Indian J Med Res 156, July 2022, pp 6-9

DOI: 10.4103/ijmr.ijmr_893_22

Editorial



Closing gaps in asthma care in India – World Asthma Day 2022

'Closing gaps in asthma care' is the theme that the Global Initiative for Asthma (GINA) has promoted for World Asthma Day 2022. We take the opportunity to shine a spotlight on a major gap in asthma care in India—the gap between the number of people with asthma in India who currently take inhaled corticosteroids (ICSs) and the number who would benefit (namely all of them).

Asthma is the most common chronic disease in children and adolescents globally and one of the most common chronic diseases in adults¹⁻⁵. Asthma is recognized by the World Health Organization (WHO) and other stakeholders as both cause and effect of poverty in low- and middle-income countries (LMICs)⁴⁻⁶. Children and adolescents with uncontrolled asthma miss out on education, adults on work opportunities and all are impacted by some combination of respiratory symptoms, adverse effects on health and well-being, costs of medication and exacerbations associated with emergency department visits, hospitalizations and even death^{4,5}. Around 96 per cent of global asthma deaths occur in LMICs⁴.

India has a high and growing burden of asthma: according to the Global Burden of Disease Report 2019, India ranks number one in the world in terms of burden, disability-adjusted life years and deaths related to asthma⁷. Over 34 million people in India have asthma, and although this reflects only 13 per cent of the world's population with asthma, 42 per cent of global asthma deaths occur in India⁸. People with asthma in India also suffer a disproportionately high burden of day-to-day symptoms, effects on quality of life and absence from school and work^{9,10}. In lowincome households, even minor levels of healthcare utilization can be financially catastrophic¹¹.

Although ICSs are the cornerstone of asthma treatment and are now recommended for all patients

with asthma because these dramatically reduce the risk of severe exacerbations and asthma-related death¹² the 2015 Asthma Insights and Management (AIM) survey found that fewer than a third of people with asthma in India used inhalers⁹. In common with the situation in other LMICs, when treatment for asthma was used. this was most likely to be oral medication including oral salbutamol, theophylline and prednisolone^{4,5,9}. Such substantial divergence from international asthma management recommendations leaves so many patients at risk of avoidable morbidity and mortality^{4,5,9}. For example, a study of 381 Indian patients receiving oral asthma medications found that almost one-third had a hospitalization due to asthma or asthma symptoms in the previous year¹³. In the 2015 AIM India survey, 89 per cent people with asthma reported using oral corticosteroids in the previous year, with an average of 10.5 courses9. Such high usage greatly increases the risk of serious long-term adverse effects such as cataract, osteoporosis and diabetes¹⁴.

Inhaled asthma treatments were developed over half a century ago and represent the safest, fastest and most effective way of delivering anti-inflammatory treatments and bronchodilators to where these are needed¹⁵. Furthermore, there is strong evidence that ICSs reduce asthma morbidity and mortality¹⁶⁻¹⁹. In turn, adequate asthma management brings economic benefits to individuals and their families²⁰. The gap between the number of people who would benefit and the number of people currently using ICS for the treatment of asthma in India is therefore of great concern. The magnitude of this gap was quantified in a recent study based on the estimated number of people with asthma in India from the Global Burden of Disease study and their estimated need for ICS, compared with the number of total ICS sales in India that were assumed to be for asthma. The authors quantified the

This editorial is published on the occasion of World Asthma Day- May 3, 2022.

gap between corticosteroid inhalers sold for asthma versus corticosteroid inhalers needed for the treatment of asthma in India as 26.4 million versus 384.2 million corticosteroid inhalers per year⁸. It seems likely that this gap explains the gap between the expected and disproportionately high death rates from asthma in India. It is a gap that needs to be filled urgently.

Apart from the social issues and the myths and misbeliefs associated with inhalation therapy²¹, one of the main reasons why people with asthma do not use ICSs in India is that primary care physicians themselves are not aware about the importance of ICS in asthma management. An estimated 90,000 modern medicine doctors graduate every year from 595 plus medical schools across India²². Unfortunately, most of the tertiary care teaching hospitals in India do not have ICS in their formularies²³, despite ICS being on the WHO's essential medicines list²⁴. Students are also not taught about the importance of inhalation therapy and are not well versed with the use of inhaler devices²⁵⁻²⁷. As a result, trainee doctors learn to prescribe oral medications for the treatment of asthma, including tablets of salbutamol and theophyllines, which then becomes the norm when they become primary care physicians. Teaching undergraduate medical students the importance of inhalation therapy and the importance of ICS in the management of asthma will help change clinical practice across India. To achieve this, every tertiary care teaching hospital must have ICS and ICS plus short-acting β_2 -agonist (SABA) or combination of ICS plus formoterol in the pharmacy for dispensing. The teaching curriculum should also focus on the importance and techniques for the use of inhaled medications. This one intervention may help bridge the gap for the underuse of ICS in asthma management and thereby help reduce asthma mortality in India.

As efforts are made to fill the gaping need for ICS for all with asthma in India, we suggest that there is a unique opportunity to do so in a way that avoids the risk of retracing the historical approach to asthma treatment in other settings. The conventional approach of starting treatment with inhaled SABA trains people with asthma to rely on its short-term symptom relief, and this reduces the likelihood that they will later be prepared, even if ICS is available and prescribed, to use it every day to address airway inflammation and reduce their risk of severe exacerbations and death²⁸. There is now robust evidence that treatment with a combination inhaler containing both ICS and formoterol [a fast-onset long-acting β_2 -agonist

(LABA)], taken whenever needed for symptom relief, reduces the risk of severe asthma exacerbations compared with using a SABA reliever, with similar levels of asthma symptom control, and with a lower overall dose of ICS; these benefits are seen both with as-needed ICS-formoterol alone, and with maintenance and reliever therapy with ICS-formoterol (MART)12. A recent systematic review and meta-analysis of trials of as-needed ICS-formoterol in patients with mild asthma found that severe exacerbations were substantially lower with ICS-formoterol compared with as-needed SABA [odds ratio (OR) 0.45; 95% confidence interval (CI) 0.34-0.60²⁹. Further, as-needed ICS-formoterol reduced the risk of emergency department visits and hospitalizations compared with daily ICS plus as-needed SABA (OR 0.63; CI 0.44-0.91)²⁹. Similarly, a systematic review and meta-analysis of trials of ICS-formoterol for maintenance and reliever therapy found that with this approach, severe exacerbations were considerably lower compared with the same maintenance dose of ICS-formoterol (relative risk 0.68; 95% CI: 0.58-0.80)³⁰. In light of this evidence and safety concerns about inhaled SABA monotherapy, GINA now recommends ICS-formoterol as the preferred reliever for adolescents and adults with asthma, with (Steps 3-5) or without (Steps 1-2) maintenance ICS-formoterol¹².

There is an urgent need to fill the gap between the number of people with asthma who take ICS and the number who would benefit in India. The same holds true for other LMICs where inhaled medicines are poorly available and largely unaffordable³¹. The ICS-formoterol inhaler-based approach has the potential to transform asthma care in India by becoming the standard of care and by avoiding SABA-only treatment. Closing the gap in actual versus expected morbidity and mortality from asthma in India would help lead the way for other LMICs where this gap is also unacceptably wide.

Financial support & sponsorship: None.

Conflicts of Interest: The first author (KM) reports advisory board fees from AstraZeneca. The second author (SSS) reports advisory board and speaker fees from GSK, Cipla Ltd. The third author (HKR) reports research funding from AstraZeneca, GlaxoSmithKline and Novartis; and honoraria from AstraZeneca, Boehringer Ingelheim, Chiesi, GlaxoSmithKline, Novartis, Sanofi and Teva, for independent medical education, advisory boards and/or consultancy.

Kevin Mortimer^{1,2}, Sundeep Santosh Salvi^{3,4} & Helen K. Reddel^{5,6*}

¹Department of Respiratory Medicine, Liverpool University Hospitals NHS Foundation Trust, Liverpool, ²Department of Medicine, University of Cambridge, Cambridge, UK, ³Department of Clinical Research, Pulmocare Research & Education (PURE) Foundation, ⁴Faculty of Health and Biological Sciences, Symbiosis International (Deemed University), Pune 412 115, Maharashtra, India, ⁵Clinical Management Group, The Woolcock Institute of Medical Research, Glebe & ⁶The Faculty of Medicine and Health, The University of Sydney, Sydney, Australia *For correspondence: helen.reddel@sydney.edu.au

Received April 17, 2022

References

- Asher MI, Rutter CE, Bissell K, Chiang CY, El Sony A, Ellwood E, et al. Worldwide trends in the burden of asthma symptoms in school-aged children: Global Asthma Network Phase I cross-sectional study. Lancet 2021; 398: 1569-80.
- García-Marcos L, Innes Asher M, Pearce N, Ellwood E, Bissell K, Chiang CY, et al. The burden of asthma, hay fever and eczema in children in 25 countries: GAN Phase I study. Eur Respir J 2022. DOI:10.1183/13993003.02866-2021.
- Mortimer K, Lesosky M, García-Marcos L, Innes Asher M, Pearce N, Ellwood E, et al. The burden of asthma, hay fever and eczema in adults in 17 countries: GAN Phase I study. Eur Respir J 2022. DOI:10.1183/13993003.02865-2021.
- 4. Meghji J, Mortimer K, Agusti A, Allwood BW, Asher I, Bateman ED, *et al.* Improving lung health in low-income and middle-income countries: From challenges to solutions. *Lancet* 2021; 397: 928-40.
- Mortimer K, Reddel HK, Pitrez PM, Bateman ED. Asthma management in low- and middle-income countries: Case for change. Eur Respir J 2022. DOI:10.1183/13993003.03179-2021.
- 6. World Health Organization. Global surveillance, prevention and control of chronic respiratory diseases: A comprehensive approach; 2007. Available from: https://www.who.int/gard/publications/GARD Manual/en/, accessed on August 17, 2020.
- 7. Global burden of Diseases. The Lancet: Latest global disease estimates reveal perfect storm of rising chronic diseases and public health failures fuelling COVID-19 pandemic. Available from: https://www.healthdata.org/gbd/2019, accessed on March 19, 2022.
- 8. Salvi S, Madas S, Ghorpade D, Gadhave S, Barne M. Is underuse of inhaled corticosteroids for asthma in India contributing to 42% of global asthma deaths? *Lung India* 2022. DOI:10.4103/lungindia.lungindia_600_21.

- Salvi SS, Apte KK, Dhar R, Shetty P, Faruqi RA, Thompson PJ, et al. Asthma insights and management in India: Lessons learnt from the Asia Pacific – Asthma insights and management (AP-AIM) study. J Assoc Physicians India 2015; 63: 36-43.
- 10. Gold LS, Thompson P, Salvi S, Faruqi RA, Sullivan SD. Level of asthma control and health care utilization in Asia-Pacific countries. *Respir Med* 2014; *108*: 271-7.
- Su TT, Kouyaté B, Flessa S. Catastrophic household expenditure for health care in a low-income society: A study from Nouna District, Burkina Faso. *Bull World Health Organ* 2006: 84: 21-7.
- 12. Reddel HK, Bacharier LB, Bateman ED, Brightling CE, Brusselle GG, Buhl R, *et al.* Global Initiative for Asthma Strategy 2021: Executive summary and rationale for key changes. *Respirology* 2022; 27:14-35.
- 13. Dhar R, Chhowala S, Lopez M, Hegde R, Madas S, Salvi S, *et al.* Assessment of asthma control in users of oral anti-asthma medications. *Int J Tuberc Lung Dis* 2021; *25*: 620-5.
- Price DB, Trudo F, Voorham J, Xu X, Kerkhof M, Ling Zhi Jie J, et al. Adverse outcomes from initiation of systemic corticosteroids for asthma: Long-term observational study. J Asthma Allergy 2018; 11: 193-204.
- Crompton G. A brief history of inhaled asthma therapy over the last fifty years. Prim Care Respir J 2006; 15: 326-31.
- 16. Suissa S, Ernst P, Benayoun S, Baltzan M, Cai B. Low-dose inhaled corticosteroids and the prevention of death from asthma. *N Engl J Med* 2000; *343*: 332-6.
- 17. Suissa S, Ernst P. Use of anti-inflammatory therapy and asthma mortality in Japan. *Eur Respir J* 2003; *21* : 101-4.
- Nunes de Melo M, Mendes Z, Martins P, Suissa S. Asthma mortality in portugal: Impact of treatment with inhaled corticosteroids and leukotriene receptor antagonists. *Treat Respir Med* 2006; 5: 143-7.
- Pauwels RA, Pedersen S, Busse WW, Tan WC, Chen YZ, Ohlsson SV, et al. Early intervention with budesonide in mild persistent asthma: A randomised, double-blind trial. Lancet 2003; 361: 1071-6.
- Franco R, Nascimento HF, Cruz AA, Santos AC, Souza-Machado C, Ponte EV, et al. The economic impact of severe asthma to low-income families. Allergy 2009; 64: 478-83.
- 21. Krishna MT, Mahesh PA, Vedanthan P, Moitra S, Mehta V, Christopher DJ. An appraisal of allergic disorders in India and an urgent call for action. *World Allergy Organ J* 2020; *13*: 100446.
- The Quint. India has largest number of medical colleges, what about good doctors? Available from: https://www.thequint. com/voices/opinion/india-has-largest-number-of-medical-col leges-what-about-good-doctors, accessed on May 11, 2022.
- Kotwani A. Access to essential medicines and standard treatment for chronic diseases. *Indian J Pharmacol* 2010; 42: 127-8.

- 24. World Health Organization. WHO model list of essential medicine. Available from: https://www.who.int/groups/expert-committee-on-selection-and-use-of-essential-medicines/essen tial-medicines-lists, accessed on March 19, 2022.
- 25. Mullerpattan JB, Udwadia ZZ, Kathar SS, Shah HD, Rastogi SA, Pandey KV, et al. Who will teach the teachers: An analysis of the inhaler technique of Indian patients and health care providers in a tertiary health care centre. Lung India 2016; 33: 493-5.
- Akhoon N, S Brashier DB. A study to monitor errors in use of inhalation devices in patients of mild-to-moderate bronchial asthma in a tertiary care hospital in Eastern India. *Perspect Clin Res* 2022; 13: 17-24.
- Padmanabhan M, Tamilarasu K, Rajaram M, Batmanabane G. Inadequate inhaler technique, an everlasting problem, is associated with poor disease control – A cross sectional study. Adv Respir Med 2019; 87: 217-25.

- Baggott C, Chan A, Hurford S, Fingleton J, Beasley R, Harwood M, et al. Patient preferences for asthma management: A qualitative study. BMJ Open 2020; 10: e037491.
- Crossingham I, Turner S, Ramakrishnan S, Fries A, Gowell M, Yasmin F, et al. Combination fixed-dose beta agonist and steroid inhaler as required for adults or children with mild asthma. Cochrane Database Syst Rev 2021; 5: CD013518.
- Sobieraj DM, Weeda ER, Nguyen E, Coleman CI, White CM, Lazarus SC, et al. Association of inhaled corticosteroids and long-acting β-agonists as controller and quick relief therapy with exacerbations and symptom control in persistent asthma: A systematic review and meta-analysis. JAMA 2018; 319: 1485-96.
- 31. Stolbrink M, Thomson H, Hadfield RM, Ozoh OB, Nantanda R, Jayasooria S, *et al*. The availability, cost and affordability of essential medicines for asthma and COPD in low-income and middle-income countries: A systematic review. Available from: http://dx.doi.org/10.2139/ssrn.4023200, accessed on March 19, 2022.