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## Prevalence and risk factors of healthcare-associated infection in Ethiopia: a systematic review and meta-analysis --Manuscript Draft--

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<b>Abstract:</b>	<p><b>Background</b></p> <p>Healthcare-related infection is a global threat because it leads to the emerging of multiple drug resistance microbial infections. Survey or surveillance data is required before or after the patient being discharged from health institutions. Different primary studies in Ethiopia showed that the magnitude of healthcare-associated infection was varying in different area at the same and/or similar study period. Hence, this systematic review and meta-analysis estimated the national prevalence and risk factors of healthcare-associated infection in Ethiopia.</p> <p><b>Methods</b></p> <p>We searched PubMed, Science Direct, Google Scholar and Addis Ababa university repository. The quality status of studies was appraised by using Joanna Brigg's Institute quality assessment scale. We employed funnel plot and Egger's regression test to assess publication bias. The national prevalence of healthcare-associated infection was estimated using a weight-inverse random-effect model meta-analysis. Finally, we did subgroup analysis to resolve the cause of statistical heterogeneity.</p> <p><b>Results</b></p> <p>A total of 18 studies with 14,240 patients were participated for prevalence estimation. The pooled prevalence of healthcare-associated infection was 16.96% (95% CI: 14.10-19.82). Those patients who had surgical procedure (AOR=3.61; 95% CI: 2.71-4.50) and diagnosis of underlying non-communicable disease (AOR=2.59; 95% CI: 1.63-3.55) were statistically significant risk factors of healthcare-associated infection.</p> <p><b>Conclusions</b></p> <p>The national prevalence of healthcare-associated infection remains high in Ethiopia. This review may help policy makers and program officers to design healthcare-associated infection preventive strategies. Infection prevention and control practice should be strengthened for patients who had surgical procedure and underlying non-communicable disease.</p>
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# Prevalence and risk factors of healthcare-associated infection in Ethiopia: a systematic review and meta-analysis

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## Abstract

**Background:** Healthcare-related infection is a global threat because it leads to the emerging of multiple drug resistance microbial infections. Survey or surveillance data is required before or after the patient being discharged from health institutions. Different primary studies in Ethiopia showed that the magnitude of healthcare-associated infection was varying in different area at the same and/or similar study period. Hence, this systematic review and meta-analysis estimated the national prevalence and risk factors of healthcare-associated infection in Ethiopia.

**Methods:** We searched PubMed, Science Direct, Google Scholar and Addis Ababa university repository. The quality status of studies was appraised by using Joanna Brigg's Institute quality assessment scale. We employed funnel plot and Egger's regression test to assess publication bias. The national prevalence of healthcare-associated infection was estimated using a weight-inverse random-effect model meta-analysis. Finally, we did subgroup analysis to resolve the cause of statistical heterogeneity.

**Results:** A total of 18 studies with 14,240 patients were participated for prevalence estimation. The pooled prevalence of healthcare-associated infection was 16.96% (95% CI: 14.10-19.82). Those patients who had surgical procedure (AOR=3.61; 95% CI: 2.71–4.50) and diagnosis of underlying non-communicable disease (AOR=2.59; 95% CI: 1.63–3.55) were statistically significant risk factors of healthcare-associated infection.

**Conclusions:** The national prevalence of healthcare-associated infection remains high in Ethiopia. This review may help policy makers and program officers to design healthcare-associated infection preventive strategies. Infection prevention and control practice should be strengthened for patients who had surgical procedure and underlying non-communicable disease.

**Keywords:** Healthcare; Infection; Meta-analysis; Ethiopia

## Background

Healthcare-associated infection (HCAI) is preventable infection that a patient can encounter in healthcare facility while receiving medical care [1, 2]. It occurs after 48 hours of hospital admission, up to 3 days after discharge, or up to 30 days after operation when someone was admitted for reasons other than infection [2].

According to World Health Organization (WHO) 2019 HCAI fact sheet report, hundred million patients were affected each year globally. This report added that point prevalence of HCAI ranges 3.5-12% and 5.7-19.1% in developed and low-and middle-income countries, respectively [3]. The burden of HCAI was also reported in Sub-Saharan Africa [4] in general and specifically, in Botswana (13.4%) [5], South African (8%) [6], and Ethiopia (13 to 22%) [7-9].

Infection acquired in healthcare settings cause prolonged hospital stay, long term disability, increased microbial resistance to antibiotics, additional financial burden for health systems, high healthcare cost for patients and their families, and substantial mortality [10]. Hence, “*clean care the safer care*” program has been launched since 2004 with WHO patient safety directive, which was aimed to reduce HCAI through improving hand hygiene practice at the center of achieving



its aim [11]. Prevention guideline and infection prevention and control policy recommendations that was initiated by WHO have been implemented in Ethiopia.

Despite these efforts, studies done at different settings of the globe revealed that admission to surgical ward and hospital type [7], chest tube placement, prolonged hospital stays, patient on mechanical ventilation, previous hospitalization [8], pediatric patients, malnutrition and length of stay in hospital >5days [9] were contributing factors of HCAI.

Varieties of studies were conducted to determine the prevalence of HCAI in Ethiopia, but it showed great variation across geographical setting and variant time periods. Scattering on this, there was a need for nationally representative data on HCAI in the country. Moreover, the effect size of risk factors of HCAI weren't reported nationwide. Consequently, this systematic review and meta-analysis was aimed to estimate the national pooled prevalence and effect size of risk factors of HCAI in ~~Ethiopian~~.

## **Methods and materials**

### **Reporting**

The results of this review were reported based on the Preferred Reporting Items for Systematic Review and Meta-analysis statement (PRISMA) guideline [12] (Supporting file 1 PRISMA-P check list).

### **Inclusion and exclusion criteria**

Cross-sectional, case-control and cohort studies were included. Those studies had reported the prevalence and/or at least one associated factor of HCAI. Studies published in English language were considered. There was no restriction of the study period. Citations without abstract and/or full-text, anonymous reports, editorials, and qualitative studies were excluded from the analysis.

### **Search strategy and information source**

PubMed, Science Direct, Google Scholar and Addis Ababa University repository were searched. The core search terms and phrases were “prevalence”, “incidence”, “epidemiology”,

“proportion”, “magnitude”, “burden”, “associated factors”, “risk factors”, “predictors”, “determinants”, “healthcare-associated infections”, “healthcare-acquired infections”, and “nosocomial infections”, “hospital acquired infections” and “Ethiopia. The search strategies were developed using different Boolean operators. Notably, to fit advanced PubMed database, the following search strategy was applied on [February 5/2020](#): [(prevalence) OR incidence[MeSH Terms]) OR epidemiology[MeSH Terms]) OR proportion[MeSH Terms]) OR magnitude[MeSH Terms]) OR burden[MeSH Terms]) AND associated factors) OR risk factors[MeSH Terms]) OR predictors[MeSH Terms]) OR determinants[MeSH Terms]) AND healthcare-associated infections) OR healthcare-acquired infections[MeSH Terms]) OR nosocomial infections[MeSH Terms]) OR hospital acquired infections[MeSH Terms]) AND (Ethiopia)].

## **Study selection**

[Retrieved studies were exported to Endnote version 8 \(Thomson Reuters, London\) reference manager software to remove duplicate studies.](#) Two independent reviewers screened the title and abstract. Disagreement was handled based on established article selection criteria. Two independent authors conducted the abstract and full-text review.

## **Quality assessment**

Two independent authors appraised the quality of studies. The Joanna Briggs Institute (JBI) quality appraisal checklist was used [13]. Disagreement was resolved by interference of a third reviewer. The following items were used to appraise cohort studies: (i) similarity of groups, (ii) similarity of exposure measurement, (iii) validity and reliability of measurement, (iv) identification of confounder, (v) strategies to deal with confounder, (vi) appropriateness of groups/participants at the start of the study, (vii) validity and reliability of outcome measured, (viii) sufficiency of follow-up time, (ix) completeness of follow-up or descriptions of reason to loss to follow-up, (x) strategies to address incomplete follow-up, and (xi) appropriateness of statistical analysis. The items used to appraise case-control studies were: (i) comparable groups, (ii) appropriateness of cases and controls, (iii) criteria to identify cases and controls, (iv) standard measurement of exposure, (v) similarity in measurement of exposure for cases and controls, (vi) handling of confounders, (vii) strategies to handle confounder, (viii) standard assessment of outcome, (ix) appropriateness of duration for exposure, and (x) appropriateness of statistical

analysis. Cross-sectional studies were appraised based on: (i) inclusion criteria, (ii) description of study subject and setting, (iii) valid and reliable measurement of exposure, (iv) objective and standard criteria used, (v) identification of confounder, (vi) strategies to handle confounder, (vii) outcome measurement, and (viii) appropriate statistical analysis. All studies which got 50% and above on the quality scale were considered low risk.

## **Data extraction**

Two independent reviewers extracted data using a structured data extraction form. Whenever variations of extracted data observed, the phase was repeated. If discrepancies between data extractors continued, third reviewer was involved. The name of the first author and year, study region, study design, target population, diagnostic methods, sample size, prevalence of HCAI, and adjusted odds ratio (AOR) of associated factors were collected.

## **Outcome measurement**

HCAI was considered, when reported as infection(s) acquired while receiving medical care based on culture confirmation [9, 14-16], or clinical and laboratory methods [7, 17-30].

## **Statistical analysis**

Publication bias was visually checked by funnel plot and objectively using Egger's regression test [31]. Heterogeneity of studies was quantified using the I-squared statistic, in which 25, 50, and 75% represented low, moderate and high heterogeneity respectively [32]. Pooled analysis was conducted using a weighted inverse variance random-effects model [33]. Subgroup analysis was done by region, study design, diagnostic method and sample size. Sensitivity analysis was employed to see the effect of single study on the overall estimation. Besides, the time-trend analysis was conducted to check whether variation through time is observed. STATA version 11 statistical software was used for meta-analysis.

## **Ethics approval and consent to participate**

Not applicable because no primary data were collected.

## Results

### Literature search

The search strategy retrieved 611 articles from PubMed, 133 from Science Direct, 19 from Google Scholar and 3 from Addis Ababa university repository. After duplication removed, 680 remained. Sixty studies were screened for full-text review. Finally, 19 studies were used for meta-analysis (Figure 1).

### The characteristics of included studies

Six studies were found in Addis Ababa [17-19, 21, 23, 30], five studies in Amhara region [7, 14, 24, 25, 27], five studies in Oromia [15, 16, 26, 28, 29], one study both in Addis Ababa and Southern Nation Nationalities and People Region (SNNPR) [20], one each in Tigray [22] and SNNPR [9]. Nine studies were done across all age group. Eight studies were done on adult population and one study was done on pediatric patients. Fourteen studies used clinical and laboratory methods to diagnosis HCAI while the remaining were culture confirmed. Four studies were conducted by using cohort study design, fourteen were cross sectional and only one was case-control study. Only six studies included > 1000 sample size (Table 1).

**Table 1:** Characteristics and quality status of the studies

First author year	Study region	Study design	Sample size	Prevalence	Quality status
Gedebu M. et al/1987 [17]	Addis Ababa	Cross sectional	2506	13.40	<u>Low risk</u>
Gedebu M. et al/1988 [18]	Addis Ababa	Cross sectional	700	17.00	Low risk
Habte-Gaber E. et al/1988 [19]	Addis Ababa	Cohort	1006	16.40	Low risk
Berhe N. et al/2001 [20]	Addis Ababa and SNNPR	Cohort	247	5.90	Low risk

Endalfer N. et al/2008 [21]	Addis Ababa	Cross sectional	854	9.00	Low risk
Tesfahun Z. et al/2009 [22]	Tigray region	Cross sectional	246	27.60	Low risk
Endalfer N. et al/2011 [23]	Addis Ababa	Cross sectional	215	35.80	Low risk
Melaku S. et al/2012 [24]	Amhara region	Cross sectional	1383	17.80	Low risk
Melaku S. et al/2012 [25]	Amhara region	Cross sectional	1254	9.40	Low risk
Mulu W. et al/2013 [14]	Amhara region	Cross sectional	294	10.90	Low risk
Sahile T. et al/2016 [26]	Oromia region	Cross sectional	500	35	Low risk
Yallew WW. et al/2016 [7]	Amhara region	Cross sectional	908	14.90	Low risk
Tolera M. et al/2018 [16]	Oromia region	Cross sectional	394	6.90	Low risk
Gashaw M. et al/2018 [15]	Oromia region	Cross sectional	1015	11.60	Low risk
Ali S. et al/2018 [29]	Oromia region	Cohort	1069	19.40	Low risk
Alemayehu T. et al/2019 [9]	SNNPR	Cross sectional	939	21.40	Low risk
Gebremeskel S. et al/2018 [30]	Addis Ababa	Cross sectional	410	19.80	Low risk
Yallew WW. et al/2017 [27]	Amhara region	Case-control	545		Low risk
Zewdu et al/2017 [28]	Oromia region	Cohort	300	14.00	Low risk

**Note:** SNNPR: Southern Nations Nationalities and Peoples Region

## Quality of studies

The JBI quality appraisal criteria established for cross-sectional, case-control and cohort studies were used. The studies included in this systematic review and meta-analysis had no considerable risk. Therefore, all the studies were considered [7, 9, 14-30] (Table 1).

## Meta-analysis

### *Publication bias*

A funnel plot showed symmetrical distribution (Figure.2). Egger's regression test p-value was 0.328, which indicated the absence of publication bias.

### *Pooled prevalence of healthcare-associated infection*

A total of 18 studies with 14,240 patients were participated for prevalence estimation. The estimated overall prevalence of HCAI is presented in a forest plot (Figure 3). The overall prevalence of HCAI was 16.96% (95% confidence interval (CI): 14.10-19.82).

### *Subgroup analysis*

The subgroup analysis based on the region, study design, diagnostic method and sample size was done. Based on this, the prevalence of HCAI was found to be 27.6% in Tigray region, 18.2% diagnosed by both clinical and laboratory method, 17.83 % in the cross-sectional studies, 18.15% in studies using < 1000 study samples (Table 2).

**Table 2:** The pooled prevalence of HCAI, 95% CI and heterogeneity estimate with a p-value for the subgroup analysis

<b>Variables</b>	<b>Characteristics</b>	<b>Pooled prevalence (95% CI)</b>	<b>I<sup>2</sup> (p-value)</b>
<b>By region</b>	Addis Ababa	18.44% (14.02-22.86)	99% (0.000)
	Oromia	17.37% (9.2-25.56)	99.5% (0.000)
	Amhara	13.27% (9.00-17.52)	98.5% (0.000)

	Tigray	27.6% (25.37-29.83)	
	SNNPR	21.4% (20.3-22.5)	
	Addis Ababa & SNNPR	5.9% (4.25-7.55)	
<b>Study design</b>	Cross sectional	17.83% (14.39-21.27)	99.3% (0.000)
	Cohort	13.96% (8.78-19.14)	98.4% (0.000)
<b>Diagnostic method</b>	Clinical and laboratory	18.2% (14.85-21.51)	99.2% (0.000)
	Culture confirmed	12.71% (6.4-19.02)	99% (0.000)
<b>Sample size</b>	<1000	18.15% (13.28-23.03)	99.3% (0.000)
	≥1000	14.66% (11.72-17.59)	98.6% (0.000)

**Note:** SNNPR: Southern Nations Nationalities and Peoples Region

### *Sensitivity analysis*

Endalafer N et al [23] and Sahile et al [26] had shown an impact on the overall estimation of the meta-analysis result (Figure 4).

### *Time trend analysis*

The time-trend analysis showed that the prevalence of HCAI is increased from 13.4% in 1983 to 19.8% in 2017. However, the pooled prevalence from year to year is not increasing significantly (p-value = 0.620) (Figure 5).

### *Associated factors*

Based on this review, HCAI in Ethiopian context is associated with socio-demographic-related, patient and healthcare-related factors.

#### *Socio-demographic-related factors*

Male sex (AOR=2.06; 95% CI: 1.01-4.22) as compared to female and age < 1 year (AOR=8.53; 95% CI:2.67-27.3) as compared to age > 56 years [30], patient age ≥ 51 years (AOR=6.38; 95% CI:1.15-35.14) as compared to age 11-20 years [14], age between 1-14 years (AOR=0.25; 95%

CI: 0.09-0.71) [27] and age between 18-30 years (AOR=0.54; 95% CI: 0.32-0.93) as compared to age <18 years [29], were more likely to had HCAI.

### *Patient and healthcare-related factors*

~~Patients taking prophylaxis (AOR=1.76; 95% CI: 1.32-2.43) as compared to not taking prophylaxis [25], catheterized patients (AOR=18.9; 95% CI: 12.2-29.37) as compared to non-catheterized patients, duration of operation 90-150 minutes (AOR=11; 95% CI: 1.98-60.83) as compared to < 30 minutes on operation [14], admission to surgical ward (AOR=2.86; 95% CI: 1.72-4.78) and admission to Felege Hiwot referral hospital Ethiopia (AOR=1.99; 95% CI: 1.36-2.93) as compared to admission to Gondar hospital Ethiopia [7] were at higher risk of developing HCAI. Patients who had central vascular catheter (AOR=6.92; 95% CI: 1.28-37.47) as compared to not having vascular catheter, taking antimicrobial (AOR=8.63; 95% CI: 3.11-23.95) as compared to not taking antimicrobial and presence of medical waste container in rooms (AOR=0.18; 95% CI: 0.03-0.97) as compared to no medical waste container in room [27] were more likely to develop HCAI. Hospital stay < 5 days (AOR=0.03; 95% CI:0.01-0.1 as compared to stayed >21 days was associated with HCAI [30]. Patients who were on mechanical ventilator (AOR=1.99; 95% CI: 1.06-3.74) as compared to patients who were not on mechanical ventilator [29] showed increased risk of HCAI.~~

~~The pooled effect of four studies [14, 23, 27, 29] showed that surgical procedure (AOR=3.61; 95% CI: 2.71-4.50) was found positively associated factor, but previous hospitalization [29, 30] and hospital stay > 5 days [9, 14] had insignificant association with HCAI. The pooled effect of four studies [14, 25, 27, 29] showed that patients who had underlying disease (AOR=2.59; 95% CI: 1.63-3.55) as compared to patients with no underlying disease were more likely to had HCAI (Figure 6).~~

## **Discussion**

The Sustainable Development Goal (SDG) goal-3 is implementing to ensure access to quality essential healthcare service in every country [34]. However, HCAI is a major challenge of the SDG of infection prevention and control in health systems, which fail to build strong principles



to reduce the risks and spread of nosocomial infection [35]. HCAI is among preventable cause of morbidity, mortality and spending health system cost.

This meta-analysis estimated the national prevalence of HCAI and contextual associated factors in Ethiopia. Accordingly, the national pooled prevalence of HCAI was 16.96% (95% CI: 14.10-19.82). This result was higher than studies done in China (3.12%) [36], Morocco (10.3%) [37], Botswana (13.54%) [5] and south Africa (7.67%) [6]. The possible reasons for high prevalence in this study might be very low hand hygiene practice by physicians, low adherence to infection prevention practice, low level of job satisfaction, morally distressed nurses, resources constraints, low implementation of nursing process and less attention given to HCAI.

~~Perform hand hygiene before patient contact is vital to prevent HCAI. However, only 7% of physicians working at two university hospitals in Addis Ababa Ethiopia performed hand hygiene before patient contact [38], as a result acquisition of HCAI from these healthcare personnel might be high. Evidence also showed that 35% of nurses were non-adherent to infection prevention practice in southwest Ethiopia [39], high HCAI could be explained by this low adherence to infection prevention. Besides, nearly 68% of health professionals were less satisfied of their work in Amhara region [40], unsatisfied health professionals are less likely to deliver quality healthcare, so nosocomial infection was inevitably high. Additionally, about 84% of nurses were morally distressed in northwestern part of the country [41], as morally less prepared nurses were unable to deliver quality nursing care, consequently HCAI was found high.~~

Resource constraints could increase HCAI in the country, because lack of hand hygiene agents and sinks were reported as hindering factors of infection prevention practice in Addis Ababa Ethiopia [38]. Implementation of nursing process was below half (49%) in northwest part of the country [42], so nursing intervention would not be planned for patients at risk of nosocomial infection, hence HCAI was found higher. Furthermore, healthcare providers, patients and/or families are more curious for primary reason of admission or healthcare visit, so less attention is given for HCAI which could justify high prevalence of HCAI in the country.

Moreover, the time trend analysis showed that HCAI was slightly increasing in Ethiopia. The possible reasons might be more emphasis given for healthcare coverage than quality, increase in

technological advancement and over utilization of invasive procedures. Evidence revealed that advance in life saving medical practices increase exposure to invasive procedures, which increase the occurrences of nosocomial infections [43]. On top of this, nurse's burnout might contribute for increasing trend. Evidence in United states (US) revealed that nurse's burnout was found as a single most important associated factor for increased nosocomial urinary tract infection (UTI) and surgical site infection (SSI) [44].

The current meta-analysis revealed that patients who had surgical procedure and underlying non-communicable disease were more likely to acquire HCAI.

Patients who had surgical procedure were associated with nearly four times increased odds of HCAI in this study. The finding is in line with previous studies done in south Africa [6] and Poland [45]. The reason for observed association could be explained by less compliance to hand hygiene practice, and high prevalence (25.22%) of surgical site infection [46] in Ethiopia. Compliance to hand hygiene practice is pivotal for prevention and control of nosocomial infection, but only 3.6% keep hand hygiene before performing aseptic procedures at Debre Birhan referral hospital, central Ethiopia [47]. The overall compliance to hand hygiene practice was 18.7% [48] at Hiwot Fana specialized and 22% [47] at Debre Birhan referral hospitals of Ethiopia in 2017. Consequently, unable to keep and maintain hand hygiene practice increases acquisition of HCAI.

The odds of patients having underlying non-communicable disease were nearly three times more at risk to develop HCAI. This finding is supported by studies that reported HCAI was positively associated with diabetes mellitus [49] and underlying renal disease [50]. The possible explanations for observed association in the current study might be due to high prevalence of underlying diseases and immunosuppression effects of these diseases. A meta-analysis studies showed that high prevalence (6.5%) of diabetes mellitus in Ethiopia [51] and immunosuppression was identified as a risk factor for HCAI [49].

## **Strength and limitation**

This systematic review and meta-analysis was the national estimation conducted in Ethiopia. It may lack national representativeness because no data were found from Benishangul Gumuz, Afar, Gambella, Somalia, Dire Dawa, and Harare regions of Ethiopia. Time-trend analysis might not reflect the exact trend because all the years didn't have reported data.

## **Conclusions**

The prevalence of healthcare-associated infection in Ethiopia remains high. This review may help policy-makers and program officers to design interventions on preventing healthcare-associated infection. The existing infection prevention and control practice for patients who had surgical procedure and underlying non-communicable disease should also be strengthened.

## **Abbreviations**

AOR: Adjusted Odds Ratio; CI: Confidence Interval; HCAI: Healthcare-Associated Infection; LMIC: Low and Middle Income Country; SNNPR: Sothern Nations Nationalities and Peoples Region; WHO: World Health Organization

## **Acknowledgments**

Not applicable

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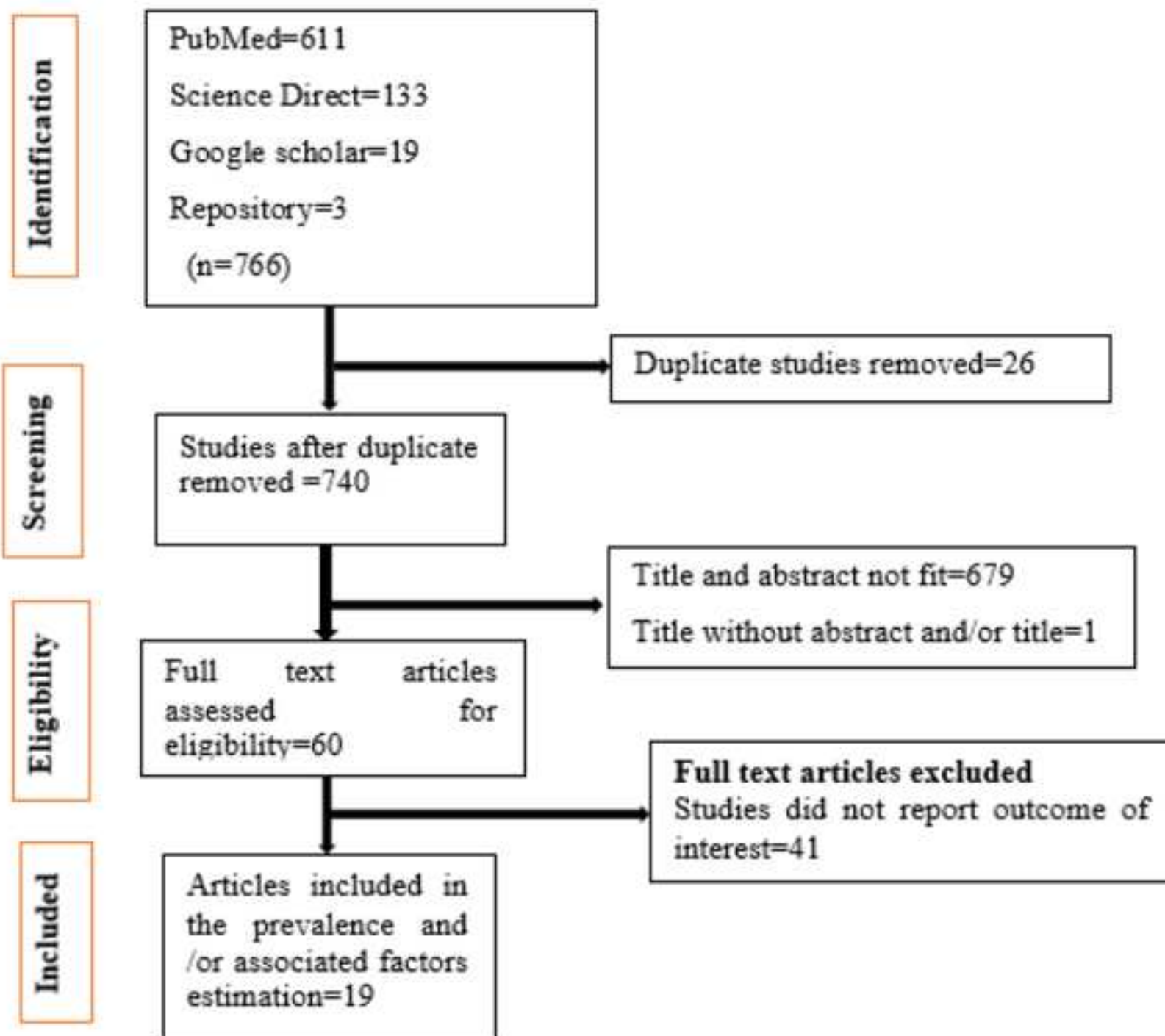
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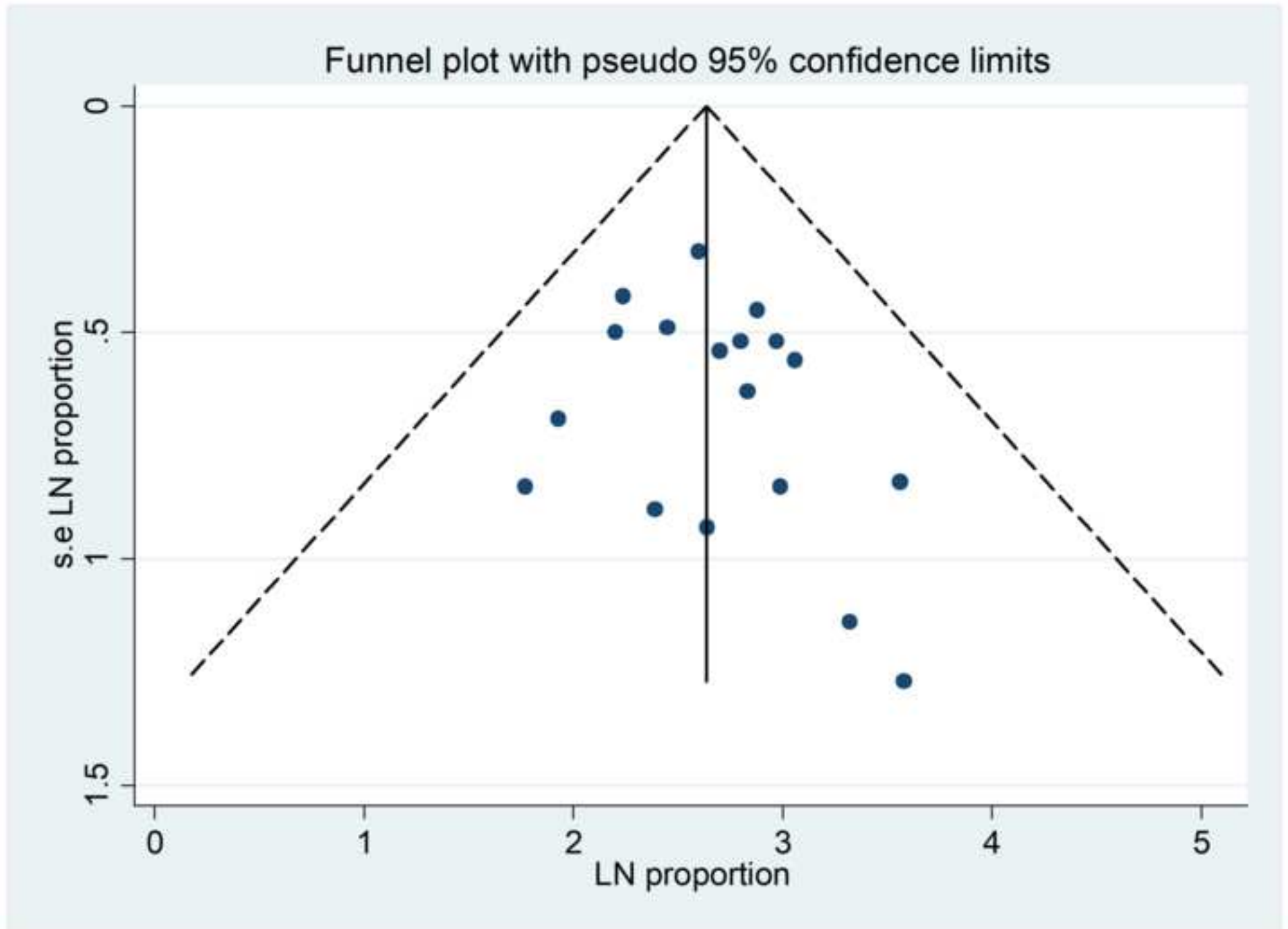
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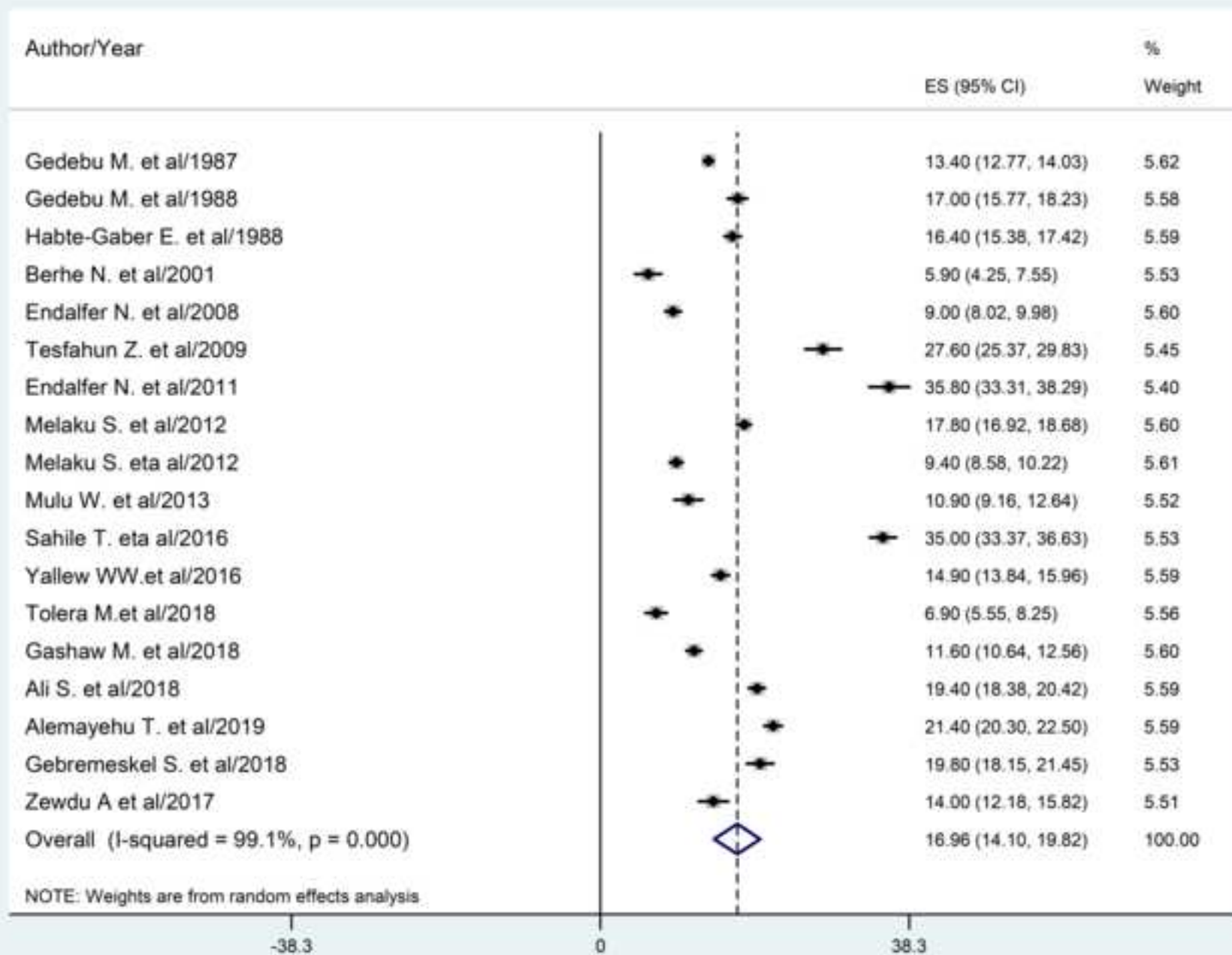
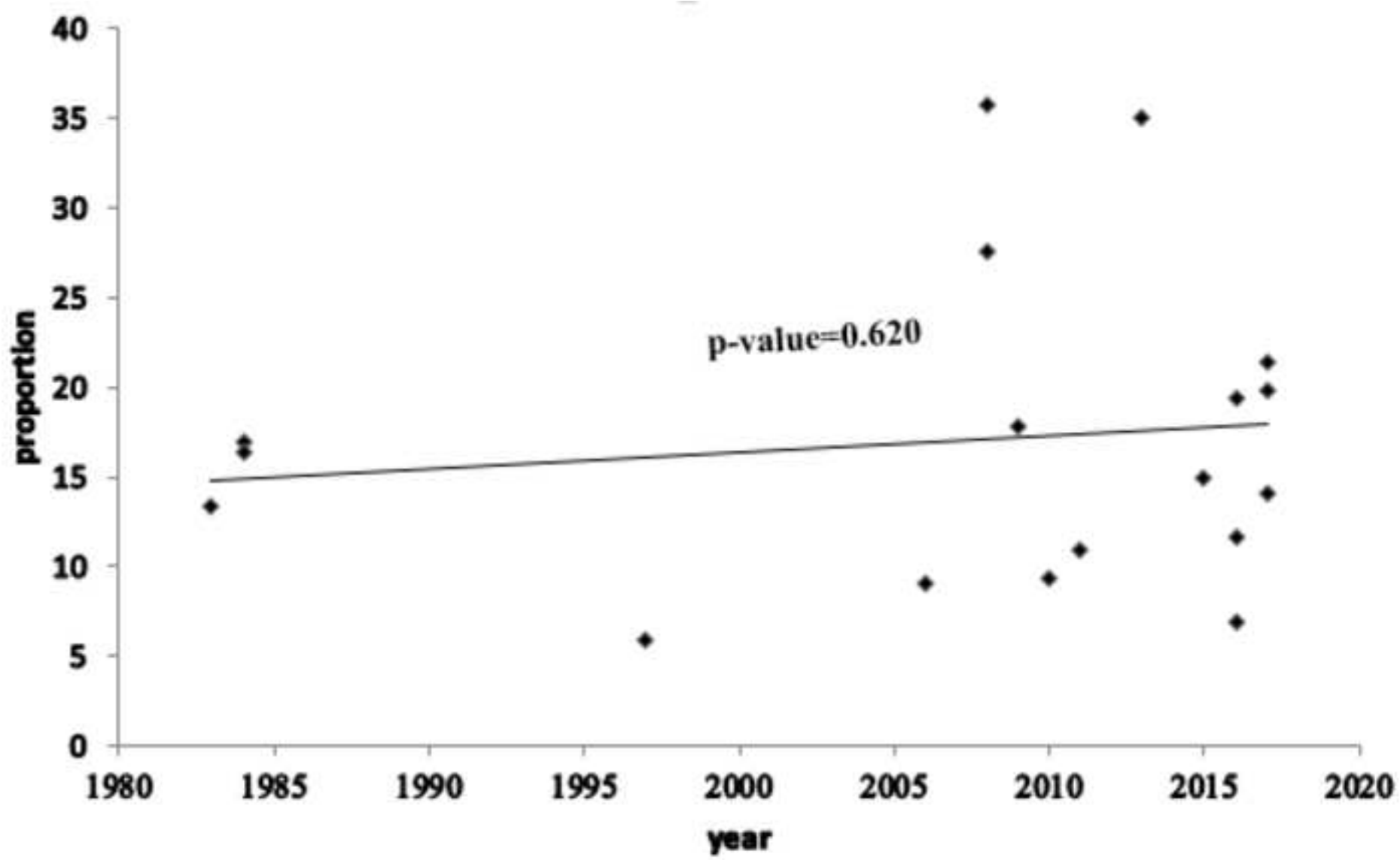
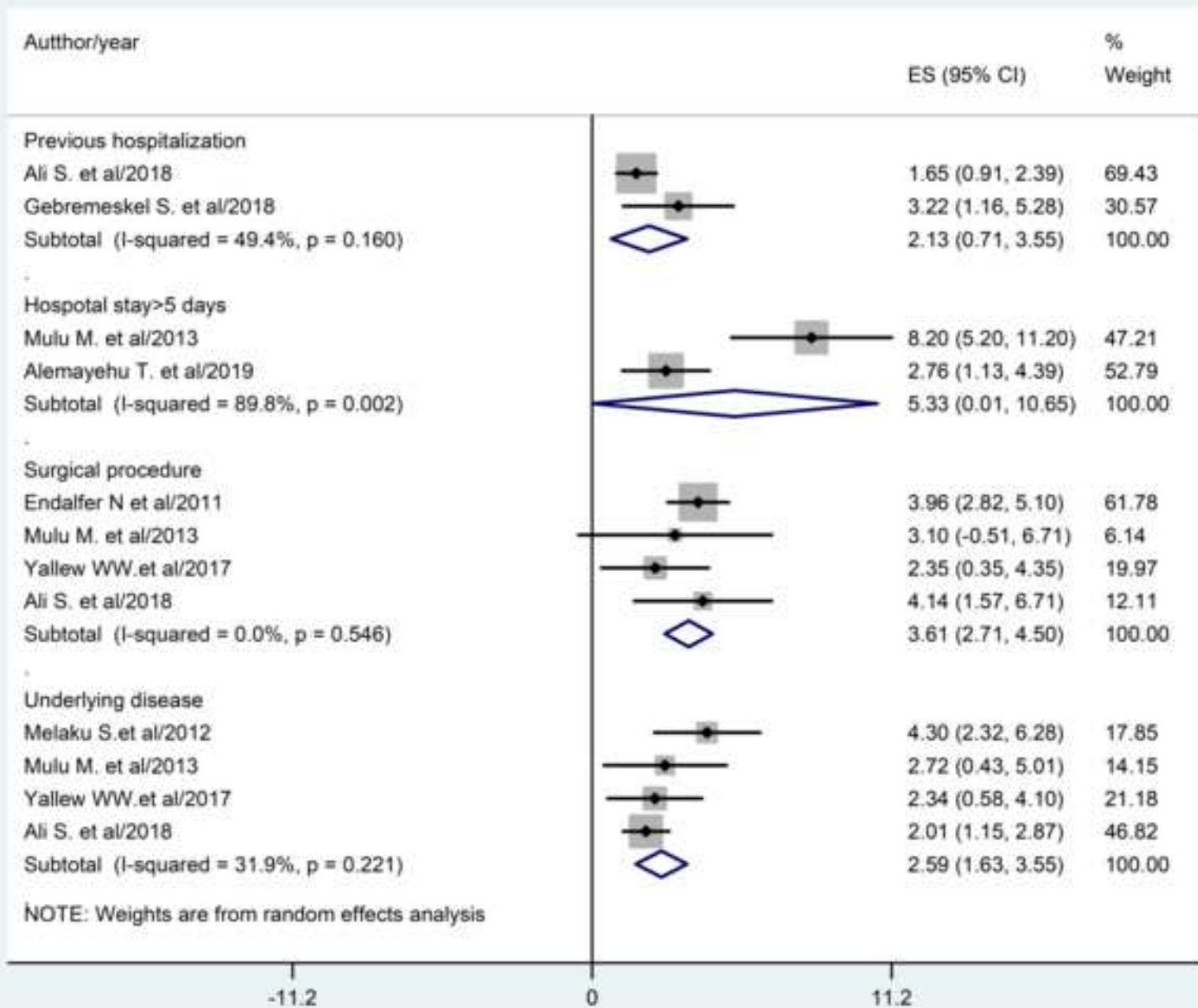




Figure 5









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