined WASH, and combined WASH plus nutrition (WSHN) <i>Control</i> s active and passive controls (no deworming)	STH infection among children aged 2 years and an older child in each household (mainly A. lumbricoides)	A. <i>lumbricoides</i> infection preva- lence was 18% lower in the water arm, 22% lower in the combined WASH arm, and 22% lower in the WSHN arm

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Setting	Design	Follow-up duration	Intervention and control	STH outcomes	Impact on STH
Bangladesh [17]	Cluster-randomized trial in 5551 compounds grouped into 702 clusters	26 months	Intervention: 6 interventions including water, sanitation, handwashing, nutrition, com- bined WASH, and combined WASH plus nutrition (WSHN) <i>Control</i> : passive controls (no deworming by study)	STH infection among children aged 2 years and up to 2 older children in each household	Water intervention reduced hookworm by 31% but did not affect other STH. Sanitation reduced <i>T. trichiura</i> by 29%, had a similar borderline effect on hookworm and no effect on A. <i>lumbricoides</i> . Combined WASH reduced hookworm by 29% and WSHN by 33%
Timor-Leste, Manufahi [19]	Cluster-randomized trial in 18 rural villages	24 months after 1st deworm- ing	Intervention: provision of water, CLTS based sanitation promotion and handwashing education, integrated with deworming of whole com- munities <i>Control</i> : deworming of whole communities	STH infection among entire communities	No additional impact of WASH interventions on STH infection compared to deworming alone
Non-randomised studies Côte d'Ivoire, south-eastern [20]	Non-randomized cluster inter- vention study in 9 villages	13 months	Intervention: CTS campaign and hygiene education plus two rounds of community-based deworming <i>Control</i> : two rounds of commu- nity-based deworming	STH infection among house- hold members (mainly hookworm present)	No statistically significant impact on STH prevalence, potentially higher hookworm ERR in intervention

Abbreviations: CLTS, community-led total sanitation; ERR: egg reduction rate; STH: soil-transmitted helminth

designed, and as such the trials were powered to detect only relatively large reductions in STH.

Further trials have evaluated the impact of mass drug administration (MDA) *versus* MDA combined with WASH. In the WASH for WORMS trial conducted in Timor-Leste, an integrated programme of WASH and community MDA was found to have no additional impact on STH infection compared to MDA alone [18, 19]. In contrast, a non-randomised intervention study in Côte d'Ivoire reported greater egg reduction rates of hookworm among communities that received a combined programme of community-led total sanitation (CLTS) and community-wide MDA compared to communitywide MDA alone [20].

Interpretation of findings

The above evidence demonstrates mixed findings. Potential explanations for such heterogeneity include that WASH interventions assessed in trials to date were either not appropriate to their study settings, were too complex resulting in limited uptake, failed to reach sufficiently high levels of coverage in the study population, or failed to achieve correct, consistent and sustained use. For instance, sanitation facilities, even those deemed "safely managed" under international monitoring standards, may present risks of user exposure inside (e.g., from unclean squatting slabs) and to the community from open sewers and untreated faecal sludge. WASH interventions have to date struggled to achieve and sustain high levels of community coverage and use, partly due to inadequate behaviour change methods to consistently achieve desired WASH practices. It is noteworthy that even in settings where high coverage of several WASH interventions is achieved, as in the Bangladesh WASH Benefits trial, the impact of interventions can still be modest.

In addition, since all communities have some access to water, some level of sanitation coverage and use and practice some personal hygiene behaviours, it is challenging to know what level of WASH intervention coverage is required to interrupt a sufficient number of exposure pathways and, in doing so, prevent reinfection. It is possible that there is a minimal required level of coverage and use for WASH interventions to have an impact, but this threshold will undoubtedly vary by intervention type, background reinfection rates, ongoing deworming or other factors not yet well defined. It is also difficult to manage contamination between arms in WASH RCTs, where control communities might have some uptake of improved water, sanitation or hygiene practices, diluting the estimated effect of the intervention.

Interestingly, the impact of WASH interventions in some trials was greatest for *A. lumbricoides* compared to hookworm. This may reflect differences in transmission (ingestion versus transdermal) or the ability of *A. lumbricoides* eggs to survive in the soil for years while hookworm larvae survive for weeks to months [21, 22]. Alternatively, the species-specific effects may reflect underlying differences in epidemiology of infection, including age patterns, within the respective study populations.

Lastly, the impact of WASH interventions seems to vary according to the underlying level of STH infection, with impact greatest at lower levels of infection. A possible explanation for this observation is that at high levels of infection and environmental contamination, WASH interventions require a longer follow-up period to see effects—most trials have follow-up periods of 1–2 years. Impact may additionally depend on the presence and length of ongoing deworming programmes. We hypothesize that the impact of WASH may be greatest after multiple years of MDA, when STH prevalence has been reduced to low levels. Here, mathematical modelling provides useful insights: using an individual-based model, Coffeng et al. show that WASH interventions have negligible short-term impact on STH infections in the context of ongoing deworming programmes, especially community-wide deworming, but that they are essential to prevent rebound of infection once deworming is stopped [23]. The dynamic interaction between intervention effort (coverage and efficacy) and infection transmission intensity is recognized in the design of vaccination programmes, and in STH population dynamic theory around MDA, but has yet to be examined seriously in the context of WASH programming. Modelling of WASH is an area which deserves more attention, using available trial data to improve the robustness of model predictions.

Implications for programmes and future research

The London meeting identified three main issues that merit further consideration: intervention design, exposure assessment, and intervention fidelity assessment.

First, there is a need to identify and evaluate contextspecific, feasible complementary interventions to support deworming and WASH programmes. Some of these complementary interventions can be remarkably simple and affordable. For example, a trial in Ethiopia showed that combined fingernail clipping with a handwashing intervention reduced STH reinfection rates among school-aged children [24]. Shoe wearing, improved flooring, food hygiene (both at the household and food-system level), household hygiene and health promotion to reduce geophagia (intentional consumption of soil) are additional interventions not typically included in WASH programmes but may reduce exposure to STH infective stages. As an illustration, an observational study in rural Bangladesh found that finished flooring was associated with lower *Ascaris* infections among children [25], highlighting the potential impact of improved household flooring. In practical terms, formative research is first needed to identify, develop, and assess the feasibility of specific intervention packages to control STH. Additional RCTs would then be needed for understanding if there are additional benefits to integrating these complementary interventions within deworming and WASH programmes.

Secondly, future work needs to improve and incorporate assessments of STH exposure from the environment, ideally using molecular assays. Measuring exposure to STHs has not typically been integrated into WASH or MDA trials. When it is, it consists of indirect measures of exposure (by STH loads in the soil) in an attempt to understand if interventions are actually interrupting or reducing environmental transmission of STH. Persistent environmental reservoirs of STH eggs may not be reduced by all types of WASH interventions, and could prevent intensive MDA programmes from achieving STH elimination. Reinfection rates six months after MDA range from 57% to 94% for STH species [26]. There are limited data on STH infective stages in the environment, as well as uncertainty regarding how long helminth eggs typically stay viable in soil, food and water and on hands and surfaces outside of a laboratory setting. Moreover, few studies have quantified the abundance and distribution of helminth eggs in soil [27], drinking water, on hands, or on surfaces. Environmental surveillance, coupled with new molecular markers of STH [28], could provide valuable insight into how environmental STH reservoirs are affected by specific interventions. Future work will also need to address how to standardise measurement of STH infective stages in the environment across matrices [29]. Taking the long view, an effective method for measuring the density of helminth eggs in the environment might provide a low-cost, less intrusive alternative to measuring human infection as a means of screening communities, as is done by screening sewage for polio virus.

Thirdly, we must improve our reporting of both intervention fidelity and measurement of WASH outcomes (i.e., intervention uptake and usage). There is a need to more clearly report on intervention fidelity with the use of standardized measures and process evaluation to understand why a programme works, and enhance the external validity and comparability of the studies [30]. Applying implementation science frameworks, as are used in other public health disciplines, could support our understanding of how to replicate and scale successful interventions [31]. We also need better measures of WASH outcomes as intermediate measures on the causal chain between interventions and STH outcomes. Evidence from recent WASH trials suggest that even potentially effective interventions often fail to achieve health impacts because of poor delivery and uptake by the target population [13, 14, 19]. Most interventions consist of a combination of technology and behaviour change. For example, improved flooring may be readily embraced by a population, but still require cleaning in order to maximize protection against STH infection; handwashing with soap requires substantial behaviour change, but compliance can be improved by hand washing stations located near latrines and food preparation areas. Sanitation interventions often have a very high uptake initially but use of latrines decreases overtime potentially due to poor construction quality. It will be important to identify a set of harmonised outcomes to help comparison across studies. Future trials also need to carefully consider the behaviour change required to improve update and usage, including understanding beliefs that could hinder behaviour change.

Conclusions

The mainstay of current STH control programmes is periodic, population-based deworming (MDA), which has been shown to be safe, scalable and cost-effective. These programmes have been shown to provide health benefits for the recipients and have demonstrated value for endemic communities today. But MDA alone has yet to be shown to provide a long-term solution: indeed, potential anthelmintic resistance, donor fatigue and other threats raise concerns about the long-term sustainability of deworming programmes alone. Historical experience in previously endemic countries, including Europe, the USA, Japan and South Korea, has shown that STH infections can be effectively and sustainably controlled in the long-term through environmental interventions. Moreover, improvements in WASH will continue to be a major policy objective beyond the goal of reducing STH, due to its wide-ranging impacts on other diseases, society and well-being. Clearly MDA needs to continue where STH infection remains high and where WASH interventions cannot have an immediate impact. What is now required is stronger evidence and policy guidance on the complementary role that WASH has for deworming programmes (especially in preventing reinfection), the specific WASH interventions that have the greatest impact on STH, the WASH coverage levels which are required to have an impact on STH, and when in a control programme cycle they should be emphasized.

Abbreviations

MDA: mass drug administration; RCT: randomized controlled trial; STH: soiltransmitted helminths; WASH: water, sanitation and hygiene.

Acknowledgements

Not applicable.

Authors' contributions

SVN, AJP and SB conceived and wrote the manuscript. All authors contributed to the manuscript. All authors read and approved the final manuscript.

Funding

The meeting was organized and supported by Bill & Melinda Gates Foundation and the Children's Investment Fund Foundation.

Availability of data and materials

Not applicable.

Ethics approval and consent to participate Not applicable.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

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Received: 31 March 2019 Accepted: 23 May 2019 Published online: 28 May 2019

References

- Pullan RL, Smith JL, Jasrasaria R, Brooker SJ. Global numbers of infection and disease burden of soil transmitted helminth infections in 2010. Parasites Vectors. 2014;7:37.
- Bethony J, Brooker S, Albonico M, Geiger SM, Loukas A, Diemert D, et al. Soil-transmitted helminth infections: ascariasis, trichuriasis, and hookworm. Lancet. 2006;367:1521–32.
- Hotez PJ, Fenwick A, Ray SE, Hay SI, Molyneux DH. "Rapid impact" 10 years after: the first "decade" (2006–2016) of integrated neglected tropical disease control. PLoS Negl Trop Dis. 2018;12:e0006137.
- WHO. Integrating neglected tropical diseases into global health and development: fourth WHO report on neglected tropical diseases. Geneva: World Health Organization; 2017.
- UNICEF W. Progress on drinking water and sanitation: 2017 update and SDG Baseline. New York and Geneva UNICEF and WHO; 2017. https:// www.who.int/mediacentre/news/releases/2017/launch-version-repor t-jmp-water-sanitation-hygiene.pdf. Accessed 15 June 2018.
- Freeman MC, Ogden S, Jacobson J, Abbott D, Addiss DG, Amnie AG, et al. Integration of water, sanitation, and hygiene for the prevention and control of neglected tropical diseases: a rationale for inter-sectoral collaboration. PLoS Negl Trop Dis. 2013;7:e2439.
- Campbell SJ, Savage GB, Gray DJ, Atkinson JA, Soares Magalhaes RJ, Nery SV, et al. Water, aanitation, and hygiene (WASH): a critical component for sustainable soil-transmitted helminth and schistosomiasis control. PLoS Negl Trop Dis. 2014;8:e2651.

- Strunz EC, Addiss DG, Stocks ME, Ogden S, Utzinger J, Freeman MC. Water, sanitation, hygiene, and soil-transmitted helminth infection: a systematic review and meta-analysis. PLoS Med. 2014;11:e1001620.
- Freeman MC, Garn JV, Sclar GD, Boisson S, Medlicott K, Alexander KT, et al. The impact of sanitation on infectious disease and nutritional status: a systematic review and meta-analysis. Int J Hyg Environ Health. 2017;220:928–49.
- Bieri FA, Gray DJ, Williams GM, Raso G, Li YS, Yuan L, et al. Health-education package to prevent worm infections in Chinese schoolchildren. New Engl J Med. 2013;368:1603–12.
- Gyorkos TW, Maheu-Giroux M, Blouin B, Casapia M. Impact of health education on soil-transmitted helminth infections in schoolchildren of the Peruvian Amazon: a cluster-randomized controlled trial. PLoS Negl Trop Dis. 2013;7:e2397.
- Freeman MC, Clasen T, Brooker SJ, Akoko DO, Rheingans R. The impact of a school-based hygiene, water quality and sanitation intervention on soiltransmitted helminth reinfection: a cluster-randomized trial. Am J Trop Med Hyg. 2013;89:875–83.
- Clasen T, Boisson S, Routray P, Torondel B, Bell M, Cumming O, et al. Effectiveness of a rural sanitation programme on diarrhoea, soil-transmitted helminth infection, and child malnutrition in Odisha, India: a clusterrandomised trial. Lancet Glob Health. 2014;2:e645–53.
- Patil SR, Arnold BF, Salvatore AL, Briceno B, Ganguly S, Colford JM Jr, et al. The effect of India's total sanitation campaign on defecation behaviors and child health in rural Madhya Pradesh: a cluster randomized controlled trial. PLoS Med. 2014;11:e1001709.
- Arnold BF, Null C, Luby SP, Unicomb L, Stewart CP, Dewey KG, et al. Cluster-randomised controlled trials of individual and combined water, sanitation, hygiene and nutritional interventions in rural Bangladesh and Kenya: the WASH benefits study design and rationale. BMJ Open. 2013;3:e003476.
- Pickering A, Njenga S, Steinbaum L, Swarthout J, Lin A, Arnold B, et al. Integrating water, sanitation, handwashing, and nutrition interventions to reduce child soil-transmitted helminth and Giardia infections: a clusterrandomized controlled trial in rural Kenya. bioRxiv pre-print. 2018; https:// doi.org/10.1101/464917.
- Ercumen A, Benjamin-Chung J, Arnold BF, Lin A, Hubbard AE, Stewart C, et al. Effects of water, sanitation, handwashing and nutritional interventions on soil-transmitted helminth infections in young children: a cluster-randomized controlled trial in rural Bangladesh. PLoS Negl Trop Dis. 2019;13:e0007323.
- Nery SV, McCarthy JS, Traub R, Andrews RM, Black J, Gray D, et al. A cluster-randomised controlled trial integrating a community-based water, sanitation and hygiene programme, with mass distribution of albendazole to reduce intestinal parasites in Timor-Leste: the WASH for WORMS research protocol. BMJ Open. 2015;5:e009293.
- Nery SV, Traub RJ, McCarthy JS, Clarke NE, Amaral S, Llewellyn S, et al. WASH for WORMS: a cluster-randomized controlled trial of the impact of a community-integrated water, sanitation, and hygiene and deworming intervention on soil-transmitted helminth infections. Am J Trop Med Hyg. 2019;100:750–61.
- Hurlimann E, Silue KD, Zouzou F, Ouattara M, Schmidlin T, Yapi RB, et al. Effect of an integrated intervention package of preventive chemotherapy, community-led total sanitation and health education on the prevalence of helminth and intestinal protozoa infections in Côte d'Ivoire. Parasites Vectors. 2018;11:115.
- Udonsi JK, Atata G. *Necator americanus*: temperature, pH, light, and larval development, longevity, and desiccation tolerance. Exp Parasitol. 1987;63:136–42.
- 22. Muller R. Chapter 5: the Nematodes. In: Muller R, editor. Worms and human disease. Wallinford: CABI; 2002.
- Coffeng LE, Vaz Nery S, Gray DJ, Bakker R, de Vlas SJ, Clements ACA. Predicted short and long-term impact of deworming and water, hygiene, and sanitation on transmission of soil-transmitted helminths. PLoS Negl Trop Dis. 2018;12:e0006758.
- Mahmud MA, Spigt M, Bezabih AM, Pavon IL, Dinant GJ, Velasco RB. Efficacy of handwashing with soap and nail clipping on intestinal parasitic infections in school-aged children: a factorial cluster randomized controlled trial. PLoS Med. 2015;12:e1001837.
- 25. Benjamin-Chung J, Nazneen A, Halder AK, Haque R, Siddique A, Uddin MS, et al. The interaction of deworming, improved sanitation, and

household flooring with soil-transmitted helminth infection in rural Bangladesh. PLoS Negl Trop Dis. 2015;9:e0004256.

- Jia TW, Melville S, Utzinger J, King CH, Zhou XN. Soil-transmitted helminth reinfection after drug treatment: a systematic review and meta-analysis. PLoS Negl Trop Dis. 2012;6:e1621.
- Gyawali P, Ahmed W, Sidhu JP, Nery SV, Clements AC, Traub R, et al. Quantitative detection of viable helminth ova from raw wastewater, human feces, and environmental soil samples using novel PMA-qPCR methods. Environ Sci Pollut Res Int. 2016;23:18639–48.
- Llewellyn S, Inpankaew T, Nery SV, Gray DJ, Verweij JJ, Clements AC, et al. Application of a multiplex quantitative PCR to assess prevalence and intensity of intestinal parasite infections in a controlled clinical trial. PLoS Negl Trop Dis. 2016;10:e0004380.

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- Collender PA, Kirby AE, Addiss DG, Freeman MC, Remais JV. Methods for quantification of soil-transmitted helminths in environmental media: current techniques and recent advances. Trends Parasitol. 2015;31:625–39.
- 30. Carroll C, Patterson M, Wood S, Booth A, Rick J, Balain S. A conceptual framework for implementation fidelity. Implement Sci. 2007;2:40.
- Proctor EK, Powell BJ, McMillen JC. Implementation strategies: recommendations for specifying and reporting. Implement Sci. 2013;8:139.

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