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7	Female Genital Tuberculosis Among Infertile Women and Its Contributions to
8	Primary and Secondary Infertility
9	A systematic review and meta-analysis
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22	
23	Abstract
24	Female genital tuberculosis (FGTB) is an infectious widespread disease among young women. This
25	meta-analysis study aimed to investigate the prevalence of Female Genital Tuberculosis among infertile
26	women and its contribution to primary and secondary infertility. A PubMed, MEDLINE, world cat log,
27	Lens.org, direct Google search, Google Scholar, and Researchgate, from 1971 to July 17, 2021, were
28	searched using the keywords; prevalence, epidemiology, urogenital tuberculosis, FGTB, infertile
29	women, infertility complaints, and FGTB testing methods. Data extracted and meta-analysis was

- performed. 42 studies were selected with a total of 30918 infertile women. Of these, the pooled
- 31 prevalence of FGTB was 20% (15-25%; 95%CI; I2 99.94%), and the prevalence of overall infertility,
- 32 primary infertility, and secondary infertility among FGTB-population were 88%, 66% and 34%,

- respectively. The proportion of FGTB is remarkable among infertile women globally. The biggest
- burden of the disease is presented in the low-income countries followed by the lower middle-income,
- 35 and upper-middle-income countries.
- *Keywords:* Female Genital Tuberculosis, Infertile women, Worldwide, Prevalence of FGTB, Infertility,
 Infertility Complaints, primary infertility, secondary infertility.
- 38

39 Introduction

- 40 Tuberculosis (TB) is an infectious disease caused by Mycobacterium Tuberculosis which is recently
- 41 listed among the top ten diseases causing death around the world. According to the World Health
- 42 Organizations (WHO), in 2019 TB was responsible for 10.0 million infections and 1.2 million people
- 43 death.¹ The two-third of this global burden presented in eight countries included; India, Indonesia,
- 44 China, the Philippines, Pakistan, Nigeria, Bangladesh, and South Africa.¹ Female Genital Tuberculosis
- 45 is commonly secondary to pulmonary TB (PTB) or extrapulmonary TB (EPTB), with the incidence rate 46 ranging between 9 to 20 and 5 to 13 among overall EPTB.^{2,3} and PTB.^{4,5} cases worldwide⁶.
- 47 respectively. Typically female genital Tuberculosis (FGTB) is known as the disease of young women
- 48 $(20-40 \text{ year-old})^{5,7}$, and it is usually diagnosed during infertility evaluations.^{2,8} A previous study
- 49 indicated that the infertility rates in women is higher compared to men⁹. Moreover, 76% of infertile
- 50 women had a history of TB^{10} , and infertility is the most frequent complaint of FGTB cases¹¹ which
- 51 occurs due to the irreversible damage to the fallopian tube.⁴ In addition to infertility, other clinical
- 52 presentations of FGTB include pelvic pain or menstrual irregularities, and its remains a major health
- problem in low-income countries^{8,12} Organs commonly affected by FGTB are the fallopian tube (90%),
- 54 ovaries (10–30%), endometrium (50%), cervix, and vagina.^{3,13} Infertile FGTB patients have been
- ⁵⁵ reported to have longer duration of infertility compared to infertility from other courses.¹⁴
- This meta-analysis study was conducted to investigate the prevalence of FGTB among infertile women
 of reproductive age and to evaluate the incidence of primary and secondary infertility among FGTB
 patients around the globe.
- 59

60 Methods

61 *Eligibility criteria*

62 Studies were eligible if they; characterized the epidemiology of FGTB among women within 63 reproductive age, if the study population were infertile women or at least indicated a proportion of 64 infertility complaints with enough explanation of epidemiology of FGTB, published in English, the 65 study published in period between 1971 to 17 July 2021, and the diagnostic methods of FGTB was 66 done based on the particular infertility centres testing protocol. Whereas studies were excluded; if

articles characterized only PTB or EPTB regardless of FGTB, and any study in which the prevalence of
FGTB reported was not that of infertile women.

69

70 Information sources

This study was carried out in accordance with the guidelines of Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA). Several electronic databases such as; MEDLINE, world cat log, Lens.org, and PubMed were used to retrieve published articles. In addition, other search engines were intensively searched including direct Google search, Google Scholar, Researchgate retrieve studies that were not indexed in PubMed. All mentioned databases were searched from their commencement in period between 1971 to July 17, 2021, for human studies published in English.

77

78 Search strategy

The Boolean search terms (AND, OR) were used to develop the research strategy to retrieve studies from PubMed and world cat log. The final search strategy included the use of Title/Abstract related to (((Female genital tuberculosis) OR (urogenital tuberculosis)) AND ((prevalence) OR (epidemiology)) AND (infertile women) OR (infertility)) taken from the study objectives. Hand intensive searches were applied in direct Google search, Google Scholar and Researchgate for the same purpose.

84

85 Study selection process

In this study, all retrieved articles were first screened by title, abstract, and full-text screened. Then 86 eligible articles exported the Mendeley citation manager software version 1.19.8, to be checked for 87 duplication. Therefore, the duplicated articles were excluded from the study. Two authors (AA, & MA) 88 screened and evaluated the remaining studies independently by a careful reading of the title and 89 abstract then full-text articles screened if the particular records mentioned the outcomes of the review 90 91 "Prevalence of Female genital tuberculosis among infertile women" in their titles and abstract. However, the screened full-text articles were considered for further evaluation based on the objectives, 92 methods, participants, and key findings. The two authors (MA & AA) independently evaluated the 93 quality of the studies against PRISMA checklist.¹⁵ any inconsistency for the included articles was 94 95 resolved through discussion, and by consulting an expert. The overall study selection process is presented using the PRISMA statement flow diagram Figure 1. 96

97

98 Data collection process

99 The relevant data from selected articles were extracted by three investigators independently (AA, MA, & SH) using a data extraction template through Microsoft word 2016. The extracted key points 100 included author name, year of publication, reference, study country, study design/setting, sample size, 101 102 FGTB proportion among infertile women, the prevalence of overall infertility, primary infertility, and secondary infertility among FGTB cases (Table 1). The data extraction accuracy was verified by 103 104 comparing the data extraction results from the second group of investigators (AB, AI, & CA), who 105 independently extracted the data in a randomly-selected subset of papers (30% of the total). The extracted quantitative data were summarized in a Microsoft Excel sheet. The prevalence of FGTB 106 among infertile women and prevalence of pooled infertility (primary & secondary) among FGTB cases 107 were conducted by STATA software version 16. 108

109

110 Data items

The main outcome of this study was the prevalence of FGTB among infertile women within 111 reproductive age worldwide, and it is measured by the direct report from the individual studies. Out of 112 these, 26 studies from India, 3 studies from Nigeria, 2 articles each from Ethiopia, South Africa and 113 Pakistan were retrieved. Also, only one each article was retrieved from Egypt, Iraq, Iran, United State 114 of America, Saudi Arabia, Sudan, and Yemen. To quantify the outcome, the investigators considered 115 studies that reported the prevalence of FGTB among infertile women and the types of tuberculosis 116 regarding FGTB among gynaecology admitted/infertile women in their statistics. The result was 117 interpreted by the proportions of the infertile population which is having any type of FGTB from the 118 total population studied. 119

120

121 Study risk of bias assessment

Inclusion criteria were appraised for all retrieved articles by using their title and abstract then, full-text 122 articles were screened to check the quality of each study before the final selection. The quality 123 assessment criteria for the studies included in the current meta-analysis and systematic review defined 124 as follows: The diagnosis of the infertility cases were performed at infertility center with consideration 125 126 that infertility is defined as a one year without conception after unprotected intercourse, the infertility was not due to male factor, the diagnosis including an infertile population who tested for FGTB 127 willingly, the diagnosis of FGTB were conducted after excluding the patients with a confirmed FGTB, 128 and finally, the sample size was representative of the population. A comprehensive search included 129 electronic database, manual and grey literature, and unpublished studies was done to manage and 130 minimize the risk of bias. Moreover, two groups of investigators (AA, MA, & SH) and (AB, AI, & CA) 131

- used Joanna Briggs Institute Quality Assessment Tool as a critical appraisal tool for the same
- 133 purpose¹⁶. The differences in the inclusion of the studies were resolved by consensus. The included
- studies were evaluated against each indicator of the tool and categorized as high-, moderate-, and low
- 135 quality. Studies with a score greater than or equal to 60% were included. The publication bias for the
- included studies was checked by both the visual inspection of the funnel plot and check the statistical
- 137 symmetry of the funnel plot using Egger's Regression Test.
- 138

139 *Summary measures*

- 140 From the standpoints of the study objectives, the proportion of FGTB among infertile women,
- 141 proportion of the type of infertility among FGTB patients were used to synthesize and present the
- 142 results for the analysis.
- 143

144 Synthesis methods

The collected data were synthesized and analysed by using the Stata software, version 16.0 (Stata Corp LLC, 77845 Texas, USA). The recommendations of the I² statistic described by Higgins et al.¹⁷ (an I² of 75/100% and above suggesting considerable heterogeneity) were used to perform this meta-analysis. The effect size, with a 95% confidence interval (CI) and standard error (SE), was used to calculate the result of this study. The effect size of this study was the prevalence of FGTB and the prevalence of the type of infertility subgroups, and they were calculated using the binomial distribution, while the SE, was calculated using the sample size (n) and the proportion of FGTB (p), and applied it one SE

- 152 formula: $\sqrt{P(1-P)} \div n$.
- 153

The potential publication bias was checked using a funnel plot, and Egger's Regression Test, and it was assumed to be significant if the P-values were less than 0.10. Subgroup analysis was applied to check the potential source of heterogeneity and possible source of bias. Any studies that had missing data and/or a risk of bias were excluded. Any study has a missing data and/or a high risk of bias were excluded. The study results were reported according to the PRISMA guidelines and the findings were presented using a narrative synthesis followed by a meta-analysis chart.

160

161 **Results**

162 Study selection

163 A total of 1203 records were identified through the major utilized databases and other relevant sources.

164 Of these 961 records were removed due to duplication and title screening, while 242 records studies

were kept for further conclusive inspection. Then another 180 records were excluded after a very

- 166 careful screening of abstracts. However, a total of 62 articles were eligible for full-text screening, 20
- articles of them were excluded due to inconsistency with the study inclusion criteria. Finally, 42
- records were fulfilled the eligibility criteria, involving 30918 participants with mainly infertility
 complaints, were included for the systematic review and meta-analysis. Figure 1 showed the selection
 process of the studies selected for the meta-analysis.
- 171

172 Study characteristics

A total of 42 studies including 30846 participants were included in the quantitative analysis for this 173 meta-analysis review study; 2 (4.8 %) were from High-income countries, 4 (9.5%) from Upper middle-174 175 income countries, 32 (76.2%) from Lower middle-income countries, and the remaining 4 (9.5%) were from the Low-income countries. Of the total included studies, 17 were cross-sectional studies, 13 were 176 prospective study design, and 12 were retrospective studies. Included studies were conducted between 177 1971 to 2021. The majority of them were hospital admitted patient settings and the most used 178 diagnostic test was only PCR or PCR combined with other relevant test methods. Table 1 showed the 179 detailed characteristics of all included studies. 180

181

182 Synthesis of results

This meta-analytical study showed that; out of 1203 retrieved records, only 42 records were included 183 and analyzed. Of these a 20% (CI 15% to 25%) pooled prevalence of FGTB among infertile women out 184 of overall study sample 30846 participants worldwide. Residual heterogeneity was high with p-value <185 .001, I^2 99.94% and ch²(2553.37). for this analysis, the random effect model was employed (Figure 2). 186 However, of 42 records only 5, 15, & 14 articles analyzed to evaluate the pooled prevalence of overall 187 infertility, primary and secondary infertility among FGTB patients respectively which were provided 188 an 88% (CI 74%-100%; I²: 99.91), 66% (CI 56%-76%; I²: 99.23), and 34% (CI 24%-43%; I²: 98.04), 189 with p-value < .001; respectively (table 2., Section A.). Also, the random effect model was applied 190 because the heterogeneity was substantially high, with P-value < .001. the publication bias was checked 191 192 by using the funnel plot of the forest plot, and the plot was visually symmetric with Egger's test (pvalue 0.25). 193

194

Due to the very high heterogeneity level presented in FGTB among infertile women analysis, a two-

subgroup analysis was performed to check the effect of the study's publication year and the World

197 Bank Economical Country Classification on the pooled prevalence of FGTB among the infertile

198 population (Table 3). The included studies were divided as the particular country classified; High

- 199 income, Upper middle-income, Lower middle-income, and Low-income countries groups. The
- analyzed data showed that the lower country economies is the highest pooled prevalence of FGTB, and

the highest income countries have the lower pooled prevalence of FGTB among infertile women. The

202 results presented as; 5.7% (I^2 78.56%), 14% (I^2 86.9%), 21% (I^2 99.95%), and 24% (I^2 99.48%) for high

- 203 income, Upper middle-income, Lower middle-income, and Low income countries, respectively (table
- 204 2., section B.)
- 205

Meanwhile the objective was to evaluate the effect of the study's publication year on the pooled
prevalence of FGTB among infertile women (table 2., Section C.) The included articles were divided
into three groups, and the results indicated an; 10%, 23%, & 22% pooled prevalence of FGTB among
infertile women for period before 2000, between 2001 to 2010, and between 2011 to 2021 study's
publication year subgroups, respectively (table 2., Section C.)

211

212 **Discussion**

Although men are significantly having the biggest burden of TB compared to women,¹⁸ in 2018, WHO estimated that 3.2 million women were infected with TB, and the disease is accompanied with severe consequences especially in women of reproductive age.¹⁸ Although, FGTB rarely occurs in developed countries³, it represent an important cause of infertility in developing countries especially in countries with high TB-incidence rates.¹⁸

218

Recently, many published studied have investigated the prevalence of FGTB among infertile women of 219 reproductive age which is showed that the lowest prevalence was 0.45% in Nigeria¹⁹ and the highest 220 prevalence was 52% in India.²⁰ Worldwide, the prevalence was 24.2% in the first published meta-221 analysis and systematic review in 2016.²¹ However, the current study finding is 20% which is slightly 222 decreased. This outcome is due to the relative progress in the availability of more sensitive TB 223 diagnosis methods such as GeneXpert and PCR in developing countries. Moreover, the relative 224 225 increase in number of TB healthcare services and many countries have adopted the WHO's END TB STRATEGY around the globe.²² 226

227

In the current comprehensive research finding, the prevalence of FGTB among infertile women
progressively increased over time to be 10%, 23%, & 22% for period before 2000, period between
2001 to 2010 and period between 2011 to 2021, respectively. This result may be due to the differences

231 in the diagnostic methods used for FGTB which have changed over times. Surprisingly, the researchers noted that the polymerase chain reaction (PCR) test was not used in studies published in period the 232 before 2000 while the same diagnosis method was used by 70% and 80.8% for period between 2001 to 233 234 2010 and 2011 to 2021, respectively. The utilized methods in currently analyzed data were histopathological examination^{23,24,25}, culture^{26,27}, acid-fast bacilli test²⁸, and laparotomy²⁵. According to 235 the literature, no standard gold test for FGTB is fixed but it depends on the facilities test protocol. 236 237 However, difference FGTB testing methods had been giving various results of the disease rate among infertile women.²⁰ The increase of the prevalence of FGTB among infertile women is due to the 238 previously mentioned reasons including utilization of TB modern diagnosis methods and adopting the 239 WHO Strategy of TB.¹⁸ Furthermore, the global funds on TB control substantially increased in recent 240 decades.29 241

242

Based on the aim of this study "to investigate the pooled prevalence of FGTB among infertile women
globally", the collected data was divided into four subgroups according to the World and Bank
Economical classification. The present study reveals that, the prevalence of FGTB is inversely
proportional to the economic situation of the country. The smallest prevalence was 5.7% in the highincome countries while, the highest prevalence was 24% in the low-income countries. The upper
middle-income and lower middle-income countries showed 14% and 22%, respectively (table 2.,
section B.).

250

Although, there was no published data to describe the rate of FGTB among infertile women in the 251 different countries based on their economic status. Many other studies have shown that female genital 252 tuberculosis is associated with PTB and EPTB as secondary infection.^{2,3} This outcome may be due to 253 the delay of TB diagnosis and other sociocultural reasons. In line with that, Getnet and colleagues 254 255 reported a 42% of PTB delayed for a varied time (a month to a year) on TB-diagnosis in low income and middle-income countries setting.³⁰ Furthermore, MacPherson et al., indicated a 4% to 38% of TB-256 patients lost the follow-up to the treatment in the same setting.³¹ In the Middle East and North Africa 257 factors such being a women and low per capita income is relatively reflected to the delay in TB-258 diagnosis.³² Although, the proportions are 1.24% and 1.26% respectively its considerable on FGTB 259 incidence. In addition, the high incidence of FGTB in low- and middle-income setting is due to factors 260 such as the higher rate of losses to follow-up with TB or EPTB treatment³³, the relatively negative 261 experiences of TB-patient and their satisfaction with healthcare system.³⁴ Moreover, poverty and the 262 high cost of the accurate diagnosis of FGTB³⁵ in developing countries has a huge negative effects on 263

- FGTB control and treatment.^{35,36} In accordance with that, D. Cazabon, et al., reported that, 32% and 46% of TB-patients had a negative experience and dissatisfaction with healthcare providers and TB services respectively.³⁴
- 267

The finding of current study reveals that the pooled prevalence of infertility among overall FGTB-268 patient was very high 88%. Of this the pooled prevalence of primary infertility was higher than that of 269 secondary infertility among FGTB-patients. Although, these results are in agreement with other meta-270 analysis findings done by Kefayat, et al., which is reported 70.7%, 75.7% and 24.3% for infertility 271 among FGTB-patient, primary infertility and secondary infertility respectively.³⁷ The present study 272 showed slight an increase in the pooled prevalence infertility and secondary infertility incidence among 273 274 FGTB compared to Kefavat, et al. study. On the other hand, the rate of primary infertility decreased over time. 275

276

To achieve the WHO End TB Strategy to eliminate catastrophic costs for TB-affected households by
2030 as Sustainable Development Goal target¹⁸, a more thorough clinical investigation should be
administrated at the level of TB and infertility clinics, particularly in low and lower-income settings.

280

281 Limitations

This review is not without limitation as articles published in languages other than English were excluded and the study population included only infertile women of reproductive age. Some grey literature may have also been omitted and regarding the incidence of FGTB among infertile women worldwide, no article however included published works from the Australian, European, and South American continents. The likelihood for publication bias is high.

287

288 Conclusions

The results of this meta-analysis found, that the pooled prevalence of FGTB among infertile women is 20%, and the pooled prevalence of overall infertility, primary infertility, and secondary infertility among FGTB patients globally, were 88%, 66%, and 34% respectively. In the last two decades, the FGTB incidence rate was increasing gradually. The biggest burden of FGTB is reported in the low- and lower-middle-income countries with a pooled prevalence of 46% globally.

294

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300 Authors' Contributions

- 301 MA, AAA, AI, CA, AB and SH conceived and designed the review. MA, AAA, and SH carried out the
- draft of the manuscript and MA is the guarantor of the review. MA, AAA, AI, CA, AB and SA
- developed the search strings. MA, AAA, and SH screened and selected studies, and extracted the data.
- AI, CA, and AB evaluated the quality of the studies. MA and AAA carried out the statistical analysis
- and interpretation. MA, AAA, AI, CA, AB and SH rigorously reviewed the manuscript.
- 306

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492	Figure 1: PRISMA Flow Diagram.
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Figure 2: Forest plot (random-effects model) for the pooled prevalence of FGTB among infertile women.
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503	Table 1: Main	characteristics of	studies included	in the meta-analysis
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Authona yoon	Study	World Bank	Count	Inf.	FGTB Testing	Proportion of	Proportion of infertility among FGTB patient		
Authors, year	Setting	Classification	ry	Рор	method	FGTB %(n)	Inf. overall	PI	SI
Chattopadhyay et al., 1986 ^{<u>38</u>}	CS/HA	High income	Saudi Arabia	945	NA	4.2 (40)	NA	NA	NA
Reshef Tal et al., 2020 ³⁹	PC/HC	High income	United States	323	QuantiFERON- TB	7.7 (25)	NA	NA	NA
Abdissa et al., 2018^{4}	CS/HA	Low income	Ethiop ia	152	PCR, CP, HE	5.3 (8)	62.5 (5)	50 (4)	12.5(1)
Abebe et al., 2004^{13}	CS/HA	Low income	Ethiop ia	25	AFB, CP, HE, PCR	64 (16)	NA	NA	NA
Abdelaziem, & Tajeldin., 2012 ⁴⁰	CS/HA	Low income	Sudan	277 8	HE	0.9 (25)	NA	NA	NA
Al eryaniet al., 2015 ⁴¹	P/O/HA	Low income	Yemen	151	AFB, PCR, CP, HE	31.1(47)	NA	NA	NA
Nezar, 2009 <u>42</u>	P/O/HA	Lower middle income	Egypt	420	Laparoscopy, HE, PCR	5.7 (24)	100 (24)	NA	NA
Kumaret al., 2008 ⁶	CS/HA	Lower middle income	India	285	PCR	39 (111)	100 (111)	NA	NA
Mohakul SK, Beela VRK, Tiru P.,2015 ⁵	P/HC	Lower middle income	India	105	PCR, Hysteroscopy	39 (41)	100 (41)	58 (24)	42 (17)
Jindal, 2006 ¹⁰	R/HC	Lower middle income	India	208 3	LAP, AFB, HE, MT, ELISA	7.2 (150)	97.3 (146)	70 (105)	27.3 (41)
Singh et al., 2008 ⁴³	R/HC	Lower middle income	India	140	MH, laparoscopy &hysteroscopy	48.5 (34)	NA	NA	NA
Sankar, 201344	R/HA	Lower middle income	India	620	AFB, PCR, CP, HE	25.5 (158)	95.5 (151)	78.8 (119)	21.2 (32)
Mahajan et al., 2016 ⁴⁵	CS/HA	Lower middle income	India	180	PCR, CP	41 (74)	NA	NA	NA
Sethi et al., 2016 ⁴⁶	CS/HA	Lower middle income	India	300	AFB, PCR, CP, HE	22.7 (68)	NA	NA	NA
Chatterjee&Basak, 2018 ²⁶	CS/HA	Lower middle income	India	120	PCR	1.7 (2)	NA	NA	NA
Gon Chowdhury R. et al., 2010 ⁴⁷	CO/HA	Lower middle income	India	517	PCR	44.5 (230)	49.7 (114)	NA	NA
Saraswat et al., 2010 ⁴⁸	CS/HA	Lower middle income	India	125	PCR, CP	20.8 (26)	NA	NA	NA
Bharti Malhotra et al., 2012 ⁴⁹	O/HA	Lower middle income	India	555	AFB, PCR, CP	25.22 (140)	NA	NA	NA
Swati B Gajbhiye et al., 2019 ⁵⁰	CS/O/HA	Lower middle income	India	50	PCR	12 (6)	83.3 (5)	80 (4)	20 (1)
Bhanothu et al., 2014^{51}	P/CC/HA	Lower middle income	India	302	PCR	28.47 (86)	NA	NA	NA
Gurjar et al., 2018 ²⁰	O/HA	Lower middle income	India	100	PCR	52 (52)	NA	NA	NA
Patil et al., 2015 ⁵²	CS/HA	Lower middle income	India	123	Gen-Probe MTD test	0.8 (1)	NA	NA	NA
Goel et al., 2013 ⁵³	R/HA	Lower middle income	India	546	PCR	3.7 (20)	NA	NA	NA
Kamal S et al, 2020 ⁵⁴	P/HA	Lower middle income	India	100	PCR, HE	27 (27)	NA	59.4 (16)	40.6 (11)
Gupta S, et al., 2021 ⁵⁵	P/HC	Lower middle income	India	59	CBNAAT, HE	3.4 (2)	NA	100 (2)	0

Meenu et al., 2020 ⁵⁶	CS/HA	Lower middle income	India	139 *	PCR	41.7 (58)	NA	NA	NA
Shende P et al., 2017 ⁵⁷	P/HA	Lower middle income	India	120	PCR	27 (32)	NA	NA	NA
Deshmukh et al., 2014 ^Z	P/HC	Lower middle income	India	218	AFB, CP, HE, PCR	39.45 (86)	NA	NA	NA
Ohri S, Patil SK, Patil A, et al., 2016 ⁵⁸	P/HA	Lower middle income	India	50	PCR	18 (9)	NA	88.9 (8)	11.1 (1)
Madkar et al., 2014 ⁵⁹	P/HA	Lower middle income	India	50	PCR	12 (6)	NA	50 (3)	50 (3)
Gupta et al., 2007 ⁶⁰	R/HA	Lower middle income	India	150	AFB, MT, PCR	26.7 (40)	NA	75 (30)	25 (10)
S Rajaram et al., 2016 ⁶¹	PC/HA	Lower middle income	India	50	HE, PCR	28 (14)	NA	NA	NA
Ojo et al., 2008 ¹⁹	R/HA	Lower middle income	Nigeri a	661	AFB, HE	0.45 (3)	NA	33.3 (1)	66.7 (2)
Ojo et al., 1971 ²³	CS/HA	Lower middle income	Nigeri a	118 96*	HE	0.7 (82)	NA	NA	NA
Emembolu, 1989 ²⁸	R/HA	Lower middle income	Nigeri a	114	AFB	16.7 (19)	NA	47.4 (9)	52.6 (10)
Gini & Ikerionwu, 1990 ^{<u>24</u>}	R/HA	Lower middle income	Nigeri a	470 0	HE	0.2 (10)	NA	NA	NA
Sughra Shahzad., 2012 ⁶²	R/HA	Lower middle income	Pakista n	150	AFB, PCR, CP	20 (30)	NA	83.3 (25)	16.7 (5)
Shaheen R, Subhan F, Tahir F., 2006 ⁶³	CS/HA	Lower middle income	Pakista n	534	CP, AFB-ZN, HE	2.43 (13)	100 (13)	NA	NA
Khan SMQ., 1985 ²⁵	R/HA	Upper middle income	Iran	91	LAP, HE	23.08 (21)	NA	71.4 (15)	28.6 (6)
Shallal et al., 2021 ⁶⁴	P-CS/HA	Upper middle income	Iraq	60	PCR, HE	10 (6)	NA	NA	NA
MARGOLIS et al, 1992 ²⁶	R/HA	Upper middle income	South Africa	650	СР	6.15 (40)	NA	40(16)	60 (24)
Oosthuizen et al., 1990 ²⁷	CS/HA	Upper middle income	South Africa	109	СР	21 (23)	NA	NA	NA

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infertility; CBNAAT = cartridge based nucleic acid amplification test; MH = menstrual history; LAP = Laparotomy.

509 *Gynaecological admitted patient including infertility. 510

511 Table 2: A: Pooled prevalence of infertility among FGTB patient; B: the pooled proportion of FGTB among 512 infertile women based on world bank country economic classification; and C. subgroup analysis of FGTB among infertile women by study's publication year 513

PC = prospective cohort study; CS = cross-sectional study; R = retrospective study; O = observational study, CC = case

histopathological examination; Inf. Pop = infertile populations; Inf = infertility; PI = primary infertility; SI = secondary

control study; HA = hospital admitted patients; HC = infertility center admitted patient; ND = no data found; PCR =

polymerase chain reaction test; AFB = acid-fast bacilli test; MT = mantoux test; CP = culture proven; HE =

Secti	Subgroups	Subgroups	No. of Studies	Total patient No.	FGTB proportion % (min-	Infertility proportion % (min-	Heterogeneity	
ion	Classification				max) 95%CI	max) 95%CI	I ² (%)	P- value
А	The type of infertility (among FGTB	Pooled infertility	5	430	-	88 (74- 100)	99.912	<.001
		Primary infertility	15	560	-	66 (56-76)	99.226	<.001
	patients)	Secondary infertility	14	558	-	34 (24-43)	98.039	<.001

в	World bank country economic Classification (among infertile patient)	High income	2	1268	5.7 (2.3- 9.1)	-	78.56	<.001
		Upper middle- income	4	910	14 (6-23)	-	86.91	<.001
		Lower middle- income	32	25562	21 (15-27)	-	99.95	<.001
		Low income	4	3106	24 (3-52)	-	99.48	.084
	Voor of	Before 2000	7	18530	10 (3-17)	-	99.96	<.001
С	publication (among infertile patient)	Between 2001 to 2010	11	7718	23 (10-36)	-	99.93	<.001
		Between 2011 to 2021	24	4623	22 (16-27)		97.98	<.001

FGTB = Female Genital Tuberculosis.