

## Authors contribution

Active participation in patient care and the search for treatment options. Writing of the clinical case and research of articles that could support the treatment and the case report

## References

- [1] Poussel G, Deschildre A, Le Bourgeois M, Cuisset JM, Catteau B, Karila C, et al. The lung is involved in juvenile dermatomyositis. *Pediatr Pulmonol.* 2013;48(10):1016–25.
  - [2] Pallo P, Shinjo S. Spontaneous pneumomediastinum in dermatomyositis: a case series and literature review. *Medical Express.* 2017;5, mo18009.
  - [3] Romanelli P, Bastos M, Zerbini C. Dermatomiosite juvenil de diagnóstico tardio com pneumotórax, pneumomediastino e enfisema subcutâneo espontâneos. *Rev Paul Med.* 1990;108(1):45–8.
  - [4] Dogra S, Suri D, Shah R, Rawat A, Singh S, Sodhi KS. Spontaneous pneumomediastinum: a rare complication of juvenile dermatomyositis. *Int J Rheum Dis.* 2012;15:e131–3.
  - [5] Le Goff B, Chérin P, Cantagrel A, Gayraud M, Hachulla E, Laborde F, et al. Pneumomediastinum in interstitial lung disease associated with dermatomyositis and polymyositis. *Arthritis Rheumatism.* 2009;61(1):108–18.
  - [6] Ma X, Chen Z, Hu W, Guo Z, Wang Y, Kuwana M, et al. Clinical and serological features of patients with dermatomyositis complicated by spontaneous pneumomediastinum. *Clin Rheumatol.* 2015;35(2):489–93.
  - [7] Sato S, Uejima Y, Nanbu M, Suganuma E, Takano T, Tanaka R, et al. Clinical analysis and outcome of interstitial lung disease complicated with juvenile dermatomyositis and juvenile polymyositis. *Modern Rheumatol.* 2016, <http://dx.doi.org/10.1080/14397595.2016.1231102>.
- Renata Lopes Francisco de Andrade <sup>a,\*</sup>,  
Ana Carolina Lopes Held <sup>a</sup>, Maria Teresa Terreri <sup>a</sup>,  
Sonia Mayumi Chiba <sup>b</sup>
- <sup>a</sup> Department of Pediatrics, Section of Rheumatology,  
Federal University of São Paulo – UNIFESP, São Paulo,  
Brazil
- <sup>b</sup> Department of Pediatrics, Section of Pulmonology,  
Federal University of São Paulo – UNIFESP, São Paulo,  
Brazil
- \* Corresponding author at: Department of Pediatrics, Section of Rheumatology, Federal University of São Paulo – UNIFESP, Borges Lagoa, St, 802. Vila Clementino, São Paulo, SP, 04083-001, Brazil.  
E-mail address: drarenatalf@gmail.com (R.L.F. de Andrade).
- 23 July 2019
- <https://doi.org/10.1016/j.pulmoe.2019.12.002>  
2531-0437/ © 2019 Sociedade Portuguesa de Pneumologia.  
Published by Elsevier España, S.L.U. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

## Severe asthma intervention in adult obese patients



Dear Editor,

Obesity and asthma are prevalent disorders, both with an important impact on public health.<sup>1</sup> A meta-analysis of seven prospective studies found a relationship between obesity and asthma: incident asthma OR in overweight and obese groups was 1.5 and 1.9, respectively, when compared to the normal weight group.<sup>1</sup> In fact, obese patients have an increased risk of asthma, and obese asthmatics have even more symptoms, frequent and severe exacerbations and worse response to asthma-specific therapies.<sup>1–4</sup> Weight loss has been proposed as essential in these patients, where up to 5–10% of weight loss is associated with improved asthma control and quality of life.<sup>1,5</sup> In this way, bariatric surgery is considered as the key option to promote a substantial weight loss.<sup>6–8</sup>

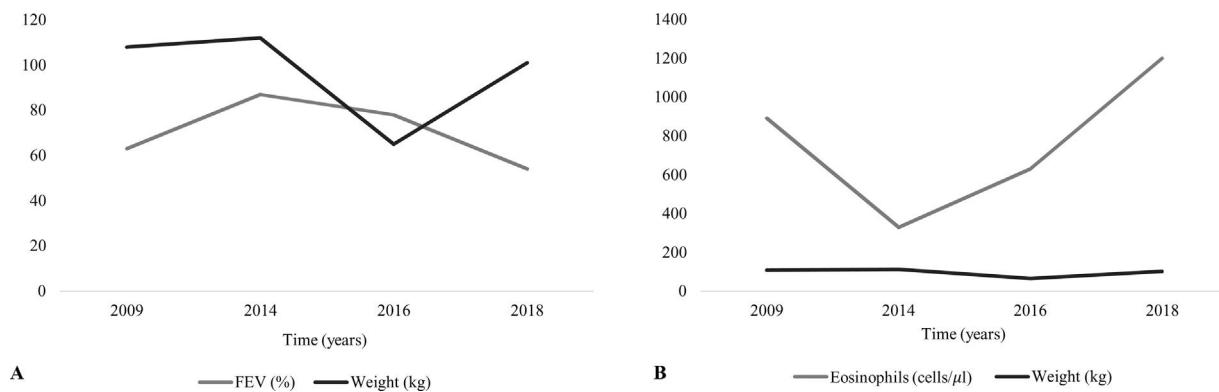
Here, we report the case of an obese adult woman who presented with severe allergic-predominant asthma, who really improved her symptoms and frequency of exacerbations following bariatric surgery. However, she developed a psychiatric disorder due to the non-acceptance of her new body image.

A thirty-eight-year-old woman was referred to a pulmonology department because of severe uncontrolled steroid-dependent asthma. She had frequent cough, shortness of breath, wheezing and recurrent exacerbations. She had other comorbidities, like obesity (BMI 38), gastroesophageal reflux disease (GERD) and chronic rhinosinusitis.

The patient's comorbidities, exposure to allergens or other harmful agents, as well as adherence and the appropriate use of current treatment were assessed. After the evaluation, weight loss was proposed and proton pump inhibitors and intranasal corticosteroids were prescribed. Asthma treatment was reviewed and optimized (with reduction of systemic corticosteroids (SC), high-dose inhaled corticosteroids (ICS) and long-acting beta 2-agonists (LABA), long-acting muscarinic antagonists (LAMA), leukotriene receptor antagonists (LTRAs)). However, the patient continued to have uncontrolled asthma.

The patient had reduced lung function (FEV1 63% pred, Tiffeneau index 61), atopy (skin prick test positive), IgE 454 kU/L and blood eosinophils count of 890 cells/l. Based on this, anti-IgE treatment (omalizumab 525 mg, SC each 2 weeks) was started.

Over the next four years (Fig. 1), the patient was asymptomatic, had a total CARAT score of 22 (9 + 13), decreased blood eosinophil count (260 cells/l), improved respiratory function tests (FEV1 87% pred), showed a clear decrease in the number of exacerbations and did not report the need for systemic therapy with corticosteroids. After that, the patient had progressively worse asthma control, despite using specific asthma therapy. Thus, the patient's comorbidities, exposure to allergens, as well as adherence and the appropriate use of current treatment were re-evaluated. The patient gained weight, reaching BMI of 40 kg/m<sup>2</sup>. She was diagnosed with moderate obstructive sleep apnea (OSA) syndrome (apnea-hypopnea index of 20.3 events/h) and automatic positive airway pressure (APAP) therapy was



**Figure 1** Variation in (A) FEV (%) and weight (kg) and (B) eosinophils count (cells/L) and weight (kg) over time following omalizumab treatment initiation.

initiated. Meanwhile, after a psychiatric evaluation, she underwent bariatric surgery. Following bariatric surgery, the patient had a rapid reduction in BMI, 65 kg in 6 months. Subsequently, the patient presented a clinical (total CARAT score of 28) and functional improvement and did not require systemic corticosteroids or omalizumab therapy. However, the patient started having depressive symptoms and lack of acceptance of her new body image, being referred for psychiatric re-evaluation. She started drug treatment and cognitive-behavioral therapy, but her weight persistently increased and asthma became uncontrolled. Thus, due to a new severe uncontrolled asthma, anti-IgE treatment was repurposed.

In fact, obesity and asthma are prevalent disorders, and there is an association between the two conditions,<sup>1,3,4</sup> where obesity is both a major risk factor and an asthma modifier. Patients with severe asthma usually have limiting symptoms, exacerbations and side effects of medications, especially with the prolonged and recurrent use of SC.<sup>2,9</sup> The adverse effects of SC include obesity, diabetes, hypertension and psychological disorders, such as depression and anxiety. Therefore, although asthma in obese patients is more commonly associated with non-type 2 inflammation,<sup>2</sup> obese patients with childhood-onset asthma tend to have higher markers of Th2 inflammation and more severe disease among obese asthmatics.<sup>4,9</sup> Studies have shown that weight loss interventions are associated with better control of asthma and respiratory function,<sup>2,10</sup> and that bariatric surgery promotes a rapid and sustained weight loss, and is generally associated with significant improvement in both asthma control and quality of life scores, reducing the risk of hospitalizations due to asthma exacerbation,<sup>7,8,10</sup> and even triggering a significant decrease in treatment step.<sup>10</sup> However, although obese asthmatic patients can be treated with specific asthma therapy and weight loss interventions, some patients still remain uncontrolled.

Obesity and asthma are frequently and independently associated with other conditions, which are also associated with worse asthma control, such as GERD, OSA and mood disorders.<sup>2-4</sup> Mood disorders, which consisted mainly of anxiety and depression, have been reported to be strongly correlated with asthma symptoms severity.<sup>11</sup> Thus, severe asthma management is difficult and needs a careful and multidimensional assessment.<sup>12</sup>

Here we report the case of an obese adult woman who presented with predominant severe allergic asthma, who truly improved her symptoms and exacerbations frequency following specific asthma therapy and bariatric surgery. However, despite all comorbidities and risk factors having been systematically assessed, their management was difficult, never allowing a longstanding asthma control. This may be due to a different type of airway inflammation, mechanical factors and other commodities that are associated with obesity.<sup>2-4</sup>

Specifically, although the evaluation of all comorbidities is extremely important, it must be emphasized that, in the case of mood disorders evaluation, this can have a negative impact if not appropriately recognized and treated.<sup>11</sup> The patient mentioned in this case started unpredictable depressive symptoms and lack of acceptance of her new body image, following a rapid reduction in BMI. Once depression was suspected, the patient was referred for psychiatric evaluation and management. She started drug treatment and cognitive-behavioral therapy, but her weight persistently increased and her asthma became uncontrolled. In conclusion, the management of this patient remains difficult, as the various comorbidities diverge and antagonize each other. Taking this into account, anti-IgE therapy was restarted.

In short, given that obesity is an important risk factor for asthma and asthma-related morbidity, weight loss interventions should be highly encouraged to ensure proper control of asthma, lung function and quality of life. However, although this aspect is extremely important, it is also essential to ensure holistic management with an appropriate approach to all comorbid conditions in obese asthmatic individuals.

## References

- Beuther DA, Weiss ST, Sutherland ER. Obesity and asthma. *Am J Respir Crit Care Med.* 2006;174:112–9.
- Global Initiative for Asthma. Global strategy for asthma management and prevention. *Glob Strateg Asthma Manag Prev* [Internet]. 2018;1–162. Available from: <http://ginasthma.org/gina-report-global-strategy-for-asthma-management-and-prevention/>.

3. Baffi CW, Winnica DE, Holguin F. Asthma and obesity: mechanisms and clinical implications. *Asthma Res Pract* [Internet]. 2015;1:1–7. Available from: <https://doi.org/10.1186/s40733-015-0001-7>
  4. Peters U, Dixon AE, Forno E. Obesity and asthma. *J Allergy Clin Immunol* [Internet]. 2018;141:1169–79. Available from: <https://doi.org/10.1016/j.jaci.2018.02.004>
  5. Dias-Júnior SA, Reis M, Carvalho-Pinto RM, de Stelmach R, Halpern A, Cukier A. Effects of weight loss on asthma control in obese patients with severe asthma. *Eur Respir J*. 2014;43:1368–77.
  6. Dixon AE, Pratley RE, Forgione PM, Kaminsky DA, Whittaker-Leclair LA, Griffes LA, et al. Effects of obesity and bariatric surgery on airway hyperresponsiveness, asthma control and inflammation. *J Allergy Clin Immunol*. 2011;124:508–15.
  7. Boulet LP, Turcotte H, Martin J, Poirier P. Effect of bariatric surgery on airway response and lung function in obese subjects with asthma. *Respir Med*. 2012;106:651–60.
  8. Hasegawa K, Tsugawa Y, Chang Y, Camargo CA. Risk of an asthma exacerbation after bariatric surgery in adults. *J Allergy Clin Immunol* [Internet]. 2015;136:288–94, e8. Available from: <https://doi.org/10.1016/j.jaci.2014.12.1931>
  9. Global Initiative for Asthma. Difficult-to-treat asthma in adolescent and adult patients: diagnosis and management – a GINA pocket guide for health professionals, vol. 2. *Global Strategy for Asthma Management and Prevention*; 2019.
  10. Santos LM, Ramos B, Almeida J, Loureiro CC, Cordeiro CR. The impact of weight loss beyond lung function: benefit with respect to asthma outcomes. *Pulmonology* [Internet]. 2019;25:313–9. Available from: <https://www.journalpulmonology.org/en-the-impact-weight-loss-beyond-articulo-S2531043719301382>
  11. Kewalramani A, Bollinger ME, Postolache TT. Asthma and mood disorders. *Int J Child Heal Hum Dev*. 2008;1:115–23.
  12. Clark VL, Gibson PG, Genn G, Hiles SA, Pavord ID, McDonald VM. Multidimensional assessment of severe asthma: a systematic review and meta-analysis. *Respirology*. 2017;22:1262–75.
- M. Serino <sup>a,\*</sup>, M. van Zeller <sup>a,b</sup>, N. Martins <sup>a,b,c</sup>,  
M. Drummond <sup>a,b</sup>  
<sup>a</sup> Pulmonology Department, Centro Hospitalar  
 Universitário São João, Porto, Portugal  
<sup>b</sup> Faculty of Medicine, University of Porto, Portugal  
<sup>c</sup> Institute for Research and Innovation in Health,  
 University of Porto, Portugal
- \* Corresponding author.  
 E-mail address: [mariana.serino@gmail.com](mailto:mariana.serino@gmail.com) (M. Serino).
- <https://doi.org/10.1016/j.pulmoe.2020.04.007>  
 2531-0437/ © 2020 Sociedade Portuguesa de Pneumologia.  
 Published by Elsevier España, S.L.U. This is an open access article  
 under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

## Strengths of breath-triggered inhalers in asthma management



Dear Editor,

Asthma affects the lives of several hundred million people around the World, across all age groups and Portugal is not an exception, with an asthma lifetime prevalence of 10.5%.<sup>1</sup> Patients, their families and the society face high direct and indirect costs, due to healthcare resources use, loss of productivity, absenteeism and presenteeism of patients. Asthma strongly influences the wellbeing and quality of life of patients. With no established curative treatment, guidelines for asthma management have identified that the primary goal of management is to achieve asthma control.<sup>2</sup> In asthma, pharmacological treatments mainly consist of inhaled drugs, which allow efficacy and significantly reduce systemic side effects, being the most efficacious treatments. However, lack of adherence to treatment in asthma occurs in more than half of all medical prescriptions. Furthermore, reduced adherence to treatment frequently allies with an incorrect inhalation technique, both considered major issues significantly impairing pharmacologic treatments effectiveness. Therefore, despite the efficacy of the available drugs, a high percentage of asthmatics are uncontrolled and have frequent exacerbations.

Difficulties in using inhalation methods are well-known problems that have been consistently maintained in recent decades, with the occurrence of several errors that affect treatment results with the use of both dry powder inhalers (DPIs) and pressurized metered-dose inhalers (pMDIs). Results from the Critikal study<sup>3</sup> analyses of the inhaler

technique assessment initiative Helping Asthma in Real-life Patients (iHARP) database have helped to identify the prevalence of critical inhaler errors (those that have a definite detrimental impact on the delivery of drug to the lungs) with different devices in patients with asthma. The most common critical errors included failure to coordinate device actuation and inhalation with pMDIs, and lack of a forceful inhalation with DPIs; overall, 89% of patients made at least one potentially critical handling error and 67% made multiple potentially critical errors.<sup>3</sup>

Breath-actuated inhalers (BAIs) represented an evolution in inhalers' design aiming to improve the management of asthma.<sup>4</sup> BAIs are aerosol devices, like pMDIs, but rather than being activated manually, they automatically release a dose of drug when the patient inhales with a sufficient inhalation flow rate. In the past, BAIs had a major limitation when chlorofluorocarbon propellants (CFCs) were used and the speed of emission of the aerosol was so high that the impact of inhaled droplets in the upper airways was still very significant; nowadays, with hydrofluoroalkane propellants (HFAs), the speed of emission is considerably lower and the aerosol may be inhaled slowly being deposited more peripherally in the airways. These new devices may offer several advantages (Table 1), and were developed to overcome the most commonly seen critical errors with other inhalers, as follows: there is no need to coordinate actuation and inhalation (which is necessary for pMDIs); as active devices, BAIs emit a propelled aerosol and patients do not need to inhale forcibly to generate respirable particles (which is required for DPIs).<sup>4,5</sup>

BAIs are intended to simplify the inhaler technique, leading to improved inhaler use by the patients and less health