Lead (Pb) contamination in drinking water in low- and middle-income countries: a systematic review and meta-

3 analysis

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- 5 Authors: Michael B. Fisher*1; Timothy Purvis¹; Zakaria Seidu²; Siddhartha Roy³; Michelle
- 6 Cawley⁴; Carrie Baldwin-SoRelle⁴; Ryan D. Cronk¹; Aaron A. Salzberg¹; Amy Guo¹; J.
- 7 Wren Tracy^{1, 5}; Emily Browning¹; Jamie K. Bartram^{*1, 6}
- 8 Author affiliations
- 9 1. The Water Institute at UNC. University of North Carolina. Chapel Hill, NC 27599, United States.
- 10 2. West African Centre for Cell Biology of Infectious Pathogens, University of Ghana, Legon, Ghana.
- 11 3. Department of Environmental Sciences, Rutgers University, New Brunswick, NJ, United States
- AUL for Health Sciences and Director, Health Sciences Library, University of North Carolina.
 Chapel Hill, NC 27599, United States.
- 14 5. ICF International, 2635 Meridian Pkwy Suite 200, Durham, North Carolina, 27713, United States
- 15 6. School of Civil Engineering, University of Leeds
- 16
- 17 *Corresponding authors: Michael B. Fisher, The Water Institute at UNC. University of North
- 18 Carolina. Chapel Hill, NC 27599, mbfisher@gmail.com or fishermb@email.unc.edu; Jamie K.
- 19 Bartram, School of Civil Engineering, University of Leeds, Woodhouse Ln, Woodhouse, Leeds
- 20 LS2 9DY, United Kingdom. jbartram@unc.edu

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- 22 Keywords: lead, Pb, heavy metal, metalloid, drinking-water, corrosion, global, quality, risk,
- 23 public health, toxic metal
- 24

25 Abstract

- 26
- 27 Lead (Pb) in drinking water causes organ damage, cardiovascular disease, cancers, and
- 28 lifelong neurological and developmental impairment, disproportionately harming infants and
- 29 developing fetuses. However, evidence on lead in drinking water in low- and middle-income
- 30 countries (LMICs) is limited and lacks robust synthesis, impeding action. To address this gap,
- 31 we conducted a systematic review and meta-analysis of peer-reviewed studies reporting lead in

32 drinking water in LMICs, according to PRISMA guidelines, to answer the following research

33 questions:

 What is the estimated prevalence of lead contamination in drinking water in LMICs at concentrations exceeding international guideline values?
 To what extent does lead contamination in LMICs vary by geography, over time, or by drinking water source type?
 Are there problematic evidence gaps with respect to the occurrence of lead in drinking water in LMICs, and if so, where is additional evidence most urgently needed?

42 We searched PubMed, Scopus, and Web of Science for studies published in English since 43 1969 reporting lead concentrations in drinking water. The decision to include English 44 language studies was made on the basis of >90% of eligible studies being published in 45 English, leading to the expectation that exclusion of non-English studies would not seriously compromise the validity of the systematic review. Study relevance was ranked using 46 47 supervised clustering and machine learning. Relevant studies were manually screened for 48 inclusion; data were manually extracted from included studies. Within-study risk of bias was 49 scored using quality items defined in this work. Between-study bias was assessed based on 50 continuity and symmetry of the (roughly lognormal) distribution of data included in the review. 51 Of approximately 40,000 search results (for both lead and other TMs), 16,868 scored 52 relevant using a trained machine learning algorithm; of these, 3,367 met inclusion criteria. Fewer than 6% of studies (n= 200) were excluded because they were unavailable in English. 53 54 Approximately one third (n=1,088) of included studies reported on lead. Central and Southern Asia accounted for 45% of included datasets, while 31% of LMICs were 55

unrepresented. Many studies (45%) had sites purposively selected for known or suspected
contamination; after excluding such "targeted" studies, metaregression indicated that >20%
(95% CI: 20-27%) of samples exceeded World Health Organization (WHO) guideline values
for lead (n=523).

Potential sources of bias within studies include sample collection and analysis limitations (lower quality studies) and purposive site selection ("targeted" studies); potential sources of bias among studies include language limitations, exclusion of grey literature, geographic heterogeneity of datasets, and potential publication bias. This work suggests that lead contaminates drinking water at levels of health concern in LMICs worldwide, and that increased collective efforts to prevent, manage, and monitor such contamination are merited.

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68

69 Introduction

Exposure to lead causes adverse health effects [1–9]: these include cardiovascular disease [2], organ damage [4–6], anemia [5] and impaired red blood cell production [7], lifelong neurological and developmental impairment [4,5,10,11], adverse reproductive and birth outcomes [4,7], cancer and/or premature death [6,12,13]. These adverse health effects are most severe for lead exposures during gestation and infancy [4,5,11]. Human exposure occurs through routes such as ingestion and inhalation in outdoor, domestic, and occupational settings [5–7,11,13,14]. As environmental exposure to lead becomes more limited through the phase-out of leaded paint,

77 gasoline, and other regulatory measures, reducing exposure through drinking water is

- 78 increasingly important [15].
- 79 Understanding the origins of toxic metals (TMs) in drinking water is important because different sources
- 80 of exposure require different management approaches [16]. Most lead in drinking water arises from
- 81 corrosion of lead- and lead-containing water system materials and components, such as pipes and
- fittings [1,2], as occurred in Flint, Michigan [3] and Washington, D.C. [4].
- 83 Several studies have reported on lead in drinking water in the USA, Canada, Europe, and other high-
- 84 income countries (HICs) [5–10]. By contrast, TM contamination in drinking water is far less extensively
- 85 characterized in low- and middle-income countries (LMICs) [11]. Where LMIC studies report occurrence
- of TM contamination in water, they may be outdated (e.g., they may predate important regulatory,
- 87 demographic, economic, and/or analytical methods changes), may be hyper-localized (e.g., study setting
- 88 may be purposively targeted based on characteristics that reduce generalizability), and/or may be
- 89 designed and reported in a way that limits internal and/or external validity concerning TM occurrence at
- 90 concentrations of public health concern in drinking water (e.g., they may not report sampling, analysis,
- 91 and quality assurance/quality control [QA/QC] methods in sufficient detail to enable meaningful
- 92 interpretation of results). As a result, the extent of TM occurrence in drinking water in LMIC settings and

93 the extent to which these occurrences go undetected are unclear.

94

95 Objectives

- 96 This review assembles and summarizes peer-reviewed evidence to address the following97 research questions:
- What is the estimated prevalence of lead contamination in drinking water in LMICs at
 concentrations exceeding international guideline values?
 - 4

100 2. To what extent does lead contamination in LMICs vary by geography, over time, or

101 by drinking water source type?

- 102 3. Are there problematic evidence gaps with respect to the occurrence of lead in
- 103 drinking water in LMICs, and if so, where is additional evidence most urgently
- 104 needed?
- 105

106 Methods

107

108 PRISMA Guidelines and Review Registration

109 The reporting of this review follows PRISMA guidelines [12] (File S1 Checklist). This work is part

of a larger systematic review of multiple toxic metals (protocol registered with PROSPERO:

111 CRD42024566116; File S1 Protocol). The current review includes those studies reporting

112 results for lead in drinking water.

113

114 Search Strategy

The search string comprised: economic terms, country names, source types, and elements. The economic terms were developed through consultations with the Health Sciences Library at UNC-Chapel Hill. The country terms included in the search string were all countries within the categories of: Low-income economies, middle-income economies as defined by the World Bank (note that the World Bank further subdivides middle-income into Lower-middle-income and Upper-middle-income economies) [13]. Water supply technology types were based on the WHO/UNICEF Joint Monitoring Programme for Water Supply, Sanitation and Hygiene (JMP) 122 classifications of water supply technology type (e.g., boreholes with handpumps, public

tap/standpipes, surface water) [14]. The list of metals and metalloids was selected based on

those with which the WHO has established guideline values (Table 1) [2].

125

126 Eligibility

127 Studies were excluded if they were published in or before 1969, unavailable in full-text in 128 English, did not present primary data (e.g., literature reviews, opinion pieces, letters to the 129 editors, commentaries), or did not report concentrations of toxic metals (larger study) or lead 130 (this work) in drinking water samples or water samples from a source likely to be used for 131 drinking water supply (e.g., a surface water or groundwater source elsewhere reported to be a 132 drinking water source for nearby populations, or a water supply technology such as a borehole 133 with handpump that is commonly used for drinking water supply; Table 2). Grey literature was 134 not included.

135

136 Search

137 The search was executed in PubMed (National Library of Medicine), Global Health 138 (EBSCOhost), Scopus (Elsevier), and Web of Science (Clarivate) databases (Table S1 in File 139 S1 Text). The initial search was conducted on November 17, 2018 and was expanded and 140 updated multiple times on December 31, 2021; October 11, 2024;, and (most recently) on 141 March 17, 2025. The initial search yielded 5,864 results that were uploaded to Covidence 142 systematic review screening software. In 2021, the search was updated and expanded, then 143 executed in PubMed, Web of Science, and Scopus, vielding a total of 33,698 results. After 144 deduplication in EndNote, the 26,646 unique results were prioritized for screening as described 145 below. The search was updated on October 11, 2024, and March 17, 2025 with expanded

- 146 geographic search terms in Scopus, yielding a total of 19,003 new results across databases, of
- 147 which 13,607 were unique additions, for a total of 46,073 unique results across all searches.
- 148 The authors added relevant results identified by hand through their previous reading,
- 149 collaborations, and review of the citations of included studies (Fig. 1).

150



Fig .1: PRISMA flow diagram for a systematic review identifying publications on lead occurrencein drinking water in LMICs

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155 Refining Search Results: Machine Learning Approach

156 To prioritize the large number of results, machine learning was used to identify results with the 157 highest likelihood of being relevant. For the original search strategy, we used supervised 158 clustering, a form of semi-supervised machine learning, with ensemble learning and for the 159 search update we used supervised machine learning (Fig. 2) as described in the literature [28-160 31] and the study protocol (File S1 Protocol). Supervised clustering uses training data in the 161 form of seed studies that are identified at random from search results. Ensemble learning 162 combines results from several individual models to improve predictions. For this project we used 163 six models and each study received an ensemble score (ES) ranging from 0 to 6. Studies with 164 an ES of 0 were not predicted relevant by any of the six models and studies with an ES = 6165 were predicted relevant by all six models. Briefly: unique search results from December 2021 166 (n=26,646) were prioritized for manual screening using supervised clustering (DoCTER [ICF, 167 Virginia, USA]), which prioritizes search results based on title and abstract text using a training 168 dataset. First, 500 randomly selected results were manually screened at the title and abstract 169 level by the first author (Fisher) to identify training data for supervised clustering. Six models 170 were run using training data and all search results and each unique study received an ES 171 ranging from 0 to 6. All studies with ES of 1 or higher were uploaded to Covidence for manual 172 screening (n=9,983). Studies with an ES of 0 (not likely to be relevant) were discarded without 173 manual review (n=16,663). Supervised machine learning was used to prioritize the unique 174 results from the search updates (n=13,607). A random selection of screening results from the 175 original search was used to train the model (DoCTER [ICF, Virginia, USA]) and each study 176 received a probability score. All studies predicted to be relevant were manually screened by the

- team (n=4,218), as well as the highest ranked 500 studies marked not likely to be relevant as an
- 178 insurance step. The remaining studies were discarded (n=8,889).
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- 180



- 181
- 182 Fig 2. Machine learning in the systematic review screening process.
- 183 To assess the accuracy of the supervised clustering and supervised machine learning tools, the
- 184 team calculated the precision (number of true relevant studies out of the pool of studies
- 185 predicted relevant by the tools) of each subset of citations after title/abstract screening. As
- 186 expected, the precision decreased as ensemble score decreased for studies from the original
- 187 search, prioritized using the supervised clustering tool (Table 3, Fig S1 in File S1 Text).

188 Similarly, for studies from the search update prioritized by supervised machine learning,

189 precision decreased as predicted relevance decreased (Table 4, Fig S2 in File S1 Text).

190

191 Manual Screening

192 Following prioritization, studies were imported into the systematic review software Covidence 193 (Veritas Health Innovation) for screening. The authors developed and documented study 194 screening and data extraction criteria, protocols, and training materials (File S1 Trainings) and 195 developed training tools, which were used to train a pool of screeners and data extractors. This 196 pool included the authors as well as multiple collaborators at The University of North Carolina at 197 Chapel Hill (Chapel Hill, North Carolina, USA), the University of Ghana, Legon (Legon, Greater 198 Accra, Ghana), and the University for Development Studies (Nyankoala, Ghana). Two trained 199 screeners from this pool then independently screened the title and abstract of each included 200 study against the inclusion and exclusion criteria (Table 2); conflicts were resolved by a third 201 screener, generally one of the study authors. At the full text review stage, studies were again 202 independently reviewed by two trained screeners and excluded if they met any exclusion criteria 203 (Table 2).

204

205 Data Extraction

Basic study data and metadata (country, region, water source type, sample collection, handling,
and analysis methods/instrumentation) were extracted from each study by trained screeners at
the full-text screening stage (Table S2 in File S1 Text). Details on the study design and setting,
including whether the setting was selected based on a known or suspected source of
contamination, were also extracted at this stage.

211 For studies included at the full-text stage, additional data extraction was conducted by trained 212 data extractors (from the same pool of collaborators described above, and trained using the 213 same set of protocols, documents, and methods described in File S1 Trainings). Extracted data 214 included sample collection, preservation, analysis details, study and setting details, quantitative 215 results, geolocation data, and other key data, as indicated in the data extraction sheet (File S1 216 Tools). Sample collection details such as the mention of a stagnation period or flushing before 217 sample collection (which affects the concentrations of TMs in water samples); the use of trace-218 metal clean and/or trace-metal grade acid-washed sampling containers (if specified, which 219 improves method precision); and whether samples were acidified or filtered before analysis (which can dissolve and preserve [acidification] or remove [filtration] particulate or sorbed 220 221 material from samples, respectively). The details of the analytical method included the type of 222 analytical instrumentation used (e.g., flame AAS vs ICP-MS; File S1 Tools). Where a study 223 reported results for substantively different sample sets (e.g., different water source categories in 224 a single setting, different settings, or different seasons [pre-versus post-monsoon], these were 225 extracted as separate "sample sets" (File S1 Trainings). Each extracted study was reviewed by 226 a different author or trained data extractor, and errors were noted and corrected as a quality 227 control measure.

228 Data were cleaned and analyzed using STATA 13 (College Station, TX). Bibliometric and 229 methodological summary statistics were compiled; summary statistics of lead occurrence data 230 were calculated, and regression analyses were performed. Bibliometric statistics were 231 disaggregated by country and SDG region [15]. Summary statistics were calculated for the 232 proportion of lead exceedances (the proportion of samples exceeding the WHO guideline value 233 for lead in drinking water [10 µg/L] [2]) across sample sets by meta-regression using the 234 "metareg" command in STATA. This command was used to ensure that studies with larger 235 numbers of samples collected were not overweighted in the analysis. For this purpose, the

236 proportion of drinking water samples exceeding the WHO guideline value (GV) for lead was 237 determined for studies in which this proportion was directly reported or could be directly derived 238 from provided lead concentration data (e.g., from provided tables, figures, or statistics). Where 239 exceedance rates were not reported and could not be derived, but summary statistics (e.g., 240 range, mean, standard deviation, and number of samples) were reported, the proportion of 241 WHO GV exceedances was estimated as follows: For each sample set, reported lead 242 concentration ranges, means, and standard deviations were used to construct a lognormal 243 distribution and n values were randomly sampled from this distribution (where n = the reported 244 number of samples in the sample set), and the proportion of these sampled values exceeding 245 the WHO GV was recorded (previous studies have reported lognormal distributions of lead and 246 TM concentrations in water) [16]. For studies reporting concentration ranges, sample numbers, 247 and measures of central tendency (e.g., mean or median) but not reporting a measure of 248 variance (e.g., variance or standard deviation), the estimation method described by Hozo et al. 249 for approximating the variance of lognormal distributions was employed [17]. Standard errors for 250 the proportion of exceedances were calculated for each sample set. Where all samples in a 251 sample set exceeded the WHO GV (proportion of exceedances =1) or all samples were below 252 the WHO GV (proportion of exceedances=0), a continuity correction was made to calculate 253 standard errors, in which the proportion of exceedances was set equal to n/(n+1) or 1/(n+1), 254 respectively (in which n = the number of samples in the sample set). This ensured that standard 255 errors would not be undefined. Estimated exceedance proportions were then calculated across 256 all sample sets. For the subset of studies that reported results with sufficient granularity for 257 reported exceedance proportions to be extracted or calculated, these "reported" exceedance 258 proportions were compared to the "estimated" exceedance proportions obtained by the methods 259 described above (Fig S3 in File S1 Text).

Exceedance proportions were also calculated for the subset of studies conducted at sites not selected based on identified contamination (abbreviated as "nontargeted" studies), and the subset of "nontargeted" studies that reported using analytical instrumentation capable of achieving method detection limits below the WHO GV for lead (e.g., ICP-MS- see definition in Table 5). Exceedance proportions were also calculated for the subset of "nontargeted" studies with high method quality scores, as described below.

266

267 Data Analysis

Fisher conducted data analyses with support from the UNC Odum Institute and other coauthors,and Purvis validated these analyses.

270 Study method quality scores were calculated for studies using the criteria shown in Table 5. 271 Method score reflects the suitability of the reported study methods for quantifying lead in 272 drinking water by adding one point for each study guality criterion met. Studies with a method 273 quality score of six or higher out of ten criteria were classified as "high quality." Estimated 274 exceedance proportions were calculated for all studies, nontargeted studies, and across 275 stratifying variables such as water system type and SDG region using metaregression. 276 Estimated exceedance proportions were compared to reported (measured) exceedance 277 proportions for the subset of studies in which such exceedances were directly reported, and 278 these comparisons were reviewed using diagnostic plots as a further validation measure. 279 Conventional tests of publication bias (e.g., funnel plots) were adapted for this work, because 280 studies conducted in different geographic locations, at various periods, and/or using other 281 methods could not reasonably be presumed to have similar population distributions of lead 282 concentrations in drinking water; thus a measurement of the symmetry of central tendencies

across studies as a function of standard error is not meaningful. However, diagnostic plots

comparing the distribution of central tendencies among studies to an expected distribution (e.g.,

lognormal distribution) can be inspected for continuity and symmetry over a region of interest-

286 where large unexplained gaps, asymmetrical features, or other deviations from the expected

287 distribution over the region of (0.1*WHO GV) – (10* WHO GV), i.e., 1-100 μg/L for lead in

288 drinking water, can be considered indicative of potential bias among studies in the meta-

analysis.

290

291 **Results**

292

293 Literature Search and Prioritization

The original 2018 search returned 5864 results, which were screened without prioritization. The expanded 2021 search returned 33,699 studies; 9,983 results were imported into Covidence following prioritization using supervised clustering. One study was added through hand searching. Search updates in 2024 and 2025 yielded an additional 13,607 unique studies. After prioritization with supervised machine learning, 4,685 studies with high expected relevance were imported into Covidence.

A total of 20,521 studies were manually screened by the team. Of these, 1,088 relevant studies reporting on lead in drinking water (out of 2169 included studies reporting on any TM of interest), comprising 1,710 sample sets and approximately 62,000 observations, were included in the final dataset after title/abstract and full-text screening. Fewer than 10% (198 out of 2169) studies were excluded on the basis of language (i.e. being unavailable in English [Fig 1]).

305

306 Bibliometric Analysis

- 307 The final set of included studies represents 66 countries across six SDG regions. 48% of
- 308 studies that included datasets for lead in drinking water were from four countries: India, China,
- 309 Pakistan, and Turkey; the 10 most represented countries account for 73% of datasets (Fig 3,
- Table 6). Less than half (66 of 137, 47%) of all LMICs were represented in the data. Fewer
- 311 studies were included from certain SDG regions (such as Latin America and the Caribbean)
- than others (such as South and Central Asia [Table 7]). Fewer than 1% of returned records
- 313 were excluded based on English language availability.

314



- 316 Fig 3: Number of studies from low- and middle-income countries reporting on
- 317 concentrations of toxic metals and metalloids in drinking water, by country

318

319 Many studies (37%) reported on samples or locations selected because of a known or 320 suspected potential source of lead contamination ("targeted" studies). Mining, agriculture, 321 industrial sources, wastewater, and landfills were the most frequently listed potential sources 322 (Table 8). 323 Inductively coupled plasma mass spectrometry, atomic absorption spectroscopy, and inductively 324 coupled plasma atomic emission spectroscopy were the most reported analytical methods 325 (Table 9). However, the analytical methods were not adequately specified in 7% of publications 326 reporting on lead. 327 Study quality varied widely regarding the suitability of methods used for quantifying lead in 328 drinking water. The overall proportion of high-quality studies was 5% (Table 10). In most cases 329 (64%), studies failed to document adequate quality assurance/quality control (QA/QC) 330 procedures. 331 Metaregression of estimated exceedance proportions indicated that more than 25% of samples 332 exceeded the WHO guideline value for lead in drinking water across all sample sets (95% CI = 333 24-28%). For the subset of studies that reported results with sufficient granularity for 334 exceedance proportions to be extracted or calculated, these "reported" exceedance proportions 335 were compared to the "estimated" exceedance proportions obtained by the methods described 336 above; reasonable agreement was obtained by the two methods (Fig S3 in File S1 Text). 337 Metaregression of exceedance estimates for all "nontargeted" studies indicated that 24% of 338 these studies (95% CI = 20-27%) exceeded the WHO GV (Table 11), and this figure decreased 339 to 23% (95% CI = 18-27%) when only studies using more sensitive instrumentation (ICP-MS)

340 from LMIC settings other than China were considered (datasets from China [PRC] had

341 systematically lower levels of lead than those from other settings ex-PRC and were therefore

342	removed for some sub-analyses Fig S4 in File S1 Text). Similarly, median lead concentrations
343	across datasets were high for all studies as well as nontargeted studies (Table 12).

344 An inspection of the concentration distribution showed broad, roughly lognormal distributions for

345 lead in drinking water for both estimated and reported sample set median and mean values (Fig

346 4, Fig S1 in File S1 Text). Overall, median lead concentration in drinking water was higher than

347 the WHO GV of 10 μ g/L across studies, with the estimated median across nontargeted studies

being 27 μg/L (95% CI 14-40 μg/L) across 317 datasets for which suitable values and standard

349 errors could be estimated (Table 12).

350 Lead occurred across source types with roughly comparable frequency except springs;

351 however, the small number of datasets reporting on lead in water from springs should be noted

352 (Fig 5, Table 13). Lead occurrence was less prevalent in East and Southeast Asia (ESEA) and

353 Latin America and the Caribbean (LAC) than in other SDG regions; however, differences

354 between ESEA and other settings were less pronounced when data from PRC were omitted

355 (Fig 6, Table 13; Table S3 in File S1 Text). Lead occurrence did not vary significantly by decade

356 of publication (Figure 7, Table S4 in File S1 Text).

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Fig 4: Diagnostic plots of the distribution of log-transformed median and mean lead
 concentrations (µg/L) for drinking water sample sets from low- and middle-income

366 countries vs normal distribution

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370 Fig 5: Distribution of mean lead concentrations for drinking water sample sets from low-

and middle-income countries, by water source type. Red line denotes WHO Guideline

372 Value.

373



Fig 6: Distribution of mean lead concentrations for drinking water sample sets from lowand middle-income countries, by SDG region (CSA: Central and Southern Asia; ENA:
Europe and North America; ESEA: East and Southeast Asia; LAC: Latin America and the
Caribbean; NAWA: North Africa and Western Asia; SSA: Sub-saharan Africa). Red line
denotes WHO Guideline Value.

381



Fig 7: Distribution of mean lead concentrations for drinking water sample sets from low and middle-income countries, by decade of study publication. Red line denotes WHO
 Guideline Value.

388

389

390 Discussion

- 391 The results suggest the widespread occurrence of lead in concentrations that raise public health
- 392 concerns about drinking water in low- and middle-income countries worldwide. Lead
- 393 concentrations in drinking water exceeded WHO GVs in a substantive minority of samples

across all studies and "nontargeted" studies, irrespective of source type, SDG region, or decadeof observation/publication.

396 The distribution of lead concentrations was roughly lognormal; where samples exceeded WHO 397 GVs, the extent of this exceedance was also roughly lognormally distributed (Fig 4, Table 13). 398 We conclude that the contamination of drinking water by lead is a widespread public health 399 concern. While there is no known safe level of lead exposure and the WHO GV for lead in 400 drinking water is set based on analytical achievability [2], it is nevertheless a relevant 401 benchmark to consider as countries work to progressively decrease the proportion of water 402 samples exceeding this threshold and generally decrease lead concentrations and occurrence. 403 Thus, reporting the proportion of samples exceeding GVs, as well as the central tendency of 404 concentrations across pooled samples, as we have done here, can be a valuable approach to 405 synthesizing and communicating large numbers of observations across diverse systems and 406 settings for analysis at a high level. Such synthesis should indicate the extent of lead 407 occurrence across systems and settings, rather than a precise or localized estimate. 408 These results go beyond previous work but are consistent with prior studies emphasizing the 409 widespread occurrence of lead contamination in drinking water and the need for additional

410 monitoring and surveillance in LMIC settings [18–20].

411

412 Limitations

413

Data availability and data quality are both limited in this study. Many countries are not
represented in any published English language studies, while others have little included
evidence. Although fewer than 1% of returned records were excluded based on English

417 language availability, excluding evidence unavailable in English language peer-reviewed articles 418 indexed in major databases was a limitation of the current review. Specifically, while more than 419 90% of recent peer-reviewed articles in science, engineering, and health fields that are indexed 420 in major databases searched in this review are available in English [21], a substantive minority 421 of such indexed studies (most notably many from China) are not, and this represents a limitation 422 of the current work. Furthermore, a substantive proportion of non-English language articles are 423 not indexed in the databases searched in this study or most systematic reviews conducted in 424 English; this proportion increases when expanded to include grey literature and other non-peer-425 reviewed evidence sources (ibid). Thus, this limitation is a feature of the inclusion criteria and 426 the search parameters. Future updates to this work will seek to include additional evidence 427 published in select additional languages and additional evidence from the grey literature, as 428 feasible.Most included studies received low method quality scores for quantifying lead in 429 drinking water (method quality score is distinct from overall study quality because many 430 included studies are adequately designed for other purposes, but do not sufficiently quantify 431 lead in drinking water at µg/L concentrations). These low scores reflect numerous limitations, 432 including failure to report sampling and/or analysis methods and failure to include and document 433 guality assurance/guality control (QA/QC) measures. While the collective evidence showed 434 modest risk of bias (Fig 4), it is helpful to note that measurement bias, selection bias 435 (concerning sampling sites), publication bias, and other biases cannot be ruled out in this work. 436 The inaccessibility of raw data or suitable (i.e., nonparametric) summary statistics for some

437 studies complicated data analysis, and parameters were estimated in some cases, reducing the 438 precision of results. Most included studies provide parametric summary statistics, whereas lead 439 concentrations in drinking water tend to be lognormally distributed [16]; nonparametric statistics 440 were therefore estimated as described in Methods. The use of meta-regression methods helps 441 provide more generalizable pooled estimates of lead occurrence in drinking water across all

LMICs and within SDG regions despite such limitations among individual studies; however,
meta-analysis of additional nationally representative sample sets would improve estimates if
and when such data become available.

445 Pooled estimates of exceedance proportions help inform decisions to undertake or enhance 446 management and monitoring of lead in drinking water at the national, regional, and global scales 447 in LMICs, but have low generalizability to specific populations or subnational locations. Many 448 included studies (45%) are "targeted," i.e., they concern settings selected because of previously 449 identified or suspected localized sources of contamination. Such studies have low 450 generalizability to locations free from such issues. Those studies that are not located in settings 451 selected based on previously identified sources of lead contamination are likely to be more 452 representative of typical exposures, and incorporation of additional high-quality "non-targeted" 453 data as they are published, particularly in understudied LMIC settings, will reduce uncertainty 454 and increase the generalizability of estimates.

Locating and including additional sample sets within unpublished grey literature, government records, and non-English language studies may also improve generalizability. Evidence may be especially challenging to obtain in contexts where governments constrain the publication of research and grey literature and where publication of water quality results may be perceived as politically problematic. Further work may seek to delineate such evidence gaps and explore alternative means of assessing potential occurrence through parallel evidence, modeling estimates, or other means.

Thus, the present estimates help inform countries' decisions on *whether* to undertake or enhance actions in response to lead in drinking water and are likely adequate to justify broad primary prevention and monitoring. In other words, it is doubtful that including more and better data will change our assessment that lead in drinking water in LMIC settings is a global issue requiring timely action. However, additional high-quality local data are needed to decide *how*

best to act and *where* to target actions most intensively to prevent optimally and progressively
remediate lead occurrence in drinking water in LMICs. In some cases, such data may be slow or
challenging to obtain, and governments may seek to apply the precautionary principle in acting
on available data while continuing to address evidence gaps as feasible. In other cases,
nationally representative monitoring and surveillance of lead in drinking water may be possible
soon, if not already underway.

473

474 Sources of Bias

475 Potential sources of bias in individual studies in this work include the study quality factors noted 476 above: 1) limitations in sample collection and analysis methods, 2) "targeted" studies, and 3) 477 incomplete reporting of methods and raw data. Potential sources of bias across studies include 478 1) language barriers (exclusion of non-English language studies); 2) exclusion of grey literature; 479 3) differential water quality monitoring and research support and capacity across settings (both 480 between and within countries and regions [e.g., rural/urban biases]); 4) potential publication bias 481 (which may include both a greater tendency to publish results with noteworthy conclusions in 482 settings in which research publication is unrestricted, and a potentially lower likelihood of 483 publishing results that may be perceived as politically problematic in contexts in which 484 governments may exert some control over publication and research support concerning political 485 priorities). It should be noted that publication bias cannot be easily assessed using statistical 486 and graphical methods that assume a common central tendency of outcomes across settings 487 (e.g., odds ratio for a given association), since the central tendency of TM occurrence data 488 cannot be assumed to be constant across study types and settings.

489

⁴⁹⁰ Implications for Policy, Practice, Research, and Monitoring

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492 These results may prompt LMICs to consider reviewing, strengthening, and/or adopting 493 additional national policies, strategies, and regulations on lead in drinking water, including 494 widespread policies and actions to prevent and monitor the occurrence of lead in drinking water 495 and the use of lead-containing parts, materials, and alloys in the construction of new water 496 systems. Such actions may include developing or updating regulations and policies to ensure 497 that supply chains for drinking water system components are free from unsuitable parts and 498 materials (e.g., brass fittings with high lead content), that new and existing water systems are 499 designed and operated with lead exposure prevention in mind (including the use of optimized 500 corrosion control treatment, where appropriate), and away from known sources of lead 501 exposure likely to contribute to such contamination (e.g., smelting activities), and are managed 502 to minimize the risk of subsequent lead contamination (e.g., monitoring and/or managing land-503 use and industrial activities likely to produce severe lead contamination impacting water sources 504 and watersheds). Implementers may likewise strengthen measures to verify that source water 505 for new water supplies does not exceed national standards for lead before new water systems 506 are commissioned. They may further wish to undertake representative water quality monitoring 507 to track progress on lead in drinking water across subnational regions and over time, if they 508 intend to demonstrate progress on national and international targets related to safely managed 509 drinking water.

510

511 Progressive remediation of existing water systems containing unsuitable components or heavily 512 impacted by nearby land use or activities may also be appropriate for achieving national water 513 quality targets in many settings. Since remediation may be more costly and slower than primary

514 prevention, risk-based prioritization of systems according to the estimated likelihood and

515 intensity of lead contamination, population served, and remaining design life could be helpful.

516 Integrating remediation with scheduled maintenance activities and other rehabilitation activities

517 to achieve economies of scale, where possible, could also be beneficial.

518

519 As part of a set of evidence-informed actions to enhance progress on access to safely managed 520 drinking water, LMICs that do not currently monitor lead in community water supplies should 521 consider initiating such monitoring and surveillance efforts at an intensity commensurate with 522 their resources and the types of water supply systems in use. Those that do conduct such 523 monitoring should periodically review their monitoring strategies to ensure that lead 524 concentrations in drinking water are quantified before and after water system installation using 525 suitable sample collection, analysis, and QA/QC methods, and that monitoring strategies are 526 consistent with available evidence and analytical methods. The resulting monitoring data inform 527 management of lead in drinking water at the national and local levels and help ensure policies, 528 implementation strategies, and targets on drinking water quality are aligned. Results should also 529 be reported to affected populations and captured in national and local management information 530 systems to enhance transparency and sector coordination (e.g., between water, environment, 531 health, and finance agencies). Sampling, monitoring, analysis, reporting, and/or knowledge 532 management capacities may benefit from additional strengthening and coordination in many 533 countries.

534

Finally, research targeting high-priority evidence gaps can further inform policies and actions to
prevent exposure to lead in drinking water. Better characterizing and understanding the root
causes of lead occurrence in water across geographies, system types, and land

538 use/demographic variables enables more effective preventive and corrective actions. Further 539 innovation, optimization, piloting, and adoption of low-cost preventive and corrective solutions, 540 monitoring tools, and approaches could facilitate better detection and management. 541 Implementation science research could also enhance the efficiency of monitoring data to inform 542 implementation. Characterizing the extent of human exposure to lead from drinking water and 543 other sources and determining the associated disease burdens could highlight the importance of 544 managing and monitoring these hazards, build institutional support for further action, and enable 545 water, environment, health, and finance decision-makers to contextualize better the hazards 546 associated with exposure to lead in drinking water among other competing priorities and 547 opportunities.

548 Where called for, additional research and monitoring can likely be undertaken in parallel with 549 timely preventive action, rather than delaying or replacing prevention activities. In our 550 estimation, the available evidence is now sufficient to justify several "no-regrets" primary 551 prevention activities in many LMIC settings. While further evidence is likely to yield more 552 accurate, more precise, and higher-resolution (e.g., more geographically disaggregated) 553 estimates of the extent of lead occurrence in drinking water of different types across LMIC 554 settings, it is unlikely to change the big-picture finding that lead occurrence in drinking water at 555 levels of public health concern is widespread among the LMIC settings included in this review, 556 and that primary prevention activities can enhance progress on safely managed water access 557 and protect vulnerable sub-populations in these settings. Timely and effective prevention, 558 monitoring, progressive remediation, and evidence-generating actions such as those described 559 above would be highly likely to reduce lead exposure, protect the health of large populations 560 across LMICs, and support progress on safely managed drinking water access under SDG 6.1.

561

562 Acknowledgements

563 The authors gratefully acknowledge USAID for financial support of this work and acknowledge 564 Cathy Zimmer and Chris Wiesen of the UNC Odum Institute for invaluable input on statistical 565 methods. The authors acknowledge the outstanding team of article screeners, data extractors, 566 and collaborators and colleagues without whom note of this work would have been possible: 567 Adelaide Henewaa; Bably Prasad; Banks Grubbs; Bayan Abdulabdah; Beatrice Asantewaa; 568 Belinda Aculley; Benjamin Ashley; Bithiah Boaitey; Brayden Priebe; Cailin Antonio; Carlos 569 Nunes; Carly Barello; Carolyn Kowalski; Chahaana Kathiravan; Claire Evans; Clay Burgess; 570 Colleen Dongarra; Cory Nixon; Daniel Appiah; David Boadi; David Dzidula Nutakor; David 571 Nutakor Dzidula : Divya Tailor; Diva Kalyanshetti; Donya Farahani; Emily Barkley; Emma 572 Woody; Fuseini Abdul-Hagg; Irene Wang; Jimmy Kruse; Jonathan Hu; Joseph Asuam Nyarko; 573 Joseph Nyarko; Joyce Liu; Kaida Liang; Karina Samuel; Kate Rodelli; Katharine Conaway; 574 Katherine Church; Keshav Srivenkatesh; Kitty Goldman; Kyle Chan; Kyle Rezek; Kylie Heilferty; 575 Lawrence Ndela; Leah Simpson; Lily Eubanks; Linh Do; Lois Bangya; Lucas Atongo; Maggie 576 Kennedy; Manasi Chaudhary; Mariah Kurtovic; Mariah Kurtovic; Medhansh Bhagchandani; 577 Mercy Adekola; Michael Tran; Mohammad Siraj; Nana Akua Koranteng; Nash; Natalie 578 Ndiforamang; Nathan Anderson; Ngan Le; Nicholas Pell; Paige Crawford; Rachel Moser; 579 Regina Menu; Rithika Jonnalagadda; Robert Murray-Gramlich; Ronald Elikem Anlonya; Sam 580 Pell; Sanjana Nalla; Sarah Hwang; Shu Chen ; Siddharth Vangara; Simrika Joshi; Stellen Li; 581 Stephen Oppong; Susan Rathbun; Sydney Martin; Syed Anjerul Islam; Taralee Cornet; Timothy 582 Purvis; Viha Patel; William Seung; Yimeng Ma; and Yukai Li.

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585 Competing Financial Interests

- 586 This work was supported in part by funds from USAID. The authors declare they have no other
- 587 actual or potential competing financial interests. The funders had no role in data analysis,
- 588 interpretation, or publication decision.

589

590 Author Contributions

591 All authors approve of the submitted manuscript, accept accountability for their contributions 592 and promise to ensure that questions related to the accuracy or integrity of any part of the work, 593 even ones in which the author was not personally involved, are appropriately investigated. 594 resolved, and the resolution documented in the literature. Michael Fisher: contributed to the 595 conception, data analysis, writing and revision processes for this manuscript. Ryan D. Cronk, 596 Timothy Purvis, Siddhartha Roy, Michelle Cawley, and Carrie Baldwin-SoRelle contributed to 597 the conception, data analysis, and revision processes for this manuscript. Zakariah Seidu, Amy 598 Guo, J. Wren Tracy, and Emily Browning: contributed to the data analysis and revision process 599 for this manuscript. Jamie K. Bartram and Aaron Salzberg contributed to the writing and revision 600 process for this manuscript.

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602

603 Table 1. Toxic metals and metalloids included in the larger systematic review of TM

604 contamination in drinking water in low- and middle-income countries; this study reports only on 605 lead

Element	Rationale (Guidelines for Drinking Water Quality, 4 th Ed.)	WHO GV (ug/L, where applicable)	Source Category (Guidelines for Drinking Water Quality, 4 th Ed.)
Antimony	Health	20	Contaminants from pipes and fittings
Arsenic	Health	10	Naturally occurring
Cadmium	Health	3	Industrial sources and human dwellings
Chromium	Health	50	Naturally occurring
Copper	Health	2000	Contaminants from pipes and fittings
Iron	Aesthetic acceptability, disinfection (c.f. Browning et al.)	300*	Naturally occurring
Lead	Health	10	Contaminants from pipes and fittings
Manganese	Acceptability, Disinfection, Health effects (c.f. Browning et al.)	100*	Naturally occurring
Mercury	Health	6	Industrial sources and human dwellings
Nickel	Health	70	Contaminants from pipes and fittings
Selenium	Health	40	Naturally occurring
Uranium	Health	30	Naturally occurring

606 *Not health-based

607

608

609 Table 2: Inclusion and exclusion criteria for study selection

Inclusion Criteria	 Based in a low- or middle-income country (LMIC, according to the World Bank's Country and Lending Groups) Provides a quantitative measure of toxic metals and/or metalloids in water sample (larger systematic review); of lead in water sample (this work) Discusses a toxic metal of interest: Antimony, Arsenic, Cadmium, Chromium, Copper, Iron, Lead, Manganese, Mercury, Nickel, Selenium, Uranium (larger systematic review); lead only (this work) Contaminant appears in a drinking water source (or a source likely to be used for drinking such as groundwater in a setting reported to rely on groundwater for drinking water supply) Peer-reviewed publication, article, or book chapter written in English since 1969
Exclusion Criteria	 Duplicate Non-English Language Study is not based in LMIC, or the setting cannot be determined Does not provide a quantitative measure of any TM of interest (larger systematic review); of lead (this work) TM/lead is not measured in drinking water sources (or sources likely to be used for drinking, such as groundwater or surface water from a water body documented as being a source of drinking water for human settlements) Example: only measured in sources or food or diet Only measured in animals, i.e., animal studies Only measured in human blood, bone, or tissue, i.e., it is a human health assessment Only measured in prepared samples or laboratory solutions to which TMs have been added (e.g., in studies focused on the testing of remediation techniques)

- 613 Table 3. Precision of studies prioritized by supervised clustering (as a function of ensemble
- 614 score)

610

611

Ensemble Score	Precision
6	55%
5	48%
4	51%
3	42%
2	34%

	1	19%
615		
616		

- 617 Table 4. Precision of studies from supervised machine learning (for results returned in all search
- 618 updates) by predicted relevance group. Group 1 studies were predicted most likely to be
- 619 relevant by supervised machine learning; Group 5 studies were predicted least likely to be
- 620 relevant.
- 621

Precision Trend (Combined Search Updates)				
Relevanc e Probabili ty from ML	Grou p	Total Studies Screene d	Total Studies Releva nt (ti/ab)	Precisi on
Highest	1	1000	856	86%
	2	1000	632	63%
	3	1000	406	41%
	4	1000	237	24%
Lowest	5	717	70	10%

622

623

- 625
 - Table 5: Criteria for calculating method quality scores for included studies

Criterion	Rationale	Definition/Detail
Analytical methods used were described	If methods are not described, their suitability cannot be confirmed	Methods are described in sufficient detail to be reproduced
Analytical methods included the use of standards, blanks, and appropriate QC procedures	Use of these measures enables differentiation between true positive samples and contamination or imprecision	Use of field or laboratory blanks is mentioned in methods; instrument calibration is mentioned in methods
Laboratory instrumentation used was described	If instrumentation is not described, its suitability cannot be confirmed	Instrumentation is described in sufficient detail to confirm its suitability for TM analysis
Laboratory instrumentation was sufficiently sensitive to	If instrumentation offers precision insufficient to	Instrumentation offers a method detection level for TM

detect TMs of interest in drinking water at µg/L concentrations	distinguish TM concentrations that are above vs below WHO GV in samples, study cannot quantify GV exceedances	of interest that is < WHO GV. ICP-MS, ICP-OES, and GF- AAS generally provide adequate sensitivity for most TMs of interest.
The type of container used to collect water samples was described	If sample collection containers are not described, their suitability for TM sampling cannot be determined	Sample container materials are described in sufficient detail to assess their suitability for TM sampling.
The type of container used was adequate	Containers that are not manufactured, cleaned, or washed in such a way that they are likely to be trace- metal free may introduce contamination, preventing quantitation of TMs in water samples.	Containers made from virgin or acid-washed PP, HDPE, PTFE, glass, or other inert material are unlikely to introduce substantive TM contamination. Reused local beverage containers or unspecified containers may introduce such contamination
The sample handling procedure was described	If sample collection procedures are not described, their suitability for TM sampling cannot be determined	Sample handling procedures are described in sufficient detail to be reproduced.
The sample handling procedure was adequate (included sample preservation with appropriate acid)	Without proper sample preservation, TMs of interest may volatilize, precipitate, or sorb to container surfaces, leading to underestimation of their concentration in water samples.	Samples are collected at the water source and preserved with trace-metal free acid prior to transport and analysis. Typically, final pH should be <=2.0
Samples were not filtered prior to analysis	Filtration can remove particulate TM contamination	Methods do not mention filtration of samples prior to analysis
Stagnation period observed	Stagnation provides an exposure profile representative of first-draw water samples, which are typically of greatest concern when characterizing water- system derived TM exposure in drinking water.	For TMs classified by the Guidelines for Drinking-Water Quality as being commonly derived from plumbing and water system components (e.g., Sb, Cu, Pb), an additional point was added for studies that reported applying

	a stagnation period of any kind before sample collection
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627

- 628 Table 6: Numbers of included sample sets in low- and middle-income countries reporting
- 629 quantitative data on the occurrence of lead in drinking water, by country

Country	Frequency
Afghanistan	1
Albania	1
Algeria	6
Argentina	0
Armenia	0
Bangladesh	50
Benin	1
Bolivia	10
Bosnia and	1
Herzegovina	
Brazil	48
Burkina	5
Faso	
Cambodia	13
Cameroon	10
China	244
Colombia	0
Costa Rica	3
Cuba	0
Djibouti	3
Ecuador	3
Egypt	41
El Salvador	1
Ethiopia	54
Ghana	45
Guatemala	9
Haiti	12
India	334
Indonesia	2
Iran	107
Iraq	29
Jordan	16
Kazakhstan	4

Kenya	23
Kosovo	8
Kyrgyzstan	2
Lebanon	12
Macedonia	1
Malawi	4
Malaysia	9
Mali	1
Mexico	82
Moldova	8
Mongolia	2
Morocco	22
Mozambique	1
Myanmar	7
Namibia	12
Nepal	10
Niger	1
Nigeria	127
North	18
Cyprus	
Pakistan	215
Palestine	6
Peru	5
Philippines	1
Rwanda	1
Serbia	8
South Africa	33
Sri Lanka	19
Sudan	2
Tanzania	12
Thailand	37
Togo	1
Tunisia	2
Turkey	148
Uganda	2
Uzbekistan	1
Vietnam	20
Yemen	3
Zambia	4
Zimbabwe	20
Total	1943

633 Table 7: Numbers of included datasets by SDG Region

SDG Region	Number of Datasets	Percent
Central and Southern		
Asia	640	38.9
Eastern and South-		
Eastern Asia	339	20.6
Sub-Saharan Africa	302	18.4
North Africa and		
Western Asia	213	13.0
Latin America and the		
Caribbean	124	7.5
Europe and Northern		
America	26	1.6
Oceania	N/A	N/A
Total	2050	100

634

636

635 Table 8. Proportion of included studies that selected study sites based on presence of

Potential Source	n	Proportion
Wastewater	206	12.0%
Mining	198	11.6%
Industrial	197	11.5%
Agriculture	186	10.9%
Landfill	169	9.9%
Geogenic	116	6.8%
Corrosion	41	2.4%
Smelting	23	1.3%
Other	40	2.3%

637

- 638
- 639 Table 9. Analytical method used across all included datasets.

Analytical Method	n	Proportion
ICP-AES	321	26.8%
ICP-MS	817	68.2%
UV-Vis	40	3.3%
Electrochemical	16	1.3%
Other	4	0.3%

640

Table 10. Study Method Quality Score ("high quality" study = a score of 7 out of 10 possible

643 method quality items.

644

Method Quality Score	n	Proportion
0	189	11.0
1	464	27.1
2	604	35.3
3	73	4.3
4	69	4.0
5	80	4.7
6	112	6.5
7	76	4.4
8	34	2.0
9	9	0.5
10	3	0.2

645

646

647 Table 11. Proportion of samples exceeding WHO Guideline Value for lead across all included 648 sample sets, for a) all included studies in which lead was quantified; b) the subset of those 649 ("nontargeted") studies in which the setting was not reported as having been purposively 650 selected due to a known or suspected specific source of contamination; c) the subset of those 651 "representative" studies that were "high-quality" ("high quality" study = a score of >=7 out 9 652 possible method study quality items); and d) the subset of those "nontargeted" studies in which 653 water samples were analyzed by ICP-MS (generally capable of differentiating between samples 654 exceeding vs conforming with GVs when used properly with suitable sample collection, 655 handling, analysis, and quality control). 656

Exceedances	Proportio	95% CI	95% CI	n
	n	(Lower)	(Upper)	
a) All studies	25.6%	23.5%	28.1%	1220
b) Nontargeted	23.7%	20.3%	27.1%	523
c) Nontargeted: high-quality	23.4%	20.1%	26.8%	521
d) Nontargeted: ICP-MS	18.9%	15.2%	22.6%	368
e) Nontargeted: ICP-MS, ex-PRC	22.7%	18.0%	27.4%	264

657

658

Table 12. Median concentration of lead across all sample sets for which values could be

660 estimated

Dataset type	Conc (μg/L)			n
a) All studies	50	36	65	854
b) Nontargeted	27	14	40	317
c) Nontargeted: high-quality	*	*	*	*

d) Nontargeted: ICP-MS	24	11	37	251
e) Nontargeted: ICP-MS, ex-PRC	32	14	51	175

661 *Insufficient usable observations to estimate

662

Table 13. Proportion of samples from nontargeted studies exceeding WHO Guideline Value for

lead* across included sample sets by A) source type and B) SDG region**, by metaregression

Source Type	Proportion	95% CI	95% CI	n
		(Lower)	(Upper)	
Borehole	0.339	0.266	0.412	148
Piped	0.318	0.187	0.448	50
Surface	0.320	0.222	0.418	85
Spring	0.556	0.358	0.753	27
Wells	0.228	0.160	0.296	130
Other	0.272	0.176	0.369	68

665

Region	Proportio	95% CI	95% CI	n
	n	(Lower)	(Upper)	
SSA	0.358	0.246	0.469	73
NAWA	0.383	0.287	0.479	93
CSA	0.389	0.324	0.454	193
SEA	0.131	0.080	0.181	143
LAC	0.127	0.0	0.256	25
EUR	N/A	N/A	N/A	N/A

* As calculated by meta-regression using the "metareg" STATA command

667 **SSA: sub-Saharan Africa; NAWA: North Africa and Western Asia; CSA: Central and Southern

Asia; SEA: Eastern and South-east Asia; LAC; Latin America and the Caribbean; EUR: Europe

669

- 671 List of Table and Figure Captions
- Table 1. Toxic metals and metalloids included in the larger systematic review of TM
- 673 contamination in drinking water in low- and middle-income countries; this study reports only on
- 674 lead
- Table 2: Inclusion and exclusion criteria for study selection
- Table 3. Precision of studies prioritized by supervised clustering (as a function of ensemblescore)
- Table 4. Precision of studies from supervised machine learning (for results returned in all search
- 679 updates) by predicted relevance group. Group 1 studies were predicted most likely to be

- relevant by supervised machine learning; Group 5 studies were predicted least likely to be
- 681 relevant.
- Table 5: Criteria for calculating method quality scores for included studies
- Table 6: Numbers of included sample sets in low- and middle-income countries reporting
 quantitative data on the occurrence of lead in drinking water, by country
- 685 Table 7: Numbers of included datasets by SDG Region
- Table 8. Proportion of included studies that selected study sites based on presence of
- 687 previously identified contamination (by source)
- Table 9. Analytical method used across all included datasets.
- Table 10. Study Method Quality Score ("high quality" study = a score of 7 out of 10 possiblemethod quality items.
- Table 11. Proportion of samples exceeding WHO Guideline Value for lead across all included
- sample sets, for a) all included studies in which lead was quantified; b) the subset of those
- 693 ("nontargeted") studies in which the setting was not reported as having been purposively
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- 695 "representative" studies that were "high-quality" ("high quality" study = a score of >=7 out 9
- possible method study quality items); and d) the subset of those "nontargeted" studies in which
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- exceeding vs conforming with GVs when used properly with suitable sample collection,
- handling, analysis, and quality control).
- Table 12. Median concentration of lead across all sample sets for which values could beestimated
- Table 13. Proportion of samples from nontargeted studies exceeding WHO Guideline Value for
- lead* across included sample sets by A) source type and B) SDG region**, by metaregression
- Fig 1. PRISMA flow diagram for a systematic review identifying publications on lead occurrencein drinking water in LMICs
- Fig 2. Machine learning in the systematic review screening process.
- Fig 3: Distribution of mean lead concentrations for drinking water sample sets from low- andmiddle-income countries, by water source type
- Fig 4: Distribution of mean lead concentrations for drinking water sample sets from low- andmiddle-income countries, by SDG region
- Fig 5: Distribution of mean lead concentrations for drinking water sample sets from low- andmiddle-income countries, by decade of study publication
- Fig 6: Distribution of mean lead concentrations for drinking water sample sets from low- and
- 715 middle-income countries, by SDG region (CSA: Central and Southern Asia; ENA: Europe and
- North America; ESEA: East and Southeast Asia; LAC: Latin America and the Caribbean;
- 717 NAWA: North Africa and Western Asia; SSA: Sub-saharan Africa). Red line denotes WHO
- 718 Guideline Value.

- 719 Fig 7: Distribution of mean lead concentrations for drinking water sample sets from low- and
- middle-income countries, by decade of study publication. Red line denotes WHO Guideline
- 721 Value.
- 722
- 723

- 724 Supporting Information Files
- 725 File S1 Checklist. PRISMA Checklist.
- File S1 Code. Code used for all calculations in this study (to be uploaded upon acceptance).
- File S1 Data. Cleaned data used for all calculations in this study (to be uploaded upon
- 728 acceptance).
- File S1 Protocol. PROSPERO protocol for this systematic review.
- 730 File S1 Studies. Table of all studies included in this systematic review.
- File S1 Text. Tables and figures presenting supporting information referenced in this study.
- Table S1. Search Strategy
- Table S2. Extracted Variables
- Table 3. Proportion of observations exceeding WHO GV for lead in drinking water by
 SDG region, including or excluding results from China.
- Table S4. Odds ratio for proportion of observations exceeding WHO GV for lead in drinking water by decade of publication (n=931)
- Fig S1. Bibliometric Results
- Fig S2. Precision of studies prioritized by supervised clustering (as a function of ensemble score)
- Fig S3. Precision of studies from supervised machine learning (for results returned in all search updates) by predicted relevance group. Group 1 studies predicted most likely to be relevant by supervised machine learning and Group 5 studies least likely to be relevant.
- Fig S4. Reported vs estimated proportion of exceedences for the subset of studies for
 which the proportion of exceedances of the WHO GV for lead was reported or could be
 extracted from available data.
- Fig S5. Diagnostic plots of the distribution of log-transformed median and mean lead
 concentrations for drinking water sample sets from low- and middle-income countries vs
 normal distribution for studies from a) median for all LMICs; b) mean for all LMICs; c)
 median for all LMICs ex-PRC; d) mean for all LMICs ex-PRC; e) median for studies
 from China (PRC); and f) mean for studies from China (PRC).
- File S1 Tools. Training tools and materials used to train screening and extraction teams in this study.
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- 757

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Fig 3





Fig 4





Fig 6





Fig. 1



Fig. 2