#### REVIEW

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# Vaccination dropout among children in Sub-Saharan Africa: Systematic review and meta-analysis

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#### ABSTRACT

Immunization is cost-effective preventive strategy for child morbidity and mortality. PubMed, Google Scholar, Scopus, Science Direct, and online institutional repository homes were searched. Data were extracted by Microsoft excel. Begg's rank test, and Egger's regression test was done. A pooled prevalence, Sub-group analysis, sensitivity analysis and meta-regression were conducted. A total of 12 articles were included in this study. The pooled prevalence of vaccination dropout was 26.06% (95% CI: 11.59, 30.53), I2 =91.2%. In sub-group analysis, Nigeria had the highest prevalence of immunization dropouts (33.59%). It was 18.01% and 29.25%, respectively, for published and unpublished research. Community-based studies and institutional-based studies also yield a prevalence of dropout 39.04% and 13.73% respectively. Dropout rate was 22.66% for sample sizes under 500 and 18.01% for sample sizes beyond 500. In Sub-Saharan Africa, the prevalence of vaccination dropout was high. Community education about vaccinations importance should be prioritized.

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children; Sub-Saharan Africa

# Introduction

Immunization is the practice of preventing an infectious pathogen from attacking the body by administering vaccines.<sup>1,2</sup> It is among the most economically advantageous public health strategies for reducing child morbidity and mortality.<sup>3</sup> Between 2010 and 2018, the measles vaccine avoided nearly 23 million fatalities.<sup>4</sup> More than half of child and young child mortality are caused by diseases that may easily be prevented by taking simple, cheap measures, like receiving vaccines.<sup>5</sup> Vaccine-preventable diseases continue to pose a public health danger in South-East Asia and sub-Saharan Africa because of insufficient and poor vaccination coverage.<sup>6</sup>

Pneumonia (12%), diarrhea (8%), injuries (6%), congenital anomalies(4%), malaria (5%) and measles (2%) are the leading causes of death in children under the age of five.<sup>7</sup> Vaccine-preventable diseases cost the lives of 8.8 million children under the age of five worldwide.<sup>8</sup> Vaccination is the most effective way to prevent infectious childhood diseases like measles, pertussis, diphtheria, tetanus, TB, meningitis, and tuberculosis in children.<sup>9</sup> The percentages of mortality brought on by pneumonia and diarrhea that can be avoided by vaccination are 59% and 29%, respectively.<sup>10</sup>

A significant number of children are not fully vaccinated despite a huge decline in the occurrence of vaccine-preventable mortality, which causes a major regional and global variation in vaccination coverage.<sup>11,12</sup> For instance, out of more over 17 million cases of measles in the world in 2017, there were

83,439 fatalities.<sup>13,14</sup> In 2018, a full three doses of the diphtheria-tetanus-pertussis (DTP) vaccine were given to 116.3 million children worldwide (86%).<sup>15,16</sup>

In 2019, 19.7 million children worldwide did not receive the third dose of the diphtheria, tetanus, and pertussis (DTP3) vaccination during the first year of life, which is a crucial sign of the effectiveness of immunization programs.<sup>17–19</sup> Of all the children who did not finish the three-dose DTP series, 6.2 million (31%) started it but never finished it.<sup>20</sup> Every year, 2–3 million children become ill and are vulnerable to diseases that can be prevented by vaccination, resulting in mortality, and about 22.4% of children fail to receive the DPT3 vaccine.<sup>21</sup> One in five infants around the world does not receive the three essential doses of the diphtheria, tetanus, and pertussis vaccine.<sup>22</sup>

Every year, 4.4 million children in sub-Saharan Africa die from communicable diseases that could be prevented by vaccination.<sup>23,24</sup> It is associated to inadequate vaccination Coverage, challenges and setup was not fully equipped in sub-Saharan Africa countries.<sup>25</sup>

The WHO's global approach known as the Immunization Agenda 2030 states that by the year 2030, every child should have received all recommended vaccinations, irrespective of their geographical region, age, socioeconomic status, or gender-related constraints.<sup>26</sup> The Expanded Program on Immunization (EPI) was established by the WHO in 1974 to control diseases that may be prevented by immunization.<sup>27</sup> After the launch of the EPI program, the

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number of fatalities among children under five remained at 5.3 million in 2018.<sup>28</sup>

Despite significant progress, the expanded immunization program still faces many obstacles. These are categorized as follows: policy, standards, and guidelines, human resources, management of vaccine, cold chain, and logistics, service delivery, communication and community partnerships; and sustainable financing.<sup>29</sup>

A review of the literature revealed various factors that were related to immunization dropout. Long travel distances to medical facilities, living in a rural region, and having a mother without a regular work were reported to be risk factors for immunization dropout in Nepal.<sup>30</sup> Children in other parts of Ghana were more likely to drop out of school if they lacked immunization records than those who did.<sup>31</sup> Last but not least, a randomized controlled experiment conducted in urban Pakistan found that receiving both a redesigned card and center-based education was expected to significantly enhance DPT3 completion compared to the control group.<sup>32</sup>

There are no data at the regional level, despite the fact that numerous primary researches have confirmed the percentage of dropout rate in Sub-Saharan Africa. Therefore, the goal of this systematic review and meta-analysis study was to identify the prevalence of dropout of vaccination among children in Sub-Saharan Africa. Clinicians and other stakeholders will be able to address gaps in immunization coverage by prioritizing and customizing immunization campaigns and operational plans based on the study's findings, which will provide them with the fundamental knowledge they need to provide every child with a life-saving vaccination.

#### Methods

#### Reporting

This systematic review and meta-analysis study was conducted to determine vaccination dropout among children in Sub-Saharan Africa using the standard PRISMA checklist guideline.<sup>33</sup> (Supplementary file 1).

# Search strategy

International online databases (Pub Med, Science Direct, Scopus, and Google Scholar) were used to search articles on the prevalence of dropout of vaccination in Sub-Saharan Africa. We also retrieved gray literature from Bangladesh University (Daffodil International University) and United States of America University (Walden University) and University of Ghana online research institutional repository. The search string was established using "AND" and "OR" Boolean operators. The following core search terms and phrases with Boolean operators were used to search related articles: ((((Dropout) OR ("Dropout" OR "failure to complete")) AND Immunization) OR ("Immunization" OR "Vaccine" OR "Vaccination" OR "Immunisation")) AND Children) OR ("Child" OR "Childhood" OR "Baby") AND Sub-Saharan Africa. Search terms were based on PICO principles to retrieve relevant articles through the databases

mentioned above. The searching period was from May 1/ 2022, to June 10/2022.

# **Outcome measurement**

The outcome variable of the current study was immunization dropout and it indicates that one has received the first recommended dose of vaccine and missed the next recommended dose. It was used for pentavalent vaccines (Diphtheria, Pertussis, Tetanus, Hepatitis B, and Hib) for the shots that require successive doses and was determined by asking yes/ no questions (1–5 doses). <sup>34</sup>

#### Inclusion and exclusion criteria

Only English language papers, including published and unpublished studies with full text available for research, and studies that took place in Africa were included in this meta-analysis. This systematic review and meta-analysis excluded research that used duplicated sources, qualitative studies, studies from developed nations, and articles without full text.

#### Quality assessment

Two authors (NAG and KDT) independently appraised the standard of the studies using the Joanna Briggs Institute (JBI) standardized quality appraisal checklist.<sup>35</sup> The disagreement raised during the quality assessment was resolved through a discussion led by the third author (GAA). Finally, the argument was solved and reached with an agreement. The critical analysis checklist has eight parameters with yes, no, unclear, and not applicable options. The parameters involve the following questions:

- (1) Where were the criteria for inclusion in the sample clearly defined?
- (2) Were the study subjects and, therefore, the setting described in detail?
- (3) Was the exposure measured result validly and reliably?
- (4) Were the main objective and standard criteria used to measure the event?
- (5) Were confounding factors identified?
- (6) Were strategies to affect confounding factors stated?
- (7) Were the results measured indeed and dependably? And (8) Was the statistical analysis suitable?. Studies were considered low risk when they scored 50% and above on the quality assessment indicators, as reported in a supplementary file (Supplementary file2).

# **Risk of bias assessment**

Two authors (NAG and GAA) independently assessed included studies for risk of bias through the bias assessment tool developed by Hoy et al,<sup>36</sup> consisting of ten items that assess four domains of bias and internal and external validity. Any disagreement raised during the risk of bias assessment was resolved through a discussion led by the third author (KDT). Finally, the argument was solved and reached with an agreement. The first four items (items 1–4) evaluate the presence of selection bias, non-response bias, and external validity. The other six items (items 5–10) assess the presence of measuring the bias, analysis-related bias, and internal validity. Therefore, studies that received 'yes' for eight or more of the ten questions were classified as 'low risk of bias.' If studies that received 'yes' for six to seven of the ten questions were classified as 'moderate risk' whereas studies that received 'yes' for five or fewer of the ten questions were classified as 'high risk' as reported in a supplementary file (Supplementary file 3).

#### **Data extraction**

Microsoft Excel spreadsheet (2016) and STATA version 11 software were utilized for data extraction and analysis. Two authors (NAG and BB) independently extracted all relevant data using a standardized Joanna Briggs Institute data extraction format. The disagreement raised during data extraction was resolved through a discussion led by the third author (GAA). Finally, the argument was solved and reached with an agreement. The data automation tool was not used due to this study's absence of the paper form (manual data). The name of the first author, year of publication, study country, study setting, study design, the prevalence of dropout of immunization, sample size, and quality of each paper was extracted.

# Data analysis

After extracting all relevant findings in a micro-soft excel spreadsheet, the data were exported to STATA software version 14 for analysis. The pooled prevalence of dropout of vaccination was computed using a 95% confidence interval. Publication bias was checked by funnel plot and more objectively through Begg and Egger's regression



Figure 1. PRISMA flow chart displays the article selection process for vaccination dropout in Sub-Saharan Africa.

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Table 1. Characteristics of the included studies in the systematic review and meta-analysis for the prevalence of vaccination dropout in Sub-Saharan Africa.

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Author/year	Country	Study setting	Study design	Sample size	prevalence	Quality
Ibrahm	Kenya	Community	Cross-sectional	216	17	Low-risk
Ayele MA	Ethiopia	Community	Cross-sectional	389	9	Low-risk
Samuel Errie Yeni	Nigeria	Institution	Cross-section	307	60.3	Low-risk
Ateudje et.al	Cameron	Institution	Cross-sectional	1430	50	Low-risk
Ayalew Kassaw	Ethiopia	Community	Cross-sectional	436	25.8	Low-risk
Robert Tewne	Ghana	Community	Cross-sectional	170	17.3	Low-risk
Bagune et.al	Ghana	Community	Cross-sectional	600	5.6	Low-risk
Chinawa et.al	Nigeria	Institution	Cross-sectional	972	6	Low-risk
Yehualashet SS	Ethiopia	Community	Cross-sectional	432	9.3	Low-risk
Powelson J	Mozambique	Community	Cross-sectional	32	19	Low-risk
Haji A	Kenya	Community	Cross-sectional	1116	12	Low-risk
Ibrahm Ali Mohamed	Somalia	Community	Cross-sectional	226	63.7	Low-risk



Figure 2. Forest plot displaying the pooled prevalence of vaccination dropout in Sub-Saharan Africa.

tests, with P < .05 indicating potential publication bias. The presence of between-study heterogeneity was checked by using the Cochrane Q statistic. This heterogeneity between studies was quantified using I<sup>2</sup>, in which a value of 0, 25, 50, and 75% represented no, low, medium, and high heterogeneity, respectively. A forest plot was used to visually assess the presence of heterogeneity, which presented at a high-level random-effect model was used for analysis to estimate the pooled estimate of dropout of vaccination. Subgroup analysis was done by sample size, country, publication, and study setting. Sensitivity analysis was executed to see the effect of a single study on the overall prevalence of the meta-analysis estimate. The findings of the study were presented in the form of text, tables, and figures

# Results

#### Search findings and study characteristics

One thousand one hundred six articles were retrieved using a search strategy about vaccination dropout in Sub-Saharan Africa through online search engines such as; PubMed, Scopus, Google Scholar, Science direct, and online research repository home. After removing duplicated studies, we obtained 701 studies selected for screening full title and abstracts. Of these, 600 studies were excluded due to title and abstracts and the remaining 101 articles were assessed for full text articles. After reviewing the full text, 89 articles were then eliminated because they contained qualitative studies, lacked full titles and abstracts, and reported findings from developed countries. Finally, 12 articles <sup>31–46</sup> with 6326 study participants were

Author/year		ES (95% CI)	% Weight
Kenya Ibrahim HA/2017 Haji A/2016 Subtotal (I-squared = 70.1%, p = 0.067)	•	17.00 (11.99, 22.01) 12.00 (10.09, 13.91) 13.94 (9.17, 18.72)	9.11 9.31 18.42
Ethiopia Ayele MA/2022 Ayalew Kassaw et.al/2022 Yehualashet SS et.al/un-pub Subtotal (I-squared = 96.1%, p = 0.000)	•	9.00 (6.16, 11.84) 25.80 (21.69, 29.91) 9.30 (6.56, 12.04) 14.57 (5.32, 23.82)	9.27 9.18 9.27 27.72
Nigeria samuel Errie Yeni/un-pub Chinawa et.al/2014 Subtotal (I-squared =80.7%, p = 0.000)	•	61.30 (55.85, 66.75) 6.00 (4.51, 7.49) 33.59 (-20.61, 87.78)	9.06 9.32 18.39
Cameroon Ateudje et.al/2020 Subtotal (I-squared = .%, p = .)	•	50.00 (47.41, 52.59) 50.00 (47.41, 52.59)	9.28 9.28
Ghana Robert Twene/un-pub Bagune et.al/2017 Subtotal (I-squared = 78.2%, p = 0.000)		17.30 (11.61, 22.99) 5.60 (3.76, 7.44) 11.13 (-0.32, 22.58)	9.04 9.31 18.35
Mozabique Powelson J./2022 Subtotal (I-squared = .%, p = .)		19.00 (5.41, 32.59) 19.00 (5.41, 32.59)	7.84 7.84
Overall (I-squared = 99.2%, p = 0.000) NOTE: Weights are from random effects analysis		21.06 (11.59, 30.53)	100.00
	<b>.0</b> 10		

Figure 3. Forest plot displaying sub-group analysis based on the country where the study was done.

included as criteria for this systematic review and metaanalysis study (Figure 1).

The cross-sectional study design was applied to all included studies. Three of these were cross-sectional studies conducted at institutions, while the remaining nine studies were community-based. Three studies conducted in Ethiopia,<sup>37,47</sup> two studies in Kenya,<sup>39,40</sup> two in Nigeria,<sup>42</sup> two in Ghana,<sup>31,43</sup> one study in Cameroon,<sup>44</sup> one study Mozambique <sup>45</sup> and one study in Somalia.<sup>46</sup> The sample size ranged from 32 to 1116. The prevalence of intention to use maternity waiting home ranged from 5.6% to 63.7%. All studies were assessed by using Joanna Briggs Institute (JBI) quality appraisal checklist and yielded low risk (Table 1).

## **Meta-analysis**

# Prevalence of dropout of vaccination in Sub-Saharan Africa

A DerSimonian and Laird random-effects model was used to determine the overall estimate of dropout of vaccination. Accordingly, the pooled prevalence of dropout of vaccination among children with a random-effects model was 21.06% (95% CI: 11.59, 30.53) with a heterogeneity index ( $I^2$ ) of 91.2% (p < 0.001) (Figure 2).

#### Subgroup analysis

Subgroup analysis was conducted based on country, setting, publication, and sample size because this meta-analysis demonstrated a notable degree of heterogeneity. Accordingly, Ghana had the lowest prevalence of vaccine dropout (11.13%; 95%CI: 0.32, 22.58;  $I^2 = 80.6\%$ ) while Nigeria had the highest (33.59%; 95%CI: 20.61, 87.78) (Figure 3). For community-based studies and institutional-based studies, the prevalence of vaccination dropout was (39.04; 95% CI:3.00, 75.09),  $I^2 = 69.7\%$ , and (13.73%; 95%CI:9.48,17.98),  $I^2 = 81.8\%$ , respectively) (Figure 4).

In studies with sample sizes higher than 500, the prevalence of vaccine dropout was 18.37% (95%CI: 1.70, 35.05);  $I^2 =$  89.7%; in studies with sample sizes less than 500, it was 22.66% (95%CI: 10.56, 34.76);  $I^2 =$  78.2% (Figure 5). In contrast to published studies, which show a rate of vaccine dropout of (18.01%), unpublished studies indicate a prevalence of



Figure 4. Forest plot displaying sub-group analysis based on study setting.

vaccination dropout of (29.25%); 95%CI: 1.69, 60.19), I<sup>2</sup> = 99.3 (Figure 6).

#### Heterogeneity and publication bias

To adjust the reported heterogeneity of this study ( $I^2 = 91.2\%$ ), we computed a Sub-group analysis based on country, study setting, sample size and publication. Univar ate meta-regression was also done to identify the source of heterogeneity using sample size and year as a covariate. It showed that no effect of sample size and year on heterogeneity between studies (Table 2).

The presence of publication bias was assessed by funnel plot visually and by Egger's test and Begg's test objectively. The funnel plot shows asymmetrical distribution of studies by visual inspection (Figure 7). Therefore, the presence of publication bias was also assessed by Egger's regression test (p = .081) and Begg's rank correlation test (p = .161) with no evidence of publication bias.

#### Leave –one-out-sensitivity analysis

A leave-one-out sensitivity analysis was carried out to detect the effect of each study on the overall prevalence of vaccination dropout by excluding one study at a time. As a result, studies omitted at a time did not show a significant change on the overall prevalence of vaccination dropout (Table 3).

# Discussion

Immunization is a cost-effective and affordable method to decrease maternal and child morbidity and mortality. Several immunogenicity studies demonstrated that pregnant women who received the vaccine developed protective antibodies against the disease.<sup>48</sup> Various evidences also showed that maternal death decline secondary to tetanus vaccine,<sup>49</sup> DTP vaccine,<sup>50-52</sup> and influenza vaccine during pregnancy.<sup>53</sup>

Pentavalent vaccines (a combination vaccine which protects against five killer diseases: diphtheria, pertussis, tetanus, hepatitis B and Hib) have been launched in by 2011.<sup>54,55</sup> The pentavalent vaccines provide a golden opportunity to curb Hib disease and hepatitis B along with (DPT) in the developing countries. If the vaccines are provided individually, the coverage of hepatitis B and Hib vaccines usually lags behind DPT coverage. This gap can be filled by using pentavalent vaccine in routine immunization programs.



Figure 5. Forest plot displaying sub-group analysis based on the sample size.

The first dosage of the pentavalent vaccine is used as a tracer indicator for subsequent doses (often the third). Low dropout rates imply good access to and utilization of vaccination services. The World Health Organization (WHO) suggested using the indicators of immunization dropout to be DTP1 to DTP3, BCG to measles-containing virus (MCV1), and MCV1 to MCV2.<sup>56</sup> A high dropout rate between Penta1 and the measles vaccination shows a service usage issue if an infant defaults to the three doses of the pentavalent vaccine.<sup>57</sup> If the dropout rate is greater than 10%, the World Health Organization (WHO) indicates that many people are not utilizing the services.<sup>58</sup>

Determining the overall prevalence of immunization dropouts among children in Sub-Saharan Africa was the goal of the current study. Thus, in Sub-Saharan Africa, the combined frequency of vaccination nonuse was 21.06% (95%.CI: 11.59, 30.53). This result is consistent with research conducted in Iraq (19.3%).<sup>59</sup> The socioeconomic similarities of the research areas where both studies were conducted in low- and middle-income countries might be a reason for the explanation.

The findings of this research is significantly higher than a study conducted in India,<sup>60</sup> which found that the dropout rates for BCG-DPT3, DPT1-DPT3, and BCG-Measles were 16.1%, 12.9%, and 8%, respectively. This might be brought on by differences in sample size, study participant makeup, health system architecture, and policy. For instance, the prior study had a total sample size of 550 individuals, but the current study had 6326 study participants. The time gap between the studies may also possibly provide an explanation for this disparity.

Comparatively, our study's findings were lower than those of research conducted in Nepal (28.5%)<sup>61</sup> and India (26.1%).<sup>62</sup> The study setting, time gap and the sample size across the studies of may be contributing factors to this variation. For instance, the current study was carried out on a sample size of 6326 whereas previous conducted on a sample size of 140 and 550 respectively.

Sub-group analysis was done based on country, study setting, sample size and publication. As a result, Ghana (11.13%) had the lowest prevalence of dropouts while Nigeria had the highest (33.6%).Studies conducted in the community (39%) revealed a higher rate of vaccination dropout than studies conducted in institutions (13.7%). This may be because women living in institutions have access to sufficient health information. Previous research that addressed attitudes and beliefs that might affect a mother's intention to vaccinate was supported by this conclusion. Healthcare professionals frequently serve as parents' major source of e2145821-8 👄 N. A. GEBEYEHU ET AL.



Figure 6. Forest plots display a sub-group analysis based on the publication.

Table
2.
Meta-regression
analysis
of
factors
affecting
between-study

heterogeneity.

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Heterogeneity source	Coefficient's	Standard error	p-value
Sample size	1.929488	1.016702	0.087
Year	-325.5629	288.3158	

vaccine information and have been demonstrated to have an impact on uptake.<sup>63,64</sup> Studies with fewer than 500 participants reported a rate of vaccination dropout of 22.7%, while studies with more than 500 participants had a prevalence of (18.37%).The prevalence of immunization dropout rate was 29.25% in unpublished research and 18.01% in published studies.

To handle a large variance that occurred in betweenstudy heterogeneity, a random-effect model was used in this research. We conducted leave-one-out sensitivity, and the results reveal that no single study had a substantial effect on the overall prevalence of incomplete immunization. We assessed the possible variability source via sub-group analysis using the study sub-region, sample size, publication and age of children. The high heterogeneity might be due to differences in the sample populations, paper qualities, or socio-cultural, ethnic, and regional differences. This study has some limitations. First, the study protocol was not registered. Second, articles were restricted to only being published in the English language. Third, all of the included studies were cross-sectional, which might affect the outcome variable because of other confounding factors. However, this research has also some strength. First, compressive electronic online international searching engines were used. Second, our review incorporated gray literature as part of the primary studies.

# Conclusion

In conclusion, Sub-Saharan Africa has a high prevalence of vaccine dropout. Vaccination dropout rates varied by country, study setting, publication, and sample size. As a result, Ghana had the lowest prevalence of vaccine dropout, compared to Nigeria's greater prevalence. The prevalence of immunization



Figure 7. Forest plot displaying the asymmetrical distribution of the included studies.

Table 3. The pooled prevalence of vaccination dropout in Sub-Saharan Africa when one study omitted from the analysis a step at a time.

Study omitted	estimate	95%Cl
lbrahim HA/2017	21.471	11.367–31.576
Ayele MA/2022	22.298	11.866-32.731
samuel Errie Yeni/un-pub	17.040	8.172-25.907
Ateudje et.al/2020	17.926	11.450-24.400
Ayalew Kassaw et.al/2022	20.585	10.515-30.655
Robert Twene/un-pub	21.438	11.375-31.500
Bagune et.al/2017	22.654	11.870-33.438
Chinawa et.al/2014	22.616	11.614–33.617
Yehualashet SS et.al/un-pub	22.269	11.792-32.746
Powelson J./2022	21.238	11.347-31.129
Haji A/2016	22.001	10.888-33.115
Ibrahim Ali Mohamed/un-pub	21.062	11.590-30.534
Combined	21.062	11.590-30.534

dropout was also shown to be greater in community-based studies and unpublished research. Studies with sample sizes under 500 also showed a higher occurrence.

To prevent immunization dropout, community education about the value of vaccinations should be prioritized.

# **Disclosure statement**

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# **Authors' contributions**

NAG conceptualized the study: NAG and KDT contributed during data extraction and analysis: NA, KA and GA, wrote result interpretation: NAG, BB and KDT, Prepared the first draft: NAG, BB and GAA contributed during the conceptualization and interpretation of results and substantial revision: NAG, KDT,BB and GAA. Revised and finalized the final draft manuscript. All the authors read and approved the final version of the manuscript.

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# Data availability statement

All relevant data are within the Manuscript and its Supporting Information files.

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