

RESEARCH ARTICLE

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Lung function in obese children and adolescents without respiratory disease: a systematic review

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Abstract

Background: Obesity in children and adolescents is associated with increased morbidity and mortality due to multisystemic impairment, including deleterious changes in lung function, which are poorly understood.

Objectives: To perform a systematic review to assess lung function in children and adolescents affected by obesity and to verify the presence of pulmonary changes due to obesity in individuals without previous or current respiratory diseases.

Methods: A systematic search was performed in the MEDLINE-PubMed (Medical Literature Analysis and Retrieval System Online), Embase (Excerpta Medica Database) and VHL (Virtual Health Library/Brazil) databases using the terms “Lung Function” and “Pediatric Obesity” and their corresponding synonyms in each database. A period of 10 years was considered, starting in February/2008. After the application of the filters, 33 articles were selected. Using the PICOS strategy, the following information was achieved: (Patient) children and adolescents; (Intervention/exposure) obesity; (Control) healthy children and adolescents; (Outcome) pulmonary function alterations; (Studies) randomized controlled trial, longitudinal studies (prospective and retrospective studies), cross-over studies and cross-sectional studies.

Results: Articles from 18 countries were included. Spirometry was the most widely used tool to assess lung function. There was high variability in lung function values, with a trend towards reduced lung function markers (FEV₁/FVC, FRC, ERV and RV) in obese children and adolescents.

Conclusion: Lung function, measured by several tools, shows numerous markers with contradictory alterations. Differences concerning the reported results of lung function do not allow us to reach a consensus on lung function changes in children and adolescents with obesity, highlighting the need for more publications on this topic with a standardized methodology.

Keywords: Lung function, Spirometry, Obesity in adolescence, Obesity in childhood, Review

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Authors summary

What is known?

(i) Obesity in children and adolescents is a risk factor to higher morbidity and mortality due to multisystemic impairment;

(ii) Obesity as a deleterious factor in lung function is poorly understood.

What is new?

(i) Spirometry was the most widely used tool to assess lung function in obesity and showed a high variability in its values, with a trend towards reduced lung function markers in children and adolescents with obesity;

(ii) Differences regarding the reported results of lung function do not allow us to reach a consensus on lung function changes in children and adolescents with obesity, highlighting the need for more publications on this topic with a standardized methodology.

Background

Obesity is a dysfunction that interferes with systems of the body and whose prevalence increases in epidemic proportion [1]. The comprehensive impact of obesity prompts researchers to reflect on its deleterious effects, which progressively worsen the quality of life of increasingly younger individuals, leading children and adolescents to suffer from impairments that had been previously observed in adults only [2, 3].

The respiratory system is one of the systems affected by obesity. Among adults, the most frequent findings in the comparison of lung function of healthy individuals versus individuals affected by obesity are the reduction in functional residual capacity (FRC) and expiratory reserve volume (ERV). One of the main reasons for these changes is the impairment of the respiratory mechanics. Excessive adipose tissue, mainly in the thorax and abdomen, causes an increase in the intra-abdominal pressure on the diaphragm and in the pressure of the fat tissue on the rib cage, hindering thoracic expansion and, consequently, lung compliance. This change leads to a reduction in lung volumes and capacities and is characteristic of a restrictive lung disease [4, 5].

Respiratory mechanics are not the only way of compromising lung function in obesity. There is also an inflammatory component that causes obstructive pulmonary disorders. Adipose tissue macrophages produce proinflammatory substances and adipocytes secrete hormones (adipokines), which reach the systemic circulation, and are able to act directly on the respiratory system or alter the immune response. The entire pathophysiological process favors the induction of bronchial hyperreactivity and may compromise pulmonary air flow [6–8].

The prevalence of obesity has increased worldwide, and although it is a relevant public health problem that

affects all age groups, the role and methods to evaluate its impact on lung function in children and adolescents is still unclear, and full understanding of the topic is still far from being attained.

As the obesity and lung function are complex phenotypes and their interaction has not been well understood, we included Fig. 1 that summarizes the mechanisms of lung function impairment due to obesity. Despite the relevance of the topic, it is still not well-established when lung function damages related to obesity start. Studies with children and adolescents diverge in their conclusions. Body changes during childhood and adolescence, variations in age range as well as in ethnic/environmental/genetic specificities make understanding of the cause-effect relationship between obesity and lung function more difficult.

Obesity and its association with lung function have been more often studied to ascertain the diagnosis of patients with asthma. Obesity and asthma have shown increasing prevalence in the last decades, and at the same time, they share common aspects, including the inflammatory process [8–10].

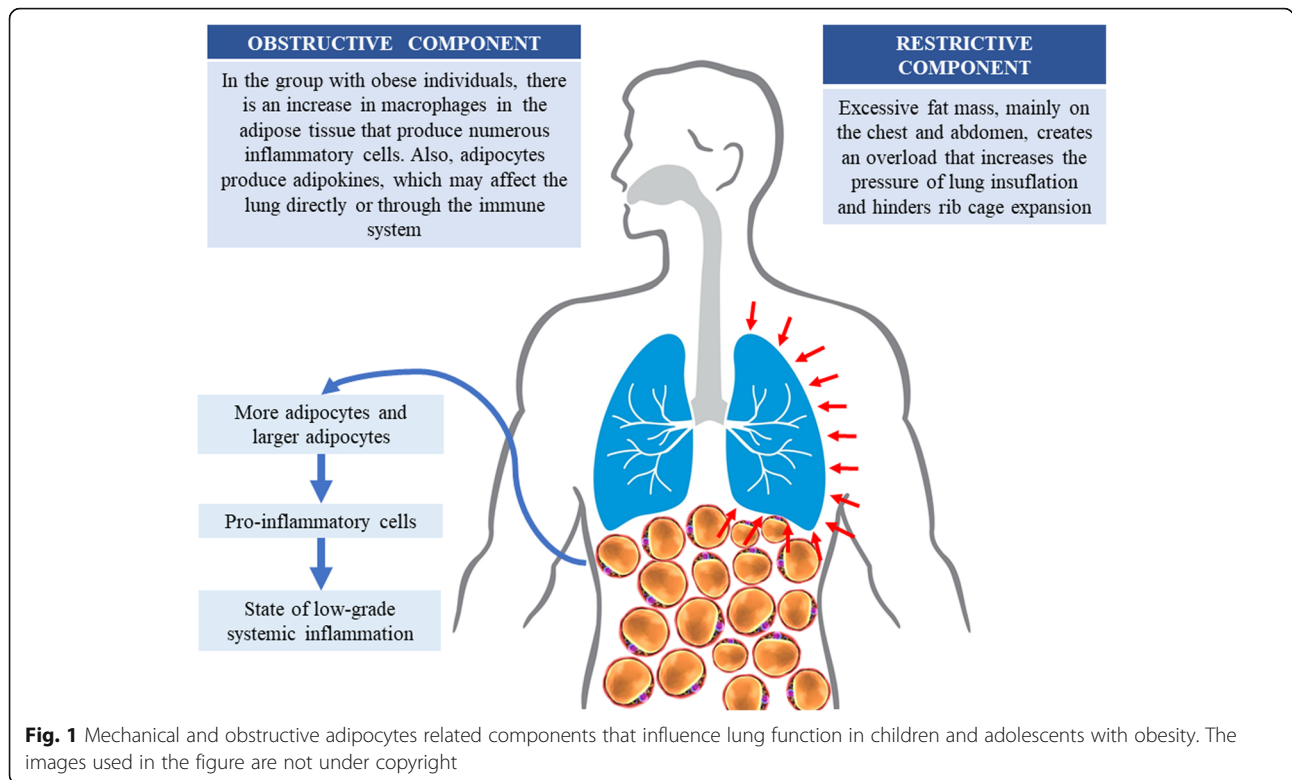
Obesity and asthma have been described as concomitant risk factors, with characteristics of a cause-effect relationship. The assessment of lung function in individuals with obesity and without asthma has yielded mixed results. Comprehensive studies are needed to understand whether mechanical and inflammatory changes are present in childhood obesity and during the growth process. Further studies should verify whether the disorders manifest differently during childhood and adolescence due to body changes throughout these periods, especially in individuals without a known lung disease, which is the focus of this systematic review [8–10].

This systematic review aimed to assess lung function in children and adolescents with obesity and to verify the presence of pulmonary restrictive or obstructive damages due to obesity in individuals without previous or current respiratory diseases, including asthma.

Methods

A systematic search was conducted in MEDLINE-PubMed (Medical Literature Analysis and Retrieval System Online, Public Medline), Embase (Excerpta Medica Database) and VHL (Virtual Health Library – Brazil) databases. Platform-specific tools were used, considering the descriptors (PubMed – MeSH Terms, VHL – DeCS and Embase – Emtree Terms) and equivalent terms, as well as excluding the descriptor “asthma” in order to achieve the objective of the study. The terms used in each database are described in Table 1.

The articles were selected in three stages, as detailed in Fig. 2. The titles were first read and the articles that were not relevant to the review were excluded (358, 1,



010 and 68 articles found in PubMed, Embase and VHL, respectively, were excluded). After the first database search filter was used, the articles were selected based on the abstracts (40, 35 and six articles found in PubMed, Embase and VHL, respectively, were excluded). In the third stage, the articles were read in full and then, carefully screened according to their relevance to the topic (10, 23 and seven articles found in PubMed, Embase and VHL, respectively, were excluded). Also, using the PICOS strategy, the following information was achieved: Patient – children and adolescents; Intervention (Exposure) – obesity; Control – healthy children and adolescents; Outcome – pulmonary function variability; Studies – randomized controlled trial, longitudinal studies (prospective and retrospective studies), cross-over studies and cross-sectional studies.

Eligibility criteria

In our study we included the following types of studies: randomized controlled trial, longitudinal studies (prospective and retrospective studies), cross-over studies and cross-sectional studies. Also, there was no restriction regarding the length of follow-up, and we considered only studies published between 2008 and 2018. We considered studies published in English, Spanish or Portuguese, and that were available in full text.

Information sources

All the studies were collected from the following databases: MEDLINE-PubMed, Embase and VHL databases. All the data was extracted directly from the studies and there was no contact with the study authors.

Search in the databases using descriptors

The following strategies were used to perform all searches in the study:

MEDLINE-PubMed.
 (((("Lung function"[Title/Abstract]) OR ((Respiratory Function Tests [MeSH Terms]) OR "Respiratory Function Tests"[Title/Abstract]) OR ((Spirometry [MeSH Terms]) OR Spirometry[Title/Abstract]))) AND (((Obesity[MeSH Terms]) OR Obesity[Title/Abstract]) OR ((Pediatric Obesity[MeSH Terms]) OR "Pediatric Obesity"[Title/Abstract]) OR (((Childhood Obesity[MeSH Terms]) OR "Childhood Obesity"[Title/Abstract]) OR Adolescent Overweight[-MeSH Terms]) OR "Adolescent Overweight"[Title/Abstract]) OR Adolescent Obesity[MeSH Terms]) OR "Adolescent Obesity"[Title/Abstract])) NOT ((Asthma[-MeSH Terms]) OR Asthma[Title/Abstract])

VHL
 ((tw:(Pediatric Obesity)) OR (tw:(Obesity, Pediatric)) OR (tw:(Childhood Obesity)) OR (tw:(Obesity, Childhood)) OR (tw:(Childhood Onset Obesity)) OR (tw:(Obesity, Childhood Onset)) OR (tw:(Child Obesity)) OR

Table 1 Descriptors used for the database search MEDLINE-PubMed (Medical Literature Analysis and Retrieval System Online-Public Medline), Embase (Excerpta Medica Database) and VHL (Virtual Health Library-Brazil)

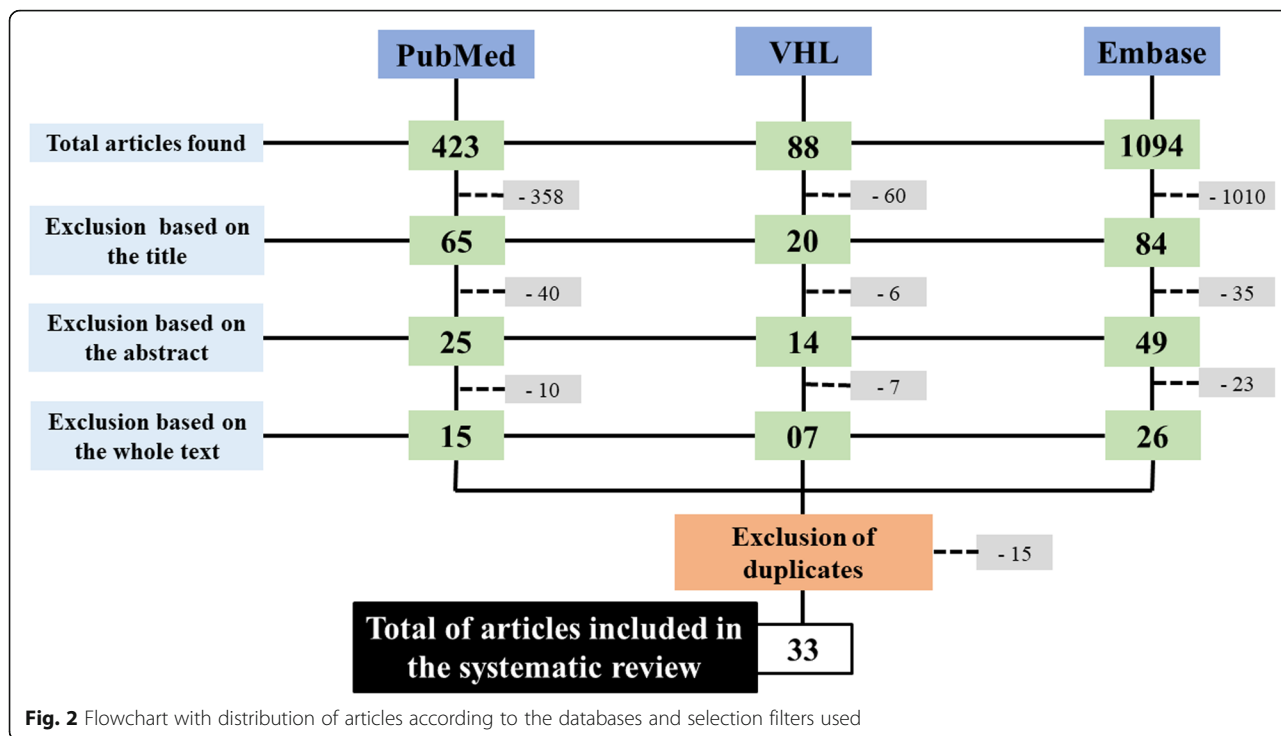
Term	PubMed (MeSH Terms)	VHL (DeCS)	Embase (Emtree Terms)
Pediatric obesity	Childhood obesity Adolescent obesity	Childhood obesity	Childhood obesity
		Obesity, pediatric	Child obesity
		Obesity, childhood	Obesity in adolescence
		Childhood onset obesity	
		Obesity, childhood onset	
		Child obesity	
		Obesity, child	
		Childhood overweight	
		Overweight, childhood	
		Obesity in childhood	
		Adolescent obesity	
		Obesity, adolescent	
		Obesity in adolescence	
		Adolescent overweight	
		Overweight, adolescent	
	Lung function	Respiratory function tests Spirometry	Function tests, pulmonary
Function test, pulmonary			Lung function test
		Pulmonary function test	Spirometry
		Test, pulmonary function	
		Tests, pulmonary function	
		Function test, lung	
		Function test, respiratory	
		Function tests, lung	
		Function tests, respiratory	
		Lung function test	
		Respiratory function test	
		Tests, respiratory function	
		Lung function tests	
		Pulmonary function tests	

(tw:(Obesity, Child)) OR (tw:(Childhood Overweight)) OR (tw:(Childhood Overweights)) OR (tw:(Overweight, Childhood)) OR (tw:(Obesity in Childhood)) OR (tw:(Infant Obesity)) OR (tw:(Obesity, Infant)) OR (tw:(Infant Overweight)) OR (tw:(Overweight, Infant)) OR (tw:(Infantile Obesity)) OR (tw:(Obesity, Infantile)) OR (tw:(Adolescent Obesity)) OR (tw:(Obesity, Adolescent)) OR (tw:(Obesity in Adolescence)) OR (tw:(Adolescent Overweight)) OR (tw:(Overweight, Adolescent)) AND ((tw:(Lung function)) OR (tw:(Function Tests, Pulmonary)) OR (tw:(Function Test, Pulmonary)) OR (tw:(Pulmonary Function Test)) OR (tw:(Test, Pulmonary Function)) OR (tw:(Tests, Pulmonary Function)) OR (tw:(Function Test, Lung)) OR (tw:(Function Test, Respiratory)) OR (tw:(Function Tests, Lung)) OR (tw:(Function Tests, Respiratory)) OR (tw:(Lung Function Test)) OR (tw:

(Respiratory Function Test)) OR (tw:(Test, Lung Function)) OR (tw:(Test, Respiratory Function)) OR (tw:(Tests, Lung Function)) OR (tw:(Tests, Respiratory Function)) OR (tw:(Lung Function Tests)) OR (tw:(Pulmonary Function Tests))) AND NOT (asthma)

Study selection

In brief, the study selection was carried out as represented in Fig. 2. Also, two authors (MFS and FALM or MFS and VLWW) decided about the eligibility before including the study in the review. In the presence of ambiguous conclusion, a third author (RTM or JDR) was contacted to perform a full consideration. Afterwards, a fourth author revised all the studies and the dataset to reach a final decision (RTM or JDR).



Data collection process

The data collection was carried out by two authors (MFS and FALM or MFS and VLWW), in this way, the data collection was performed twice for each study. Also, after the data extraction, the study was described as Table 2 and both authors included a summary using both datasets generated in the individual data collection. In the presence of ambiguous information, a third author (RTM or JDR) was contacted to perform a full consideration.

Results

As described in the methods, in brief, the articles were selected in three stages. A total of 48 articles were selected. After the exclusion of duplicates, 33 articles were included in the systematic review.

Table 2 shows a detailed informative and descriptive summary of the articles in this review: authorship, year of publication, place of study (country), study objective, presence or absence of respiratory disease, type of study, type of evaluation of lung function and type of posture and markers used in the analysis, main results and conclusions.

A wide age range (5 to 18 years) could be observed in the studies, with a higher prevalence between 11 and 13 years, which was an inclusion criterion in 69.7% (23/33) of the articles [11, 13–15, 17–22, 24–27, 29, 30, 32, 33, 35, 36, 38–40].

It is important to highlight the exclusion of studies that included participants with respiratory diseases, since

the focus was to evaluate the pulmonary changes resulting exclusively (or with the least possible influence of other factors) from obesity. Therefore, including children with previous respiratory diseases could evolve into sampling (selection of individuals), confounding (proven impact on outcomes) or information bias (previous knowledge of an existing problem). In this context, 66.7% (22/33) [12–15, 17–21, 25, 26, 29, 32–37, 39–42] of the studies excluded individuals with respiratory diseases, 27.3% (9/33) [16, 22–24, 27, 28, 30, 31, 38] did not mention respiratory diseases as a factor of exclusion or non-inclusion, and 6% (2/33) [11, 43] excluded only individuals with a history of smoking. Among the studies that excluded previous respiratory diseases, several exclusion criteria could be observed: some authors excluded only individuals with exacerbation of asthma or cough; others excluded any respiratory conditions that might impair the evaluation; and others used standardized instruments such as the ISAAC questionnaire (The International Study of Asthma and Allergies in Childhood). This demonstrates the variability of methods adopted by the authors of the different studies, low standardization of exclusion/inclusion criteria, and difficulties/limitations to evaluate, diagnose and exclude patients with possible respiratory diseases in some specific cases among the evaluated children and adolescents [44].

Additionally, in the studies analyzed, the inclusion of healthy controls (HC), without obesity, was described as a criterion to compare lung function. In this context, 78.8% (26/33) [11, 13, 14, 16–20, 24–38, 41–43] of the

Table 2 Descriptive analysis of the articles included in the systematic review

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Yao et al. [10] (2017) - China	To evaluate the effect of excess weight on lung function and FeNO in Asian children with a focus on changes in atopy	Smoking (Analysis was performed including and excluding individuals with asthma, but this was not an initial exclusion criterion)	Prospective cohort	1717 Asian children aged 5 to 18 years	Spirometry (Spiro Lab II, Medical International Research, Roma, Italy) FeNo (CLD 88 NO analyzer, Evo Medics, Duernten, Switzerland)	Spirometry, ATS - position not mentioned FeNO: ATS and ERS - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, PEF, FEF _{25-75%} , FeNO	There were associations (+) of z-score of BMI with FVC, FEV ₁ , PEF and FEF _{25-75%} and (-) with FEV ₁ /FVC and FeNO. The associations occurred for the variables in the analysis with the entire group and excluding individuals with asthma	Excess weight changes lung volume and flow in a disproportionate manner, which reflects in the increase in FVC, FEV ₁ , PEF and FEF _{25-75%} , and reduction in FEV ₁ /FVC
Peng et al. [11] (2016) - China	To evaluate whether weight index is associated with high BP, reduced FVC, dental caries and low vision, as well as whether nutritional status can predict diseases in schoolchildren	Yes (individuals with chronic or infectious diseases, e.g., cardiovascular, renal, hepatic, diarrhea, pneumonia, upper respiratory tract infection and influenza)	Cross-sectional	12,297 children aged 6 to 18 years	Spirometry (model not mentioned)	Standardized instrument not mentioned - standing	FVC	Group 6 to 12 years - males compared to females: higher weight, WC, BMI, SBP, FVC, low weight, overweight and obesity, FVC/weight; and lower SAH and low vision. Group 13 to 18 years - males: higher weight and height, WC, BMI, SBP, FVC/weight, prevalence of underweight, overweight, obesity, abdominal obesity and poor FVC/weight; and lower DBP, SAH, caries and low vision. Group of individuals with underweight had the lowest value of WC, SBP, FVC and visual acuity; and better FVC/weight. Group of individuals affected by obesity presented higher value of WC, SBP, DBP and FVC; and lower number of caries and FVC/weight. Children with overweight and obesity were at increased risk of high BP and poor FVC/weight when	Inadequate nutritional status was associated with BP, FVC, caries and visual acuity. The prevalence of common diseases in school-aged children is greater in children with altered weight. Thus, weight index is a potential marker to predict some diseases, reinforcing the importance of maintaining weight in the prevention of diseases in schoolchildren. However, the causal relationship and physiological mechanisms to explain the changes need to be further studied

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
LoMauro et al. [12] (2016) - Italy	To verify whether the thoracoabdominal volume of male adolescents with obesity during exercise has specific characteristics to deal with the increasing ventilatory demand and to investigate whether a short period of multidisciplinary program for weight loss, including respiratory muscle resistance training, can modify the geometry and volume of the rib cage	Not mentioned	Prospective and intervention	11 male adolescents (Tanner 3 to 5), with standard deviation of BMI > 2, in relation to the Italian standards	Spirometry (MedGraphics CPXD, Medical Graphics Corp., Saint Paul, Minn., USA) and OEP (Smart System BTS, Milan, Italy)	Spirometry: ERS – standing OEP: Standardized instrument and position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, PEF, total and compartmental volume in FRC and TLC, TV, RR, MV	<p>compared to children with normal weight, while undernourished children were at higher risk of caries, and in both groups, there was a higher risk of low vision</p> <p>FEV₁/FVC was greater than 80% predicted in the individuals, indicating the absence of OVD, but with a possible indication of a restrictive pattern. After the intervention, there was an improvement in the absolute and predicted values of FVC, and a reduction in IC of the lung and abdominal rib cage was observed</p>	<p>Hyperinsufflation of the abdominal rib cage occurs during incremental exercise from moderate intensity to peak intensity in order to recruit lung volume, being an adaptation of the ventilatory dynamics to deal with the overload of the chest wall due to obesity, optimizing the synergism between the diaphragm and the abdominal musculature. The system starts to function at high volume to optimize lung compliance. After training, there was reduction in abdominal load, pulmonary recruitment and thoracic cavity volume, improvement of physical performance, reduction in dyspnea and delay in dynamic hyperinsufflation of the abdominal thoracic cavity without ventilatory and metabolic</p>

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Faria et al. [17] (2014) - Brazil	To investigate lung function response during exercise in adolescents with non-morbid obesity and without respiratory diseases	Yes (history of acute or chronic respiratory diseases, thoracic or skeletal deformity, heart diseases and congenital diseases)	Cross-sectional	92 adolescents aged 10 to 17 years - 47 with obesity (23 males) and 45 HC (21 males)	Spirometry (GPFS/D - MedGraphics Saint Paul, Minnesota, USA), FMR (Gerar*)	Spirometry: ATS and ERS - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, IC, ERV, VC, MVV, MIP, MEP	Baseline BP and HR were higher among individuals with obesity, while SpO ₂ was lower. MVV, FVC and FEV ₁ were lower in males with obesity when compared to HC. IC in the group of females + obesity was higher than in the group of females + control. ERV was lower in both sexes among individuals with obesity when compared to controls. There were no differences in lung function before and after exercise. RMS showed differences between the sexes, but not between individuals with obesity and HC	
Davidson et al. [18] (2014) - Canada	To investigate the relationship between age, sex and BMI and lung volume of healthy	Yes (children with cardiorespiratory or ribcage diseases, asthma and individuals with	Retrospective	327 healthy individuals divided into 4 groups: underweight (pBMI < 5), normal weight	Spirometry, Body	Plethysmography [SensorMedics (Northridge, CA) Vmax 22 system with volume	Manuvacuometry: Standardized instrument not mentioned - sitting	FEV ₁ , FVC, FEF _{25-75%} , TLC, VC, FRC, ERV, RV, D _{lco}	ATS - position not mentioned

The distribution of body fat alters lung function in a sex-dependent manner among individuals with obesity and does not change after exertion

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
	individuals aged 6 to 17 years	reversible obstruction in spirometry)		(pBMI between 5 and 85), overweight (pBMI between 85 to 95) and individuals with obesity (pBMI ≥95)			measurement by the 6200 auto box and Vmax Legacy Plethysmography software (Viasys, Yorba Linda, CA)		
Individuals with obesity showed lower values in the predicted % of FRC and ERV. RV was lower in the groups of individuals with overweight and obesity. Individuals with low weight had lower FVC and RV. In the group of individuals with obesity, there was lower FEV ₁ /FVC. Additionally, there was a (+) linear association of the BMI z-score with the % of predicted of FVC, VC and DLCO and (-) linear association of the BMI z-score with the % predicted of FRC, ERV, RV and absolute value of FEV ₁ /FVC	Obesity was related to lower lung volume in children and adolescents. Changes in lung function may result in worsening respiratory symptoms and reduced functional capacity. Thus, there is a need to develop and implement effective strategies to prevent and manage obesity in childhood and adolescence		Cross-sectional and prospective	26 individuals with obesity and 25 HC aged 10 to 18 years	Spirometry (SensorMedics, Yorba Linda, CA, USA), multi-breath nitrogen wash-out (Vmax 29, hardware and software - SensorMedics)	Spirometry. ATS - position not mentioned	FVC, FEV ₁ , FEF _{25-75%} , FRC, TLC, RV	There was no difference between the groups for TLC, FVC, FEV ₁ and FEF _{25-75%} . However, the individuals affected by overweight and obesity presented lower z-scores of the FRC and RV. The expiratory flow during the submaximal exercise was associated	Young individuals with overweight and obesity may perform submaximal tests, and they tend to have a higher limitation of expiratory flow during submaximal exercise than healthy children. The use of compensatory
Gibson et al. [19] (2014) - Australia	To evaluate (i) whether children and adolescents with overweight or obesity can be submitted to submaximal exercise; (ii) respiratory limitations during exercise in children and adolescents with overweight and obesity	Yes (children with chronic cardiorespiratory problems or if it was not safe to exercise due to medical and/or musculoskeletal conditions)							

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
	compared to the control group							with the variables of weight (z-score BMI, weight and % of fat mass) and FEF _{25-75%}	breathing strategies allows overweight individuals to exercise at this intensity without feeling short of breath
Rio-Camacho et al. [20] (2013) - Spain	To investigate the left ventricular mass (echocardiography) of children with obesity, with and without metabolic syndrome; to evaluate the association between the level of adipokine and circulating cytokine and the alteration of left ventricular mass and Spirometry; and to determine the best variable to predict cardiovascular risk	Yes (Children with chronic diseases and/or Tanner = 5 (to avoid sexual dysmorphism of adipokines analyzed at this stage)	Cross-sectional and descriptive	41 individuals with obesity and over 8 years old (20 with metabolic syndrome criterion)	Spirometry (Frow Screen - Jaegger)	ATS - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, FEF _{25-75%}	MCP-1, LAR and CRP were higher in the presence of metabolic syndrome. There were no differences between the groups, with and without metabolic syndrome, for Spirometry and left ventricular mass	Obesity with metabolic syndrome has a higher degree of inflammation, and CRP is the best predictor of vascular risk. However, left ventricular mass and Spirometry were not influenced by the chronic inflammatory state in children and adolescents with obesity
Berntsen et al. [21] (2011) - Norway	To evaluate whether lung function measured in standing position is higher in children with overweight and obesity, when compared to the sitting position	The study does not mention respiratory conditions, only organic causes or diseases that may lead to obesity, may restrict the ability of being physically active and the use of medication that acts on growth or weight gain	Randomized and cross-over	115 individuals with overweight and 92 individuals with obesity aged 7 to 17 years	Spirometry (Vmax Series, SensorMedics, Yorba Linda, CA, USA)	ERS - sitting and standing	FVC, FEV ₁ , FEF _{50%} , FEV ₁ /FVC, PEF	15% of the patients had asthma. Females, when compared to males, showed higher value of FEV ₁ and FVC. FEV ₁ , FVC and FEF _{50%} were higher in Spirometry performed in the sitting position when compared to the evaluation in the standing position. In the linear regression analysis, the % of BMI, diagnosis of asthma, use of corticosteroids and sex were associated with changes in FVC, FEV ₁ , FEF _{50%} and PEF	FVC, FEV ₁ and FEF _{50%} were higher in the sitting position when compared to the standing position. However, the increase had little clinical significance. In this way, the sitting position is the most appropriate posture to perform forced expiratory flow-volume maneuver
Chen et al. [22] (2009) - Canada	To evaluate WC as a predictor of lung	Not mentioned (8 participants were	Cross-sectional	718 individuals aged 6 to 17 years	Spirometry (MedGraphics)	ATS - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC	WC had a (+) correlation with FVC	The association (-) of WC and FEV ₁ /

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Kalhoff et al. [23] (2011) - Germany	function markers and to compare it with BMI in children and adolescents	excluded because they presented reduced lung function value and low reproducibility)	Cross-sectional	518 preschoolers aged 6 years	System CPFS - Medical Graphics Corporation, St. Paul, MN, 1992)	ATs and ERS - position not mentioned	Airway resistance at 5 Hz and pulmonary reactance at 5 Hz	and FEV ₁ , and (-) correlation with FEV ₁ /FVC. On average, 1 cm increase in WC was associated with an increase of 7 mL in FVC and 4 mL in FEV ₁ . The assessment of height showed no changes in the association between WC and lung function. However, by adding body weight, the 1 cm increase in WC was associated with an increase of 4 mL in FVC and 2 mL in FEV ₁	FVC may be related to adiposity and/or lower predictability of FEV ₁ in relation to FEV ₁ /FVC in children. Thus, obesity is likely to be associated with reduced lung function in childhood
Kalhoff et al. [23] (2011) - Germany	To investigate whether overweight or obesity are associated with abnormalities in IOS in a random sample of pre-school children aged 6 years	Not mentioned	Cross-sectional	518 preschoolers aged 6 years	IOS (MasterScreen IOS - CareFusion, Höchberg, Germany)	ATs and ERS - position not mentioned	Airway resistance at 5 Hz and pulmonary reactance at 5 Hz	The study found no differences in resistance and reactance at 5 Hz in children with high BMI	In children aged 6 years, abnormalities in IOS were not associated with increased BMI. IOS requires little cooperation to have the test performed, unlike Spirometry. Therefore, this technique enables the analysis of pulmonary development with the age by measurements in series, from childhood to adolescence
Gundogdu et al. [24] (2011) - Turkey	To evaluate the effects of obesity on lung function and to define the relation of BMI as independent variable and PEF as dependent	Yes (children who had major dysfunctions - cardiac, respiratory, renal or hematological or those with asthma symptoms)	Cross-sectional	1439 children aged 6 to 14 years	PEF (Mini Wright Peak Flow)	GINA, 2005 - standing	PEF	Simple multiple linear regressions showed reduced PEF associated with an increase in BMI category. PEF was lower in the group of individuals affected by obesity when compared to	PEF was lower in children with obesity than in children with normal weight. As PEF is an indicator of pulmonary airflow resistance, there was an increase in

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Ferreira et al. [25] (2017) - Brazil	To evaluate lung function of children and adolescents with obesity (without asthma) by Spirometry and VC to compare them to HC of the same age group	Yes (Children with a history of respiratory diseases - asthma, obstructive sleep apnea or chronic obstructive pulmonary disease)	Cross-sectional	38 individuals with obesity and 39 HC aged 5 to 17 years	Spirometry (CPFS/D - Medical Graphics Corp., MN, USA and software Breeze PF 3.8 - Medical Graphics Corp., MN, USA) and VolC (CO2SMO - Dixtal, São Paulo, Brazil)	Spirometry: ATS and ERS - position not mentioned VolC: Standardized instrument not mentioned - sitting	FVC, FEV ₁ , FEV ₁ /FVC, FEV _{75%} , FEV _{75%} /FVC, ERV, MV, MV _{alv} , TV, TV _{alv} , DSV, DSV/TV, IC ETCO ₂ , VCO ₂ , RR, SpO ₂ , SIp ₂ , SIp ₃ , SIp ₂ /TV, SIp ₃ /TV	individuals without obesity	respiratory resistance in children with obesity. The association of high BMI with reduced PEF indicated that obesity is a risk factor for reduced airflow and lung function. Reduced prevalence of asthma may be a result of the patients' awareness and obesity prevention, i.e., prevention of obesity can reduce respiratory symptoms
								The lowest z-score of FEV ₁ /FVC, FEV _{75%} and FEV _{75%} /FVC occurred in the group of individuals with obesity, and 36.8% of individuals affected by obesity had FEV _{25-75%} lower than 70% (OVD by flow), and changes did not occur in the control group. In CV, the group of individuals with obesity showed lower DSV/TV and SIp ₃ /TV. In the linear regression, the BMI z-score influenced FVC, FEV ₁ /FVC, FEV _{75%} /FVC, FEV _{25-75%} /TV _{alv} , DSV/TV, VCO ₂ , SIp ₃ and SIp ₃ /TV. There was no response to BD among individuals with obesity. In the division by age group (5 to 11 years or > 11 years) there were changes of FEV ₁ /FVC	Even without the diagnosis of asthma by clinical criteria and without response to BD, individuals with obesity show lower FEV ₁ /FVC and FEV _{75%} /FVC, indicating an obstructive process. In CV, in the group with individuals with obesity, there was higher TV _{alv} with no alteration in ventilation homogeneity, suggesting that these individuals have altered flow, but no changes in lung volumes

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Del-Rio Navarro et al. [26] (2013) - Mexico	To compare bronchial hyperreactivity by the methacholine challenge testing in Mexican children with normal weight. In addition, to associate the group with normal weight with children with obesity or morbid obesity	Yes (underweight children, chronic respiratory diseases – including asthma and rhinitis, acute respiratory infection in the last month, endocrine diseases, dysmorphic exposure to tobacco)	Cross-sectional	229 children aged 10 to 18 years (40 – normal weight, 116 – with obesity and 73 – with morbidly obesity)	Spirometry (Vmax, Sensor Medics, Anaheim, CA), methacholine challenge testing (provococholine, 100 mg, Methaparm, Inc., Coral Springs, FL) performed with dosimeter (Mark Salter Labs, Arvin, CA)	Spirometry, ATS - sitting	FVC, FEV ₁ , FEF _{25-75%} , PEF	in both groups and only in the older individuals for expiratory flow and pulmonary volume In the group with obesity or morbid obesity, there was higher FVC and lower FEF _{25-75%} when compared to children with normal weight. Individuals with obesity, when compared with morbidly obese ones, had lower FEF _{25-75%} . PEF was higher among children with obesity when compared to children with normal weight or with morbid obesity. During the methacholine challenge testing, FEV ₁ was lower among children with obesity than in children with morbid obesity, starting from a dose of 0.25 mg/mL up to a dose of 16 mg/mL. In the comparison of group of individuals with normal weight with the group of individuals with obesity, there was higher value of FEV ₁ during methacholine challenge testing with 0.25 and 1 mg/mL of methacholine and lower with 4 and 16 mg/mL in the control group	Obesity did not change aerobic responsiveness due to the use of methacholine and studies should be performed to confirm the findings

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Spathopoulos et al. [27] (2009) - Greece	To evaluate the effect of obesity on lung function in a cohort of children aged 6 to 11 years and to associate obesity, atopy and asthma	Yes [high or low respiratory infection, exacerbation of asthma in the last 3 weeks, uncontrolled asthma (GINA), congenital heart abnormality, thoracic deformity or neuromuscular diseases]	Cohort	2715 children aged 6 to 11 years, (1978 – normal weight, 357 – with overweight and 300 – with obesity)	Spirometry (Vitalograph 2120)	AIS and ERS - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, FEF _{25-75%}	Among overweight individuals, FVC, FEV ₁ , FEF _{25-75%} and FEV ₁ /FVC levels were lower when compared to controls. Although the diagnosis of atopy and asthma is frequent in children with overweight and obesity, there was no difference in lung function in individuals with and without asthma. High BMI was an independent variable to predict reduction in Spirometry (mainly for FEF _{25-75%}) and a risk factor of asthma and atopy. When separated by sex, high BMI was associated with FVC in females and FEV ₁ /FVC in males	High BMI is a marker of obesity in children that can be easily measured and can determine the reduction in Spirometry measures, risk of atopy (both sexes) and asthma among females
Jeon et al. [28] (2009) – South Korea	To evaluate the factors that influence lung function in female adolescents, focusing on the hormonal factors of the menstrual cycle and obesity	Not mentioned	Cross-sectional	103 Korean high school children aged 15 to 18 years	Spirometry (Super Spiro, Micro Medical LTD, Kent, UK)	Standardized instrument and position not mentioned	FVC, FEV ₁ , FEF _{25-75%} , FEV ₁ /FVC	FEV ₁ /FVC was lower in females with obesity, when compared to HC of the same gender. The individuals who were evaluated in the menstrual period had lower FEV ₁ , FEV ₁ /FVC and FEF _{25-75%}	The literature is scarce on the study of asthma, lung function and puberty. In the study, there was a limitation of airflow associated with obesity, allergy, menstrual cycle and sensitization by inhaled allergens. Studies should be conducted to evaluate the relationship between gender hormones, leptin, lung function and asthma

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
He et al. [29] (2009) - China	To evaluate the relationship of obesity and asthma, asthma symptoms and lung function of Chinese schoolchildren using the definition of overweight and obesity of a Chinese group	Not mentioned	Cross-sectional	2179 children (1138 boys and 1041 girls) aged 8 to 13 years	Spirometry (Minato AS-505 portable electric spirometer - Minato Ltd., Tokyo, Japan)	ATS - sitting	FVC, FEV ₁ , FEF _{25-75%} , FEF _{75%} , FEF _{25-75%}	2% of the sample had asthma. Overweight children had higher FVC than in HC. Men with overweight and women with obesity had higher FEV ₁ than controls	Lung function was not altered by obesity; however, there was a higher prevalence of respiratory symptoms in individuals with overweight or obesity. Longitudinal studies need to assess the cause-effect relationship between overweight, obesity and lung function
Silva et al. [30] (2011) - Brazil	To assess the onset of EIB in children and adolescents, without asthma and overweight	Yes (acute and chronic lung diseases, cardiopathy, diabetes, musculoskeletal deformity and pain, steroid and non-steroid anti-inflammatory medication, symptoms of viral infection (cold or flu) in the last 6 weeks and FEV ₁ /FVC < 80%, FEV ₁ and PEF < 70% of predicted)	Cross-sectional	69 school children aged 8 to 15 years (39 children with obesity without asthma and 30 HC without respiratory diseases)	Spirometry (EasyOne model 2001 - Zurich, Switzerland) and PFE (Peak flow meter Healthscan® Personal Best)	ATS - position not mentioned	FEV ₁ , PEF, FVC, FEV ₁ /FVC, FEF _{25-75%}	The prevalence of EIB was 62% in the group of individuals with obesity and 16% in the control group. There was no difference in Spirometry between groups, except for PEF, which was lower in the group of individuals with obesity	PFE was important in EIB diagnosis. Possibly, different etiologies are related to EIB and studies of pathophysiology of the central and peripheral airways and the onset of EIB in children and adolescents with excess weight should be performed
Bekkers et al. [31] (2013) - The Netherlands	To associate WC and BMI with lung function in 8-year-old children	Not mentioned	Cohort	1,106 children aged 7.4 to 9.2 years	Spirometry (Jaeger)			pneumotachograph - Viasy Healthcare, San Diego, CA)	ATS and ERS - sitting
FVC, FEV ₁ , FEV ₁ /FVC	Children with lower or higher WC showed lower FVC and FEV ₁ than those with normal WC. Children with low or high BMI had lower FVC and FEV ₁ when compared to normal BMI.	In patients aged ~ 8 years, higher BMI or increased WC were not associated with FEV ₁ or FVC, demonstrating that this association may change over the course of childhood to							

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
	<p>were determined. However, IOS in the group of individuals with obesity presented higher value than in the control group for: absolute value of Z5, and absolute value and % of predicted of R5, Fres and AX. The group with individuals with overweight presented higher value than the control group for % of predicted value of R5, Fres and AX and for absolute Fres</p>								
Cibella et al. [33] (2015) - Italy	<p>To investigate the effects of weight on lung function of healthy children in a sample registered in 2 cross-sectional surveys with selected age group</p>	<p>Yes (history of wheezing, night cough or cough due to exercise)</p>	Cross-sectional	<p>2,393 Caucasian individuals aged 10 to 17 years (51.1% boys)</p>	<p>Spirometry (Microloop, Miro Medica, Chatham Maritime, Kent, UK)</p>	<p>ATS and ERS - position not mentioned</p>	<p>FVC, FEV₁, FEV₁/FVC, FEF_{25-75%}, FVC/FEF_{25-75%}</p>	<p>In the control of the variables weight, height, age and sex in the multiple linear regression, the weight corrected for height. However, B coefficient was (+) for FVC and FEV₁, being higher for FVC, and (-) for FEV₁/FVC and FEF_{25-75%}/FVC. In the division by age group (< 11, 12, 13 and > 14 years), there was a (+) association of the B weight coefficient with FVC and FEV₁, and a (-) association with FEV₁/FVC and FEF_{25-75%}/FVC (association of FEF_{25-75%} and B weight coefficient did not occur in the group < 11 years). Among individuals with obesity and overweight, the % of predicted for FVC and</p>	<p>FVC and FEV₁ were positively associated with weight, when corrected for height. However, due to a different magnitude in the effect of weight on FVC and FEV₁, FEV₁ showed a disproportionately smaller growth with weight gain when compared to FVC. Therefore, in individuals with a high BMI, there is a reduction of FEV₁/FVC and FEF_{25-75%}/FVC, and this change does not depend on respiratory symptoms</p>

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Silva et al. [34] (2015) - Brazil	To evaluate the effects of posture on thoracoabdominal kinematics of children with obesity and to compare them with a control group with normal weight	Yes (pulmonary or neuromuscular diseases)	Cross-sectional	35 children aged 8 to 12 years (18 with obesity and 17 with normal weight)	Spirometry (Micromedical Microloop MK8, Kent, England), RMS (digital manometer - MVD Globalmed 300, São Paulo, Brazil), OEP (OEP - BTS Bioengineering, Italy)	Spirometry, ATS and ERS - sitting; maximum respiratory pressure - sitting; OEP: Aliverti and Pedotti, 2003 - sitting and supine	FVC, FEV ₁ , FEV ₁ /FVC, MEP, MIP, TV variation, VTRCp, VTRCa, VTAB, VTRCp%, VTRCa%, VTAB%, Ti, Te, MV, RR, TV and θ	FEV ₁ was higher and the absolute value of FEV ₁ /FVC and FEF _{25-75%} /FVC was lower than in HC MIP, MV and TV were higher among individuals with obesity. The posture influenced TV (total and compartmental). There was higher TV, VTRCp and VTRCa in the sitting position, while VTAB was higher in supine position among children with obesity. TV was more influenced by the compartments VTRCp% and VTRCa% in the sitting position, while VTAB% was higher in the supine position. In addition, VTAB% was higher among individuals with obesity	The study demonstrated that the thoracoabdominal kinematics of children with obesity is influenced by the supine position, with an increase in abdominal contribution and reduction in the contribution of the rib cage to ventilation, suggesting that supine areas of pulmonary hypoventilation may occur. However, the thoracoabdominal kinematics was not different in the sitting position between the groups. Sitting posture is recommended during therapeutic procedures to achieve better distribution of regional rib cage volume and pulmonary ventilation
Turun et al. [35] (2014) - Turkey	To compare lung function in children with normal weight, overweight, obesity or morbid obesity and to	Yes (atopy or chronic lung diseases, asthma or family history of asthma, atopic dermatitis, food	Cross-sectional	170 individuals (30 - with overweight, 34 - with obesity, 64 - with morbid obesity and 42 - with normal	Spirometry (MIR, Spirolab III colour, Roma, Italy)	Standardized instrument and position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, FEF _{25-75%} , PEF	Overweight, obesity and morbid obesity showed lower FEF _{25-75%} and PEF, when compared to the group of individuals	The study considered FEV ₁ /FVC < 80% of predicted as OVD. Thus, despite the difference, the

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
	evaluate the effects of degree of obesity on lung function	intolerance or syndrome		weight) aged 9 to 17 years				with normal weight	study did not identify an obstructive abnormality in the group of individuals with obesity or morbid obesity individuals, when compared to controls with normal weight individuals and pointed out that longitudinal studies should investigate the effect of obesity degree and weight loss on lung function among individuals with obesity
Khan et al. [36] (2014) - Canada	To associate anthropometric measures and lung function in children	Not mentioned	Cross-sectional	1583 children aged 6 to 17 years (males: 573 – with normal weight, 216 – with obesity; females: 626 – with normal weight and 168 – with obesity)	Spirometry (Koko)	ATS and ERS - sitting	FVC, FEV ₁ , FEV ₁ /FVC, FEV _{0.75}	There was higher FVC, FEV _{0.75} and FEV ₁ in males than in females, and the opposite occurred in FEV ₁ /FVC. When the variable was adjusted according to the sex of the participants, there was association of BMI and WC with residual FVC in males and FVC and residual FEV ₁ in females. Both sexes had an inverse correlation of BMI with residual FEV ₁ /FVC. In the division by body mass, in the individuals with normal weight, there was a (+) effect of the BMI on FVC, FEV _{0.75} and FEV ₁ , and a (-) effect on FEV ₁ /FVC. WHR had a (+) correlation with FVC and FEV ₁ and a (-) correlation with FEV ₁ /FVC.	In males, there was worsening of lung function with overweight. Lung function was altered by abdominal and subcutaneous fat, and skinfolds were more sensitive to measure adiposity when compared to anthropometric data. The best indicator of adiposity in the analysis of lung function in males was the triceps skinfold

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Rosa et al. [37] (2014) - Brazil	To evaluate RMS by maximum respiratory pressure in healthy schoolchildren with overweight and obesity, and to identify whether the anthropometric and respiratory variables are related to the outcomes	Yes (SAAC)	Cross-sectional	90 school children aged 7 to 9 years (30 – with obesity, 30 – with overweight and 30 – HC	Spirometry (Piko-1, Spire Health, USA) and RMS (one-way valve digital		manovacuometer (MVD 300; G-MED, Brazil)	FVC. The WHR presented a (–) correlation with FEV ₁ /FVC. In children with overweight and obesity, there was a (–) association of WC and WHR with FVC and FEV ₁ . In this group, there was a (–) correlation of the skinfold of the triceps, biceps, iliac crest and medial calf with FVC, FEV _{0.75} and FEV ₁ , and the same was observed for the subscapular fold and the sum of all folds, adding the association with FEV ₁ /FVC. In HC, there was a correlation between: (i) triceps skinfold and FVC; (ii) iliac crest fold and FEV ₁ /FVC; (iii) sum of the 5 folds (triceps, biceps, subscapular, iliac crest and medial calf) and FEV ₁ /FVC	FEV ₁ , MIP, MEP
There was higher MIP in HC when compared to the others. The correlation of age	Obesity and overweight were associated with lower MIP when compared to HC.							Spirometry: ATS/ERS - sitting FMR: ATS - sitting	

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
with FEV ₁ , MIP and MEP was (+), and of MIP with BMI (-). MIP and MEP correlated with FEV ₁ , mainly, each other and, with less intensity, with the FEV ₁ . MEP had a (+) correlation with height and FEV ₁ . In the individual analysis of the groups, there was correlation of age with weight and height, except in the group of individuals with overweight; weight with height and BMI; MIP with age in the HC group, FEV ₁ in the group of individuals with obesity and MEP in the 3 groups; MEP with age and height in the HC group and FEV ₁ in the 3 groups; FEV ₁ with age in the 3 groups, and weight and height in the HC group and in the group of individuals with overweight	To describe pulmonary functional alterations in asymptomatic and overweight children and adolescents	Yes (history of wheezing, cough, chest pain, or known lung diseases)	Cross-sectional and descriptive	59 individuals aged 8 to 18 years (4 – with overweight, and 27 – with morbid obesity)	Spirometry (Koko Digidoser - Ferraris Respiratory, Louisville, CO, USA) and helium washout (mass flow sensor Vmax 21) (Masys Healthcare, Palm Springs,	Spirometry, ATS and Brazilian Society of Pulmonology - Phthysiology - position not mentioned helium washout - Standardized instrument and position not	FVC, FEV ₁ , RV, TLC, FEV ₁ /FVC; FEF _{25-75%}	30.3% of individuals had TLC < 80% of predicted and 3.5% TLC > 120%. In the sample, 25.5% of the individuals had a (+) response to BD in FEV ₁ , most of them with morbid obesity. Individuals with (+) response to BD had FEV ₁ /FVC < than LLN,	Asymptomatic respiratory individuals with excess weight had a high prevalence of ventilatory disorders, predominantly OVD. Additionally, there was a (+) response to the BD, higher than that

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Van de Griendt et al. [39] (2012) – The Netherlands	To evaluate the effects of weight reduction on lung function in children with morbid obesity aged 8 to 18 years	Yes (asthma or regular use of inhaled corticosteroids)	Longitudinal	112 children aged 8 to 18 and with BMI ≥ 30 Kg/m ² with comorbidities or BMI ≥ 35 Kg/m ²	CA, USA	mentioned		therefore, OVD. Regarding the use of BD and FVC, 2 individuals had a (+) response. Other findings were: 32.2% of individuals with OVD (15.2% - with overweight or obesity and 16.9% - morbid obesity) 25.4% with RVD (11.8% - with overweight or obesity and 13.5% - morbid obesity), and 6.7% with MVD (3.3% - with overweight or obesity and 3.3% - with morbid obesity). In addition, there was a (-) correlation between BMI with WC and FEV ₁ /FVC in the MVD group	reported in the literature, most frequently in morbid obesity
After 6 months of treatment to reduce weight, there was an increase of 3.08% in FuncVC, 2.91% in FEV ₁ , 2.27% in TLC and 14.8% in ERV. WC had a (-) correlation with ERV. Changes in BMI score correlated with ERV					Spirometry and Body		Plethysmography (MasterScreen PFT + body box - Jaeger Viasy, Wuertzburg, Germany)	Standardized instrument and position not mentioned	FEV ₁ , FEF _{50%} , ERV, FRC, TLC and FuncVC
Alghadir et al. [40] (2012) – Saudi Arabia	To investigate the relationship between severity of obesity and parameters of lung	Not mentioned (an interview and questionnaire about the medical history of lung	Cross-sectional	60 male individuals aged 6 to 13 years (20 in each group: with obesity, with overweight and		Spirometry (Pony FX - COSMED, Italy	FEV ₁ , FVC, FEV ₁ /FVC	The more fat a child has, the more compromised the lung function will be. Saudi children had	Lung function of male Saudi Arabians with obesity or overweight was

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Paralikar et al. [41] (2012) - India	function, comparing lung function in Saudi men with overweight and obesity with individuals with normal weight, and to compare the value found with the reference values for Caucasian Individuals	infection were conducted, but did not mention this factor as exclusion criterion)	Cross-sectional	with normal weight)	60 male individuals aged 12 to 17 years (30 – with obesity and 30 – HC)	Spirometry (MEDI; SPIRO - Maestros Medline Systems Ltd., Navi Mumbai, India)	FEV ₁ , FVC, FEV ₁ /FVC; PEF; FEV _{25-75%} ; MW	lower value than predicted for height and age. In the group of individuals with obesity and overweight, there was a lower (predicted) value for FVC and FEV ₁ and higher for FEV ₁ /FVC. Lung function of children with obesity was lower than that of the other groups, and the difference between the % of the measured value and the % of the predicted showed higher value in obesity for FVC and FEV ₁ than in HC or individuals with overweight	lower than that of children of the same age range in the HC group. The difference in relation to the predicted for their ages may indicate restriction in thoracic expansion and affect exercise capacity
	To evaluate lung function in adolescents with obesity in the city of Baroda, Gujarat	Yes (cough)				Spirometry, ATS and ERS - sitting		The mean % of predicted FEV ₁ was lower in the group of individuals affected by obesity, as well as the mean absolute value and % of predicted FEV ₁ /FVC and MW values. However, no individuals had OVD. Weight, BMI and WC had a (-) correlation with FEV ₁ /FVC, MW and FEV _{25-75%} and WHR with MW and FEV _{25-75%}	Lung function among individuals with obesity was lower than that of the HC, being obesity a health risk in the evaluated age group. Despite the difference between groups, no individual had OVD or RDV. Longitudinal studies are needed to understand the relationship between increased body weight and lung function

Table 2 Descriptive analysis of the articles included in the systematic review (Continued)

Authors (years) - country	Objective	Exclusion of respiratory diseases?	Type of study	Population	Lung function	Standardized instrument and position for assessment	Marker of lung function (Intervention)	Main results (Outcome)	Conclusions
Supriyatno et al. [42] (2010) - Indonesia	To determine the prevalence of abnormalities in lung function among Indonesian male adolescents and young people with obesity	Yes (Children with exacerbated asthma)	Cross-sectional	110 children with obesity aged 10 to 12 years	Spirometry (P57 Spirometer)	Spirometry; Polgar, 1971 - position not mentioned	FVC, FEV ₁ , FEV ₁ /FVC, FEF _{25%} , FEF _{50%}	In the sample, there was history of 29.1% asthma, 41.8% allergic rhinitis, 58.2% abnormality in lung function (30% MVD (obstructive and restrictive), 25.5% RVD and 2.7% OVD)	Abnormalities in lung function occur in obesity in early adolescence, with the most frequent change being MVD. There was no correlation between BMI and lung function. Studies are needed to assess the association of the degree of obesity and abnormalities in lung function with more accurate measures to assess body fat and with HC

FeNo Fraction of exhaled nitric oxide, ATS American Thoracic Society, ERS European Respiratory Society, FVC Forced vital capacity, FEV₁ Forced expiratory volume in the first second of forced vital capacity, FEV₁/FVC Relation between forced expiratory volume in the first second and forced vital capacity, PEF Peak expiratory flow measured by spirometry, FEF_{25-75%} Forced expiratory flow between 25 and 75% of forced vital capacity, BMI Body mass index (weight/height²), HC Healthy controls, BP Blood pressure, WC Waist circumference, SBP Systolic Blood Pressure, FVC/weight Forced vital capacity index by weight, Tanner Pubertal developmental stage according to Tanner's criteria, OEP Optoelectronic plethysmography, FRC Functional residual capacity, TLC Total lung capacity, TV Tidal volume, RR Respiratory rate, MV Minute volume, δ MWT Six-minute walk test, FEF_{25%} Forced expiratory flow at 25% of forced vital capacity, FEF_{50%} Forced expiratory flow at 50% of forced vital capacity, FEF_{75%} Forced expiratory flow at 75% of forced vital capacity, ERV Expiratory reserve volume, RV Residual volume, RV/TLC Ratio of residual volume and total lung capacity, IC Inspiratory capacity, HOMA-IR Homeostasis model assessment of insulin resistance, HDL High density lipoprotein, MMV Maximum voluntary ventilation, MIP Maximum inspiratory pressure, MEP Maximum expiratory pressure, WHR Waist hip ratio, HR Heart rate, SpO₂ Peripheral oxygen saturation, ρ BMI, BMI percentile, D_{LCO} Diffusing capacity of the lungs for carbon monoxide, MCP-1 Monocyte Chemoattractant Protein-1, LAR Leptin to adiponectin ratio, CRP C-reactive protein, IOS Impulse oscillometry, VolC Volumetric capnography, tot Total, alv Alveolar, DSV Dead space volume, DSV/TV Relation between Dead space volume and tidal volume, SIp₂ Slope of phase 2, SIp₃ Slope of phase 3, SIp₂/TV Relation between slope of phase 2 and tidal volume, SIp₃/TV Relation between slope of phase 3 and tidal volume, EtCO₂ End-tidal carbon dioxide, VCO₂ Volume of exhaled carbon dioxide, CI Capnography index [(SIp₂/SIp₃)x 100], Hz Hertz, EIB Exercise-induced bronchospasm, Z5 Respiratory impedance, R5 Total resistance, R20 Central airway resistance, X5 Reactance at 5 Hz, AX Reactance area, Fres Resonant frequency, VTRCp Tidal volume of the pulmonary rib cage, VTRCa Tidal volume of the abdominal rib cage, VTAB Tidal volume of the abdomen, VTRCp% Percentage of contribution of the tidal volume in the rib cage to total tidal volume, VTRCp% Percentage of contribution of the tidal volume in the abdominal rib cage to the total tidal volume, VTAB% Percentage of abdominal tidal volume contribution to tidal volume, T_e Expiratory time, θ Phase transition between 2 compartments, FuncVC Functional vital capacity, LLN Lower limit of normal, OVD Obstructive ventilatory disorder, RVD Restrictive ventilatory disorder, MVD Mixed ventilatory disorder, FEV_{0.75} Forced expiratory volume at 0.75 s, SAH Systemic arterial hypertension, DBP Diastolic blood pressure, BD Bronchodilator, PFE Peak expiratory flow measured by Peak Flow Meter, GINA Global Initiative for Asthma

studies compared individuals affected by obesity with HC.

The articles included were produced in 18 countries, with a predominance of European (8) [21–23, 31, 34, 35, 40], South American (8) [17, 18, 26, 32, 33, 37, 39, 41] and Asian (7) [11, 13, 15, 27, 30, 31, 42] countries. Also, 4 studies from North America were included [19, 24, 38, 43], as well as one from Central America [29], one from Oceania [20] and four from intercontinental countries (three Euroasians [14, 25, 36] and one from Asia and Oceania [12]).

In the evaluation of lifestyle habits, 15.2% (5/33) [20, 24, 30, 41, 42] of the studies assessed the participation of individuals in physical activities. Also, 9.1% (3/33) [16, 20, 42] of the studies assessed screen time of the participants and one [24] study mentioned the use of a lifestyle habits questionnaire but did not detail the assessed variables.

There was no uniform definition of obesity among studies: 27.3% (9/33) [11, 14, 15, 20, 35–37, 40, 42] used references by Cole et al. [45–47]; 21.2% (7/33) [12, 17, 19, 26, 29, 38, 43] used the criteria established by the Center for Disease Control and Prevention (CDC); 15.1% (5/33) [13, 18, 33, 39, 41] used the definition of the World Health Organization (WHO); 12.1% (4/33) [16, 24, 32, 34] did not mention any references for the definition of obesity; and 27.3% (9/33) [14, 21–23, 25, 27, 28, 31, 36] used references according to their countries of origin. Two [14, 36] of the studies mentioned above, used references by Cole et al. [45–47] in addition to references according to their countries of origin.

Inflammatory markers were assessed and correlated with lung function in just 6% (2/33) [11, 21] of the studies, which is a quite low percentage, considering the systemic and complex nature of obesity, which requires an interdisciplinary approach. Among the studies that assessed inflammatory process, one study evaluated fraction of exhaled nitric oxide (FeNO) [11] and another serum levels of C-reactive protein (CRP), adiponectin, leptin, interleukin 6 (IL-6), tumor necrosis factor (TNF- α), monocyte chemoattractant protein-1 (MCP-1), visfatin and retinol binding protein 4 [21].

Spirometry was the most commonly used tool to assess lung function, i.e., in 93.9% (31/33) [11–24, 26, 27, 29–43] of the studies. Next, body plethysmography and measurement of the respiratory muscle strength (RMS) were the most used tools in 12.1% (4/33) [15, 19, 40, 43] and 9.1% (3/33) [18, 37, 41] of the studies, respectively. Optoelectronic plethysmography (OEP) [23, 37], impulse oscillometry (IOS) [28, 33] and peak expiratory flow measured by peak flow meter (PFE) were included in the analyses of 6.1% (2/33) [25, 32] of the studies. Other tools were used in only one study each: nitrogen [20] and helium [39] washout, FeNO [11], volumetric capnography (VolC) [26] and methacholine challenge testing [29].

Among the spirometry variables, forced expiratory volume in the first second (FEV₁) of the forced vital capacity (FVC) was the most prevalent marker in the studies, included in 90.9% (30/33) [11, 12, 14–24, 26, 27, 29–43] of the analyses, followed by FVC and the FEV₁/FVC ratio, used in 87.9% (29/33) [11–24, 26, 27, 29–39, 42, 43] and 72.7% (24/33) [11, 12, 14–18, 21–24, 26, 27, 31–39, 42, 43] of the studies, respectively. Forced expiratory flow between 25 and 75% of FVC (FEF_{25–75%}) was also a widely assessed marker, being analyzed in 57.6% (19/33) [11, 14, 15, 17–21, 29–36, 39, 42, 43] of the studies. In addition, PEF and PFE were included in 33.3% (11/33) [11, 14, 17, 22, 23, 25, 29, 32, 36, 42] of the studies.

Among the variables analyzed using other tools besides spirometry, total lung capacity (TLC) and FRC could be observed in 21.2% (7/33) [15, 19, 20, 23, 39, 40, 43] and 18.2% (6/33) [15, 19, 20, 23, 40, 43] of the studies, respectively.

The comparison of the studies with a control group that showed comparative values or significant associations is described in Table 3. In this context, a great number of studies for each variable could be observed due to the selection of the various markers evaluated. No clear pattern emerged, as regards FEV₁ and FVC, with about half of the studies reporting no association between obesity and lung function parameters [11, 14, 16–20, 26, 27, 29–38, 41–43]. A clearer pattern emerged as regards FEV₁/FVC and FEF_{25–75%}, as most studies found either a negative association or no association between obesity and lung function [11, 14, 16–20, 26, 27, 29–38, 41–43], with only one study reporting a positive association (Table 3) [13].

Discussion

Effects of growth and development on lung function

Childhood and adolescence are characterized by major changes in the structure and functions of the human body systems. The physiological processes that influence lung function in 6-year-old children are different from those influencing 15-year-old adolescents, even if we disregard other multiple factors, such as gender, ethnicity, environment and genetics. Changes in lung function of adults with obesity are related to increased intra-abdominal pressure due to the deposition of fat in this region, which compromises the efficiency of diaphragmatic mobility, as well as the deposition of fat on the rib cage, which reduces its compliance. However, such aspects are insufficient to understand the influence of obesity on lung function of children and adolescents [48, 49].

Figure 3 lists some factors that influence lung function in children and adolescents and that should be considered in the discussion about the variability of the findings in this systematic review. The first factor is the increase in lung volume and surface for gas exchange

Table 3 Descriptive analysis of the markers evaluated in lung function in individuals aged 5 to 18 years

Variable	Association			Total
	+	-	No difference	
FVC	8	5	10	23
FEV ₁	4	6	13	23
FEV ₁ /FVC	1	10	7	18
FEF _{25-75%}	1	7	8	16
Peak expiratory flow measured by spirometry	2	4	3	9
Expiratory reserve volume	-	3	2	5
Functional residual capacity	-	3	-	3
Total lung capacity	-	-	3	3
Residual volume	-	3	-	3
Inspiratory capacity	2	-	-	2

+, studies that showed comparative markers with greater value in obesity or positive associations with variables that are indicative of obesity; -, studies that showed comparative markers with lower value in obesity or with negative association with variables that are indicative of obesity; FVC, forced vital capacity; FEV₁, forced expiratory volume in the first second of forced vital capacity; FEV₁/FVC, forced expiratory volume in the first second and forced vital capacity ratio; FEF_{25-75%}, forced expiratory flow between 25 and 75% of forced vital capacity

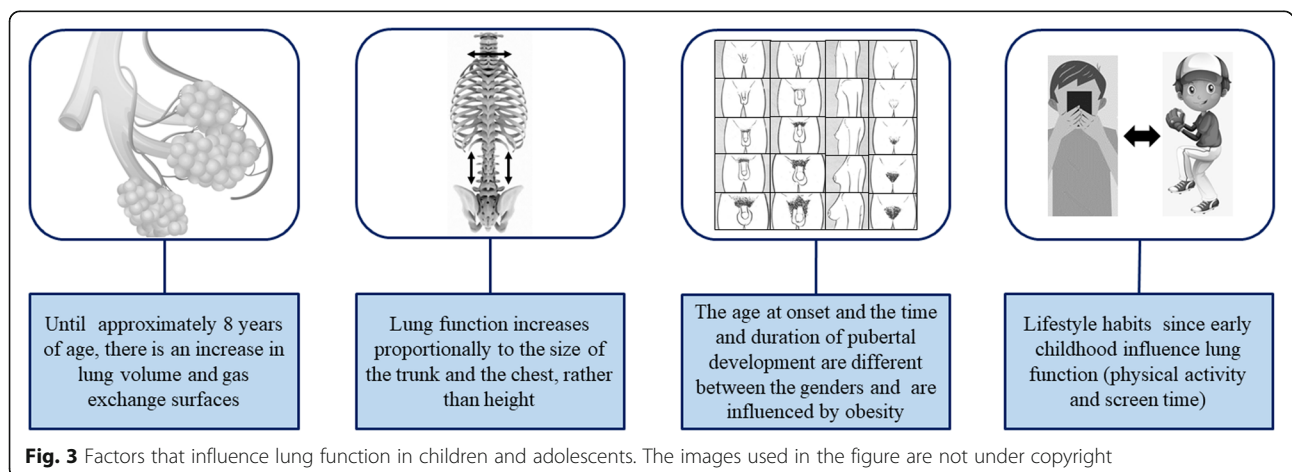
until approximately eight years of age. In this context, 45.5% (14/33) [11, 13, 16, 17, 22, 24–28, 33, 34, 38, 41] of the studies in this review included individuals under 8 years old – a period characterized by airway growth and development. Among these studies, 13 [11, 13, 16, 17, 22, 24–27, 33, 34, 38, 41] included, at the same time, individuals over eight years of age – a period, in which the respiratory tract is anatomically formed.

According to the literature, until the end of the pre-school age, the respiratory tract growth follows a dysnaptic pattern, i.e., the growth of the airways is slower than the growth of the lung parenchyma. After this period, the growth is isometric, showing greater homogeneity. This pattern of growth up to early childhood may lead to increased airway resistance and higher risk of obstructive processes in these individuals, especially in males, who have proportionally smaller airways than females during this maturation period. Thus, the findings regarding lung function in children and adolescents

should be analyzed considering the lung growth phase and the child’s development [49, 50].

The second and third factors described in Fig. 3 are inter-related and associated with changes arising from growth. The onset of puberty marks the beginning of a process of maturation, characterized by body and psychological changes. The changes vary according to gender, and in women, the pubertal development begins approximately two years earlier than in men. Moreover, hormonal changes may directly influence lung function: at first, by growth spurts, followed by increase in trunk height and ribcage diameter, which influence the increase in lung capacity and volumes. Another example is the increase in the production of male testosterone during puberty, which triggers a muscle growth peak, which includes the respiratory muscles and favors the increase of FVC and respiratory flows [51, 52].

Among individuals affected by obesity, the changes mentioned in the previous paragraph tend to occur at an earlier stage. Although the mechanisms are not yet well-



established, studies have shown a relationship between insulin resistance and increased serum levels of leptin. Thus, comparing individuals of the same age group, at different stages of pubertal development and different genders, may be a bias in the analysis of lung function. Of the studies included in the systematic review, 21.2% (7/33) [14, 16, 21, 23, 32, 38, 43] analyzed pubertal development [53, 54].

The last item described in Fig. 3 is fundamental to understand lung function in children and adolescents and is associated with physical activities and sedentary lifestyle habits. In this context, we should emphasize that the comprehensive knowledge of the studied sample is of utmost importance. Today, the majority of the population is sedentary, and sedentary behaviors are not exclusively associated to the group of individuals affected by obesity. So, if the sample is composed of individuals with obesity under treatment/follow-up and HC with sedentary lifestyle habits, there may be a bias in relation to cardiorespiratory conditioning, which may affect lung function. Therefore, in the analysis of lung function in children and adolescents with obesity, information about physical activities and screen time is fundamental [55, 56].

Interestingly, despite the importance of this data, only 15.1% (5/33) [20, 24, 30, 41, 42] of the studies in this systematic review assessed participation in physical activities, 9.1% (3/33) [16, 20, 42] included the assessment of screen time, and 1 [24] study mentioned lifestyle habits without detailing the assessment items used.

Measurement tools to define obesity

Body mass index (BMI) was the most commonly used tool to determine obesity in children and adolescents. However, the criteria to define obesity varied among the studies. The option of using country-specific standards of normality or those published by the CDC or WHO, determined the lack of homogenization of the samples. Also, the cut-off points for the definition of obesity within the selected references were different. Therefore, the use of comparable criteria between the studies would allow a more precise definition regarding the presence of obesity in children and adolescents. Comprehensive references, including data collection at a global level, are best indicated as they analyze normality patterns, taking ethnic differences into account.

The amounts of body fat and lean mass make the assessment of lung function in individuals with obesity more reliable. This can be explained because BMI, which is the most commonly used indicative of obesity, shows some limitations. BMI measures excess weight rather than excess fat and the variability due to gender, age, ethnicity and lifestyle habits may act as modifiers. Thus, well-trained individuals with high lean mass indexes are classified with obesity [57, 58].

In this systematic review, only 12.1% (4/33) [15, 18, 23, 38] of the studies used instruments that allowed the quantification of body fat. The first study used bioimpedance (BIA) [23], which estimates fat mass, fat-free mass and total body water. The second study [18] used BIA and the skinfold measurement test, which estimates the amounts of fat in each segment. However, the literature reports that this method has its limitations for the assessment of individuals with obesity [59]. The third study [38] only analyzed the skinfolds, and the fourth [15] used BIA and dual energy X-ray absorptiometry (DXA), which is a more accurate tool, as DXA assesses the amount and distribution of fat and lean body mass. The analysis of the distribution of body fat is an important tool to detect alterations caused by impaired respiratory mechanics, which is greater with the increase in fat in the thorax and abdomen.

Obesity epidemic reflected on the diversity of the studied populations

The inclusion of studies with individuals from almost all continents, except for Africa, is relevant for the analysis of the findings. They reflect a global epidemic of obesity among children and adolescents. According to the WHO, the prevalence of overweight and obesity among children aged 5 to 19 years increased from 4% in 1975 to 18% in 2016. Currently, more than 124 million children and adolescents are affected by excess of weight. Weight gain trends include both developed and underdeveloped countries, and currently, overweight and obesity are more prevalent and more often associated with causes of death than underweight, except in some parts of Africa (especially sub-Saharan Africa) and Asia [60, 61].

Trends related to lung function in children and adolescents with obesity

The variability of the results showed an inability to establish lung function changes in children and adolescents with obesity. However, some trends have been detected and are discussed as follows.

Among the variables analyzed, the comparison of FEV₁/FVC between individuals with obesity and HC was observed in 54.5% (18/33) [11, 12, 14, 16–18, 26, 27, 31–38, 42, 43] of the studies and of those, 55.6% (10/18) [11, 12, 16, 17, 26, 31, 34, 35, 38, 42] found lower value or negative association of the variable with obesity. The described changes may be an indication of the obstructive disorder in individuals affected by obesity during childhood and adolescence. The obstruction is related to pro-inflammatory activity of the adipose tissue, which could trigger bronchial hyperreactivity. Prior to our study, a review described similar findings regarding FEV₁/FVC [62].

The analysis of inflammatory markers to identify systemic impacts caused by obesity was reported in only 6%

(2/33) [11, 21] of the studies. One study [21] did not make any references to comparisons with the HC group. But, in another study [11], a positive association of FeNO with BMI was determined, suggesting that inflammation was more often detected in individuals affected by obesity and that these inflammatory changes should be considered in the clinical evaluation.

Despite the changes in FEV₁/FVC, in the 22 [11, 14, 16–20, 26, 27, 29–38, 41–43] studies that analyzed FEV₁ in children and adolescents with obesity and HC, there were discrepancies in the findings to confirm the presence of obstructive ventilatory disorder: (i) 56.5% (13/23) [17, 19, 20, 26, 29, 31–34, 36, 37, 41, 43] found no differences or associations between the groups; (ii) 26.1% (6/23) [14, 16, 18, 27, 38, 42] found lower value or a negative association between FEV₁ and obesity; (iii) 17.4% (4/23) [11, 24, 30, 35] found higher value or a positive association in individuals with obesity. Thus, FEV₁ was not associated with lung function impairment in overweight individuals.

FEF_{25–75%} should also be considered, as it is a marker of obstructive ventilatory disorder. Some studies indicate that this tool is more sensitive than FEV₁, and it can detect early ventilatory changes, especially in the small airways [63, 64]. However, as with most variables, there was also variability in the results. In total, 48.5% (16/33) [11, 14, 17, 19, 20, 26, 29–36, 42, 43] of the studies analyzed FEF_{25–75%} of individuals with obesity and HC and of those: (i) 43.8% (7/16) [19, 20, 30–33, 43] found no differences and associations between indicators of obesity and FEF_{25–75%}; (ii) 46.7% (8/16) [14, 17, 26, 29, 34–36, 42] found lower value or a negative association of FEF_{25–75%} with obesity; (iii) one (6.6%) [13] found a positive association between FEF_{25–75%} and obesity in children and adolescents.

In short, FEV₁/FVC was the spirometry marker with the greatest sensitivity to identify a possible obstructive process due to obesity in children and adolescents. However, the variability of this marker – and, even more of other indications of obstruction – was high, regardless of the study. Thus, the development of cohort studies on indicatives of growth stages and body development is of utmost importance, since these factors are different between individuals affected by obesity and HC and also influence lung function. Pubertal development or age cohorts considering the time of dysanaptic growth and isometric growth would considerably reduce confounding factors and allow better understanding of lung function changes due to obesity in children and adolescents.

In the evaluation of FVC, which indicates restrictive respiratory disorder, 66.7% (23/33) [11, 13, 14, 16–20, 24, 26, 27, 29–38, 42, 43] of the studies included comparison with HC and 43.5% (10/23) [14, 17, 20, 31–33, 36, 37, 42, 43] of them did not find any differences

between individuals with or without obesity. Only 21.7% (5/23) [16, 18, 27, 34, 38] found a negative association or lower values of FVC in children and adolescents with obesity. In a systematic review conducted in 2012, the authors concluded that the literature references demonstrated an association between reduced FVC and FEV₁ with obesity in children and adolescents, in disagreement with our findings [65].

Besides the spirometry variables, some other measures contributed to the analysis of lung function in children and adolescents with obesity. TLC, FRC and residual volume (RV) were markers used in only 9.1% (3/33) [19, 20, 43] of the studies comparing individuals with obesity and HC. All these studies reported that individuals affected by obesity showed lower value or a negative association of FRC and RV with obesity. However, there were no differences in relation to TLC. If TLC – which is the sum of the inspiratory capacity (IC), and FRC (FRC = RV + ERV) – does not present a difference between individuals with obesity and HC, and if FRC and RV are reduced, it can be assumed that IC should be higher in individuals affected by obesity. Only 2 [18, 43] studies analyzed this variable and found a higher value or a positive association of IC with obesity.

These results are in agreement with a systematic review published in 2016, which evaluated the effects of obesity on lung volume and capacity in children and adolescents, and found a reduction in some markers in obesity, especially the reduction in FRC, ERV and RV [66].

Issues on respiratory physiology and biomechanics requiring further investigation

The values for respiratory mechanics of individuals with obesity are incoherent. Some hypothesis and questions can be raised, namely:

- (i). Are individuals with obesity actually “stronger” and are, therefore, able to inspire more air?
- (ii). However, if there is an increase in RMS, considering the 9 [11, 14, 17, 25, 29, 32, 33, 36, 42] studies that analyzed peak expiratory flow measured by spirometry (individuals with obesity x HC), why did only 2 [11, 29] studies find a positive association with obesity? Why did 4 [17, 25, 32, 36] studies find lower value or a negative association between obesity and peak expiratory flow measured by spirometry? And, why did 3 [14, 33, 42] studies find no differences between groups?
- (iii). Are lung function changes in individuals with obesity due to the differences between males and females, since in females, there is a pattern of gynoid obesity, with higher fat concentration on the hip and legs; whereas in males, an android pattern

occurs, with more volume of fat in the chest and abdomen?

- (iv) It is well-known that individuals affected by obesity tend to initiate pubertal development earlier than healthy individuals. So, is there greater muscle development among individuals with obesity, which tends to be balanced at the end of puberty? Can impairment of lung function in individuals with obesity be clearly observed after puberty?

Confounding biases to clarify the mechanisms that interfere with lung functions in children and adolescents affected by obesity

Obesity is a multisystemic dysfunction, and therefore it is difficult to control the variables in order to understand the damage caused to lung function. For this reason, we found high variability in the results. Given the studies included in this systematic review, we are not able to establish which ventilatory changes are due to obesity in children and adolescents, even excluding data whose focus was the influence of asthma on obesity, which is a bias in this analysis. There are mechanisms that correlate both dysfunctions and the causal relationship between them may hinder perception of what is actually a consequence of obesity and/or asthma [8–10].

The findings of this review, although inconclusive, may give us a direction for future research. The strategies include: greater sampling control; reduction of confounding variables; conducting interdisciplinary and longitudinal studies with individuals with obesity versus HC; detailed analysis of environmental and social aspects; validation of findings among different populations; larger sample size; inclusion of measurements of lean mass and fat mass in order to unify and establish better criteria to define obesity; and future studies aiming to associate different genetic aspects, with predisposition to variability for weight gain, as well as for the individual nuance of lung function.

Thus, the inclusion and analysis of lung function in children and adolescents have become fundamental. Pubertal staging should be considered in order to avoid the influence of early maturation of individuals affected by obesity on the overestimation of lung capacity. It is important to analyze fat distribution, considering the concentration of abdominal and thoracic fat as factors that directly influence lung function. For this analysis, the use of instruments, such as DXA, may help determine the influence of the distribution and amount of body fat on lung function.

Assessing RMS with manuvacuometry or the distribution of lean mass with DXA, or even amount of lean mass using BIA, also favours the understanding of the physical conditions that influence lung function in children and adolescents. In order to analyze physical

conditions, it is also important to include the evaluation of programmed and non-programmed physical activities as well as the screen time.

It is essential to exclude previous respiratory conditions that may influence lung function and control variables that are indicative of inflammation, including CRP, erythrocyte sedimentation rate, FeNO or serum levels of leptin, adiponectin, IL-6 and TNF- α , which will allow precise determination of the influence of overweight in lung function of children and adolescents with obesity.

Meta-analysis

Meta-analysis is the gold standard in order to interpret a specific topic such as the importance of lung function in cases of obesity in the pediatric population. However, as described in our data there is no standardization in the studies about lung function in children and adolescents with obesity. To perform a meta-analysis, a minimum of standardization should be applied in the data acquisition. However, looking for the data included in Table 2, the studies were performed using different methodologies and/or lung function tools and/or lung functions measures (markers). Moreover, the age range was not equal, and the objectives were different among the studies. In brief, articles [11, 13–17, 19, 20, 22, 25, 26, 28–30, 32–38, 40–42] were described as cross-sectional studies on children-adolescents without respiratory diseases, where spirometry markers (such as FVC, FEV₁ and FEV₁/FVC) were assessed, and the relation between lung function and body mass, expressed as either BMI or norm weight/overweight/obesity, was estimated. Those studies did not allow us to perform a meta-analysis because there is a disparity of study objectives, population type (age range, sex distribution), origin of the population, presence of other lung function measurement, obesity as an independent variable and the exercise analysis. The information about the disparities among the studies is shown in Table 2 and Table 4.

Conclusion

The different results observed for lung function in children and adolescents with obesity show that there is no consensus on the impairment in such individuals in the literature. Considering the influence of growth and development on the function of all systems, it is fundamental to control the variables to reduce sampling, information and confounding biases, as well as to enable the analysis of the deleterious effects of obesity. In this context, new studies should require greater control of variables that influence growth and development to better understand the influence of obesity on lung function of children and adolescents.

Table 4 Disparities between the markers of cross-sectional studies on children-adolescents without respiratory diseases, where spirometry markers were assessed, and the relation between lung function and body mass, expressed as either body mass index or norm weight/overweight/obesity preventing the performance of meta-analysis

Authors (year)	Objective	Spirometry measurement					Other lung function measurement			Obesity as an independent variable	Exercise
		FVC	FEV ₁	FEV ₁ /FVC	FEF _{25-75%}	PEF	ERV				
		Yes	No	Yes	No	Yes	No	Yes	No		
Peng et al. [11] (2016) - China	To evaluate whether weight index is associated with high blood pressure, reduced FVC, dental caries and low vision, as well as whether nutritional status can predict diseases in schoolchildren	Yes	No	No	No	No	No	-	No	No	
Özgen et al. [13] (2015) - Turkey	To evaluate the relation between lung function tests and functional capacity during exercise in children with obesity	Yes	Yes	Yes	Yes	Yes	No	-	Yes	Yes	
Kongkiattkul et al. [14] (2015) - Thailand	To evaluate the correlation between obesity indexes (anthropometry and bioimpedance) and lung function parameters and to identify whether the indexes correlate with abnormalities in lung function of children and adolescents with obesity	Yes	Yes	Yes	Yes	No	No	+ Body Plethysmography	Yes	No	
Ferreira et al. [15] (2014) - Brazil	To assess the influence of obesity on physical and lung function of children and adolescents with obesity and to associate the variables with a control group	Yes	Yes	Yes	Yes	Yes	Yes	-	Yes	Yes	
Rastogi et al. [16] (2014) - United States of America	To investigate the association between total fat, trunk fat and metabolic abnormality with lung function of a sample of minority urban adolescents	Yes	Yes	Yes	Yes	No	Yes	+ Body Plethysmography	No	No	
Faria et al. [17] (2014) - Brazil	To investigate lung function response during exercise in adolescents with non-morbid obesity and without respiratory diseases	Yes	Yes	Yes	Yes	No	Yes	+ Respiratory Muscular Force	Yes	Yes	
Rio-Camacho et al. [20] (2013) - Spain	To investigate the left ventricular mass (echocardiography) of children with obesity, with and without metabolic syndrome; to evaluate the association between the level of adipokine and circulating cytokine and the alteration of left ventricular mass and spirometry; and to determine the best variable to predict cardiovascular risk	Yes	Yes	Yes	Yes	No	No	-	Yes	No	
Gibson et al. [19] (2014) - Australia	To evaluate (i) whether children and adolescents with overweight or obesity can be submitted to submaximal exercise; (ii) respiratory limitations during exercise in children and adolescents with overweight and obesity compared to the control group	Yes	Yes	No	Yes	No	No	+ Multi-breath Nitrogen Wash Out	Yes	Yes	
Chen et al. [22] (2009) - Canada	To evaluate waist circumference as a predictor of lung function markers and to compare it with BMI in children and adolescents	Yes	Yes	Yes	No	No	No	-	No	No	

Table 4 Disparities between the markers of cross-sectional studies on children-adolescents without respiratory diseases, where spirometry markers were assessed, and the relation between lung function and body mass, expressed as either body mass index or norm weight/overweight/obesity preventing the performance of meta-analysis (Continued)

Authors (year)	Objective	Spirometry measurement					Other lung function measurement			Obesity as an independent variable	Exercise
		FVC	FEV ₁	FEV ₁ /FVC	FEF _{25-75%}	PEF	ERV				
Ferreira et al. [25] (2017) - Brazil	To evaluate lung function of children and adolescents with obesity (without asthma) by Spirometry and volumetric capnography and to compare them to healthy control of the same age group	Yes	Yes	Yes	Yes	No	Yes	+ Volumetric Capnography	Yes	No	
Del-Rio Navarro et al. [26] (2013) - Mexico	To compare bronchial hyperactivity by the methacholine challenge testing in Mexican children with normal weight. In addition, to associate the group with normal weight with children with obesity or morbid obesity	Yes	Yes	No	Yes	Yes	No	+ Methacoline Challenge Testing	Yes	No	
Jeon et al. [28] (2009) - South Korea	To evaluate the factors that influence lung function in female adolescents, focusing on the hormonal factors of the menstrual cycle and obesity	Yes	Yes	Yes	Yes	No	No	-	No	No	
He et al. [29] (2009) - China	To evaluate the relationship of obesity and asthma, symptom of asthma and lung function of Chinese schoolchildren using the definition of overweight and obesity of a Chinese group	Yes	Yes	No	Yes	No	No	-	No	No	
Silva et al. [30] (2011) - Brazil	To assess the onset of exercise-induced bronchospasm in children and adolescents, without asthma and overweight	Yes	Yes	Yes	Yes	Yes	No	+ Peak Expiratory Flow Meter	Yes	Yes	
Assumpção et al. [32] (2017) - Brazil	To compare IOS parameters of children with normal weight, overweight and obesity	Yes	Yes	Yes	Yes	Yes	No	+ Impulse Oscillometry	Yes	No	
Cibella et al. [33] (2015) - Italy	To investigate the effects of weight on lung function of healthy children in a sample registered in 2 cross-sectional surveys with selected age group	Yes	Yes	Yes	Yes	No	No	-	No	No	
Silva et al. [34] (2015) - Brazil	To evaluate the effects of posture on thoracoabdominal kinematics of children with obesity and to compare them with a control group with normal weight	Yes	Yes	Yes	No	No	No	+ Respiratory Muscular Force + Optoelectronic Plethysmography	Yes	No	
Torun et al. [35] (2014) - Turkey	To compare lung function in children with normal weight, overweight, obesity or morbid obesity and to evaluate the effects of degree of obesity on lung function	Yes	Yes	Yes	Yes	Yes	No	-	Yes	No	
Khan et al. [36] (2014) - Canada	To associate anthropometric measures and lung function in children	Yes	Yes	Yes	No	No	No	-	Yes	No	
Rosa et al. [37] (2014) - Brazil	To evaluate respiratory muscular strength by maximum respiratory pressure in healthy,	No	Yes	No	No	No	No	+ Respiratory Muscular Force	Yes	No	

Table 4 Disparities between the markers of cross-sectional studies on children-adolescents without respiratory diseases, where spirometry markers were assessed, and the relation between lung function and body mass, expressed as either body mass index or norm weight/overweight/obesity preventing the performance of meta-analysis (Continued)

Authors (year)	Objective	Spirometry measurement				Other lung function measurement			Obesity as an independent variable	Exercise
		FVC	FEV ₁	FEV ₁ /FVC	FEF _{25-75%}	PEF	ERV	ERV		
Assunção et al. [38] (2014) - Brazil	schoolchildren with overweight and obesity, and to identify if the anthropometric and respiratory variables are related to the outcomes	Yes	Yes	Yes	Yes	No	No	+ Helium Washout	Yes	No
Alghadir et al. [40] (2012) - Saudi Arabia	To describe pulmonary functional alterations in asymptomatic and overweight children and adolescents	Yes	Yes	No	No	No	-	-	Yes	No
Paralikar et al. [41] (2012) - India	To investigate the relationship between severity of obesity and parameters of lung function, comparing lung function in Saudi men with overweight and obesity with individuals with normal weight, and to compare the value found with the reference values for Caucasian individuals	Yes	Yes	Yes	Yes	Yes	No	-	Yes	No
Supriyatno et al. [42] (2010) - Indonesia	To evaluate lung function in adolescents with obesity in the city of Baroda, Gujarat	Yes	Yes	Yes	No	No	-	-	No	No

ERV Expiratory reserve volume, FVC Forced vital capacity, FEV₁ Forced expiratory volume in the first second of forced vital capacity, FEV₁/FVC forced expiratory volume in the first second and forced vital capacity ratio, FEF_{25-75%} Forced expiratory flow between 25 and 75% of forced vital capacity, PEF Peak expiratory flow measured by spirometry, + Studies that showed comparative markers with greater value in obesity or positive associations with variables that are indicative of obesity, - Studies that showed comparative markers with lower value in obesity or with negative association with variables that are indicative of obesity

However, studies on individuals with obesity describe a trend towards lower FEV₁/FVC, FRC, ERV and RV, suggesting that both mechanical and inflammatory impairments influence lung function throughout childhood and adolescence.

Studies on pubertal development would be significant for a standard comparison including hormonal and structural changes in this period and the onset and duration of maturation. The quantification and distribution of body fat and the analysis of lifestyle habits would promote coherence and standardization on this subject, favoring the clinical approach to individuals.

The prevalence of obesity has increased worldwide, and although it is a relevant public health problem that affects all age groups, the role and methods to evaluate its impact on lung function in children and adolescents have not been established yet, and full understanding of the topic is still far from being attained.

Abbreviations

-: Studies that showed comparative markers with lower value in obesity or with negative association with variables that are indicative of obesity;
+: Studies that showed comparative markers with greater value in obesity or positive associations with variables that are indicative of obesity; 6MWT: Six-minute walk test; alv: Alveolar; ATS: American Thoracic Society; AX: Reactance area; BD: Bronchodilator; BIA: Bioimpedance; BMI: Body mass index (Weight/Height²); BP: Blood pressure; CDC: Center for Disease Control and Prevention; CI: Capnography index [(Slp₂/Slp₃) × 100]; CRP: C-reactive protein; DBP: Diastolic blood pressure; D_{LCO}: Diffusing capacity of the lungs for carbon monoxide; DSV: Dead space volume; DSV/TV: Relation between dead space volume and tidal volume; DXA: Dual energy X-ray absorptiometry; EIB: Exercise-induced bronchospasm; Embase: Excerpta Medica Database; ERS: European Respiratory Society; ERV: Expiratory reserve volume; EtCO₂: End-tidal carbon dioxide; FEF_{25–75%}: Forced expiratory flow between 25 and 75% of forced vital capacity; FEF_{25%}: Forced expiratory flow at 25% of forced vital capacity; FEF_{50%}: Forced expiratory flow at 50% of forced vital capacity; FEF_{75%}: Forced expiratory flow at 75% of forced vital capacity; FeNO: Fraction of exhaled nitric oxide; FEV_{0.75}: Forced expiratory volume at 0.75 s; FEV₁: Forced expiratory flow in the first second of forced vital capacity; FEV₁/FVC: Relation between forced expiratory volume in the first second and forced vital capacity; FRC: Functional residual capacity; Fres: Resonant frequency; FuncVC: Functional vital capacity; FVC: Forced vital capacity; FVC/weight: Forced vital capacity index by weight; GINA: Global Initiative for Asthma; HC: Healthy controls; HDL: High density lipoprotein; HOMA-IR: Homeostasis model assessment of insulin resistance; HR: Heart rate; Hz: Hertz; IC: Inspiratory capacity; IL-6: Interleukin 6; IOS: Impulse oscillometry; ISAAC: The International Study of Asthma and Allergies in Childhood; LAR: Leptin to adiponectin ratio; LLN: Lower limit of normal; MCP-1: Monocyte chemoattractant protein-1; MEDLINE: PubMed, Medical Literature Analysis and Retrieval System Online - Public Medline; MEP: Maximum expiratory pressure; MIP: Maximal inspiratory pressure; MV: Minute volume; MVD: Mixed ventilatory disorder; MVV: Maximum voluntary ventilation; OEP: Optoelectronic plethysmography; OVD: Obstructive ventilatory disorder; pBMI: BMI percentile; PEF: Peak expiratory flow measured by spirometry; PFE: Peak expiratory flow measured by Peak Flow Meter; R20: Central airway resistance; R5: Total resistance; RMS: Respiratory muscle strength; RR: Respiratory rate; RV: Residual volume; RV/TLC: Ratio of residual volume and total lung capacity; RVD: Restrictive ventilatory disorder; SAH: Systemic arterial hypertension; SBP: Systolic blood pressure; Slp₂: Slope of phase 2; Slp₂/TV: Relation between slope of phase 2 and tidal volume; Slp₃: Slope of phase 3; Slp₃/TV: Relation between slope of phase 3 and tidal volume; SpO₂: Peripheral oxygen saturation; Tanner: Pubertal developmental stage according to Tanner's criteria; Te: Expiratory time; Ti: Inspiratory time; TLC: Total lung capacity; TNF-α: Tumor necrosis factor; tot: Total; TV: Tidal volume; VC: Vital capacity; VCO₂: Volume of exhaled carbon dioxide; VHL: Virtual Health Library (Brazil); VolC: Volumetric capnography; VTAB: Tidal

volume of the abdomen; VTRCa: Tidal volume of the abdominal rib cage; VTRCa%: Percentage of contribution of the tidal volume in the abdominal rib cage to the total tidal volume; VTAB%: Percentage of abdominal tidal volume contribution to tidal volume; VTRCp: Tidal volume of the pulmonary rib cage; VTRCp%: Percentage of contribution of the tidal volume in the rib cage to total tidal volume; WC: Waist circumference; WHO: World Health Organization; WHR: Waist hip ratio; X5: Reactance at 5 Hz; Z5: Respiratory impedance; θ: Phase transition between two compartments

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The authors declare that there is no competing of interest to declare;

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