

Update on Discontinuing TB Isolation

Appendix B, Provincial TB Manual

Updates reflect Canadian TB Standards, 8th ed, Appendix B recommendations

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Key message

Prolonged airborne isolation is associated with patient harms and increased costs and is often not necessary to lower risk of spread.

Potential Harms of Prolonged Isolation

- TB Isolation is associated with anxiety, fear, and mood disorders
- Patients under isolation often have less visitors than other patients
- Recent qualitative studies in patients with MDR-TB and their family indicate a strong preference for early de-isolation
- Providers may feel obligated to “intensify therapy” to facilitate smear conversion (even if this may not improve cure rates)
- Providers may perform more frequent sputum sampling to detect smear conversion ASAP, increasing costs

Ref: Harms of Isolation in TB, 1-5

Key message

Studies suggest the infectiousness of people with pulmonary TB disease on effective therapy is related to the efficacy and duration of TB treatment. People with pulmonary TB quickly become non-infectious once effective TB therapy starts, usually within days.

Pre-Clinical Studies

- Guinea pig data demonstrate that human-to-guinea pig transmission nearly stops within 1-2 days of effective therapy. (Ref 1-4)
- Cough aerosol sampling show that once treatment starts, production of cough aerosols with infectious bacilli ends in days (Ref 5-9)
- Transcriptomic studies demonstrate that the *M. tuberculosis* transcriptome rapidly changes once effective therapy has started, which may in turn impact infectiousness (Ref 9-11)

Clinical Studies

- RCT data: one RCT in Madras >50 years ago: no difference in TB incidence between household contacts of people treated in sanatorium versus those “de-isolated” at home (Ref 1-2)
- Observational data:
 - TST positivity in contacts does not differ by sputum status at time of de-isolation (3)
 - No conversions in contacts of people de-isolated while smear positive (4)

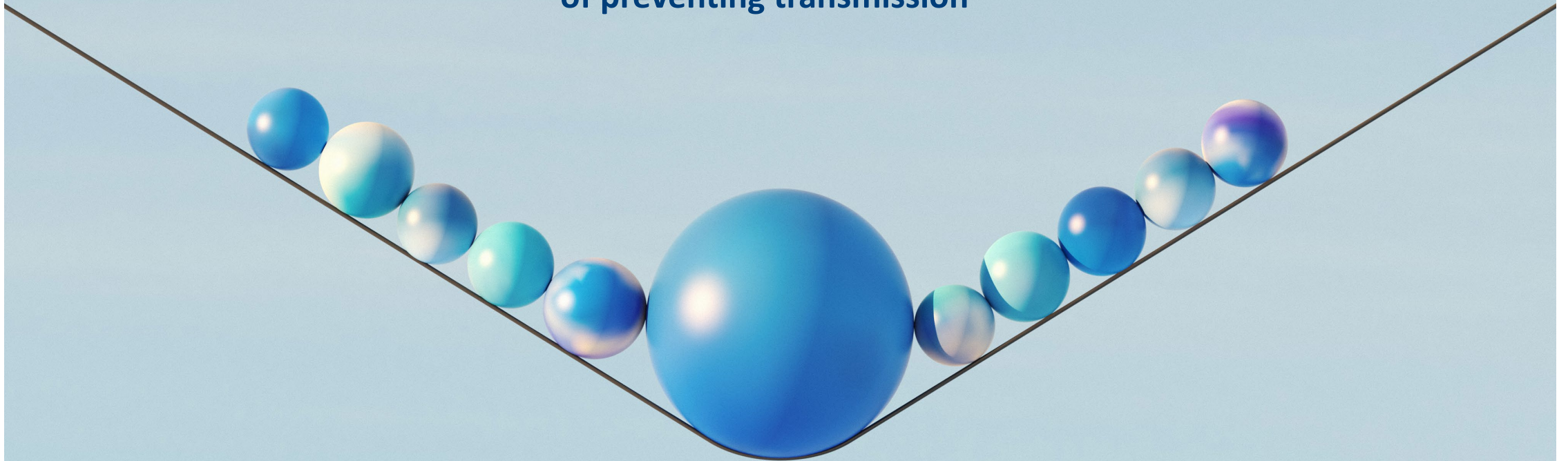
Practice Implications

- Smear and culture status are helpful indicators of infectiousness pre-treatment.
- Once effective TB therapy is started, microbiological results are poor indicators of infectiousness.
- De-isolation of people with pulmonary TB should focus on treatment efficacy and duration.



Aim of TB de-isolation recommendations

Balance the real, tangible harms of prolonged isolation to person with TB with responsibility of preventing transmission



De-Isolation Recommendation



Discontinue isolation for **rifampin-susceptible*** TB results when the following are true:

At diagnosis: AFB smear-**negative**/culture-positive

- Evidence of clinical improvement
- 2 weeks of effective, multidrug therapy is completed

AFB smear-**positive**/culture-positive

- Evidence of clinical improvement
- 2 weeks of effective, multidrug therapy is completed
- 3 consecutive AFB-negative sputum-smears

Note: No change from previous recommendations

*De-Isolation recommendations are specific to rifampin-susceptible TB organisms since at many sites the available rapid resistance testing will be completed via a GeneXpert. Further confirmation at the BCCDC Public Health Lab may take longer than 2 or 4 weeks.

De-Isolation Recommendation



Discontinue isolation for **rifampin-susceptible** TB results when the following are true:

Persistent* AFB smear-**positive**/culture-**positive**

- Evidence of clinical improvement
- 4 weeks of effective, multidrug therapy is completed

This update reflects that a **maximum duration** of isolation has been established.

*persistent means smears continue to be positive after 2 weeks effective therapy

De-Isolation Recommendation



Discontinue isolation for **rifampin-resistant** TB results when the following are true:

Rifampin-resistant pulmonary TB

- Case review with TB Services
- Evidence of clinical improvement
- 4 weeks of effective, 2nd line multidrug therapy is completed
- 2nd line sensitivity results are available
- If initially AFB smear positive, three consecutive AFB smears must be negative



This update reflects that confirmation of culture conversion is not needed to de-isolate.



Further points

- Emphasis on home-isolation wherever possible, and ASAP
 - Emphasis on strategies to support strong adherence
- US guidelines de-isolate within as few as 5 days post-effective therapy
 - The practicality of these recommendations and rollout have yet to be tested

References – Guidance Documents

1. Cooper R. Appendix B: De-isolation review and recommendations. Canadian TB Standards, 8th edition. *Can J Resp Crit Care Med*; Vol 6, 2022.
2. Shah M. National Tuberculosis Coalition of America (NTCA) Guidelines for Respiratory Isolation and Restrictions to reduce transmission of Pulmonary Tuberculosis in Community settings. *Clin Inf Dis* 18 April 2024.

References – Harms of Isolation in TB

1. Gammon J. The psychological consequences of source isolation: a review of the literature. *J Clin Nurs*. 1999 Jan;8(1):13–21.
2. Basham CA, Karim ME, Cook VJ, Patrick DM, Johnston JC. Tuberculosis-associated depression: a population-based cohort study of people immigrating to British Columbia, Canada, 1985-2015. *Ann Epidemiol*. 2021 Nov;63:7–14.
3. Sprague E, Reynolds S, Brindley P. Patient Isolation Precautions: Are They Worth It? *Can Respir J*. 2016;2016:5352625.
4. Moffatt J, Mayan M, Long R. Sanitoriums and the Canadian colonial legacy: the untold experiences of tuberculosis treatment. *Qual Health Res*. 2013 Dec;23(12):1591–9.
5. Horter S, Stringer B, Reynolds L, Shoaib M, Kasozi S, Casas EC, et al. “Home is where the patient is”: a qualitative analysis of a patient-centred model of care for multi-drug resistant tuberculosis. *BMC Health Serv Res* [Internet]. 2014 Feb 21 [cited 2025 Jan 9];14:81. Available from: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3943511/>

References – Pre-Clinical Evidence

1. Riley RL, Mills CC, O’Grady F, et al. Infectiousness of air from a tuberculosis ward. Ultraviolet irradiation of infected air: comparative infectiousness of different patients. *Am Rev Respir Dis*. 1962;85(9):511–525. doi:10.1164/arrd.1962.85.4.511.
2. Dharmadhikari AS, Mphahlele M, Venter K, Stoltz A, Mathebula R, Masotla T, et al. Rapid impact of effective treatment on transmission of multidrug-resistant tuberculosis. *Int J Tuberc Lung Dis Off J Int Union Tuberc Lung Dis*. 2014 Sep;18(9):1019–25. Escombe AR *PLoS Med*. 2008;5(9):1387–1396
3. Escombe AR, Moore DAJ, Gilman RH, Pan W, Navincopa M, Ticona E, et al. The Infectiousness of Tuberculosis Patients Coinfected with HIV. *PLOS Med [Internet]*. 2008 Sep 16 [cited 2025 Jan 9];5(9):e188. Available from: <https://journals.plos.org/plosmedicine/article?id=10.1371/journal.pmed.0050188> Fennelly KP *AJRCCM* 2004 and 2012;186(5):450–457 6.
4. Escombe AR, Oeser C, Gilman RH, Navincopa M, Ticona E, Martínez C, et al. The detection of airborne transmission of tuberculosis from HIV-infected patients, using an in vivo air sampling model. *Clin Infect Dis Off Publ Infect Dis Soc Am*. 2007 May 15;44(10):1349–57.
5. Fennelly KP, Jones-López EC, Ayakaka I, Kim S, Menyha H, Kirenga B, et al. Variability of infectious aerosols produced during coughing by patients with pulmonary tuberculosis. *Am J Respir Crit Care Med*. 2012 Sep 1;186(5):450–7.
6. Fennelly KP, Martyny JW, Fulton KE, Orme IM, Cave DM, Heifets LB. Cough-generated aerosols of *Mycobacterium tuberculosis*: a new method to study infectiousness. *Am J Respir Crit Care Med*. 2004 Mar 1;169(5):604–9.

References – Pre-Clinical Evidence

7. Jones-López EC, Acuña-Villaorduña C, Ssebidandi M, Gaeddert M, Kubiak RW, Ayakaka I, et al. Cough Aerosols of *Mycobacterium tuberculosis* in the Prediction of Incident Tuberculosis Disease in Household Contacts. *Clin Infect Dis Off Publ Infect Dis Soc Am*. 2016 Jul 1;63(1):10–20.
8. Jones-López EC, Namugga O, Mumbowa F, Ssebidandi M, Mbabazi O, Moine S, et al. Cough Aerosols of *Mycobacterium tuberculosis* Predict New Infection. A Household Contact Study. *Am J Respir Crit Care Med* [Internet]. 2013 May 1 [cited 2025 Jan 9];187(9):1007–15. Available from: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3707366/>
9. Acuña-Villaorduña C, Ayakaka I, Schmidt-Castellani LG, et al. Host determinants of infectiousness in smear-positive patients with pulmonary tuberculosis. *Open Forum Infect Dis*. 2019;6(6):ofz184–9. doi:10.1093/ofid/ofz184.
10. Theron G, Limberis J, Venter R, Smith L, Pietersen E, Esmail A, et al. Bacterial and host determinants of cough aerosol culture positivity in patients with drug-resistant versus drug-susceptible tuberculosis. *Nat Med*. 2020 Sep;26(9):1435–43. Shaikh A. et al. *Sci Rep* 2021;11:17812.
11. Shaikh A, Sriraman K, Vaswani S, Oswal V, Rao S, Mistry N. Early phase of effective treatment induces distinct transcriptional changes in *Mycobacterium tuberculosis* expelled by pulmonary tuberculosis patients. *Sci Rep* [Internet]. 2021 Sep 8 [cited 2025 Jan 9];11(1):17812. Available from: <https://www.nature.com/articles/s41598-021-96902-7>
12. Walter ND, Born SEM, Robertson GT, Reichlen M, Dide-Agossou C, Ektnitphong VA, et al. *Mycobacterium tuberculosis* precursor rRNA as a measure of treatment-shortening activity of drugs and regimens. *Nat Commun* [Internet]. 2021 May 18 [cited 2025 Jan 9];12(1):2899. Available from: <https://www.nature.com/articles/s41467-021-22833-6>

References – RCT and cohort studies

1. Kamat SR, Dawson JJY, Devadatta S, et al. A controlled study of the influence of segregation of tuberculous patients for one year on the attack rate of tuberculosis in a 5-year period in close family contacts in South India. *Bulletin of the World Health Organization*. 1998;76(2):109–124.
2. Devadatta S, Dawson JJ, Fox W, et al. Attack rate of tuberculosis in a 5-year period among close family contacts of tuberculous patients under domiciliary treatment with isoniazid plus PAS or isoniazid alone. *Bull World Health Organ*. 1970;42(3):337–351.
3. Gunnels J. J., Bates J. H., Swindoll H. Infectivity of sputum-positive tuberculous patients on chemotherapy. *Am Rev Respir Dis*. 1974 Mar;109(3):323–330. doi: 10.1164/arrd.1974.109.3.323.
4. Riley RL, Moodie AS. Infectivity of patients with pulmonary tuberculosis in inner city homes. *Am Rev Respir Dis*. 1974;110(6):810–812. doi:10.1164/arrd.1974.110.6P1.810.