RESEARCH ARTICLE

How to improve TB outpatient service in a TB low-endemic country during SARS-CoV-2 pandemic

NICCOLÒ RICCARDI^{1,2}, PAOLA RODARI², SILVIA DURANTI^{1,2}, SIMONE VILLA^{1,3}, LUCA ROSARIO ASSANTE⁴, PAOLA CASTELLOTTI^{1,5}, MAURIZIO FERRARESE^{1,5}, MARIANO MARTINI^{6,1}, TERESA ZUPPINI⁷, ROBERTO TESSARI⁷, CRISTINA CALZA⁷, LUIGI CODECASA^{1,5}, GIORGIO BESOZZI¹, ANDREA ANGHEBEN²

 ¹ StopTB Italia, Milan, Italy; ² Department of Infectious, Tropical Diseases & Microbiology, IRCCS Sacro Cuore Don Calabria Hospital, Negrar di Valpolicella, Verona, Italy; ³ Centre for Multidisciplinary Research in Health Science, University of Milan, Italy;
⁴ Division of Pulmonology and Thoracic Endoscopy, IRCCS Sacro Cuore-Don Calabria Hospital, Negrar-Verona, Italy; ⁵ Regional TB Reference Centre, Istituto Villa Marelli, Niguarda Hospital, Milan, Italy; ⁶ Department of Health Sciences, University of Genoa, Italy; ⁷ Service of Hospital Pharmacy, IRCCS Sacro Cuore Don Calabria Hospital, Negrar di Valpolicella, Verona, Italy

Keywords

Tuberculosis • Outpatient care • Italy • TB-care • Medical ethics

Summary

Despite notable progresses in the recent decades, tuberculosis (TB) continues to remain a public health concern even in Europe. TB prevention and care should be people-centred, improving ambulatory models across countries, in order to expand access to diagnosis and treatment for both TB infection and disease. Even more, during emergencies such as the cur-

Despite notable progress in the recent decades [1, 2], tuberculosis (TB) remains a public health concern in the World Health Organization (WHO) European Region, which includes Italy, currently being the leading cause of death from a single microorganism [3, 5].

The emergence of drug-resistant (DR)-TB in TB lowendemic countries is a consequence of failings in the health care system that does not provide easy access to TB care to those in need, fuelling the TB epidemic and slowing the progress to TB elimination [6, 7].

A milestone of TB prevention and care is a peoplecentred approach, which implies overall improving ambulatory models across countries, in order to expand access to diagnosis and treatment of both TB infection and disease [6]. In fact, hospitalization of TB patients is needed only for case of severe localizations (e.g. TB meningitis, TB pericarditis) and/or when complications arise because of the treatment (e.g. severe allergies) and/ or if the patient has comorbidities that require close monitoring during anti-TB and TB preventive treatment (TPT) [8].

Otherwise, independently from the spectrum of resistance of the *Mycobacterium tuberculosis* (Mtb) strain, all TB continuum of care, from screening and diagnosis to treatment and post-treatment follow-up, should be carried on in specialized outpatient services [8]. Even more during pandemic times, where seeking of TB care has been replaced by the fear of coronavirus disease 2019 (COVID-19) and outpatients services have been hampered by the health emergency status, TB patients care is fundamental [9, 10].

rent pandemic, when seeking of TB care has been replaced by the fear of coronavirus disease 2019 (COVID-19), TB patient's care is fundamental. In this short communication, we document how was possible to implement a TB outpatient service meanwhile a local outbreak of SARS-CoV-2 transmission was ongoing.

.....

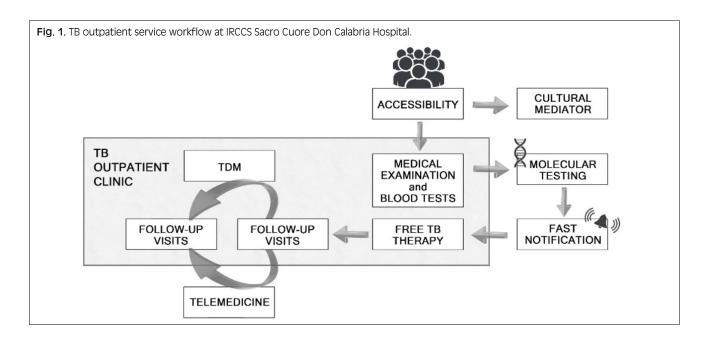
Aim of this paper is to describe how a TB outpatient service was implemented in a referral centre for Infectious and Tropical diseases in Northern Italy when the COVID-19 pandemic was raging in the area.

Following the example of the Regional TB Reference Centre of Lombardy Region, Villa Marelli Institute/ ASST Niguarda Ca' Granda (Milan, Italy), which works as an outpatient reference centre for drug-susceptible and DR-TB, serving a population of > 10 million people and managing ~250 patients per year (3% with DR-TB) [7], we created, at the IRCCS Sacro Cuore Don Calabria Hospital (Negrar di Valpolicella, Italy), an integrated outpatient TB service with a multidisciplinary team composed by infectious disease specialists, pulmonologists, pharmacists, microbiologists, radiologists, gastroenterologists, and geriatrics.

Moreover, *on demand* cultural mediators allow to further improve the service, guaranteeing the patient's full understanding of the disease, treatment, scheduled follow-ups visits as well our full understanding of the patient's complaints. In case of other organ involvement (e.g. genitourinary tract) other specialists were called on demand for consultations.

The outpatient TB service was based on the following major pillars against the most common barriers:

- 1. easy and direct access for patients (reducing barriers to access);
- 2. rapid molecular diagnosis and drug-resistance profiling (reducing risk of suboptimal treatment or resistance induction and optimizing treatment duration);



- 3. instant notification of molecular and microbiological tests' results to the treating physicians (timely adjustment of treatment);
- 4. tailored treatment regimens mindful of patients' comorbidities and at-risk behaviours (like unhealthy diet, comorbidities, drug-drug interactions etc.);
- 5. direct distribution of anti-TB drugs to the patient (delivered to patients at time of visit, in order to increase adherence to treatment);
- consultation vie e-mails with patients between follow-ups (essential during COVID-19 lock-down period);
- 7. scheduled follow-up visits with the possibility of convenient specialists' consultation (to give comprehensive care to patients);
- 8. therapeutic drug-monitoring of anti-TB drugs (increasing adjustment of treatment and tolerance) [11];
- 9. track patients' adherence using electronic tools to reduce dropouts (increasing compliance);
- 10. infectious (e.g. hepatitis B virus, hepatitis C virus, HIV, aspergillosis) and tropical (e.g. schistosomiasis, strongyloidiasis) diseases screening for patients from endemic areas;
- 11. discussion of difficult-to-treat cases with experts from the Villa Marelli Institute (Fig. 1).

Applying the model to the current context of the COVID-19 ongoing pandemic, we established to screen all patients with rapid antigen test for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) before entering the hospital and, if the results are either positive or pending, adequate personal protective equipment is worn by both patients and health care workers.

Since the beginning of the outpatient service in July 2020, 30 tuberculosis cases have been diagnosed, 28 (93,3%) and 2 (6,7%), DS and DR-TB, respectively.

Twenty-three were diagnosed pulmonary TB and 7 extra-pulmonary TB. Overall, 6 patients were born in

Italy and 24 were foreigners. Just one patient (3.3%) was lost to follow-up.

In conclusion, starting from a people-centred approach, our outpatient service for TB patients care was organized to be nearer and more accessible to the people who need it the most [12].

Overall, it points to a model easily replicable in other low-endemic-high income countries in order to increase patient's adherence to treatment and to answer to patients' necessities [5, 13].

Furthermore, hospitalization can be restricted only to those with severe disease, with reduction of isolation stay and related costs; leaving hospital-bed free for other infective-pathologies such as the current numerous cases of COVID-19 pneumonias [5].

Finally, our contribution represents a step forward in building up extended regional and national networks for TB outpatients. Ensuring quality care, sharing information on diagnosis, management and outcomes, and addressing social and cultural aspects by employing professional mediators may help to increase treatment success, reduce drop out and, ultimately, contribute to the WHO goal of ending global TB epidemic by the year 2035 [14].

Acknowledgements

Funding sources: this research did not receive any specific grant from funding agencies in the public, commercial, or not-for-profit sectors.

The authors wish to thank Dr. Luca De Santis for Graphic Support.

.....

Conflict of interest statement

The authors declare no conflict of interest.

Authors' contributions

NR, AA and GB conceived the study; NR, PR, SD, SV, MM drafted the manuscript; NR, PR, LRA, PC, MF, MM, TZ, RT, CC, LC revised the manuscript; NR, PR, SD, SV, PC, MF, MM performed a search of the literature; NR, PR, SD, MM, LC, GB and AA revised critically the manuscript. All authors read and approved the last version of the manuscript.

References

- [1] Villa S, Riccardi N, Canetti D, Alagna R, Castellotti P, Ferrarese M, Cirillo D, Barberis I, Bragazzi NL, Gazzaniga V, Ricucci V, Codecasa LR, Besozzi G, Martini M. From the past, a long way to future challenges for a greater control of tuberculosis. Tuberculosis 2020;123:101948. https://doi.org/10.1016/j. tube.2020.101948
- [2] Martini M, Besozzi G, Barberis I. The never-ending story of the fight against tuberculosis: from Koch's bacillus to global control programs. J Prev Med Hyg 2018;59:E241-E247. https:// doi.org/10.15167/2421-4248/jpmh2018.59.3.1051
- [3] Martini M, Riccardi N, Giacomelli A, Gazzaniga V, Besozzi G. Tuberculosis: an ancient disease that remains a medical, social, economical and ethical issue J Prev Med Hyg 2020;61:E16-E18.
- [4] Global tuberculosis report 2019. Geneva: World Health Organization 2019.
- [5] A people-centred model of tuberculosis care: a blueprint for eastern European and central Asian countries, 1st edition. Copenhagen: WHO Regional Office for Europe 2017.

- [6] https://www.euro.who.int/__data/assets/pdf_file/0004/388039/ ambulatory-care-eng.pdf?ua=1
- [7] Riccardi N, Pontarelli A, Alagna R, Saderi L, Ferrarese M, Castellotti P, Viggiani P, Cirillo D, Besozzi G, Sotgiu G, Codecasa L; StopTB Italia Onlus Group. Epidemiology and treatment outcome of MDR and pre-XDR TB in international migrants at two reference centers in the North of Italy: a cross-sectional study coordinated by Stop TB Italia Onlus. Public Health 2020;180:17-21. https://doi.org/10.1016/j.puhe.2019.10.022
- [8] Guiding principles to reduce tuberculosis transmission in the WHO European Region. Copenhagen: WHO Regional Office for Europe 2018.
- [9] Mantica G, Riccardi N, Terrone C, Gratarola A. Non-COV-ID-19 visits to emergency departments during the pandemic: the impact of fear. Public Health 2020;183:40-1. https://doi. org/10.1016/j.puhe.2020.04.046
- [10] Migliori GB, Thong PM, Akkerman O, et al. Worldwide Effects of Coronavirus Disease Pandemic on Tuberculosis Services, January-April 2020 [published online ahead of print, 2020 Sep 11]. Emerg Infect Dis. 2020;26(11):10.3201/eid2611.203163. https://doi.org/10.3201/eid2611.203163
- [11] Riccardi N, Villa S, Alagna R, Giacomelli A, Saderi L, Cirillo DM, Besozzi G, Sotgiu G, Codecasa L. Advantages and challenges of tailored regimens for drug-resistant tuberculosis: a StopTB Italia look into the future. Infect Drug Resist 2020;13:2795-800. https://doi.org/10.2147/IDR.S257480
- [12] Morgan S, Yoder LH. A concept analysis of person-centered care. J Holist Nurs 2012;30:6-15.
- [13] Strengthening integrated, people-centred health services. Geneva: World Health Organization 2016.
- [14] Martini M, Barberis I, Gazzaniga V, Gazzaniga V, Icardi G. The fight to end tuberculosis: a global challenge in strong partnership. J Prev Med Hyg 2020;61:E1-E2.

Received on January 27, 2021. Accepted on May 17, 2021.

......

Correspondence: Mariano Martini, Department of Health Sciences, University of Genoa, largo R. Benzi 10 Pad. 3, 16132 Genoa, Italy - Tel./ Fax: 010.353.85.02 - E-mail: mariano.yy@gmail.com / mr.martini@unige.it

How to cite this article: Riccardi N, Rodari P, Duranti S, Villa S, Assante LR, Castellotti P, Ferrarese M, Martini M, Zuppini T, Tessari R, Calza C, Codecasa L, Besozzi G, Angheben A. How to improve TB outpatient service in a TB low-endemic country during SARS-CoV-2 pandemic. J Prev Med Hyg 2021;62(Suppl. 1):E50-E52. https://doi.org/10.15167/2421-4248/jpmh2021.62.1S3.2008

© Copyright by Pacini Editore Srl, Pisa, Italy

This is an open access article distributed in accordance with the CC-BY-NC-ND (Creative Commons Attribution-NonCommercial-NoDerivatives 4.0 International) license. The article can be used by giving appropriate credit and mentioning the license, but only for non-commercial purposes and only in the original version. For further information: https://creativecommons.org/licenses/by-nc-nd/4.0/deed.en