

Open Peer Review on Qeios

What is the impact of COVID-19 era on annual tuberculosis notifications in India? A comparative study (2017-2022)

Dr Piyush Kumar¹, Advocate Anupama², Harshika Singh³

1 B. R. Ambedkar Bihar University

2 Bihar State Bar Council 3 University of Glasgow

Funding: No specific funding was received for this work

Potential competing interests: No potential competing interests to declare.

Abstract

Globally Tuberculosis (TB) is 13th leading cause of mortality and the 2nd leading infectious killer after COVID-19, 1.6 million people died in 2021 from TB (including 187 000 people with HIV). The emergence of COVID-19 pandemic situations on the already overburdened health care system of India may have affected new TB cases notification as well as treatment, setting back the previous achievements made to eliminate TB by 2025. Lockdown-induced under-reporting (80% reduction in TB notification rates) of active TB will affect TB epidemiology as well as NTEP (National Tuberculosis Elimination Program) elimination goal by 2025. There are significant studies which found that COVID-19 is promoting the growth of active tuberculosis in a patient with dormant/latent tuberculosis infection. In the COVID-19 epidemic, diagnosis and treatment of tuberculosis, or tuberculosis and the co-infection with COVID-19, may be compromised as TB and COVID-19 both present with respiratory symptoms that are similar to each other. According to the 2019 India TB Report, 2, 24,000 incident cases per year of paediatric TB are estimated accounting for 22% of global burden. In spite of many similarities and differences, there is still a lack of clarity about the exact epidemiological relationship between COVID-19 and TB. This was a cross-sectional, retrospective, quantitative; research study aimed to know the impact of COVID-19 era on TB (tuberculosis) notifications in India (includes all 36 states and UTs of India). All the 36 participants' states and UTs newly detected tuberculosis cases notifications were followed from 1st January 2017 to 31st December 2022. The total number of newly detected tuberculosis cases notifications from 1st January 2017 to 31st December 2022 was 12290180 (n) including all 36 participants. The (period prevalence) total number of newly detected tuberculosis cases notifications during pre-COVID-19 era (2017-2019) was 5914621 and during the COVID-19 era (2020-2022) were 637559. The total annual newly detected tuberculosis cases notifications (including public and private healthcare facilities) in India including all 36 participants from beginning of year 1st January till 31st December during a year were 1525045, 1997873, 2391703, 1810834, 2145678, 2419047, for years 2017, 2018, 2019, 2020, 2021, and 2022 respectively. The mean of newly detected tuberculosis cases notifications from all public and private healthcare facilities during Pre-COVID-19 era (2017-2019) was 164295 (Std. Err. - 37404.92; [95% Conf. Interval] -88359.01- 240231; Std. Dev. - 224429.5). The mean of newly detected tuberculosis cases notifications from all public and private healthcare facilities during COVID-19 era (2020-2022) was 177098.9 (Std. Err. - 43081.17; [95% Conf. Interval] - 89639.43 -264558.3; Std. Dev. - 258487). The incidence rate of newly detected tuberculosis cases notifications per lakh population was 112.62; 145.94; 172.92; 129.68; 152.44; and 169.63 respectively for years 2017, 2018, 2019, 2020, 2021, 2022. This research study revealed that the incidence rate of newly detected tuberculosis cases notifications per lakh population kept on increasing during the pre-COVID-19 era and it decreased abruptly during the first COVID-19 year 2020 by 25.00 percent (%) compared to previous year of pre-COVID-19 era 2019.

Key Author

Dr Piyush Kumar, M.B.B.S. E.M.O.C., P.G.D.P.H.M.¹

¹ Senior General Medical Officer, Health Department, Government of Bihar, India, drpiyush003@gmail.com; ORCID: https://orcid.org/0000-0001-9857-478X.

Co-Authors

Advocate Anupama²

 $^2 \ Senior \ Lawyer- \ Bar-council, \ Patna, \ Bihar, \underline{advocateanupama007@gmail.com}, \ ORCID: \ \underline{https://orcid.org/0000-0003-3651-2705}$

Harshika Singh

³ Masters Student of Epidemiology of Infectious Disease and Antimicrobial Resistance, University of Glasgow UK<u>harshika1437@gmail.com</u>

Impact of COVID-19 era on annual tuberculosis notifications in India

Public Health / Original Research

Keywords: COVID-19, Tuberculosis, Incidence, Impact, Health, annual, notification.

Introduction

Background/rationale

Globally Tuberculosis (TB) is 13th leading cause of mortality and the 2nd leading infectious killer after COVID-19, 1.6 million people died in 2021 from TB (including 187 000 people with HIV) [1]. The emergence of COVID-19 pandemic situations on the already overburdened health care system of India may have affected new TB cases notification as well as treatment, setting back the previous achievements made to eliminate TB by 2025 [2]. The COVID-19 have affected several routine healthcare services deliveries even in well to do countries with advanced technologies and ample human resource [3]. Worldwide, TB incidence is reducing at the rate of 2%per year and between 2015 and 2020 the total decrease was 11% only



which was only half way to achieve the goal to eliminate TB Strategy milestone by achieving 20% reduction between 2015 and 2020 [1]. The most obvious impact of COVID-19 was alarge global reduction in the reported number of newly diagnosed TB which fell from 7.1 million in 2019, to 5.8 million in 2020 (–18%) and India is among the three countries which accounted for most of the reduction in 2020 besides Indonesia and the Philippines (67% of the global total) [4] India have remarkably high annual TB incidence at 210/100,000 in 2021 as per data of World Bank [5].

An essential element of the global effort to eradicate tuberculosis (TB) is curbing TB in India. Approximatel \$25\$ per cent of the world's TB cases are estimated to have occurred in India [6]. By 2025, five years before the global deadline in 2030, India has committed to meeting the Sustainable Development Goal targets of 80% TB incidence reduction and 90% TB death reduction [7]. The key author previous research work (is available at WHO COVID-19 research database) has found that with phase-wise unlocking beginning on 1 June 2020, a nationwide lockdown was implemented in India in response to the COVID-19 pandemic for 68 days (from 25 March to 31 May 2020) which disrupted several routine essential healthcare services delivery in India as [8]. It is anticipated that lockdown-induced poverty and under nutrition will worsen the TB cases situation by downgrading poor population health status, pushing those living in vulnerable economic circumstances below poverty line, and that lockdown-induced under-reporting (80% reduction in TB notification rates) of active TB will affect TB epidemiology as well as NTEP (National Tuberculosis Elimination Program) elimination goal by 2025 [9, 10]. There are significant studies which found that COVID-19 is promoting the growth of active tuberculosis in a patient with dormant/latent tuberculosis infection [11]. One study found that high mortality (11%) is associated with COVID-19/TB co-infection, additionally, M. tuberculosis co-infection also resulted in the more severe COVID-19 and a faster progression of the disease [12]. The World Bank determined that \$3.2 a day is a poverty line for lower-middle-income countries, and the economic downturn due to COVID-19 may push 104 million additional Indians into poverty [13].

In the COVID-19 epidemic, diagnosis and treatment of tuberculosis, or tuberculosis and the co-infection with COVID-19, may be compromised as TB and COVID-19 both present with respiratory symptoms that are similar to each other [14]. COVID-19 often affects children less severely, whereas 1.2 million children fell ill with TBin 2021, globally and TB is at present the leading cause of death for children of all ages globally dying from infectious diseases[15, 16]. According to the 2019 India TB Report, 2, 24,000 incident cases per year of paediatric TB are estimated accounting for 22% of global burden [17]. As India rank fourth in COVID-19 cases and mortality, there is a possibility that the number of TB cases and deaths might increase in the future [18]. In spite of many similarities and differences, there is still a lack of clarity about the exact epidemiological relationship between COVID-19 and TB [19].

Hence to know the real situation this study was carried out in order to alert policy maker for needful action.

Methods

This was a cross-sectional, retrospective, quantitative; research study aimed to know the impact of COVID-19 era on TB (tuberculosis) notifications in India(includes all 36 states and UTs of India). This version of study is having objective to assess the new TB cases (tuberculosis) notifications of India at country level and theext version of this study will discuss performance of different states and UTs on individual levels. As the first case of COVID-19 was notified in January 2020 hence for this studyperiod before 1st January 2020 (1st January 2017 to 31st December 2019) is considered as pre COVID-19 era and from 1st January 2020 till end of this study was considered as COVID-19 era(1st January 2020 to 31st December 2022), see table-1 and 2 [20].

Table 1. Annual pre COVID-19 era TB case notifications across 36 states and UTs of India from 1st January 2017 onward till 31st December 2019



| State Wise Total Notified From 01/01/2017 To: 31/12/2019 | 2017 | 2017 | 2018 | 2018 | 2019 | 2019 | PRE-COVID ERA |
|--|-------------------------------|--------------------------------|-------------------------------|--------------------------------|-------------------------------|--------------------------------|------------------------------------|
| State | 2017-Total Public Notified | 2017-Total Private Notified | 2018-Total Public Notified | 2018-Total Private Notified | 2019-Total Public Notified | 2019-Total Private Notified | Total Public & Private Notified |
| Andaman & Nicobar Islands | 615 | 13 | 544 | 18 | 573 | 7 | 1770 |
| Andhra Pradesh | 62731 | 5787 | 65645 | 19142 | 76239 | 22222 | 251766 |
| Arunachal Pradesh | 3297 | 2 | 3045 | 4 | 2908 | 36 | 9292 |
| Assam | 36066 | 618 | 36929 | 4115 | 40612 | 7982 | 126322 |
| Bihar | 55906 | 9298 | 66337 | 32282 | 77922 | 44240 | 285985 |
| CHANDIGARH | 4225 | 85 | 5124 | 176 | 6457 | 496 | 16563 |
| Chhattisgarh | 31280 | 3764 | 29901 | 8736 | 31536 | 11679 | 116896 |
| Dadra and Nagar Haveli and Daman and Diu | 1282 | 59 | 1244 | 83 | 1357 | 130 | 4155 |
| Delhi | 57255 | 1667 | 69911 | 9731 | 79743 | 27852 | 246159 |
| Goa | 1409 | 116 | 1833 | 431 | 1921 | 463 | 6173 |
| Gujarat | 97002 | 17115 | 106001 | 36750 | 104921 | 53865 | 415654 |
| Haryana | 35060 | 3350 | 47836 | 11727 | 50895 | 21835 | 170703 |
| Himachal Pradesh | 13477 | 542 | 15181 | 1244 | 15785 | 1560 | 47789 |
| Jammu & Kashmir | 8218 | 571 | 10499 | 764 | 10493 | 904 | 31449 |
| Jharkhand | 36155 | 1991 | 38487 | 6884 | 43677 | 12544 | 139738 |
| Karnataka | 63636 | 5705 | 67316 | 11942 | 71861 | 19619 | 240079 |
| Kerala | 17160 | 3671 | 20926 | 3227 | 20659 | 4884 | 70527 |
| Ladakh | 274 | 2 | 355 | 57 | 360 | 29 | 1077 |
| Lakshadweep | 49 | 0 | 19 | 0 | 15 | 0 | 83 |
| Madhya Pradesh | 116574 | 6067 | 120681 | 29650 | 139013 | 47445 | 459430 |
| Maharashtra | 123919 | 18877 | 139098 | 52196 | 143187 | 82282 | 559559 |
| Manipur | 1626 | 304 | 2165 | 413 | 2004 | 551 | 7063 |
| Meghalaya | 3171 | 222 | 4025 | 573 | 4709 | 724 | 13424 |
| Mizoram | 2457 | 41 | 2561 | 36 | 2939 | 40 | 8074 |
| Nagaland | 2439 | 275 | 3678 | 471 | 4149 | 696 | 11708 |
| Odisha | 48319 | 1933 | 45807 | 2683 | 48889 | 4479 | 152110 |
| Puducherry | 1890 | 5 | 3485 | 30 | 4564 | 72 | 10046 |
| Punjab | 36252 | 3251 | 42232 | 8211 | 43891 | 13940 | 147777 |
| Rajasthan | 85440 | 8161 | 111042 | 37519 | 121574 | 51374 | 415110 |
| Sikkim | 1199 | 2 | 1507 | 11 | 1427 | 24 | 4170 |
| Tamil Nadu | 77815 | 8708 | 75392 | 19359 | 82290 | 27229 | 290793 |
| Telangana | 37009 | 1713 | 42087 | 8922 | 50554 | 20551 | 160836 |
| Tripura | 2036 | 0 | 2624 | 19 | 2719 | 46 | 7444 |
| Uttar Pradesh | 235505 | 18069 | 303728 | 82555 | 326306 | 159771 | 1125934 |
| Uttarakhand | 13393 | 3496 | 16951 | 3872 | 19744 | 6157 | 63613 |
| West Bengal | 82118 | 3306 | 86951 | 12893 | 84897 | 25185 | 295350 |
| Total | 1396259 | 128786 | 1591147 | 406726 | 1720790 | 670913 | 5914621 |

In India, all the TB (tuberculosis) new case notifications from 36 states and UTs of India are electronically transmitted / reported / communicated through the Government of India established web based NIKSHAY platform [21]. We collected data from the NIKSHAY platform for annual case notifications across 36 states and UTs of India from 1st January 2017 onward till 31st December 2022, to compare three equal pre-pandemic years with three years of COVID-19 erato ascertain the positive / negative impact of COVID-19 on the TB notification ratessee table-1 and 2.

Table 2. Annual COVID-19 era TB case notifications across 36 states and UTs of India from 1st January 2020 onward till 31st December 2022



| State Wise Total Notified From 01/01/2020 To: 31/12/2022 | 2020 | 2020 | 2021 | 2021 | 2022 | 2022 | COVID-19 ERA |
|--|-----------------------|------------------------|-----------------------|------------------------|-----------------------|------------------------|---------------------------------|
| State | Total Public Notified | Total Private Notified | Total Public Notified | Total Private Notified | Total Public Notified | Total Private Notified | Total Public & Private Notified |
| Andaman & Nicobar Islands | 481 | 0 | 506 | 6 | 510 | 24 | 1527 |
| Andhra Pradesh | 46868 | 17235 | 62124 | 24765 | 62010 | 30106 | 243108 |
| Arunachal Pradesh | 2522 | 1 | 2750 | 16 | 2715 | 141 | 8145 |
| Assam | 29271 | 6202 | 29729 | 8385 | 36662 | 10972 | 121221 |
| Bihar | 52317 | 47521 | 62365 | 70260 | 78619 | 82052 | 393134 |
| CHANDIGARH | 3767 | 537 | 4245 | 506 | 5653 | 399 | 15107 |
| Chhattisgarh | 20981 | 8400 | 23695 | 8882 | 26739 | 11684 | 100381 |
| Dadra and Nagar Haveli and Daman and Diu | 869 | 81 | 947 | 65 | 1293 | 105 | 3360 |
| Delhi | 59631 | 27086 | 68274 | 35256 | 76790 | 29757 | 296794 |
| Goa | 1340 | 326 | 1631 | 381 | 1612 | 455 | 5745 |
| Gujarat | 77184 | 43305 | 92852 | 51853 | 100903 | 50911 | 417008 |
| Haryana | 41472 | 21482 | 45620 | 23546 | 51152 | 24493 | 207765 |
| Himachal Pradesh | 12194 | 1273 | 13062 | 1515 | 14439 | 1630 | 44113 |
| Jammu & Kashmir | 7943 | 881 | 9503 | 1395 | 9995 | 1783 | 31500 |
| Jharkhand | 30516 | 15349 | 35455 | 17155 | 43573 | 13612 | 155660 |
| Karnataka | 48716 | 17262 | 52972 | 19745 | 59446 | 20840 | 218981 |
| Kerala | 15076 | 5822 | 15437 | 6628 | 16725 | 6590 | 66278 |
| Ladakh | 231 | 6 | 282 | 10 | 311 | 9 | 849 |
| Lakshadweep | 18 | 0 | 12 | 0 | 11 | 0 | 41 |
| Madhya Pradesh | 104682 | 33284 | 111154 | 55870 | 130140 | 55364 | 490494 |
| Maharashtra | 95748 | 64693 | 110116 | 90320 | 133590 | 102293 | 596760 |
| Manipur | 1151 | 433 | 1256 | 544 | 1627 | 917 | 5928 |
| Meghalaya | 3455 | 689 | 3278 | 892 | 4054 | 920 | 13288 |
| Mizoram | 1991 | 131 | 1486 | 273 | 1696 | 388 | 5965 |
| Nagaland | 2906 | 689 | 2971 | 741 | 3341 | 750 | 11398 |
| Odisha | 40435 | 5264 | 45156 | 7369 | 50331 | 9967 | 158522 |
| Puducherry | 2684 | 88 | 3407 | 49 | 3731 | 102 | 10061 |
| Punjab | 34694 | 11720 | 36737 | 14482 | 43218 | 11684 | 152535 |
| Rajasthan | 95855 | 41374 | 102979 | 46507 | 126480 | 42712 | 455907 |
| Sikkim | 1158 | 180 | 1305 | 102 | 1282 | 108 | 4135 |
| Tamil Nadu | 54008 | 16522 | 64534 | 18611 | 71842 | 21909 | 247426 |
| Telangana | 40540 | 22703 | 41488 | 19311 | 52264 | 20556 | 196862 |
| Tripura | 1995 | 73 | 2448 | 111 | 2860 | 156 | 7643 |
| Uttar Pradesh | 242722 | 125317 | 315422 | 140843 | 372651 | 149230 | 1346185 |
| Uttarakhand | 14292 | 5798 | 17356 | 5610 | 21157 | 6328 | 70541 |
| West Bengal | 61905 | 17489 | 69644 | 21476 | 76751 | 23927 | 271192 |
| Total | 1251618 | 559216 | 1452198 | 693480 | 1686173 | 732874 | 6375559 |
| | | | | | | | |

We then performed a linear regression of existing data of pre pandemic years for counterfactual analysis to find out the TB cases which may be notified if the natural intervention COVID-19 had not occurred. The data were collected and analysed with Microsoft office and stata 15.1software.

Data sources/measurement

The data is available at link given below:

https://reports.nikshay.in/Reports/TBNotification

Bias

To reduce the bias three equal years before and during pandemic were included in this study.

Results

This six year retrospective novel cross-sectional research study included all 36 numbers of different states and union territories (UTs) of India to assess the overall country status regarding the title mentioned research question. All the states and UTs were potentially eligible due to the fact that they (36 states and UTs) continuously and regularly register/communicate/transmit all the of newly detected tuberculosis cases notifications electronically through the NIKSHAY platform of Government of India. NI-KSHAY-(Ni=End, Kshay=TB) is the online web based patient management system in India exclusively for TB (Tuberculosis) control under the NTEP-2020 (National Tuberculosis Elimination Programme, previously RNTCP) with the key aim to eliminate TB by 2025. All the 36 participants' states and UTs newly detected tuberculosis cases notifications were followed from ft January 2017 to 31st December 2022 and analyzed to get an answer to the title mentioned research question. All participants participated from beginning to end of this study.



There are wide variations in demographic, clinical, social characteristics of above mentioned 36 participants evident from NITI AAYOG annual reports [22]. Information on exposures and potential confounders of different participants is not taken into account as this is a quantitative research study based on secondary data. There were no missing data.

The total number of newly detected tuberculosis cases notifications from1st January 2017 to 31st December 2022 was 12290180 (n) including all 36 participants. The(period prevalence) total number of newly detected tuberculosis cases notifications during pre-COVID-19 era (2017-2019) was 5914621 and during the COVID-19 era (2020-2022) were 637559. The total annual newly detected tuberculosis cases notifications (including public and private healthcare facilities) in India including all 36 participants from beginning of yeaf st January till 31st December during a year were 1525045, 1997873, 2391703, 1810834, 2145678, 2419047, for years 2017, 2018, 2019, 2020, 2021, and 2022 respectively, see table-1 and 2

During the base year 2017 of this research study the total newly detected tuberculosis cases notifications from public and private healthcare facilities were 1396259 and 128786 respectively while for consecutive years 2018, 2019, 2020, 2021, 2022 of study it was (1591147, 406726), (1720790, 670913), (1251618, 559216,) (1452198,693480,) and (1686173,732874) respectively, see table-1 and 2.

2017

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2017 was 38784.97 (Std. Err.-8202.51; [95% Conf. Interval]-22132.99 -55436.95; Std. Dev.-49215.05) whereas for private healthcare facilities during 2017 it was 3577.39 (Std. Err.-860.98; [95% Conf. Interval] - 1829.51-5325.26; Std. Dev. - 5165.85). Among public healthcare facilities Lakshadweep reported minimum 49 newly detected tuberculosis cases notifications whereas maximum 235505 was reported from UP (Uttar Pradesh) during 2017. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 18877 was reported from Maharashtra during 2017, see table-3.

2018

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2018 was 44198.53 (Std. Err.-9905.34; [95% Conf. Interval]- 24089.62-64307.44; Std. Dev. - 59432.04) whereas for private healthcare facilities during 2018 it was 11297.94 (Std. Err.- 2985.37; [95% Conf. Interval] - 5237.322- 17358.57; Std. Dev. - 17912.22). Among public healthcare facilities Lakshadweep reported minimum 19 newly detected tuberculosis cases notifications whereas maximum 303728 was reported from UP (Uttar Pradesh) during 2018. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 82555 was reported from UP during 2018, see table-3.

2019

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2019 was 47799.72 (Std. Err. - 10636.6; [95% Conf. Interval] - 26206.28 -69393.16; Std. Dev. - 63819.58) whereas for private healthcare facilities during 2019 it was 18636.47 (Std. Err. - 5203.06; [95% Conf. Interval] - 8073.698- 29199.25; Std. Dev. - 31218.36). Among public healthcare facilities Lakshadweep reported minimum 15 newly detected tuberculosis cases notifications whereas maximum 326306 was reported from UP (Uttar Pradesh) during 2019. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 159771 was reported from UP during 2019, see table-3.

The mean of newly detected tuberculosis cases notifications from all public and private healthcare facilities during Pre-COVID-19 era (2017-2019) was 164295 (Std. Err. - 37404.92; [95% Conf. Interval] - 88359.01- 240231; Std. Dev. - 224429.5). Among all public and private healthcare facilities Lakshadweep reported minimum 83 newly detected tuberculosis cases notifications whereas maximum 1125934 was reported from UP (Uttar Pradesh) during Pre COVID-19 era (2017-2019), see table-3.

| Table 3. Statistical Analysis of tuberculosis cases notifications of Pre-COVID era. | | | | | | | | |
|---|-----|----------|-----------|---|--------------|-----|---------|--|
| Variable | Obs | Mean | Std. Err. | Confidence Interval-Mean [95% Conf. Interval] | Std. Dev. | Min | Max | |
| 2017-Total Public Notified | 36 | 38784.97 | 8202.51 | 22132.99 -55436.95 | 49215.05 | 49 | 235505 | |
| 2017-Total Private Notified | 36 | 3577.39 | 860.98 | 1829.51- 5325.26 | 5165.85 | 0 | 18877 | |
| 2018-Total Public Notified | 36 | 44198.53 | 9905.34 | 24089.62-64307.44 | 59432.04 | 19 | 303728 | |
| 2018-Total Private Notified | 36 | 11297.94 | 2985.37 | 5237.322- 17358.57 | 17912.22 | 0 | 82555 | |
| 2019-Total Public Notified | 36 | 47799.72 | 10636.6 | 26206.28 -69393.16 | 63819.58 | 15 | 326306 | |
| 2019-Total Private Notified | 36 | 18636.47 | 5203.06 | 8073.698- 29199.25 | 31218.36 | 0 | 159771 | |
| Pre-COVID-19 Total Public & Private Notified | 36 | 164295 | 37404.92 | 88359.01- 240231 | 224429.5 | 83 | 1125934 | |

2020

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2020 was 34767.17 (Std. Err. - 7838.52; [95% Conf. Interval] - 18854.13-50680.21; Std. Dev. - 47031.11) whereas for private healthcare facilities during 2020 it was 15533.78 (Std. Err. - 4142.46; [95% Conf. Interval] - 7124.13 -23943.43; Std. Dev. - 24854.78). Among public healthcare facilities Lakshadweep reported minimum 18 newly detected tuberculosis cases notifications whereas maximum 242722 was reported from UP (Uttar Pradesh) during 2020. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 125317 was reported from UP during 2020, see table-4



2021

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2021 was 40338.83 (Std. Err. - 9743.34; [95% Conf. Interval]- 20558.81 -60118.86; Std. Dev. - 58460.02) whereas for private healthcare facilities during 2021 it was 19263.33 (Std. Err. - 5064.32; [95% Conf. Interval] - 8982.22-29544.45; Std. Dev. - 30385.93). Among public healthcare facilities Lakshadweep reported minimum 12 newly detected tuberculosis cases notifications whereas maximum 315422 was reported from UP (Uttar Pradesh) during 2021. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 140843 was reported from UP during 2021, see table-4

2022

The mean of newly detected tuberculosis cases notifications from public healthcare facilities during 2022 was 46838.14 (Std. Err. - 11485.15; [95% Conf. Interval] - 23522.05-70154.23; Std. Dev. - 68910.89) whereas for private healthcare facilities during 2022 it was 20357.61 (Std. Err. - 5426.21; [95% Conf. Interval] - 9341.82 - 31373.41; Std. Dev. - 32557.27). Among public healthcare facilities Lakshadweep reported minimum 11 newly detected tuberculosis cases notifications whereas maximum 372651 was reported from UP (Uttar Pradesh) during 2022. Among private healthcare facilities Lakshadweep reported minimum 0 newly detected tuberculosis cases notifications whereas maximum 149230 was reported from UP during 2022, see table-4

| Table 4. Statistical Analysis of tuberculosis cases notifications of COVID-19 era | | | | | | | | |
|---|-----|----------|-----------|---|--------------|-----|---------|--|
| Variable | Obs | Mean | Std. Err. | Confidence Interval-Mean [95% Conf. Interval] | Std. Dev. | Min | Max | |
| 2020-Total Public Notified | 36 | 34767.17 | 7838.52 | 18854.13- 50680.21 | 47031.11 | 18 | 242722 | |
| 2020-Total Private Notified | 36 | 15533.78 | 4142.46 | 7124.13 -23943.43 | 24854.78 | 0 | 125317 | |
| 2021-Total Public Notified | 36 | 40338.83 | 9743.34 | 20558.81 -60118.86 | 58460.02 | 12 | 315422 | |
| 2021-Total Private Notified | 36 | 19263.33 | 5064.32 | 8982.22-29544.45 | 30385.93 | 0 | 140843 | |
| 2022-Total Public Notified | 36 | 46838.14 | 11485.15 | 23522.05- 70154.23 | 68910.89 | 11 | 372651 | |
| 2022-Total Private Notified | 36 | 20357.61 | 5426.21 | 9341.82 -31373.41 | 32557.27 | 0 | 149230 | |
| COVID-19-Total Public & Private Notified | 36 | 177098.9 | 43081.17 | 89639.43 -264558.3 | 258487 | 41 | 1346185 | |

The mean of newly detected tuberculosis cases notifications from all public and private healthcare facilities during COVID-19 era (2020-2022) was 177098.9 (Std. Err. - 43081.17; [95% Conf. Interval] - 89639.43 -264558.3; Std. Dev. - 258487). Among all public and private healthcare facilities Lakshadweep reported minimum 41 newly detected tuberculosis cases notifications whereas maximum 1346185 was reported from UP (Uttar Pradesh) during COVID-19 era (2020-2022).

Other analyses

Total newly detected tuberculosis cases notifications in a specified year x 100000

Incidence Rate (newly detected tuberculosis cases notifications) per lakh population = Population (forecasted value as real census is not done) at risk (we considered whole population at risk)

| Table | Table 5. Incidence Rate of newly detected tuberculosis cases notifications in a specified year per lakh population | | | | | | | |
|-------|--|---|------------------------|--|--|--|--|--|
| Year | Population | Newly detected tuberculosis cases notifications from all public and private | Incidence Rate/lakh | | | | | |
| 2017 | 1354195680 | 1525045 | 112.62 | | | | | |
| 2018 | 1369003306 | 1997873 | 145.94 | | | | | |
| 2019 | 1383112050 | 2391703 | 172.92 | | | | | |
| 2020 | 1396387127 | 1810834 | 129.68 | | | | | |
| 2021 | 1407563842 | 2145678 | 152.44 | | | | | |
| 2022 | 1426099353 | 2419047 | 169.63 | | | | | |

Population Data Source- The World Bank available at -https://data.worldbank.org/indicator/SP.POP.TOTL?locations=IN

The populations for year 2022 was forecasted with the help of Microsoft excel by utilizing the previous all year available data.

The incidence rate of newly detected tuberculosis cases notifications per lakh population was 112.62; 145.94; 172.92; 129.68; 152.44; and 169.63 respectively for years 2017, 2018, 2019, 2020, 2021, 2022, see figure-1.



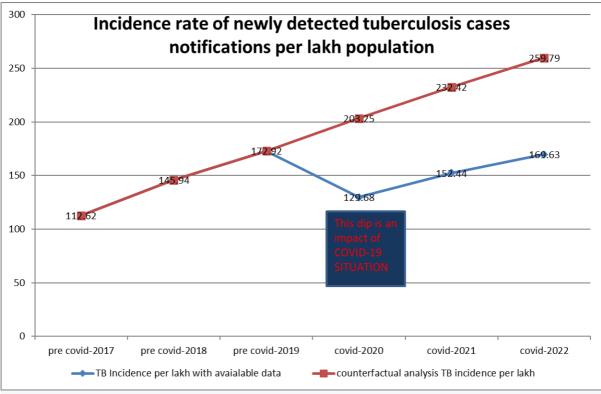


Figure 1. Comparison of newly detected tuberculosis cases notifications per lakh population with counterfactual analysis

Discussion

This research study revealed that the incidence rate of newly detected tuberculosis cases notifications per lakh populationkept on increasing during the pre-COVID-19 era and it decreased abruptly during the first COVID-19 year 2020 by 25.00 percent (%) compared to previous year of pre-COVID-19 era 2019 and 11.84 percent, 1.90 percent for subsequent years 2021, 2022. The incidence rate during pre-COVID-19 era increased by 29.59 and 18.49 percent compared to previous year data considering 2017 as base year.

The incidence rate during COVID-19 era also increased by 17.55 and 11.28 percent compared to previous year data considering 2020 as base year. The trend shows that before COVID-19 era as well as in COVID-19 era the incidence rate is increasing. In coming years it is expected to cross the pre-COVID-19 era 2019 highest incidence rate as well as there are more chances of abrupt increase in incidence of tuberculosis due to missed cases during COVID-19 era. We all know that the disease cannot be eradicated in one night and the big drop in incidence rate of newly detected tuberculosis cases notifications is most probably a negative impact of prevailing COVID-19 situations/restrictions. As compared to pre- COVID-19 eraIndia made partial recoveries in 2021, 2022 but still reduction is seen compared with 2019.

The statement of PIB (press information bureau), Government of Indiabased on Global TB Report 2022 released by WHO (World Health Organization) Posted On: 28 OCT 2022 6:22PM by PIB Delhi is questionable in the light of findings of this novel research study[23]. Without considering and analysing COVID-19 impact on the incidence rate of newly detected tuberculosis cases by visualizing pre- COVID-19 trends it seems that this above statement was made. This novel research study is intended to ALERT the policy makers that – THE incidence rate/lakh population of newly detected tuberculosis cases WILL KEEP INCREASING FOR FEW YEARS as it seems that a large number of TB new cases have been missed due to COVID-19 situations. (NTEP BY 2025 SEEMS TO BE IMPOSSIBLE IN LIGHT OF THIS RESEARCH STUDY- AVAILABLE SOON)

Counterfactual analysis

Considering COVID-19 as a natural intervention in routine healthcare activities we have analyzed the situation in the absence of the COVID-19 intervention, by comparing counterfactual outcomes to data observed under the intervention. Through simple linear regression we have forecasted the incidence rate of newly detected tuberculosis through the previous year's data of this study.

| Table 6. Counterfactual analysis results | | | | | | | | | |
|--|------------------------|---|------------------------|---|--|--|--|--|--|
| Public forecast without COVID-19 newly detected tuberculosis cases | new tuberculosis cases | Private forecast without COVID-19 newly detected tuberculosis cases | new tuberculosis cases | Counterfactual analysis incidence rate/lakh population- COVID-19 ERA | | | | | |
| 2020 | 1893930 | 2020 | 944268.7 | 203.25 | | | | | |
| 2021 | 2056195 | 2021 | 1215332 | 232.42 | | | | | |
| 2022 | 2218461 | 2022 | 1486396 | 259.79 | | | | | |



This counterfactual analysis suggest that 1027364, 1125849 and 1285810 new tuberculosis cases were missed during 2020, 2021, 2022 respectively as a probable negative impact of COVID-19 in India, see figure-1 and Table-6.

Strength and Limitations

There are several studies done on this research question but this is the first study which also considered year 2022. We know that the first two years of pandemic was quite harsh and it has shattered several routine healthcare services. The COVID-19 is not over and it is quite important to assess the situation of year 2022 in which the cases and mortality from COVID-19 decreased as well as to know that are the healthcare facilities are returning to normal functions? This research study revealed that the notification rates for TB has improved a lot during 2022 which may be a sign to say that healthcare facilities are returning to normal functions. This is the only study which has taken into account 6 years with equal distribution among pre-COVID-19 and COVID-19 period. Lack of more details about data on reported cases, including socio-demographic, **old non-notified cases missed during COVID-19 years**, residence wise data (rural vs. urban, etc.) is one major limitation.

Conclusion and Recommendation

The finding of this research study suggests that about over 3439023 new TB cases were not notified in India that would have been expected in the absence of the COVID-19 pandemic. Where are these cases? Factors which may have affected TB cases notifications may be:

- Decreased mobility and increased COVID-19 related hospital admissions per 100,000 population
- Disruption in TB diagnostic services, the impact of mask use
- Further research is needed to clarify this association
- · Identification of other key contributors to the observed gap in TB case notification in India during the pandemic.

Other information

Abbreviations

Tuberculosis (TB); COVID-19- Coronavirus Disease 2019; HIV- Human Immunodeficiency Virus; NTEP (National Tuberculosis Elimination Program).

Declarations

- This version of paper has not been previously published in any peer reviewed journal and is not currently under consideration by any journal. The document is Microsoft word with English language & 2500 words excluding reference and declaration etc. (5292 words Total including all).
- Ethics approval and consent to participate: Not applicable. This study has not involved any human or animals in real or for experiments. The submitted work does not contain any identifiable patient/participant information.
- Consent for publication: The author provides consent for publication.
- Availability of data and materials: Electronic records from Nikshay, India,
- Conflicts of Interest/ Competing Interest: There are no conflicts / competing of interest
- Funding:
 - The author declares that no funds are taken from any individual or agency-institution for this study.
 - This is the first version of this work and next versions will evolve in future with more information and analysis.
- Authors' contributions: The whole work is done by the Author Dr Piyush Kumar, M.B.B.S., E.M.O.C., P.G.D.P.H.M., -Senior General Medical Officer- Bihar Health Services- Health Department- Government of Bihar, India, Advocate Anupama-Senior Lawyer, Bar Council, Patna and Harshika Singh, Masters Student of Epidemiology of Infectious Disease and Antimicrobial Resistance, University of Glasgow UK,
- Acknowledgements: I am thankful to Advocate Anupama my wife and daughters Aathmika-Atheeva for cooperation.
- Author information: The author is currently working as Senior General Medical Officer for the government of Bihar and Advocate Anupama-Senior Lawyer, Bar Council, Patna.
- Correspondence: Department of Health, Government of Bihar, MOBILE +919955301119/+917677833752, Email drpiyush003@gmail.com

References

- World Health Organization Home/Newsroom/Fact sheets/Detail/Tuberculosis available at -https://www.who.int/news-room/fact-sheets/detail/tuberculosis#:~:text=A%20total%20of%201.6%20million.with%20tuberculosis(TB)%20worldwide.
- 2. World Health Organization Home/News/Detail/States and UTs accelerate action to end TB by 2025 available at https://www.who.int/india/news/detail/09-11-2021-states-and-uts-



accelerate-action-to-end-tb-by-2025

- 3. World Health Organization Home/News/COVID-19 continues to disrupt essential health services in 90% of countries available at https://www.who.int/news/item/23-04-2021-covid-19-continues-to-disrupt-essential-health-services-in-90-of-countries
- 4. World Health Organization Global Tuberculosis reports-1997- 2022- available at -https://www.who.int/teams/global-tuberculosis-programme/tb-reports
- 5. The World Bank Incidence of tuberculosis (per 100,000 people) India- available at https://data.worldbank.org/indicator/SH.TBS.INCD?locations=IN
- Aggarwal, Ashutosh Nath; Agarwal, Ritesh; Dhooria, Sahajal; Prasad, Kuruswamy Thurai; Sehgal, Inderpaul Singh; Muthu, Valliappan. Impact of COVID-19 pandemic on tuberculosis notifications in India. Lung India 39(1):p 89-91, Jan–Feb 2022. | DOI: 10.4103/lungindia_lungindia_604_21
- 7. NATIONAL STRATEGIC PLAN FOR TUBERCULOSIS ELIMINATION 2017–2025 available at -https://tbcindia.gov.in/WriteReadData/NSP%20Draft%2020.02.2017%201.pdf
- 8. WHO COVID-19 Research Database available at https://search.bvsalud.org/global-literature-on-novel-coronavirus-2019-ncov/?lang=en&q=au:%22Dr%20Piyush%20Kumar%22
- 9. Khandelwal, S. (2022). Malnutrition and COVID-19 in India. In: Pachauri, S., Pachauri, A. (eds) Health Dimensions of COVID-19 in India and Beyond. Springer, Singapore. https://doi.org/10.1007/978-981-16-7385-6_9
- 10. Husain, A. A., Monaghan, T. M., & Kashyap, R. S. (2021). Impact of COVID-19 pandemic on tuberculosis care in India. Clinical microbiology and infection: the official publication of the European Society of Clinical Microbiology and Infectious Diseases, 27(2), 293–294. https://doi.org/10.1016/j.cmi.2020.08.014
- 11. COVID-19 promoting the development of active tuberculosis in a patient with latent tuberculosis infection: A case report; Mohammed Khayat, Hanan Fan, Yusuf Vali, https://doi.org/10.1016/i.rmcr.2021.101344
- 12. TB/COVID-19 Global Study Group (2022). Tuberculosis and COVID-19 co-infection: description of the global cohort. The European respiratory journal, 59(3), 2102538. https://doi.org/10.1183/13993003.02538-2021
- 13. Databank files- THE WORLD BANK available at -https://databankfiles.worldbank.org/public/ddpext_download/poverty/33EF03BB-9722-4AE2-ABC7-AA2972D68AFE/Global POVEQ SDN.pdf
- On tuberculosis and COVID-19 co-infection-Marina Tadolini, José-María García-García, François-Xavier Blanc, Sergey Borisov, Delia Goletti, Ilaria Motta, Luigi Ruffo Codecasa, Simon Tiberi, Giovanni Sotgiu, Giovanni Battista Migliori; European Respiratory Journal Aug 2020, 56 (2) 2002328; DOI: 10.1183/13993003.02328-2020
- 15. Zimmermann P, Curtis N Why is COVID-19 less severe in children? A review of the proposed mechanisms underlying the age-related difference in severity of SARS-CoV-2 infections Archives of Disease in Childhood 2021;106:429-439.
- 16. UNICEF Tuberculosis is now the leading cause of death from infectious diseases for children of all ages globally available at https://data.unicef.org/topic/child-health/tuberculosis/
- 17. India TB report-2019- available at -https://tbcindia.gov.in/WriteReadData/India%20TB%20Report%202019.pdf
- 18. Rath, R. S., Dixit, A. M., Koparkar, A. R., Kharya, P., & Joshi, H. S. (2020). COVID-19 pandemic in India: A Comparison of pandemic pattern in Selected States. Nepal journal of epidemiology. 10(2), 856–864. https://doi.org/10.3126/nie.v10i2.28960
- 19. Utomo, B., Chan, C. K., Mertaniasih, N. M., Soedarsono, S., Fauziyah, S., Sucipto, T. H., Aquaresta, F., Eljatin, D. S., & Adnyana, I. M. D. M. (2022). Comparison Epidemiology between Tuberculosis and COVID-19 in East Java Province, Indonesia: An Analysis of Regional Surveillance Data in 2020. Tropical medicine and infectious disease, 7(6), 83. https://doi.org/10.3390/tropicalmed7060083
- 20. Andrews, M. A., Areekal, B., Rajesh, K. R., Krishnan, J., Suryakala, R., Krishnan, B., Muraly, C. P., & Santhosh, P. V. (2020). First confirmed case of COVID-19 infection in India: A case report. The Indian journal of medical research, 151(5), 490–492. https://doi.org/10.4103/ijmr.IJMR 2131 20
- 21. Ni-kshay Reports Available at -https://reports.nikshay.in/Reports/TBNotification
- 22. NITI AAYOG REPORTS available at-https://niti.gov.in/documents/reports
- 23. Press Information Bureau GoI- available at -https://pib.gov.in/PressReleasePage.aspx?

 PRID=1871626#:~:text=India's%20TB%20incidence%20for%20the.the%20qlobal%20average%20of%2011%25