



Research Article

Sexually Transmitted Infections among Key Populations in India: Systematic Review with Spatiotemporal Distribution

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Abstract

In the developing world, sexually transmitted infections (STIs) are among the key sources of health and financial adversities, contributing significantly to morbidity, death, and stigma. Goal of the current systematic review is to identify the geographic distribution and current of STIs among the Indian key population. A protocol was prepared and registered in the PROSPERO, with the registration number CRD42022357425 and published as a Systematic Review Protocol. The data on frequency and prevalence of four treatable STIs— syphilis, chlamydia, gonorrhea and trichomoniasis, focused on the MSM (men sex with men) and FSW (female sex workers) populations, followed by H/TG (Hijras with Transgenders) and PWID (those who inject drugs), was gathered and analyzed from different geographical regions in India. However, it was found that most of the research used aetiological diagnosis to report prevalence and were belongs to the western and southern regions of India. Few studies from the northern and north eastern regions were also being found. In the light of present findings and with the identified limitations, it can be concluded that existing HIV surveillance system under NACP (National AIDS Control Programme) in India, can be utilized with additional bio-specimen collection to determine STI prevalence among high risk populations.

Keywords: Sexually transmitted infections; Key population; Syphilis; Gonorrhea; Chlamydia; Trichomoniasis; India

Introduction

In the developing parts of the world, sexually transmitted infections (STIs) are among the key sources of health, and financial adversities, causing extensive morbidity, mortality and stigma [1]. A remarkable variation of frequency and prevalence of the four curable STIs, which are syphilis, chlamydia, gonorrhea and trichomoniasis are observed with the spatiotemporal variation [2]. Studies suggest that the prevalence of these four STIs among general people has a range of 0-3.9% in India [3], but the STI burden is considerably higher among key populations having high-risk behaviour like men who have sex with men (MSM), female sex worker (FSW) hijras with transgender, and people who inject drugs (PWID). However, there are inadequate literature till date to report STI burden in Indian key populations and all of the information have not been unified to enlighten the general spatiotemporal tendencies of STI infections in various

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key or high-risk population. To gain insight of the status of the prevalence and spatiotemporal distribution of four STIs viz. syphilis (by *Treponema pallidum*), gonorrhoea (by *Neisseria gonorrhoeae* or NG), chlamydia (by *Chlamydia trachomatis* or CT), and trichomoniasis (by *Trichomonas vaginalis* or TV) among different key populations such are, men who have sex with men (MSM), female sex workers (FSW), hijras with transgenders (H/TG) and people who inject drugs (PWID) in India, present systematic review has been initiated using existing published evidence [3-4].

Materials and Methods

A study protocol was developed according to the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) guidelines. All available peer reviewed published articles are extracted using suitable search terms [5] in PUBMED, MEDLINE, EMBASE, Cochrane Library, Science Direct, Google Scholar and Psycinfo along with published re-ports (as grey literature) from reliable sources, comprising all studies from January 2000 to December 2023.

A protocol for the systematic review was prepared and registered in the PROSPERO, having the registration number CRD42022357425 and published subsequently [5].

Evaluation of the methodological quality

Evaluation of selected studies was performed through title, text and abstract prior to the accumulation of it into the decisive analysis. Evaluation was performed with the help of modified Newcastle - Ottawa Quality Assessment tool [6]. Evaluation was completed before collection of information.

Geographical distribution with spatiotemporal attributes

Geographical distributions of previously mentioned STIs in Indian key population were presented using heat map and choropleth maps, generated with the help of Quantum GIS (Ver. 3.18 Zurich) [7]. An administrative map of India containing the latest information of states and union territories is georeferenced and the resulting polygon was used as the base map layer. State-level STI prevalence data were added as comma delimited layers. Prevalence data from accepted studies were added as spatiotemporal attributes in separate layers respectively. Finally, maps were generated for overall included studies for each key population with prevalence of four curable STIs in histogram format in a separate layer. Multiple points were used to generate heat map of STI prevalence. Data based on regional distribution of STIs among key population has been entered into the newly pre-pared maps as non-spatial attributes [8].

Results

The present findings would be vital to understand the STI status among key population and to design evidence-based strategies for STI prevention in key population in India. There are total of 8919 publications are selected initially. The abstract and titles were screened for eligibility and duplicates were removed. A total of forty articles were included from peer reviewed journals. Among the 70 reports (grey literature)-excluding Sankalak Reports (two documents), published by several competent national, international, and state agencies, any reports could not be included based on the screening tool. Details of the inclusion and exclusion criteria, along with the, number of each category of articles are described through PRISMA flow Diagram [Figure 1].

Search of relevant databases (PUBMED, Google Scholar, Mendeley, Cochrane Library, Scopus, Science Direct and EMBASE) yielded 40 articles which met the inclusion criteria set in the protocol for the present systematic review. However, it is found that published articles were not available from all geographical regions in India. Most of the studies were conducted in southern and western part. However, four studies were also reported from north-eastern part of India. A few studies from northern India was also conducted in Delhi and Lucknow (Figure 2).

Maximum numbers of studies reported on STI prevalence, were belongs to MSM and FSW population followed by H/TG and PWID and mostly based on syphilis seroprevalence. Data on chlamydia and gonorrhoea were less available, trichomoniasis was least reported STI. While synthesizing data, articles which included more than one key population was counted separately. Eighteen articles reported STI prevalence among FSWs [1-4,8-23]. These eighteen articles included twenty-six different studies based on prevalence data of different STIs, different FSW subgroups and diverse geographical region. All data related to FSW was presented through two different diagnostic methods, aetiological [1-4,8-20] (Table 1) and syndromic [20,23] (Table 2). STI prevalence among MSM was reported in nineteen articles [3,15,24-40]. Based on different STIs and geographical regions total forty studies were recorded for this review. Relevant extracted data on MSM was recorded under aetiological diagnosis [3,15,24-40] (Table 3) and syndromic [33,40] (Table 4) diagnosis. Four articles on H/TG included four studies on STI prevalence among H/TG based on different STI and geographical region [18,29,31,41]. Data on H/TG under aetiological diagnosis was presented in Table 5. There was no data available for syndromic diagnosis. Two articles [15,42] were available for STI prevalence among PWID. These two articles comprise six studies on prevalence of different STIs among PWID population in India (Table 6).

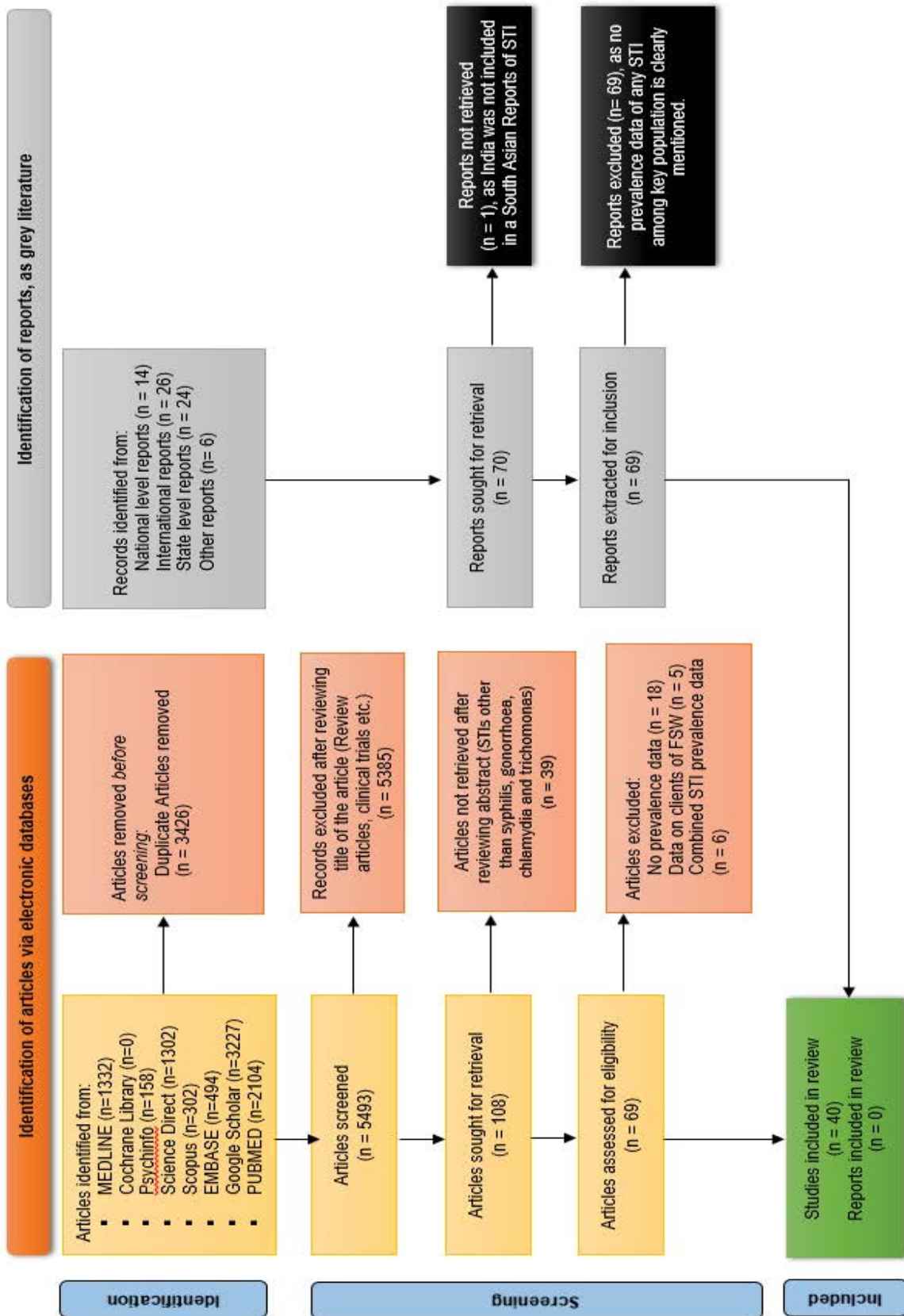


Figure 1: PRISMA Flow Diagram.

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Table 1: Aetiological diagnosis and prevalence of four curable STIs among FSW from available published articles from different parts of India.

Study	Study period	Study design	Location	FSW Sub group	Sample size	Study setting	Sampling Technique	Specimen	Specimen source	Syphilis		Gonorrhoea		Chlamydia		Trichomoniasis				
										Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence			
Reza-Paul et al. [1]	2004-2006	CS	Mysore	NR	429	Facility	TLC	Blood for Syphilis and urine for remaining three	NR	RPR + TPHA	24.7	PCR	5.4	PCR	10.8	PCR	32.9			
Shethwala et al. [2]	2005-2006	CS	Surat	NR	300	Community	NR	Blood for syphilis and chlamydia, cervix-swab for Trichomoniasis	NR	RPR + TPHA	6.66	NR	NR	NR	NR	Wet mount microscopy	2			
Gutierrez et al. [3]	2003-2007	CS	Andhra Pradesh	NR	1750	Facility	NR	Blood	NR	NA	13	NR	NR	NR	NR	NR	NR			
Ramesh et al. [4]	2004-2006	CS	Karnataka	NR	2312	Community	CCS	Blood for Syphilis and urine for gonorrhoea, chlamydia	NR	NR	10.2	NAAT	3.5	NAAT	6.5	NR	NR			
Hemalatha et al. [9]	Dec. 2005-Oct. 23006	CS	Andhra Pradesh	NR	3223	Community	CCS	Urine	NR	NR	NR	NAAT	2	NAAT	3.4	NR	NR			
Das et al. [10]	Oct. 2008-May 2009	CS	Mumbai & Hyderabad	NR	416	Facility	NR	Blood	NR	RPR + TPHA	10.1	NA	NA	NA	NA	NA	NA	NA		
					417						Swab	Cervix	NA	NA	NAAT	14.1	NAAT	16.1	NA	NA
					399										NA	NA	NA	NA	PCR	31.1
Gupte et al. [11]	Dec.2007-Feb.2008	CH	Mumbai	Brothel	2858	Community	NR	Blood	NR	RPR	6.6	NR	NR	NR	NR	NR	NR			
Gupte et al. [11]	Dec.2007-Feb.2008	CH	Mumbai	Bar	9296	Community	NR	Blood	NR	RPR	1.3	NR	NR	NR	NR	NR	NR			
Gupte et al. [11]	Dec.2007-Feb.2008	CH	Mumbai	Street	1199	Community	NR	Blood	NR	RPR	2.9	NR	NR	NR	NR	NR	NR			

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Gupte et al. [11]	Dec.2007- Feb.2008	CH	Mumbai	Home	4512	Community	NR	Blood	NR	RPR	1.8	NR	NR	NR	NR	NR	NR
Mainkar et al. [12]	Mar.2006- Dec 2006	CS	Maharashtra	NR	2525	Facility	RDS	Blood for Syphilis and urine for gonorrhoea, chlamydia	NR	RPR + TPHA	15.8	NAAT	7.4	NAAT	8	NR	NR
Rachakulla et al. [14]	Nov 2005- Dec 2006	CS	Andhra Pradesh	NR	3271	Facility	Cluster	Blood for Syphilis and urine for gonorrhoea, chlamydia	NR	RPR + TPHA	10.8	NAAT	2.2	NAAT	3.5	NR	NR
Ghosh et al. [15]	NR	CS	West Bengal	NR	45	Facility	NR	Blood for syphilis and chlamydia, cervix-swab for Trichomoniasis	NR	VDRL + TPHA	28.8	NR	NR	Serological	15.5	Kupferburg culture	17.7
Medhi et al. [16]	Feb. 2006 - Apr. 2006	CS	Nagaland	NR	423	Community	RDS	Blood for Syphilis and urine for gonorrhoea, chlamydia	NR	RPR + TPHA	15.3	NAAT	4.2	NAAT	22.2	NR	NR
Das et al. [17]	Oct 2008- Nov 2009	CH	Mumbai & Hyderabad	NR	417	Facility	NR	Blood for syphilis and Cervix-swab for remaining three	NR	RPR + TPHA	5.8	NAAT	14.2	NAAT	16.1	NAAT	31.2
Boily et al. [18]	Aug. 2004- Feb 2005	CS	Mysore	NR	429	Facility	TLC	Blood	NR	RPR + TPHA	24.9	NR	5.4	NR	10.8	NR	NR
Boily et al. [18]	Aug.2005- Aug. 2006	CS	Shimogga	NR	393	Facility	TLC	Blood	NR	RPR + TPHA	4	NR	1.3	NR	5.6	NR	NR
Boily et al. [18]	Nov. 2005- May 2007	CS	Belgaum	NR	366	Facility	TLC	Blood	NR	RPR + TPHA	8	NR	4.7	NR	6.5	NR	NR
Boily et al. [18]	Nov. 2005- Feb 2007	CS	Bellari	NR	423	Facility	TLC	Blood	NR	RPR + TPHA	5.2	NR	2.7	NR	4.1	NR	NR
Boily et al. [18]	Jul.2006- Jun. 2007	CS	Bangalore	NR	673	Facility	TLC	Blood	NR	RPR + TPHA	12.6	NR	3.6	NR	6.5	NR	NR

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Jayaraman et al. [19]	Jul.2006- Jun. 2007	CS	Bangalore	NR	673	Facility	TLC	Blood	NR	RPR + TPHA	12.6	NAAT	NR	NAAT	NR	NR	NR
Sarna et al. [20]	Jan 2010 -Dec 2010	CH	Nellore	NR	529	Community	RDS	Blood for syphilis and Cervix-swab for remaining three	NR	RPR + TPPA	2.8	PCR	0.7	PCR	2.2	Pap Smear	15.5
Beksinska et al. [22]	July and August 2011	CS	Karnataka	NR	1092	Community	CCS	Blood	NR	RPR + TPHA	3.2	NR	NR	NR	NR	NR	NR

CH- Cohort study; CS- Cross-sectional study; TLC- Time Location Cluster, CCS- Conventional Cluster Sampling, NR- Not Reported; TPHA- Treponema Pallidum Heamagglutination Test, RPR Rapid Plasma Reagin; NAAT- Nucleic Acid Amplification Test

Table 2: Syndromic diagnosis and prevalence of four curable STIs among FSW from available published articles from different parts of India.

Study	Study period	Study design	Location	FSW Sub group	Sample size	Study setting	Sampling method	Urethral discharge	Genital Ulc Herpetic	Cervical discharge	Genital Ulcer Non Herpetic	PSSw	Inguinal Bubo	Vaginal discharge	Ano-rectal discharge
Sarna et al. [20]	Jan 2010 -Dec 2010	C1H	Nellore	NR	529	Community	RDS	NR	NR	5.3	NR	NR	NR	50.7	NR
Shukla et al. [23]	Aug 2012- Jul.2013	CS	Lucknow	Street based	83	Community	Random	NR	10.8	30.1	2.4	NR	1.2	30.1	0
Shukla et al. [23]	Aug 2012-Jul.2013	CS	Lucknow	Home based	205	Community	Random	NR	1.5	20.5	1.5	NR	1	20.5	1.5

CH- Cohort study; CS- Cross-sectional study; NR- Not reported; TLC- Time Location Cluster, RDS- Respondent Driven Sampling

Table 3: Aetiological diagnosis and prevalence of four curable STIs among MSM from available published articles from different parts of India.

Study	Study period	Study design	Location	MSM Sub group	Sample size	Study setting	Sampling Technique	Specimen	Syphilis		Gonorrhoea		Chlamydia		Trichomoniasis	
									Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence
Setia et al. [24]	2001	CS	Mumbai	NR	122	Facility	CCS	Blood	TPHA	17.2	NR	NR	NR	NR	NR	NR
Gupta et al. [25]	1998-2002	CH	Pune	NR	708	Facility	NR	Blood for syphilis and glans penis swab for gonorrhoea	RPR + TPHA	3	Gram Stain	4.1	NR	NR	NR	NR

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Gupta et al. [25]	1993-1997	CH	Pune	NR	708	Facility	NR	Blood for syphilis and glans penis swab for gonorrhoea	RPR + TPHA	8.2	Gram Stain	4.5	NR	NR	NR	NR
Brahmam et al. [26]	Mar.2006-Apr.2007	CS	Tamilnadu	NR	2025	Facility	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR	14	NAAT	0.1	NAAT	0.6	NR	NR
Brahmam et al. [26]	Mar.2006-Apr.2007	CS	AAAndhra Pradesh	NR	1621	Facility	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR	13	NAAT	0.6	NAAT	1.6	NR	NR
Brahmam et al. [26]	Mar.2006-Apr.2007	CS	Karnataka	NR	298	Facility	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR	11.9	NAAT	0.6	NAAT	1.6	NR	NR
Brahmam et al. [26]	Mar.2006-Apr.2007	CS	Maharashtra	NR	653	Facility	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR	8.4	NAAT	0.3	NAAT	4	NR	NR
Gutierrez et al. [3]	2003-2007	CS	Andhra Pradesh	NR	1106	Facility	NR	Blood	NR	20	NR	NR	NR	NR	NR	NR
Kumta et al. [27]	Jan.2003-Dec.2004	CH	Mumbai	NR	831	Facility	NR	Blood	VDRL	6.5	NR	NR	NR	NR	NR	NR
Solomon et al. [28]	Oct.2008-Nov 2008	CS	Tamil Nadu	NR	721	Community	RDS	Blood	RPR, TPPA	8.04	NR	NR	NR	NR	NR	NR
Sahastrabudhe et al. [29]	1993-2002	CS	Pune	NR	641	Facility	RDS	Blood	TPHA	5.8	NR	NR	NR	NR	2rrrr	NR
Ghosh et al. [15]	NR	CS	West Bengal	NR	26	Facility	NR	Blood	TPHA	7.7	NR	NR	NR	NR	NR	NR
Narayanan et al. [30]	2006-2009	CS	Mumbai and Hyderabad	NR	149	Facility	NR	Blood for syphilis and rectal swab for gonorrhoea and chlamydia	TPHA, RPR	4	PCR	14.8	PCR	4.7	NR	NR

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Subramanian et al. [31]	2006	CS	Tamilnadu	NR	1621	Community	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR + TPHA	14.3	NR	0.07	NAAT	0.68	NR	NR
Goswami et al. [32]	2006	CS	Andhra Pradesh	NR	1218	Facility	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR + TPHA	8.1	NAAT	0.3	NAAT	1.1	NR	NR
Nagarajan et al. [33]	2005-2007	CS	Andhra Pradesh, Maharashtra and Tamilnadu	NR	3895	Community	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR + TPHA	12.8	NAAT	1.2	NAAT	0.3	NR	NR
Ramanathan et al. [34]	Jan. 2005-Mar. 2009	CS	Maharashtra	NR	653	Community	TLC	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR + TPHA	8.8	NAAT	0.3	NAAT	3.7	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Hyderabad	NR	998	Community	RDS	Blood	TPHA	3.9	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Vijaywada	NR	1000	Community	RDS	Blood	TPHA	4.4	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Vizag	NR	1000	Community	RDS	Blood	TPHA	2.1	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Bangalore	NR	1000	Community	RDS	Blood	TPHA	4	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Belgaum	NR	1003	Community	RDS	Blood	TPHA	0.8	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Mangalore	NR	1000	Community	RDS	Blood	TPHA	1.4	NR	NR	NR	NR	NR	NR

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Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Chennai	NR	1000	Community	RDS	Blood	TPHA	1.1	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Coimbatore	NR	1000	Community	RDS	Blood	TPHA	3	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Madurai	NR	1001	Community	RDS	Blood	TPHA	1.8	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Bhopal	NR	998	Community	RDS	Blood	TPHA	1.3	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Delhi	NR	997	Community	RDS	Blood	TPHA	3.5	NR	NR	NR	NR	NR	NR
Solomon et al. [35]	Sep.2012-Jun. 2013	CS	Lucknow	NR	1000	Community	RDS	Blood	TPHA	4.1	NR	NR	NR	NR	NR	NR
Mayer et al. [36]	NR	CS	Mumbai	NR	307	Community	RDS	Blood for syphilis , oro pharyngeal + rectal swab and urine for gonorrhoea and chlamydia	RPR, TPHA	4.02	Gen Probe	3.26	Gen Probe	17	NR	NR
Ramakrishnan et al. [37]	2009-2010	CS	Andhra Pradesh, Maharashtra and Tamilnadu	MSMO	2396	Community	TLC	Blood	RPR + TPHA	5.5	NR	NR	NR	NR	NR	NR

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Ramakrishnan et al. [37]	2009-2010	CS	Andhra Pradesh, Maharashtra and Tamilnadu	MSMW	1343	Community	TLC	Blood	RPR + TPHA	7	NR	NR	NR	NR	NR	NR
Prabakar et al. [38]	Jun.2015-May 2016	CH	Chennai	NR	1215	Facility	NR	Blood	VDRL + TPHA	4.53	NR	NR	NR	NR	NR	NR
Safren et al. [39]	NR	CS	Chennai	NR	304	Facility	NR	Blood for syphilis and urine +rectal and oro-pharyngeal swab for gonorrhoea and chlamydia	TPHA	16.1	NAAT	8.3	NAAT	11.3	NR	NR
Safren et al. [39]	NR	CS	Mumbai	NR	304	Facility	NR	Blood for syphilis and urine +rectal and oro-pharyngeal swab for gonorrhoea and chlamydia	TPHA	14.8	NAAT	16	NAAT	18.2	NR	NR
Swamiappan et al. [40]	July 2016- Jun. 2019	CH	Chennai	NR	489	Facility	NR	Blood	RPR	8.8	NR	NR	NR	NR	NR	NR

CH- Cohort study; CS- Cross-sectional study; TLC- Time Location Cluster, NR- Not Reported; TPHA- Treponema Pallidum Heamagglutination Test, RPR Rapid Plasma Reagin; NAAT- Nucleic Acid Amplification Test

Table 4: Syndromic diagnosis and prevalence of four curable STIs among MSMs from available published articles from different parts of India.

Study	Study period	Study design	Location	MSM sub group	Sample size	Study setting	Sampling method	Urethral discharge	Genital Ulcer Herpetic	Genital Ulcer Non-Herpetic	PSSw	Inguinal Bubo	Ano-rectal discharge
Nagarajan et al. [33]	2005-2007	CS	Andhra Pradesh, Maharashtra and Tamilnadu	NR	3895	Community	TLC	0.1	NR	NR	0.3	NR	NR
Swamiappan et al. [40]	July 2016- Jun. 2019	CH	Chennai	NR	489	Facility	NR	2	3.1	2	NR	NR	NR

CH- Cohort study; CS- Cross-sectional study; NR- Not Reported, ; TLC- Time Location Cluster, PSSw- Painful Scrotal Swelling

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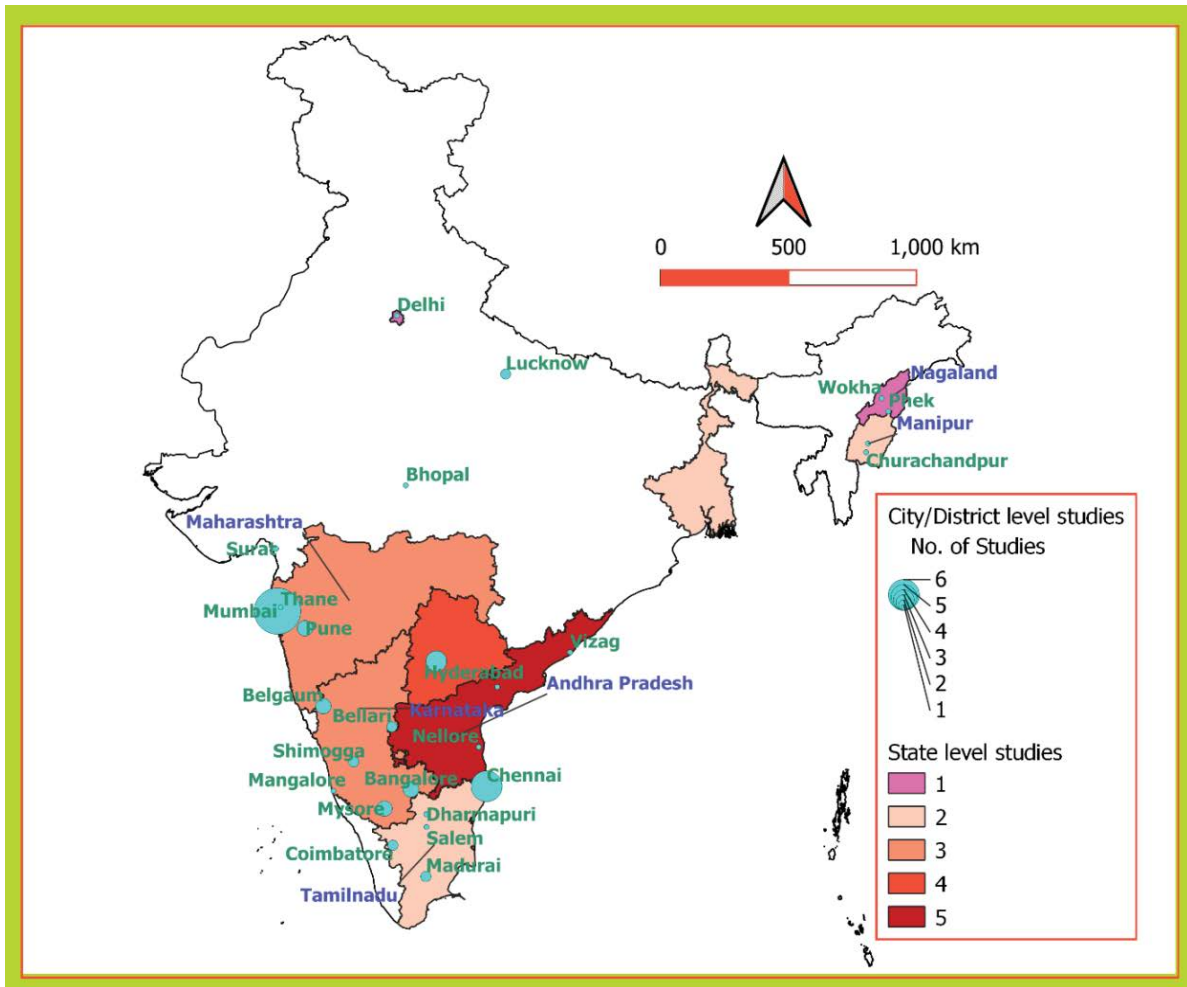


Figure 2: Map of India indicating the regions where included studies were conducted.

Findings among FSW

As per the search, there are eighteen articles comprises of twenty-six studies, which were find out, tell us out about the different STIs among FSW (Table 1 and 2), most of these studies were conducted in Andhra Pradesh, Telangana, Karnataka, Maharashtra, Uttar Pradesh, West Bengal, and Nagaland. Almost half of the studies were conducted in facilities like STI clinics or tertiary care hospitals (Figure 2). Out of 26 studies, most of the studies reported prevalence through aetiological diagnosis (Table 1). Most of the studies reported on single STI, namely syphilis. Among the published articles, an article from Beksinska et al. was published in 2018 [22], however, the study period was 2011. There was very little published article on STI prevalence estimation among FSW after 2011. This cross-sectional study on syphilis among FSW performed in Karnataka and the estimated prevalence was 3.2. Screening was done through RPR followed by TPFA.

Out of all available studies on FSW, a facility level cross-sectional study with a little lower sample size of 45, estimated

very high syphilis prevalence (28.8) in West-Bengal [15]. A study conducted in Mumbai during the period of 2007- 2008. [11] and published in 2011, estimate prevalence of syphilis among two groups of sex workers, brothel based FSW had highest syphilis prevalence of 6.6 and the bar-based sex workers had lowest prevalence of 1.3. Trichomoniasis was the least examined and hence least reported STI among FSW (Figure. 3). A study, which based on syndromic diagnosis, was conducted in Nellore, Andhra Pradesh during the period of January to December, 2011, unveiled genital herpetic and non-herpetic ulcer, cervical discharge, Inguinal buboes vaginal discharge and ano-rectal discharge in FSWs (Table 2). The highest reported symptom was vaginal discharge (50.7%) [20]. There was a single reported study each from Nagaland, West Bengal, and Maharashtra, two from Karnataka and Tamil Nadu and three studies from Andhra Pradesh. Most of the district level studies were reported from Karnataka and Andhra Pradesh. Four district level or small studies were reported from Mumbai and two each from Bangalore and Mysore (Figure 3).

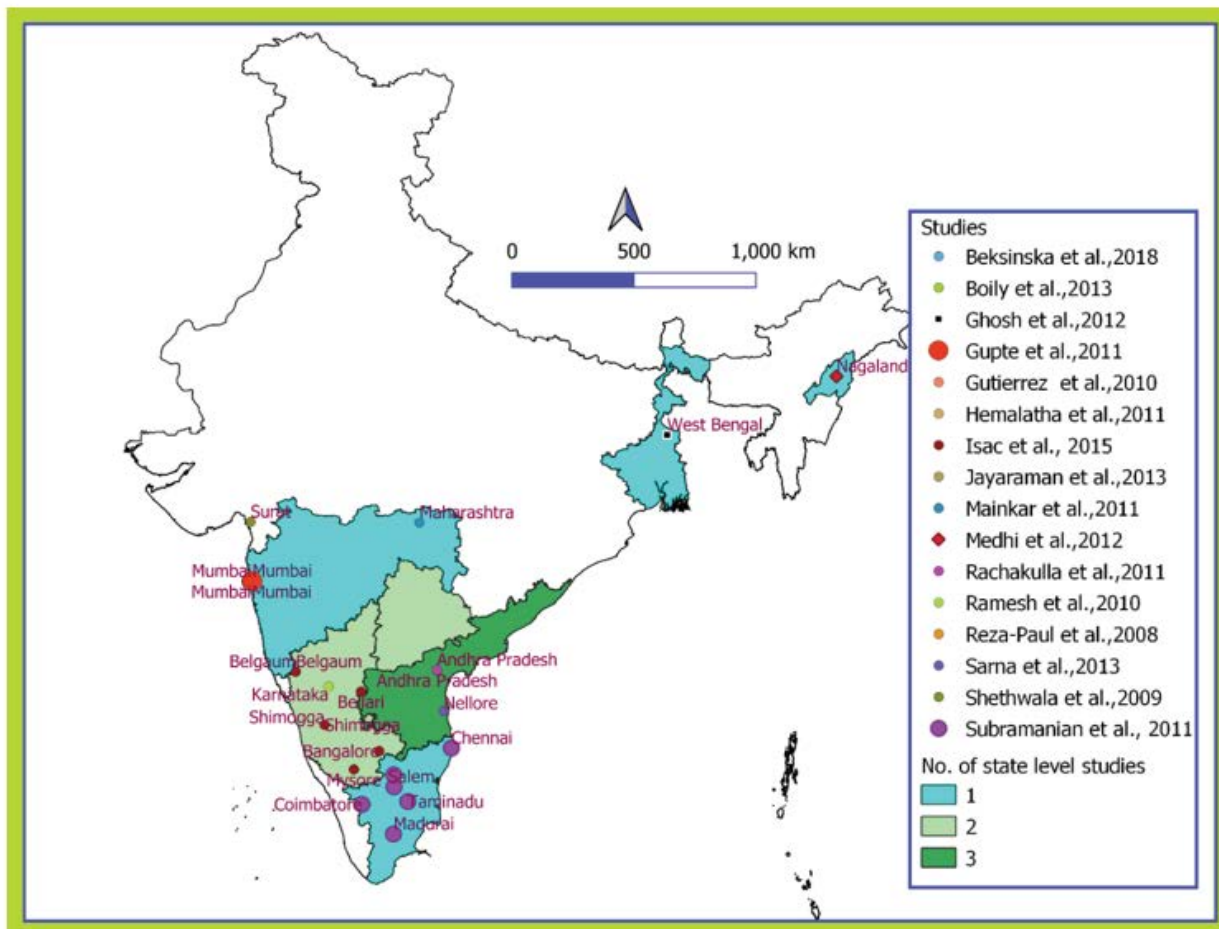


Figure 3: Representation of published articles on four curable STIs among FSWs in India.

Findings among MSM

Nineteen articles [3,15,24-40] on MSM were found out through literature search. Most of them were conducted in Tamil Nadu, West Bengal, Andhra Pradesh, Telangana, Karnataka, and Maharashtra (Figure 5). Almost half of the studies reported from facility level such as STI clinics or tertiary care hospitals. Most of the studies reported prevalence through aetiological diagnosis (Table 3). The most recent available publication reported through literature search, was by Swamiappan et al. [40] and by Safren et al. [39] and both published in 2020. The study conducted in Chennai during July 2016 to June 2019 [40] was reported prevalence of syphilis (8.8) through RPR without mentioning any confirmatory test. Along with the aetiological diagnosis, this study [40] reported symptoms of Urethral Discharge (2%), Genital herpetic Ulcer (3.1%) and Non-herpetic Ulcer (2%). Another cross-sectional study conducted in Mumbai and Chennai without specifying any study period (Table 4), reported prevalence of syphilis, gonorrhoea, and chlamydia as 16.1, 8.3 and 11.3 respectively in Chennai and 14.8, 16, 18.2 respectively in Mumbai [39]. Most of the studies reported Syphilis as none of the studies reported Trichomoniasis.

Gonorrhoea and chlamydia were also examined by many of these studies. For aetiological diagnosis RPR screening test followed by TPHA confirmatory test were administered for syphilis and PCR and NAAT for gonorrhoea and chlamydia. Similarly, a cross-sectional community-based study [3] in Andhra Pradesh, estimate a very high syphilis prevalence of 20.0, during the study period of 2003 to 2007. In the same state, Andhra Pradesh a cross-sectional study was conducted in Vijayawada and Vizag, during 2012-2013 and had estimated prevalence of 4.4, and 2.1 respectively [28]. As per the literature search, there are single study from each state of Karnataka, West Bengal and Maharashtra was reported whereas; two studies were reported from Andhra Pradesh and Tamil Nadu. A considerable number of studies were reported from Mumbai, Chennai, and Pune. A few sporadic studies were also evident from Delhi, Lucknow, Bhopal, and Hyderabad. There was no published article from North-eastern India. Studies from Mumbai, Chennai and Pune showed higher STI prevalence. Similar to FSW population, prevalence of syphilis was reported in most of the studies, whereas a fewer study reported studies on trichomonas (Figure 4).

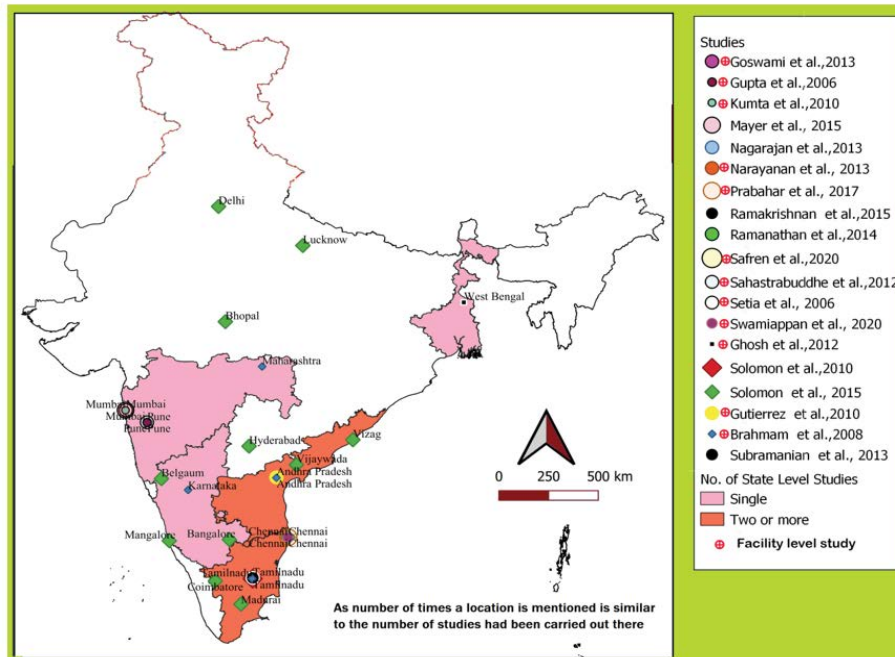


Figure 4: Representation of published articles on four curable STIs among MSMs in India.

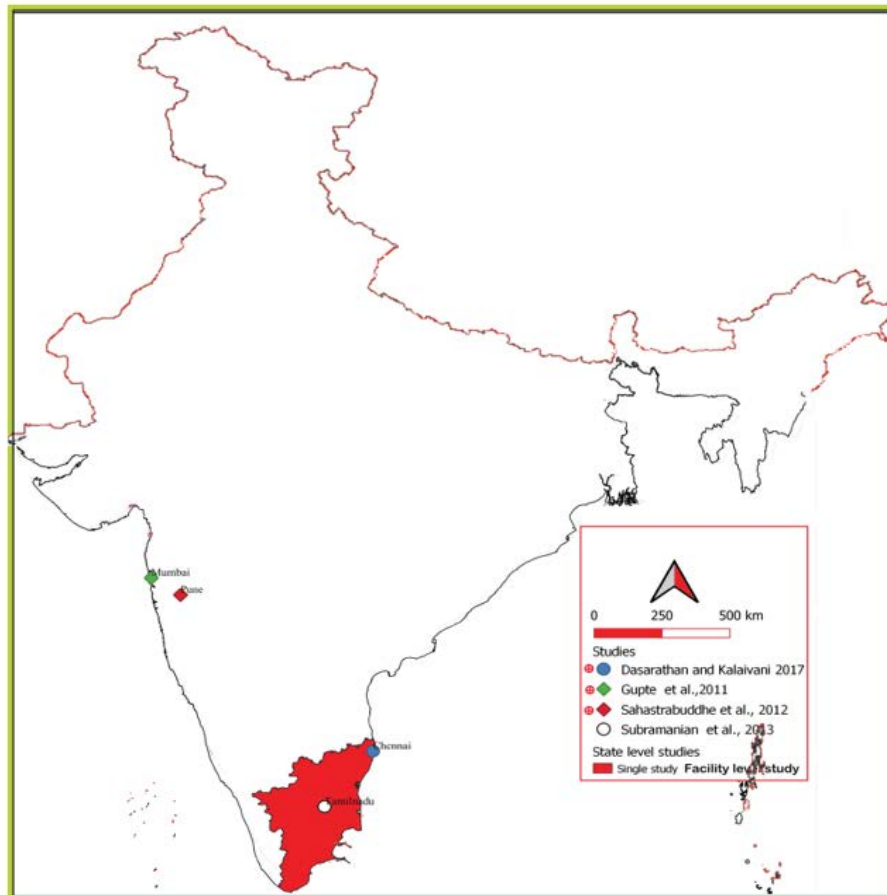


Figure 5: Geographical representation of published four curable STIs among H/TGs in India.

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Table 5: Aetiological diagnosis and prevalence of four curable STIs among H/TG people from available published articles from different parts of India.

Study	Study period	Study design	Location	Sample size	Study setting	Sampling Technique	Specimen	Syphilis		Gonorrhoea		Chlamydia		Trichomoniasis		
								Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	
Gupte et al. [11]	Dec.2007-Feb.2008	CH	Mumbai	620	Facility	NR	Blood	RPR	13.1	NR	NR	NR	NR	NR	NR	NR
Sahastrabudhe et al. [29]	1993-2002	CS	Pune	68	Facility	RDS	Blood	TPHA	10.3	NR	NR	NR	NR	NR	NR	NR
Subramanian et al. [31]	2006	CS	Tamilnadu	404	Community	CCS	Blood for syphilis and urine for gonorrhoea and chlamydia	RPR + TPHA	16.6	NAAT	0	NAAT	0	NR	NR	NR
Dasarathan and Kalaivani [41]	2011-2014	CH	Chennai	82	Facility	RDS	Blood	VDRL + TPHA	20.7	NR	NR	NR	NR	NR	NR	NR

CH- Cohort study; CS- Cross-sectional study; TLC- Time Location Cluster, NR- Not Reported; TPHA- Treponema Pallidum Heamagglutination Test, RPR Rapid Plasma Reagin; NAAT- Nucleic Acid Amplification Test

Findings among H/TG

From the literature search it was revealed that among the all STIs, syphilis was only re-reported among H/TGs in India. Present search finds out. four articles [11,29,31,41]. The recent published article on facility level study [41], which was conducted in a cohort of a sub group of males to female transgenders estimates prevalence of syphilis was

20.7 (Figure 5). Other four studies reveal from the present search, are reported from Tamilnadu, Mumbai, Chennai, and Pune. Although current finding shows a poor regional representation of H/TG on STI prevalence (Table 5).

Findings among PWID

There were only four articles reported STI prevalence among PWID [11,42-44], found chlamydia in India. A cross-

Table 6: Aetiological diagnosis and prevalence of four curable STIs among PWID from available published articles from different parts of India.

Study	Study period	Study design	Location	Sample size	Study setting	Sampling Technique	Specimen	Syphilis		Gonorrhoea		Chlamydia		Trichomoniasis	
								Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence	Methods of diagnosis	Prevalence
Mahanta et al. [42]	NR	CS	Bishnupur	420	Community	RDS	Blood	RPR + TPHA	5.7	NR	0.3	NR	1.7	NR	NR
Mahanta et al. [42]	NR	CS	Churachand	419	Community	RDS	Blood	RPR + TPHA	0.9	NA	NA	NA	NA	NR	NR
				417			Urine	NA	NA	NR	0	NR	2.1		
Mahanta et al. [42]	NR	CS	Phek	440	Community	RDS	Blood	RPR + TPHA	7.4	NA	NA	NA	NA	NR	NR
				430			Urine	NA	NA	NR	0.6	NR	11.4		
Mahanta et al. [42]	NR	CS	Wokha	420	Community	RDS	Blood	RPR + TPHA	19.5	NA	NA	NA	NA	NR	NR
				409			Urine	NA	NA	NR	1.6	NR	11		
Mahanta et al. [42]	NR	CS	Mumbai & Thane	355	Community	RDS	Blood	RPR + TPHA	4.9	NR	NR	NR	NR	NR	NR
				373			Urine	NA	NA			NR	0.7		
Ghosh et al. [15]	NR	CS	West Bengal	58	Facility	NR	Blood	TPHA	0	NR	NR	NR	NR	NR	NR
Armstrong et al. [43]	2009	CS	Nagaland	1705	Community	RDS	Blood for syphilis, urine for chlamydia	RPR+TPHA	13.4	NR	NR	NAAT	9.1	NR	NR
Armstrong et al. [43]	2009	CS	Manipur	1657	Community	RDS	Blood for syphilis, urine for chlamydia	RPR+TPHA	3.9	NR	NR	NAAT	1.8	NR	NR
Goswami et al. [44]	2014	CS	Manipur	1657	Community	RDS	Blood for syphilis, urine for chlamydia	RPR+TPHA	4	NR	NR	NAAT	0.2	NR	NR
Goswami et al. [44]	2014	CS	Nagaland	1705	Community	RDS	Blood for syphilis, urine for chlamydia	RPR+TPHA	13	NR	NR	NAAT	1	NR	NR

CH- Cohort study; CS- Cross-sectional study; TLC- Time Location Cluster, NR- Not Reported; TPHA- Treponema Pallidum Heamagglutination Test, RPR Rapid Plasma Reagin; NAAT- Nucleic Acid Amplification Test

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sectional study on PWID published on 2008 carried out in Manipur, Nagaland and Maharashtra (Mumbai, and Thane), which reported seroprevalence of syphilis and gonorrhoea as 19.5% and 1.6 respectively at Wokha, and 11.4% as chlamydia for Phek Nagaland [42]. without mentioning any specific study period. Another article comprised of data from North-eastern India viz. Phek, Wokha, Churachandpur, Bishnupur along with Mumbai-Thane [43] reported on prevalence of syphilis and chlamydia. Representation from North-eastern region was reported during present literature search (Figure 6), but no STI prevalence was reported from West Bengal. The data also implied a weak regional representation of STI prevalence among PWID (Table 6) in India.

Key findings

Forty articles were included in this systematic review after screening for inclusion criteria. Ghosh et al., 2012 reported a very high syphilis prevalence is 28.8 among FSW in West Bengal for a very small sample size of 45. A very low syphilis prevalence 0.8 was reported [28] among MSM in Belgaum, where the sample size was 1003. A facility based cross-sectional study was conducted in Mumbai and Hyderabad for a sample size of 149 [30] reported a very high prevalence for Gonorrhoea (14.8) among MSM. However,

a community based cross sectional study conducted in Tamilnadu [31] reported a very low gonorrhoea prevalence (0.07) among MSM with the sample size of 1621. Only nine articles reported Chlamydia among key population. A very high chlamydia prevalence (22.2) was reported by Medhi et al. [16] a community based cross-sectional study conducted on FSW in Nagaland with a sample size of 423. A similar community based cross-sectional study conducted in Tamilnadu [33] reported a very low chlamydia prevalence, 0.3 for MSM, Trichomoniasis was reported only by 1/10th [n=40] of the reviewed articles. From the available data of the included studies, it was found that STI prevalence was high when samples were taken from facility setting irrespective of region and key populations.

Quality of the selected studies

The modified Newcastle - Ottawa Quality Assessment Scale was applied to determine quality of included studies and the result was presented pictorially (Figure 7). Here, it must be noted that, quality of all the included studies calculated and presented here is only to accomplish the demands of present systematic review, except this author have no purposeful intention (Figure 7).

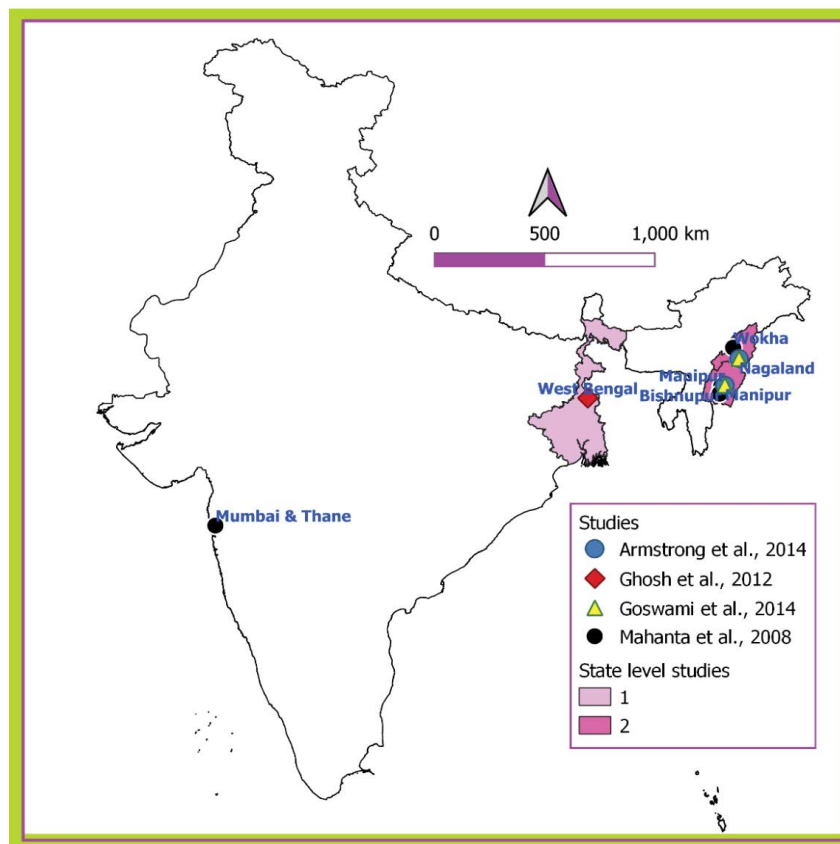


Figure 6: Representation of published articles on four curable STIs among PWIDs in India.

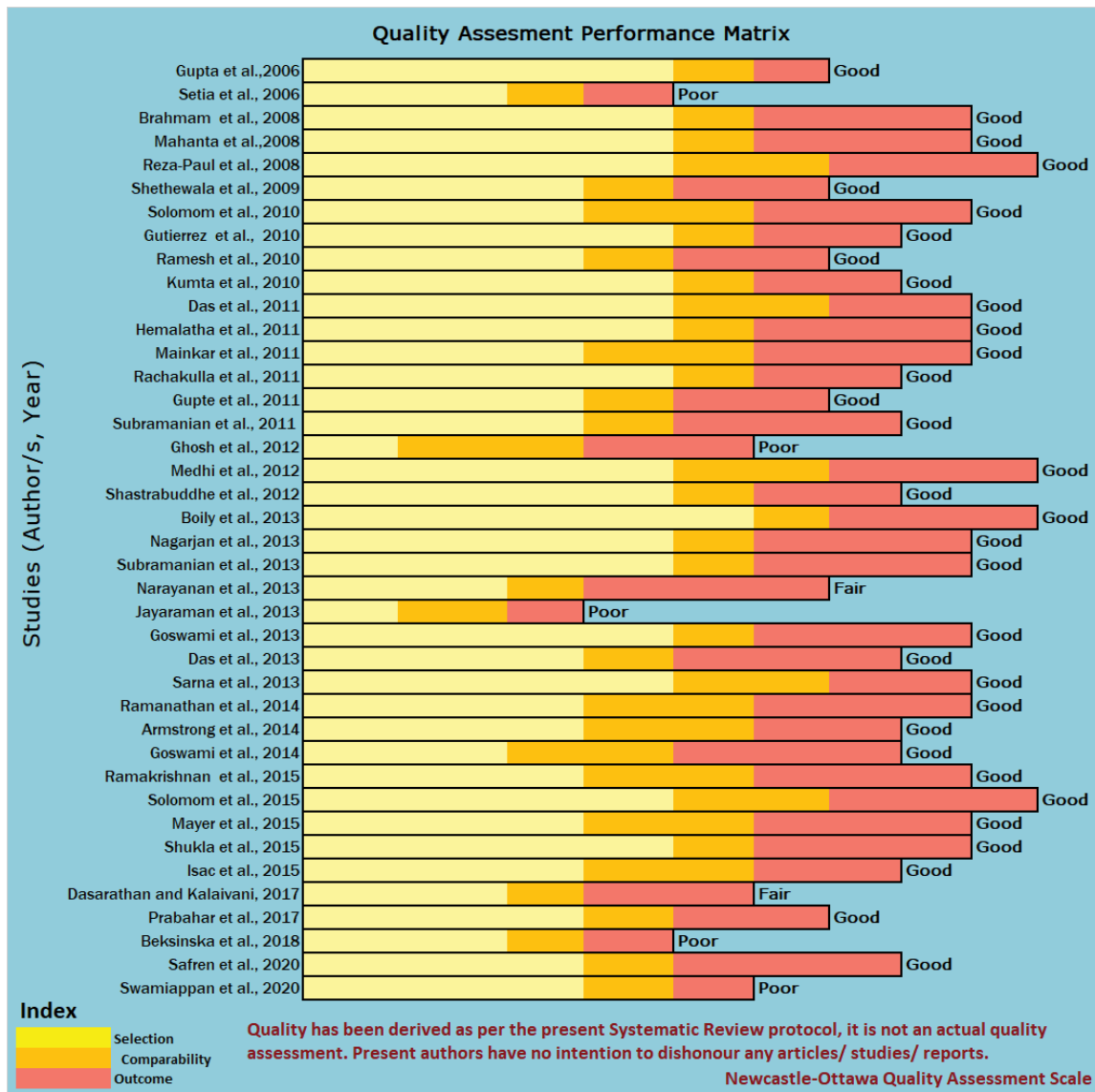


Figure 7: Quality Assessment Performance Matrix.

Strength and limitations

The prevalence of STIs shows considerable heterogeneity by geographical setting and key population group. At the same time, limited numbers of study reported STI prevalence, sampling heterogeneity across studies prevent definite conclusions and how the prevalence of STIs varies among key populations. All the included studies in the present systematic review used probability-based sampling and both in facility as well as community-based setting. However, there were limited data points from all regions in India. The variation among included studies highlights a limitation of the present review, as the findings can vary based on the electronic databases and selected search terms. However, present article summarizes of all study that allows pre-defined acceptance criteria and qualify the quality assessment tool (Tables 1-6).

Discussion

Forty articles were included in this systematic review after screening for inclusion criteria. State-wise distribution revealed that Tamilnadu, Andhra Pradesh, Karnataka, Maharashtra were study locations for 3/4th of the included articles [n=40]. Two articles determined prevalence of the four STIs among PWID in Manipur and Nagaland. Three articles with the prevalence of STIs among FSW and MSM in the cities of central and Northern India. One article was found from West Bengal. There were three articles that reported aetiological as well as syndromic prevalence among key population. One article reported only syndromic prevalence for FSW. Remaining articles reported the prevalence of four STIs among key populations through aetiological diagnosis. RPR and TPHA tests were conducted for diagnosing

syphilis; NAAT or Gen Probe APTIMA were used for diagnosing gonorrhoea and chlamydia; kupferburgs culture, wet mount microscopy tests were conducted for diagnosing trichomoniasis. Over the last two decades, the National AIDS Control Program of India (NACP) implemented by National AIDS Control Organization (NACO) which, undertakes prevention of STI as one of the key strategies. NACO through its network of designated STI or RTI (reproductive tract infection) clinics (situated at government health care facilities located mostly at the district level and above) is providing free standardized STI/RTI services. These clinics have been branded as “Suraksha Clinics” and provide sexual and reproductive health services [45]. Function of these Centre is to provide validation of syndromic case management by doing etiologic testing, antibiotic susceptibility testing for Gonococci, EQAS for syphilis, and conducting operations research providing evidence to the programme. [46]. Still, there is a limited literature till date to report pooled prevalence among key populations across India and these data have not been integrated to inform the overall geographical and temporal trends of STI infections among various key populations in India.

Conclusion

The present systemic review had generated on the prevalence and spatiotemporal distribution of the four STIs in Indian key population. Prevalence of STIs showed extensive heterogeneity through spatiotemporal setting and people practicing high-risk behaviour in India. A national STI surveillance cum prevention programme is essential among key populations in India. In the light of present findings and with the identified limitations, it can be concluded that existing HIV surveillance system under NACP, can be utilized with additional bio-specimen collection to determine STI prevalence among high risk populations. However, in future, with the availability of the comparable data from most of the regions in India, it will be possible to conduct a systemic review with pooled prevalence on the spatiotemporal distribution of the four curable STIs in the general population of India.

Supplementary Materials:

The following supporting information can be downloaded at: www.mdpi.com/xxx/s1, Table S1: search terms and search strategy; Table S2 PRISMA chart.

Author Contributions:

MB, SB, AR, BTV, BSB, TLNP, LS, ND and SD Conceptualize and design the study. Data curation was done by MB, UG, PG, PB, and SB. Investigation MB, UG, PG, PB, SA, and SB. Methodology MB, UG, PB, SA, SB, AKD, DC, CB, GKM, BSB, FD and SD. Formal analysis UG, PG, MB, PGG, PK, BSB, GKM and PB. Funding acquisition AR, RA,

and SD. Project administration MB, TLNP, LS, ND, SB, RA, and SD. Supervised by BTV, TLNP, LS, MB, ND, PR, CB, RA, AKD, GKM and DC. Resources DC, AKD, AR, RA, and FD. Writing – original draft MB, UG, PK, PB, and SB; and Writing – review & editing MB, AM, TLNP, PR, PK, PGG, BSB, SA, SB, and SD. All authors read the manuscript before they have given the final approval for publication.

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