

Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.



Available online at www.sciencedirect.com

ScienceDirect

journal homepage: http://www.journals.elsevier.com/ indian-journal-of-tuberculosis/

Letter to the Editor

Indeterminate mycobacterium tuberculosis QuantiFERON post Moderna mRNA Covid-19 vaccination

Keywords:
Mycobacterium tuberculosis
QuantiFERON
Moderna
mRNA Covid-19 vaccination

ABSTRACT

We report an interesting case of an indeterminate MTB QuantiFERON for a 26-year-old healthy soldier planned for a routine field exercise to Brunei. Further medical history revealed that the patient had a Moderna mRNA Covid-19 vaccine the day before his MTB QuantiFERON test. The patient was subsequently asked to repeat a T-spot test which was non-reactive, there were no longer any issues with the positive control for the T-spot test.

Current Covid-19 research suggests that infection causes a dysregulation of the immune system, perhaps this might also be extrapolated where a Covid-19 vaccine might provoke an immune response which might interfere with some immunological assays. In summary there should be more research invested into the immunological interactions that the newly developed Covid-19 vaccinations have with our existing immunological tests such as QuantiFERON tests which forms a key cornerstone in our fight against tuberculosis.

© 2022 Tuberculosis Association of India. Published by Elsevier B.V. All rights reserved.

Dear Editor

We read with interest the recently published article positive QuantiFERON test and the severity of COVID-19 disease: A prospective study. It was interesting to note that a negative mycobacterium tuberculosis (MTB) QuantiFERON test was a strong predictor of mortality in severe Covid-19 diseases.

In response, the authors would like to report an interesting case of an indeterminate MTB QuantiFERON for a 26-year-old healthy soldier planned for a routine field exercise to Brunei. As part of the pre-departure health screening, soldiers in Singapore are supposed to be screened for MTB, of which the preferred method is the MTB QuantiFERON assay. This patient was referred by the military medical officer due to the indeterminate (MTB) QuantiFERON result. The reason for the indeterminate result was due to the positive control being not interpretable as the mitogen minus tube did not have a sufficient amount of activation. The patient was asked multiple screening questions regarding recent infective symptoms, recent illness/Covid-19 infection, chronic medication or complementary medication usage. A thorough history of sexually transmitted infections, risky sexual behavior and risky sexual behavior was also performed. All of the above questions were negative. Further questioning about recent vaccinations did reveal that the patient had a Moderna mRNA Covid-19 vaccine the day before MTB QuantiFERON test. The patient was subsequently asked to repeat a T-spot test which was non-reactive, there were no longer any issues with the positive control for the T-spot test.

霐

TUBERCULOSIS

This is an interesting and likely first reported case of an indeterminate MTB QuantiFERON result post mRNA vaccination in an otherwise healthy patient. Currently the Centre for Disease Control in the United States of America stipulates that a QuantiFERON test be delayed 4–6 weeks if a patient has been vaccinated with a live attenuated virus.¹ As current evidence shows that vaccination with live viruses (such as the MMR vaccine) can cause mild immune system suppression. This may reduce the reactivity of the tuberculin skin test and possibly causing a false–negative reaction.² While inactive vaccines do not interfere with TB test results. Currently there is little data regarding mRNA vaccines in the setting of a QuantiFERON test. There have been studies

regarding QuantiFERON in the setting of severe Covid-19 infection. Current evidence suggests that there is an increased rate of indeterminate QuantiFERON results in critically ill COVID-19 patients with most showing huge reduction in mitogen stimulus thus suggesting gross general unresponsiveness of T cells.^{3–5} It seems that Covid-19 infection causes a dysregulation of the immune system, perhaps this might also be extrapolated where a Covid-19 vaccine might provoke an immune response which might interfere with some immunological assays.

In summary there should be more research invested into the immunological interactions that the newly developed Covid-19 vaccinations have with our existing immunological tests such as the tuberculin skin tests, QuantiFERON tests and T-spot tests which forms a key cornerstone in our fight against tuberculosis.

Author contributions

All authors contributed to (1) concept or design, (2) acquisition of data, (3) analysis or interpretation of data, (4) drafting of the manuscript, and (5) critical revision for important intellectual content.

All authors had full access to the data, contributed to the study, approved the final version for publication, and take responsibility for its accuracy and integrity.

Conflicts of interest

The authors have none to declare.

REFERENCES

- Lewinsohn DM, Leonard MK, LoBue PA, et al. Official American thoracic society/infectious diseases society of America/centers for disease control and prevention clinical practice guidelines: diagnosis of tuberculosis in adults and children. Clin Infect Dis. 2017;64(2):111–115.
- Kroger A, Bahta L, Hunter P. General best practice guidelines for immunization. In: Best Practices Guidance of the Advisory Committee on Immunization Practices (ACIP). 2021.
- **3.** Gupta A, Sural S, Gupta A, et al. Positive QuantiFERON test and the severity of COVID-19 disease: a prospective study. *Indian J Tubercul.* 2021;68(4):474–480.
- Shier KL, Tang Y-W. Elevated rates of indeterminate results on QuantiFERON-TB gold plus in COVID-19 patients. J Clin Microbiol. 2021;59(10):e01414-e01421.
- Ward JD, Cornaby C, Schmitz JL, Tang Y-W. Indeterminate QuantiFERON gold plus results reveal deficient interferon gamma responses in severely ill COVID-19 patients. J Clin Microbiol. 2021;59(10):e00811–e00821.

Samuel S.Y. Wang Tuberculosis Control Unit, Tan Tock Seng Hospital, Singapore, Singapore E-mail address: samuel.wang@mohh.com.sg

> 1 February 2022 Available online 31 March 2022

0019-5707/\$ — see front matter © 2022 Tuberculosis Association of India. Published by Elsevier B.V. All rights reserved. https://doi.org/10.1016/j.ijtb.2022.03.019