INTRODUCTION

Tuberculosis (TB) is caused by the airborne infectious pathogen, Mycobacterium tuberculosis. It primarily infects the lungs causing pulmonary TB and can invade other organs causing extrapulmonary TB (EPTB). TB is prevalent in all countries and distributed in various age groups. Of note, 10% of healthy individuals who are infected with M. tuberculosis develop TB within two years. In addition, the incidence is higher among people with human immunodeficiency virus (HIV) infections.

One-third of world’s population is infected with M. tuberculosis, which is ranked among top 10 causes...
of deaths worldwide surpassing HIV. It is also denoted as the major cause of deaths from a single infectious agent. In 2019, where about 10 million people developed TB and 1.4 million died. This is equivalent to 130 cases/100,000. The World Health Organization (WHO) indicated that the incidence of TB globally is declining very slowly in recent years and similarly the Saudi Ministry of Health (MOH) stated a slow decline in TB. However, we would like to notify the readers that this declining rate is variable between different Saudi cities, particularly in the capital city of Riyadh and in holy cities (Makkah and Madinah) during Umrah and Hajj seasons. Regarding Saudi Arabia, the number of reported TB cases was 3336 in 2014. Saudi Arabia is a global destination where the influx of pilgrims throughout the year confers highly dynamic nature to the population as well as the large proportion of expatriates in the country’s population.

Despite of the major measures to prevent and control the spread of tuberculosis, the disease continues to be a serious global public health concern, particularly in the developing countries. In this respect, the top six TB burdened countries are India, Indonesia, China, Nigeria, Pakistan, and South Africa where infections in these countries account for 60% of new cases worldwide. India and China account for half of all global TB cases. It was also reported that pulmonary TB is the most prevalent form of the disease in the northern part of Iran. Another Middle East country with a high incidence rate as reported in 2014 is Turkey, with 22% of all reports out of 58252 recorded cases in the Middle East. Also, the global TB report that was published in 2016 ranked the gulf countries based on the incidence rate of TB, where Kuwait came first with an incidence of 200 cases/million, followed by Saudi Arabia (89/million), and then United Arab Emirates with the least incidence rate (6.8/million). Coming to the developed countries, the rates in the United States has declined remarkably to three cases/100,000, which could be attributed to the advanced healthcare system in the country in comparison to the developing countries.

Regarding mortality, it increases with the presence of comorbidities. For example, HIV and TB co-infection is an important cause of mortality in HIV patients. In addition, diabetes mellitus (DM) is associated with an increased mortality rates following TB infections. In this regard, DM is an established comorbidity that accelerates TB complications and derail its treatment. Likewise, smoking increases the risk of TB as well as its related mortality rates. To sum up, the mortality predictors include older age, low body weight, rural living, discontinued treatment, EPTB, immunocompromisation, comorbidities, and coinfections, especially HIV infections.

The objective of the current retrospective study was to analyze the reported cases of pulmonary TB to comprehend the epidemiology and ascertain its association with the risk factors that may aid the National Tuberculosis Program (NTP) of the Saudi Arabia to spot high risk groups and figure out the success of infection control measures to revise their policies and measures to combat the disease.

METHODS

This is a retrospective study that was conducted to analyze official pulmonary TB figures in Saudi Arabia cities that were reported by the MOH for the year 2018 (https://data.gov.sa/Data/en/organization/ministry_of_health). The official data figures of pulmonary TB cases were obtained and comprised the figures of 13 provinces and seven districts for Saudi and non-Saudi nationals as well as incidence amongst males and females. Cases were categorized into seven age groups, which were as follows: < 15, 15–25, 25–35, 35–45, 45–55, 55-65, and ≥ 65 years old. Also, cases were stratified according to gender, nationality, and region to compare the incidence rates between different strata of the study sample. Of note, the status of non-Saudi cases in the MOH reports was not indicated as being visitors or residents.

Statistical Analysis: The data was tabulated in Microsoft Office Excel and the descriptive statistics such as mean; sum and percent distribution were calculated. The comparison between the incidence rates in different age groups and gender for Saudi and non-Saudi nationals were computed by Chi Square test using IBM SPSS (Version 23; IBM Corp., Armonk, N.Y., USA). P-value of ≤ 0.05 was considered statistically significant.

Ethical Approval: This is a secondary data analysis-based study, which does not require to be ethically reviewed and approved as all data are publicly available through the Ministry of Health portal without any reference to personal identifying information of patients, for this reason, IRB approval is not applicable.

RESULTS

The rate of incidence: The number of cases of pulmonary TB reported by the MOH in Saudi Arabia cities in 2018 is presented in Table-I. The
results indicated that the highest number of cases amongst Saudi nationals was recorded in the capital city of Riyadh where most subjects being males (79.2%), which was also the case for Jeddah where males represented most cases (81.2%). Overall, the occurrence of infections in Saudi males was higher than that in females, except in Jazan, wherein the percentage of infections in females was 58.2%.

Regarding the incidence of infection in non-Saudi nationals, the highest incidence rate was recorded in Jeddah, followed by Riyadh. Also, the incidence rate was significantly higher (p ≤0.001) in males than in females representing 73.6% and 71.3%, respectively. The incidence of infections in males amongst the non-Saudi nationals was higher than that in females in all cities without exceptions. The map in Fig.1 depicts the incidence of pulmonary TB in different cities; the symbols code indicates the magnitude of incidences and classifies them into four different rates.

**Assessment of Risk Factors:** The distribution of infections in different age groups in males and

| Region          | Saudi Male | Saudi Female | Saudi Total | Non-Saudi Male | Non-Saudi Female | Non-Saudi Total | Total Cases |
|-----------------|------------|--------------|-------------|----------------|-----------------|-----------------|-------------|
| Riyadh          | 255        | 67           | 322         | 241            | 97              | 338             | 660         |
| Makkah          | 60         | 27           | 87          | 66             | 41              | 107             | 194         |
| Jeddah          | 243        | 56           | 299         | 332            | 119             | 451             | 750         |
| Taif            | 14         | 06           | 20          | 16             | 05              | 21              | 41          |
| Madinah         | 42         | 19           | 61          | 43             | 17              | 60              | 121         |
| Qaseem          | 13         | 06           | 19          | 15             | 05              | 20              | 39          |
| Eastern province| 56         | 20           | 76          | 146            | 28              | 174             | 250         |
| Al-Ahsa         | 06         | 02           | 08          | 16             | 05              | 21              | 29          |
| Hafr Al-Baten   | 10         | 02           | 12          | 10             | 04              | 14              | 26          |
| Aseer           | 15         | 14           | 29          | 34             | 03              | 37              | 66          |
| Bishah          | 08         | 05           | 13          | 05             | 05              | 10              | 23          |
| Tabouk          | 03         | 03           | 06          | 15             | 03              | 18              | 24          |
| Hail            | 06         | 02           | 08          | 06             | 03              | 09              | 17          |
| Northern        | 03         | 01           | 04          | 06             | 00              | 06              | 10          |
| Jazan           | 38         | 53           | 91          | 87             | 18              | 105             | 196         |
| Najran          | 08         | 05           | 13          | 17             | 07              | 24              | 37          |
| Al-Bahah        | 09         | 00           | 09          | 10             | 03              | 13              | 22          |
| Al-Jouf         | 02         | 01           | 03          | 03             | 00              | 03              | 06          |
| Qurayyat        | 04         | 01           | 05          | 10             | 00              | 10              | 15          |
| Qunfudah        | 05         | 01           | 06          | 10             | 01              | 11              | 17          |
| Total           | 545        | 291          | 836         | 1,088          | 364             | 1,452           | 2,543       |

* Number of cases per 100,000.
females, Saudi and non-Saudi nationals. Table-II

Amongst male Saudis, the highest percentages of infections were reported in the age range <15, 15-25 and 25-35 in order. In Saudi females, the highest percentage of infections were in age groups 45-55, followed by 35-45. In this respect, there was a statistically significant difference between the number of infections in males and females in the above-mentioned age groups ($p=0.001$).

The highest number of infections amongst male non-Saudi nationals was recorded in the age groups 25-35 and 35-45, followed by 45-55. On the other hand, the highest female infections were in ≥65 then 55-65 age groups. Likewise, the difference was statistically significant between the number of infections in male and female non-Saudi nationals in these age groups ($p=0.001$).

### DISCUSSION

In the current study, the overall rate of incidence of pulmonary TB found to be 7.61/100,000 in Saudi Arabia for the year 2018. Globally, the TB incidence rate is falling by roughly 2% a year.7,18 In 2019, the incidence of TB in Saudi Arabia was 9.9 cases/100,000. The incidence of tuberculosis in Saudi Arabia fell gradually from 19 cases/100,000 in 2000, to 9.9 cases/100,000 in 2019.6 The difference in the incidence of TB between genders could be attributed to sex hormones that may be responsible for modulating the immune response, which is necessary for fighting the pathogen and other infectious diseases.19,20 Recently, the mortality rate of TB was reported to be 22/1000 person-years being higher in males and elderly. Also, it has been suggested that the reactivation of latent MTB infection in elderly age could be due to immunosenescence and co-morbidities.21,22

In the current study, the number of cases of TB in males amongst the non-Saudi nationals was found to be higher than that in Saudi nationals; most likely because most of them came from countries with a high burden of TB.6 The highest number of infections amongst male non-Saudi nationals was recorded in the age groups 25-35 and 35-45, followed by 45-55; this might be explained by the socioeconomic status, where in these age-groups there are a lot of unskilled workers who tend to live in crowded shelters, with poor sanitation and hygiene.23 These conditions make them susceptible for reactivation of latent TB. Expatriates arriving to the kingdom for work are not screened for TB, except for house workers and health professionals.24 The highest incidence rate of TB amongst non-Saudis, which was recoded in Jeddah, followed by Riyadh, may be associated with the higher proportion of non-Saudis in urban areas and industrial cities. Jeddah receives more than ten million foreign visitors every year for Hajj and Umrah.25

Another Saudi study reported low prevalence of TB and multidrug resistant TB over 17-year period in a single center.26 Having indicated that Saudi Arabia is a global destination and with continuous influx and outflux of people in and out from the country, there might be a chance of Saudi Arabia to become a spread point of infection in case TB incidence rises in a steady state. Therefore, we have to notify the readers that gulf health council (GHC) members that include Saudi Arabia have the highest prevalence of diabetes in the world, where the International Diabetic Federation (IDF) ranked five members of the GHC amongst the top 10 countries with high prevalence of diabetes. In this context, Saudi Arabia ranked third in the world by the IDF. Since diabetes has been found to increase TB risk, the high prevalence of diabetes in Saudi Arabia should be a warning sign.9
CONCLUSION

There was a significant difference between males and females regarding contracting the TB infection amongst both the Saudi and the non-Saudi nationals suggesting an association between gender and the increased risk of infection. Likewise, analysis of infection in different age groups suggested that the incidence rates in Saudis were recorded in younger age groups than in the non-Saudis, which is surprising. Importantly, we do not know whether non-Saudis are residents or visitors as it was not mentioned in the MOH database. In this respect, there should be a proper screening program for high-risk groups as well as a plan to investigate the risk factors that might be associated with the increased incidence of pulmonary TB. Altogether, these analyses and approaches would help health authority decision makers to establish policies and procedures to limit the infection and apply appropriate precautionary measures. To this end, this study analyzed the data of pulmonary TB incidence rates in 2018 where the data for other years were not reported on the MOH data repository. It would be more informative if we could obtain data for a longer period for the study to be more conclusive.

Conflict of Interest: None.

Funding: None.

Availability of data and materials: The data used to support the findings of this study are available freely on public domain of The Ministry of Health, Saudi Arabia. Open Data Portal. https://data.gov.sa/Data/en/organization/ministry_of_health

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