

## Introduction

Giovanni Battista Migliori<sup>1</sup>, Graham Bothamley<sup>2,3,4</sup>, Raquel Duarte<sup>5,6,7</sup> and Adrian Rendon<sup>8</sup>

@ERSpublications

Physicians feel TB is either very easy to treat with standard regimens or is too complex. This book: provides trainees with basic TB management knowledge; offers insight into addressing complexities in individual patients; is a useful resource for experts. http://ow.ly/zroQ30mipoT

On 26 September 2018, the United National General Assembly agreed to take concerted action on TB. With over 10 million new TB cases (90% in adults and 9% in HIV co-infected individuals) and 1.6 million deaths (300000 in HIV coinfected persons) in 2017, TB is a global health priority [1]. Of particular concern for both clinicians and national TB programmes is MDR-TB: in 2017, WHO were notified of 558000 new rifampicin-resistant cases and 460000 confirmed MDR-TB cases [1]. Heads of state and government agreed to mobilise US\$13 billion a year by 2022 in order to ensure that TB care is received by 40 million people and preventive treatment is given to 30 million people [2].

Although the "white plague" has historically been studied in an extensive manner (it was "the" respiratory disease in the pre-antibiotic era), there is still much to learn about its prevention, diagnosis and treatment. TB is therefore a hot topic in respiratory medicine, attracting an increasing amount of interest from clinicians, scientists, public health officers and the pharmaceutical industry, given that new drugs are finally available after more than 40 years of neglect. The High-Level Meeting on the Fight to End Tuberculosis also agreed to fund a US\$2-billion research agenda [2].

With this *Monograph*, our aim is to provide an accessible resource that will help the young physician in training to recognise and treat TB in all its manifestations, as well as address a need for help with other mycobacterial diseases which might become apparent during the diagnostic process. Primarily providing clinical support, this book will also act as a reference resource for difficult TB. It will introduce topics of interest and scientific advances in TB that can be investigated by the interested reader at their leisure. Many of the chapters also indicate where TB management is going.

Copyright ©ERS 2018. Print ISBN: 978-1-84984-099-6. Online ISBN: 978-1-84984-100-9. Print ISSN: 2312-508X. Online ISSN: 2312-5098.

<sup>&</sup>lt;sup>1</sup>Istituti Clinici Scientifici Maugeri IRCCS, Tradate, Italy. <sup>2</sup>Homerton University Hospital, London, UK. <sup>3</sup>Blizard Institute, Barts and The London School of Medicine and Dentistry, Queen Mary University of London, London, UK. <sup>4</sup>London School of Hygiene and Tropical Medicine, London, UK. <sup>5</sup>Departamento de Pneumologia, Centro Hospitalar Vila Nova de Gaia, Vila Nova de Gaia, Portugal. <sup>6</sup>Departamento de Ciências da Saúde Pública e Forenses e Educação Médica, Faculdade de Medicina, Universidade do Porto, Porto, Portugal. <sup>7</sup>ISPUP-EPIUnit, Faculdade de Medicina, Universidade do Porto, Porto, Porto, Porto, Barta de Medicina, University Hospital of Monterrey, Monterrey, Mexico.

Correspondence: Giovanni Battista Migliori, Istituti Clinici Scientifici Maugeri IRCCS, Via Roncaccio 16, Tradate, Varese, 21049, Italy. E-mail: giovannibattista.migliori@icsmaugeri.it

This *Monograph* will discuss the main issues related to TB, with an innovative approach, beginning with a patient's perspective [3]. The role of patients is very important, given the burden of the disease on healthcare systems globally.

A chapter on the history of TB discusses recent advances in human and TB genetics and presents historical vignettes that are relevant to the current introduction of new treatments [4]. Social determinants are included in the chapter on epidemiology, so that the measures to control and eventually eliminate TB are more holistic [5]. Molecular biology has made significant advances since the last ERS Monograph on TB was published in 2012 [6], and clinically relevant material has been included in a number of chapters [4, 7-12]. The diagnostic aspects (clinical diagnosis, laboratory diagnosis, imaging, bronchoscopy and other invasive procedures) are reiterated and updated so that they are accessible to the physician in training [5, 9, 13, 14]. The treatment of drug-susceptible and drug-resistant cases, new and repurposed drugs, adverse events and the role of surgery are discussed, together with broad principles, so that physicians can apply these to the likely rapid changes in this area [15–18]. Specific patient groups (children, pregnant women and the elderly) are addressed [19]. Comorbidities have become an increasing problem in the management of TB, and diabetes, chronic renal impairment, liver disease and transplantation are addressed, in addition to coinfection with HIV [20]. The modern TB physician has to work with a team to manage homelessness, alcohol and opiate addictions, poverty and malnutrition and the disruptions caused by migration and fleeing war zones and persecutions [21].

A later chapter includes both treatment and therapeutic drug monitoring, noting that the latter will become increasingly important in personalised treatments regimens [22]. Rehabilitation after TB has become an important topic and receives its own chapter [23]. NTM have been included, as they are frequently diagnosed when TB is considered their increasing importance may merit an entire *Monograph* in the not-too-distant future [24]! Preventive issues have come to the fore, especially with the End TB Strategy and there are chapters on vaccines [10], infection control [25] and latent TB infection management [11]. Looking to the future, there is a chapter on research priorities [12], and one addressing the needs of the physician training [26].

Lastly, to emphasise the realities of managing TB, there are some clinical cases drawn from the experience of early career members with expertise in managing MDR-TB [27].

The developments and challenges over the last 6 years, since the publication of the first TB *Monograph* [6], have exceeded our expectations. We expect the recent pledges of world leaders to defeat TB will be met by a mixture of attention to patients' needs and scientific advances, in addition to those we have outlined in this *Monograph*. We hope the *Monograph* will encourage TB physicians and basic scientists to see the gaps and fill these with their own excellent research for the next TB *Monograph*.

## References

- 1. World Health Organization. Global Tuberculosis report 2018. WHO/CDS/TB/2018.20. Geneva, World Health Organization, 2018. http://apps.who.int/iris/bitstream/handle/10665/274453/9789241565646-eng.pdf?ua=1
- UNGA High Level Meeting on Tuberculosis, 26 September 2018. https://www.un.org/pga/73/event-detail/fight-toend-tuberculosis/.

- 3. Spita G, Clegg H, Dumitru M, *et al.* The patients' perspective. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 1–7.
- 4. Loddenkemper R, Murray JF, Gradmann C, et al. History of tuberculosis. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 8–27.
- Duarte R, Santos JV, Santos Silva A, *et al.* Epidemiology and socioeconomic determinants. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 28–35.
- 6. Lange C, Migliori GB, eds. Tuberculosis. Sheffield, European Respiratory Society, 2012.
- Barreira-Silva P, Torrado E, Nebenzahl-Guimaraes, et al. Aetiopathogenesis, immunology and microbiology. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 62–82.
- 8. Zellweger J-P, Sousa P, Heyckendorf J. Clinical diagnosis. *In*: Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 83–98.
- 9. Tagliani E, Nikolayevskyy V, Tortoli E, *et al.* Laboratory diagnosis. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 99–115.
- Ruhwald M, Andersen PL, Schrager L. Towards a new vaccine. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 343–363.
- Rendon A, Goletti D, Matteelli A. Diagnosis and treatment of latent tuberculosis infection. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 381–398.
- 12. Bothamley G. What next? Basic research, new treatments and a patient-centred approach. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 414–429.
- 13. Chesov D, Botnaru V. Imaging for diagnosis and management. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 116–136.
- 14. Bhowmik A, Herth FJF. Bronchoscopy and other invasive procedures for diagnosis. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 137–151.
- Caminero JA, Scardigli A, van der Werf T, *et al.* Treatment of drug-susceptible and drug-resistant tuberculosis. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 152–178.
- 16. Krutikov M, Bruchfeld J, Migliori GB, *et al.* New and repurposed drugs. *In*: Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 179–204.
- Caminero JA, Lasserra P, Piubello A, et al. Adverse anti-tuberculosis drug events and their management. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 205–227.
- Olland A, Falcoz P-E, Guinard S, *et al.* Surgery as a treatment. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 228–233.
- 19. Repossi A, Bothamley G. Pregnancy and the elderly. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 263–275.
- 20. Magis-Escurra C, Carvalho ACC, Kritski AL, *et al.* Comorbidities. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 276–290.
- Viney K, Wingfield T, Kuksa L, et al. Access and adherence to prevention and care for hard-to-reach groups. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 291–307.
- 22. Alffenaar J-WC, Akkerman OW, Bothamley G, *et al.* Monitoring during and after treatment. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 308–325.
- 23. Muñoz-Torrico M, Cid S, Galicia-Amor S, *et al.* Sequelae assessment and rehabilitation. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 326–342.
- 24. Zweijpfenning S, Hoefsloot W, van Ingen J, *et al.* Nontuberculous mycobacteria. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 399–413.
- 25. Nardell E, Volchenkov G. Transmission control: a refocused approach. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 364–380.
- 26. Casalini C, Matteelli A, Komba A, *et al.* Opportunities for training and learning. *In:* Migliori GB, Bothamley G, Duarte R, *et al.*, eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 430–445.
- 27. Tiberi S, Payen MC, Manika K, et al. Clinical cases. In: Migliori GB, Bothamley G, Duarte R, et al., eds. Tuberculosis (ERS Monograph). Sheffield, European Respiratory Society, 2018; pp. 446–460.

Disclosures: None declared.