





Karnataka Paediatric Journal

Oscillometry – The future of estimating pulmonary functions

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Review Article

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Received : 21 October 2020 Accepted : 23 November 2020 Published : 25 January 2021

DOI 10.25259/KPJ_25_2020

Quick Response Code:



ABSTRACT

The prevalence of asthma is increasing rapidly, worldwide, due to changing gene-environment interactions. The rate of rise is more in resource poor nations due to lack of knowledge and non-availability of expertise. Monitoring of lung functions is mandatory for diagnosis and further management of asthmatic patients. Spirometry, the widely available investigation, is the gold standard test used for mapping pulmonary dynamicity. It has got its own limitations in the form of operational difficulties in children, the elderly, and in those with neuromuscular or behavioral issues. In the current era of COVID-19 pandemic, the utility of spirometry has been further restricted to selected cohort only, due to potential risk of viral transmission during the procedure. Oscillometry technique has been used previously, to monitor lung functions, with promising results. Ultrasonic waves of various frequencies accompany the tidal breath of patients and respiratory impedance is calculated by measured pressure and flow signals from exhaled breath. The results are interpreted in the form of resistance, reactance, resonant frequency, and reactance area. Various manufacturers have developed different mechanical models with slight variation in impulse pattern till date. There are certain distinct advantages of oscillometry over spirometry. Being tidal breath-based maneuver, it is more child friendly. People with neuromuscular weakness, cognitive limitations, and the elderly can easily perform it with only minor understanding and effort. Oscillometry is more sensitive than spirometry in detecting peripheral airway diseases. Post-bronchodilator reversibility can be evaluated by comparing with the baseline respiratory characteristics. Their utility in restrictive diseases and vocal cord dysfunction has also been explored. Less aerosol generation during the normal tidal breath is another advantage of oscillometry, over spirometry needing forceful efforts, which makes it more suitable for use in viral pandemic situations for monitoring patients with both asthma and pneumonia. More research is needed, in various geographic locations and heterogeneous populations, to devise the normative data of oscillometric parameters. Simultaneously, there is an urgent need for standardization of available machines at global platform.

Keywords: Oscillometry, Spirometry, Pulmonary function test, Respiratory impedance, COVID-19

INTRODUCTION

Allergic disorders are increasing worldwide with substantial affliction toward developing countries.^[1] The current global prevalence of diagnosed asthma cases is 300 million.^[1] In India, about 30% of the population is affected by some form of allergy.^[2] Asthma prevalence has increased many folds over the past 6 decades and is currently influencing 15% of people in Indian capital.^[3] As per a recent estimate, approximately half of the children present to a physician with at least one episode of wheezing by their sixth birthday.^[4] These figures are underestimate of real quantity due to lack of awareness among health-care providers and non-availability of reliable diagnostic modalities in resource poor nations.^[3] Demonstration of reversible airflow obstruction is desired in any patient with features of chronic airway inflammation (such as recurrent wheeze, breathlessness, chest tightness, and cough of variable intensity) for making a diagnosis of asthma.^[5] Spirometry

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is considered gold standard technique for demonstrating airway reversibility in suspected cases.^[6] Although universally accepted, this modality has practical difficulties such as requirement of patient cooperation and forceful respiratory efforts.^[6] It is cumbersome in young children, the elderly, patients with neuromuscular weakness, post-cardiothoracic surgery, and those with learning problems.^[7,8] Finkelstein et al., in a multicentric survey, demonstrated that only 21% of 671 primary care physician used spirometry, due to operational dilemmas.^[9] About 28% of people were either missed or overdiagnosed in another study using this tool.^[10] Inability to reliably diagnose peripheral airway and parenchymal diseases are other shortcomings of this age-old technique.^[10] Spirometry is not dependable in under-five wheezers.^[5] With the emerging global problem of asthma, we urgently require a competent method which can overcome many of these shortcomings. An ideal lung function test should be possible at any age, safe, simple to perform, reproducible in different circumstances, and sensitive enough to detect minor changes in respiratory mechanics.^[11]

Any technique based on tidal breathing would be ideal, especially in pediatric age group, where the cooperation expected from the subjects is minimal. Among the various available diagnostic modalities, some are oscillometry, interrupter technique, and body plethysmography.^[12] Oscillometry seems a reasonable option to determine pulmonary mechanics in patients who are unable to perform spirometry.^[13] It works on the principle of moving sound waves over tidal breath during the respiratory cycle. Resistance and reactance are calculated from measured changes in pressure and flow at different frequencies. Oscillometry is better in children and can provide additional information in adults for monitoring lung functions compared to spirometry.^[14] Oscillation principle has been used previously in both pre-school and school-aged children to assess lung functions.[15] Gupta et al. have recently demonstrated airway reversibility using this technique in children as young as 2 years of age.^[16] Patients with physical and cognitive limitations were also able to perform this maneuver convincingly.^[17] Komarow et al. have explored its utility in identifying vocal cord dysfunction.^[18] Another key highlight is reduced aerosol generation in oscillometry, which makes it a safer alternative in COVID-19 pandemic.^[19]

HISTORICAL PERSPECTIVE

Oscillations were first used to quantify the mechanical behavior of respiratory system by Dubois *et al.* in 1956.^[13] The approach commonly known as forced oscillation technique (FOT), in which airway characteristics in the form of impedance were monitored using sound waves of various frequencies. There have been several modifications in FOT over the past 6 decades with regard to configuration, oscillation type, frequencies, and assessment of airway parameters.^[20] One such development, known as impulse oscillometry (IOS), was demonstrated by Michaelson *et al.* in 1975 using multiple frequencies at one point of time.^[21] Oscillometry provides a detailed description of pressure-flow relationships over discrete frequencies. This provides a better insight about resistance and reactance of respiratory system than conventional spirometry.^[8]

TYPES

Depending on the type of oscillation signals used, this technique can be classified as: $^{\left[22\right] }$

- Monofrequency using single sinusoidal pressure waveform
- Pseudorandom noise (PRN) where impulses of several frequencies are simultaneously applied
- IOS in which recurrent impulses (square waveform) applied at a fixed frequency of 5 Hz.

Single frequency impulses are useful for monitoring patients with sleep apnea or those on mechanical respiratory support (ventilation or continuous positive airway pressure). PRN FOT impulses are widely used for monitoring various obstructive (asthma, bronchitis, and emphysema) and restrictive (interstitial lung disease, pulmonary fibrosis, and thoracic wall abnormalities) diseases. Intrabreath changes are better defined using recurrent impulses. Dandurand *et al.* have demonstrated that devices with PRN FOT signals are better in measurement of peripheral lung characteristics, than others, when subjected to higher mechanical load.^[23]

The commonly used commercially available machines working on PRN impulses are Wave Tube, TremoFlo C-100, MostGraph-02 prn, and Resmon Pro, whereas Master Screen IOS and MostGraph-02 imp work with recurrent impulses.^[23] The two most commonly available FOT models, Master Screen IOS and Resmon Pro, use 5, 10, and 20 Hz and 5, 11, and 19 Hz frequencies, respectively, to study the airway mechanical properties.^[23]

PRINCIPLE

Oscillations of different frequencies are used to study respiratory impedance in FOT. The technique requires only passive cooperation from the patients for the evaluation of lung functions.^[24] Respiratory impedance (Z) is a measure of resistive, inertial, and elastic forces of lungs and thoracic cage.^[25] Sound waves of multiple frequencies, generated by a loud speaker, are superimposed over tidal breath of subjects through the respiratory system [Figure 1].^[8]

These sound waves, being the mini pressure waves, cause subtle pressure changes in the airway which leads to change in airflow.^[24] Smaller frequencies (2–4 Hz) can travel till the depth of lung peripheries, whereas higher frequencies (>20 Hz)

reach to proximal conducting airways only.^[26] Middle range frequencies (5–20 Hz) are commonly used in clinical practice to determine respiratory characteristics [Figure 2] using fast Fourier transform technique.^[14] Frequencies less than 5 Hz get easily altered by harmonics of the normal breathing.^[26,27] whereas larger frequencies (>30 Hz) cause subjective discomfort and are affected by shunting properties of upper airways.^[26,28]

The sound wave signals of pressure and flow are separated from the breathing pattern, by signal filtering mechanism, while returning from lungs during exhalation. The complex ratio of sinusoidal pressure (P) and flow (Q) for individual sound wave frequency, as determined by the pressure and flow transducers (pneumotachograph) at the mouthpiece, informs about the impedance of various segments of the respiratory system [Figure 1].^[25] Impedance is calculated at discrete frequencies by ohm's law.

$$Z(\omega) = P(\omega)/Q(\omega)$$
$$\omega = 2.\pi f$$

Where, *Z* – Impedance, *P* – Pressure, *Q* – Flow, ω – oscillation frequency function, *f* – frequency

Measured impedance (Z) is the sum of opposing forces in the respiratory system, resistance (R), and reactance (X). Z, calculated at individual frequency, informs about the mechanical properties of respective portion of airways [Figure 2]. during its travel through the respiratory system. It can be represented by a combination of resistance (R), a real force, and reactance (X), an imaginary component. Resistance and reactance are associated with energy dissipation and storage, respectively^[25]

$$Z(\omega) = R(\omega) + jX(\omega)$$

R – Resistance, *j* – unit imaginary number defined as $\sqrt{-1}$, and *X* – Reactance

- Resistance (R) It is a measure of opposition to airflow. It is directly proportional to length and inversely proportional to fourth power of radius of conducting tubes. Resistance measured at a particular frequency (f) is labeled as R_f , for example, R_5 is resistance detected at 5 Hz. R_f includes the resistance of oropharynx, larynx, trachea, large and small airways, lungs, and chest wall tissue
- Reactance (X) Reactance can be understood as rebound resistance produced by distensible airways.^[29] It is an imaginary component of impedance, which is defined by balance between inertance (I) (positive force) of conducting airway and capacitance (C) (negative force) of pulmonary parenchyma. Capacitance is in inverse correlation with elastic properties of lung

$$X(\omega) = \omega . I - E/\omega$$
$$X(\omega) = \omega . I - 1/\omega . C$$
$$X \alpha I \qquad X \alpha C$$

Where, X – Reactance, I – Inertance, E – Elastance, and C – Capacitance

• Reactance measured at a particular frequency (f) is labeled as X_6 for example, X_5 is reactance at 5 Hz. At lower

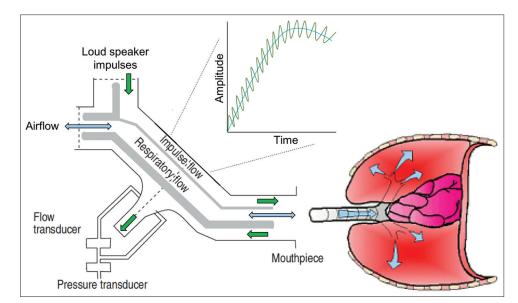


Figure 1: Principle of oscillometry. Sound waves (green) are superimposed over the normal tidal breath (blue) during respiration. Transducers measure the pressure and flow during exhalation for individual frequency which is utilized to calculate impedance of the respiratory system.

TERMINOLOGY

• Impedance (Z) – It is defined as sum total of all the resistive, inertial, and elastic forces of respiratory system which a pressure impulse has to encounter

frequency, reactance is negative due to predominant capacitative forces, whereas it becomes positive toward higher frequencies, with major contribution from inertial forces [Figure 3]

- Resonant frequency (Fres) It is the arbitrary frequency number at which capacitative and inertial forces equalize and reactance becomes zero [Figure 3]. Elastic forces dominate below Fres, whereas airway inertance plays major role above Fres.^[29] Fres is usually higher in children and reduces with age
- Reactance area (Ax) It is the triangular area limited by Fres (on right side) and reactance at 5 Hz (on left side) [Figure 3]. Ax provides information regarding peripheral airways and lung parenchyma. It is more sensitive parameter than Fres and X₅ (in descending order) for the detection of small airway obstruction as well as to document the bronchodilator reversibility
- Coherence It is a quality control parameter which reflects the reliability of oscillometry maneuver. The value depends on relative comparison between input (flow) and output (reflected pressure) in the respiratory

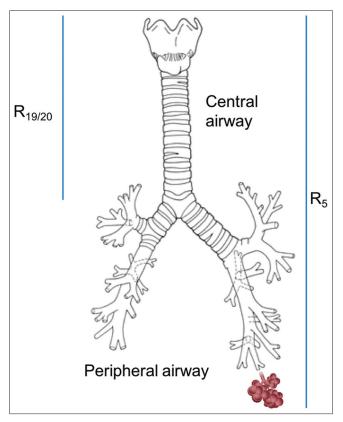


Figure 2: Impulses of various frequencies traveling through airway. Smaller frequency sound waves (5 Hz), being more energetic, travel till the farthest point of respiratory system, that is, alveoli. Higher frequency (20 Hz) remains in the central airways depicting the characteristic of respective part.

system.^[30] Coherence values of >0.8 at 5 Hz and >0.9–1 at 20 Hz are considered satisfactory in adult subjects but cutoffs are yet to be validated for children.^[15,31] The potential problem in relying on coherence is varied approaches used for calculations leading to different values by different manufacturers. Coherence values are reduced in pathological conditions or due to improper technique including swallowing, glottis closure, tongue causing airflow obstruction, and irregular breathing during the oscillometry maneuver.^[32] High coherence values cannot rule out measurement errors or artifacts.^[32]

 Coefficient of variation (CoV%) – This should be used over coherence to determine quality control, whenever available. CoV should be ≤10% in adults and ≤15% in children for two sets of R₅.^[32]

TECHNIQUE

Calibration of the machine, at least once a day, is desired with the external resistor or as per the manufacturer's specifications.^[11] Bronchodilator medications (short-acting β -2 agonist for 4 h and long-acting β -2 agonist for 24 h) should be stopped before the procedure. After explaining the procedure (preferably by recorded video demonstration) to patient and attendants, anthropometric measurements (weight, height, and body mass index) are documented. Demographic (name, age, gender, area of residence, and identification number) and anthropometric parameters are entered in the machine after calibration. Patient is asked to sit on an examination stool/chair, with uncrossed legs to reduce the influence of extrathoracic pressure with straight

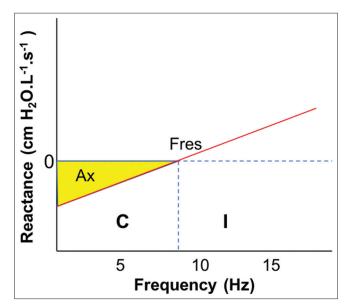


Figure 3: Changes in reactance with frequency. Fres: Resonant frequency, Ax: Reactance area, C: Capacitance, I: Inertance.

back [Figure 4].^[30] He/she is asked to hold the mouth piece with his/her teeth and to make a tight seal around it with lips to prevent any air leak during the FOT maneuver. A nose clip is applied to occlude both the nostrils and he/she is asked to breathe through mouth piece. Height of sitting stool/chair and/or mouth piece is adjusted to achieve a comfortable position for the patient with slight neck extension. As FOT is based on small pressure oscillations and a little change in resistance or air leak can affect the interpretations significantly, it is important to ensure adequate seal around nose and mouth piece. The cheeks, most compliant part of respiratory system in children, should be supported firmly either by patient him/herself or attendant to minimize wobbling [Figure 4].^[20] After appropriate positioning, patient is asked to perform normal tidal breathing in a relaxed manner. An average of 10 respiratory efforts or 1 min, whichever is earlier, is required to assess the respiratory characteristics during any maneuver.^[12] Acquisitions of minimum 30 s for adults and 16 s for children (<12 years of age) with at least three acceptable breaths are recommended.^[32] Respiratory efforts meeting acceptability criteria [Table 1] are considered valid, whereas maneuvers with artifacts such as airflow obstruction by tongue or glottic closure, irregular breathing, coughing, crying, swallowing, and improper technique will be discarded by the machine. A maximum of three acceptable maneuvers are recorded and checked for coherence or CoV. Mean respiratory impedance (resistance and reactance), resonant frequency, and reactance area are documented. The procedure is repeated 15 min after inhaled short-acting β -2 agonist to identify any postbronchodilator reversibility.

NORMAL VALUES

• Impedance – Several studies have provided reference values for respiratory characteristics till date.^[20] The



Figure 4: Technique of performing oscillometry.

references might vary as per the ethnicity and the oscillation technique used in different machines.^[26] The commonly used regression equations, by the machine, for calculating R and X based on height (H) in meters, weight (W) in kilograms, and age (A) in years are as follows:^[33]

for men -

R_{men} = -0.2454. H+0.001564. W-0.00055. A+0.5919

X_{men} = 0.1479. H-0.000402. W-0.00022. A-0.1721

and for women -

 $R_{women} = -0.4300. H+0.00165. W-0.00070. A+0.9312$

 $X_{women} = 0.2487. H-0.001700. W-0.00053. A-0.2158$

Normative values from Indian populations need to be devised. Gupta *et al.* have recently demonstrated a negative correlation between oscillometry parameters with height followed by body mass index in Indian children, while evaluating airway reversibility in asthmatic patients.^[16] There was no gender influence observed on any of the parameter.

- Resonant frequency The normal values of Fres varies in between 6 and 12 Hz in healthy adults^[8,34] and it tends to be more in children
- Reversibility The recommended cutoffs for significant bronchodilator response in both adults and children are -40% in R₅, +50% in X₅, and -80% in Ax.^[32] These values might vary with severity of disease and more studies are required before considering them as benchmark
- Degree of bronchoconstriction Cutoffs for X₅ have varied from 50 to 80% and more studies are needed to provide reference values for specific populations.^[32]

INTERPRETATION

 Resistance (R) – Total, large/central, and small/ peripheral airway resistances are represented as R₅, R_{19/20}, and R₅-R_{19/20}. R₅ is always higher than R_{19/20}. This difference is practically negligible in adults, whereas

Table 1: Acceptability criterias – All of the following are required.				
Domain	Criterias			
Patient position	Sitting position at comfortable height Back straight with slightly extended neck Legs uncrossed Nose clip on Cheek firmly supported			
Patient-machine interface	Tight seal around mouth piece No artefact due to irregular breathing efforts, tongue obstruction, speaking or coughing during maneuver, glottis closure or swallowing			

it increases in younger children due to significant contribution by peripheral airway resistance. In peripheral airway obstruction, R_5 will increase with normal $R_{19/20}$ (and higher $R_5-R_{19/20}$) making frequency dependent airway resistance (R α 1/f).^[24] In larger airway obstruction, both R_5 and $R_{19/20}$ will rise equally (with normal $R_5-R_{19/20}$), which will be frequency independent [Figure 5]. There will not be any change in restrictive lung diseases [Table 2]

- Spiky pattern in inspiration, demonstrated by >2 standard deviation variation in subsequent efforts at 5 Hz, may suggest vocal cord dysfunction.^[18] The finding needs to be supported with further research
- Reactance (X) It is usually measured at 5 Hz and becomes more negative in both peripheral airway obstruction and lung parenchymal disease [Figure 5]. It is not affected by large airway obstruction
- Resonant frequency (Fres) It increases (shift to right) in both restrictive and peripheral airway obstructive diseases.^[24] It is not affected by large airway obstruction
- Reactance area (Ax) It increases in both small airway obstruction and restrictive diseases.^[8] It is not affected by central airway problems.

COMPARISON WITH SPIROMETRY

Oscillometry is more sensitive for the detection of peripheral airway obstruction and restrictive diseases affecting lung parenchyma.^[11,12] [Table 3] highlights the salient differences between spirometry and oscillometry.^[8]

Use of spirometry has been restricted in current COVID-19 pandemic due to reasons of enhanced risk of disease transmission by potential aerosol generation.^[35] Forced breathing maneuver causes more aerosol generation due to "airway reopening phenomenon."^[36] Breathing till residual volume will reopen the collapsed alveoli causing increased air turbulence, leading to more production and release of smaller particles.^[36] A small volume tidal breath, as used in oscillometry, will not cause much disturbance in the internal milieu and thus safeguarded in situations of active infections (such as influenza and corona). Gupta *et al.* have highlighted this concept recently, which suggest oscillometry procedure safer than spirometry in viral pandemic situations.^[19]

CLINICAL APPLICATIONS OF OSCILLOMETRY

 Provides practically useful information regarding the subtle changes in airways with greater sensitivity, in both

Table 2: Changes in oscillometry parameters during pathological conditions.							
Conditions	R_5	R _{19/20}	$R_5 - R_{19/20}$	\mathbf{X}_5	Ax	Fres	
Peripheral obstruction	$\uparrow \uparrow \uparrow$	Ν	$\uparrow \uparrow$	More negative	$\uparrow\uparrow$	$\uparrow\uparrow$	
Central airway obstruction	$\uparrow\uparrow$	$\uparrow\uparrow$	Ν	Ν	Ν	Ν	
Combined airway obstruction	$\uparrow\uparrow\uparrow$	$\uparrow\uparrow$	\uparrow	More negative	$\uparrow\uparrow$	$\uparrow\uparrow$	
Restrictive lung disease	Ν	Ν	Ν	More negative	$\uparrow \uparrow$	$\uparrow\uparrow$	

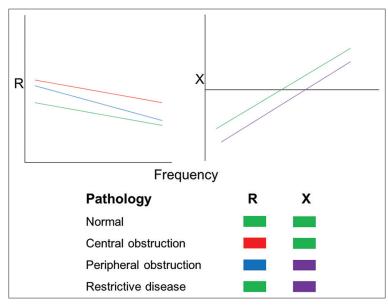


Figure 5: Respiratory characteristics in health and disease. R – Resistance (in cm H₂O.L⁻¹.s⁻¹), X – Reactance (in cm H₂O.L⁻¹.s⁻¹). During normal healthy conditions, the resistance and reactance are at baseline (green). Various combinations of changes in these parameters will help in determining the nature and location of pathology.

children and adults, when compared to spirometry^[37,38]

- Useful to assess abnormal distal airway function, in case of clinical suspicion with normal spirometry^[38,39]
- Bronchodilator reversibility can be demonstrated with short-acting β-2 agonists and ipratropium^[40,41]
- Good potential in diagnosis and monitoring of restrictive lung diseases such as bronchopulmonary dysplasia,^[42] cystic fibrosis,^[43] and interstitial lung disease^[44]
- Feasible option in children, the elderly, and those with neuromuscular diseases and impaired intellect^[24,26]
- Potentially useful in patients on mechanical ventilation^[45] and during sleep^[11]
- Safer than spirometry during viral pandemic situations (e.g., influenza and corona) due to less aerosol generation.^[19] Oscillometry can be used to reliably diagnose and monitor patients with asthma and COVID-19 pneumonia.

[Figure 6] shows an algorithmic approach to a patient with oscillometric lung function assessment.

Table 3: Comparative analysis between spirometry and oscillometry.					
Parameter	Spirometry	Oscillometry			
Principle	Measures flow rates and lung volumes	Calculates impedance by measuring flow and pressure of sound waves			
Parameters measured	FEV ₁ , FVC, PEFR, FEF _{25-75%}	Z, R, X, Fres, Ax			
Type of breath required	Forced maneuver	Tidal breath			
Patient cooperation needed	High	Minimal			
Can be performed in children	>7 years	>2 years			
Patients with neuromuscular weakness, intellectual disability, post cardiothoracic surgery	Procedure cannot be done	Can be done			
Sensitivity for detection of peripheral airway obstruction	Low	High			
Aerosol generation	High	Very low			
Standardization of method	Yes	Yet to be done			
References value	Available	Need more studies			

FEV₁ – Forced Expiratory Volume in 1 second, FVC – Forced Vital Capacity, PEFR – Peak Expiratory Flow Rates, FEF_{25-75%} - Forced Expiratory Flow at 25-75% of FVC, Z – Impedance, R – Resistance, X – Reactance, Fres – Resonant frequency, Ax – Reactance area

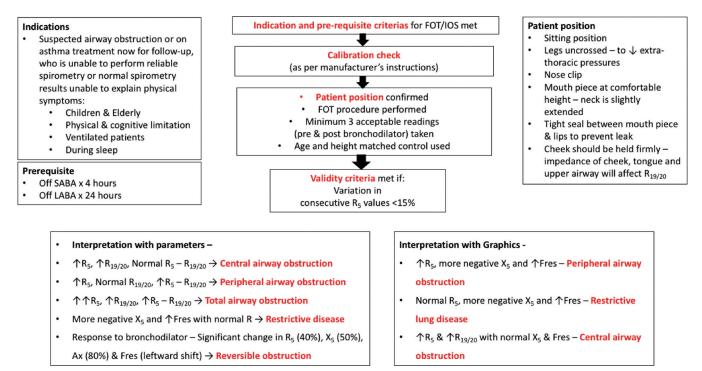


Figure 6: Approach to a patient with oscillometric assessment. FOT: Forced oscillation technique, IOS: Impulse oscillometry, SABA: Shortacting $\beta 2$ agonist, LABA: Long-acting $\beta 2$ agonist, R: Resistance (in cm H₂O.L⁻¹.s⁻¹), X: Reactance (in cm H₂O.L⁻¹.s⁻¹), Fres: Resonant frequency (in Hz), Ax: Reactance area.

Limitations

- Although this technique is tidal breath based, still a minimum amount of cooperation is needed from patients
- Standardization of the available machines with different manufactures is needed
- Reference values for different populations are not available
- Reference cutoff values for bronchodilator reversibility need to be validated with more studies
- Poor cheek support can reduce the resistance values^[46]
- More research required in restrictive diseases, ventilated, and/ or sedated patients and patients with vocal cord dysfunction.

CONCLUSION

Oscillometry, being a tidal breath-based technique, can be a real privilege to physicians and their patients for monitoring lung functions. It is more sensitive in detecting small airway pathologies than conventional spirometry. Limited aerosol generation could be another reason for its use in viral pandemics for monitoring lung functions. More research is required for identifying regional reference values and standardization of machines.

Declaration of patient consent

Patient's consent not required as patients identity is not disclosed or compromised.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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How to cite this article: Gupta N, Sachdev A, Gupta D, Gupta S. Oscillometry – The future of estimating pulmonary functions. Karnataka Paediatr J 2020;35(2):79-87.