Review Article

Electrical impedance tomography

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Abstract: Continuous assessment of respiratory status is one of the cornerstones of modern intensive care unit (ICU) monitoring systems. Electrical impedance tomography (EIT), although with some constraints, may play the lead as a new diagnostic and guiding tool for an adequate optimization of mechanical ventilation in critically ill patients. EIT may assist in defining mechanical ventilation settings, assess distribution of tidal volume and of end-expiratory lung volume (EELV) and contribute to titrate positive end-expiratory pressure (PEEP)/tidal volume combinations. It may also quantify gains (recruitment) and losses (overdistention or derecruitment), granting a more realistic evaluation of different ventilator modes or recruitment maneuvers, and helping in the identification of responders and non-responders to such maneuvers. Moreover, EIT also contributes to the management of life-threatening lung diseases such as pneumothorax, and aids in guiding fluid management in the critical care setting. Lastly, assessment of cardiac function and lung perfusion through electrical impedance is on the way.

Keywords: Acute lung injury; critical care; electric impedance; physiologic monitoring; respiratory distress syndrome

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Introduction

Continuous assessment of respiratory status is one of the cornerstones of modern intensive care unit (ICU) monitoring systems. Lung imaging techniques have experienced overwhelming progress: we currently not only base our knowledge on chest X-ray or computed tomography, but use lung ultrasound, positron emission tomography, electrical impedance tomography (EIT) or magnetic resonance imaging to improve our understanding of the disease and the effect of therapeutic strategies (1-3).

EIT uses electrical currents to assess the conductivity distribution within the body from voltage measurements of its surface. This concept was first described as a method to explore subterranean mineral deposits in the early twentieth century. Medical use of EIT was developed in the 1980s by

Barber and Brown at the Department of Medical Physics and Clinical Engineering in Sheffield (UK). A broad spectrum of possible applications in medicine were suggested, from gastric emptying to brain function monitoring, and then from breast imaging to interpretation of lung function. Since the first publication of an EIT image of the human thorax in 1985 by Brown et al., multiple advances have seen the light in this field of study, driven by the clinical need for monitoring of lung ventilation and assessment of regional lung function at the bedside (4). Several commercial EIT devices are currently available and being used on a day-today basis in different ICUs around the world: Pulmovista® 500 (Dräger Medical GmbH, Lündbeck, Germany), Goe-MF[®] II (CareFusion - Becton, Dickinson and Company, San Diego, United States), Mark 1[®] and Mark 3.5[®] (Maltron International Ltd, Rayleigh, United Kingdom),

BB[®] (Swisstom AG, Landquart, Switzerland) and Enlight[®] (Timpel S/A, Sao Paulo, Brazil) (5).

Principles of EIT

EIT is a non-invasive, radiation-free, real-time imaging of ventilation. Regional electrical bioimpedance measurements require an electrode belt of 16 electrodes (or 32 in the current models by Swisstom AG and Timpel S/A manufacturers) placed around the chest wall, including a reference electrode connected to a central body, preferably to the abdomen (the reference electrode ensures that all measurements of the different electrode pairs are related to the same electrical potential). The 16-electrode belt is placed in one transverse plane, usually between the 4th and 5th intercostal space (it is not recommended to place the belt lower than the 6th intercostal space because the diaphragm may enter the measurement plane).

EIT images are determined by the distribution of intrathoracic bioimpedance. Bioimpedance is defined by the specific composition of tissues, taking into account water, electrolytes, fat, etc. High content of extracellular water, high concentration of electrolytes, large cells and a high number of cell connections reduce impedance, whilst fat accumulation, bone and air increase impedance. Therefore, pathological changes of the tissue composition (such as pleural effusion, lung fibrosis, alveolar fluid, etc.) entail a change in regional bioimpedance. Lung imaging is therefore an ideal application of EIT; large changes in lung volume (and therefore in conductivity) can be easily detected due to the closeness of the lungs to the surface (6-12).

Small alternating high frequency and low amplitude electrical currents are applied though a pair of electrodes while the resulting surface potentials (or voltages) are measured by the remaining electrodes. Applying Ohm's law, the bioelectrical impedance between the injection and measurement electrode pairs is established from the known applied current and the measured voltages. Subsequently, the adjacent electrode pair is used for the next current application, and the other 13 measurements of voltage are carried out. The current applications and voltage measurements through adjacent electrode pairs imply a complete rotation around the thorax, creating voltage profiles at the 16 electrode positions, each composed of 13 voltage measurements. The resulting 208 values, also called frames (or raw images), are used to reconstruct a transverse EIT image; current EIT devices offer scan rates (number of frames acquired per second) of about 50 images per second.

In humans, an inspiratory maneuver from residual volume to total lung capacity amplifies the regional bioimpedance by approximately 300% (13). Cardiac activity and perfusion cause a change in thoracic bioimpedance, from diastole to systole, in a range of 3% (14). Extravascular lung water, body movement and electrode resistance in the skin may also change thoracic bioimpedance. In summary, an increase in intrapulmonary gas volume decreases conductivity, while an increase in blood or fluid volume, or disruption of cellular barriers, increase conductivity.

To be able to provide a high-resolution image, EIT requires a complex system of processors and software that compiles and analyzes all the information given by the electrodes and compares the recollected data with interindividual variations, adjusting to the thoracic geometry. A reconstruction algorithm will reject measurements at poorly connected electrodes, or ignore redundancy of data; they implicitly make assumptions about the level of random noise or interference in the raw data, and vary depending on the manufacturer: Sheffield backprotection (used by Goe-MF II and Mark 1 and 3.5), FEMbased Newton-Raphson method (used by PulmoVista and Enlight) and GREIT (used by BB).

EIT devices use the relative approach and thus calculate difference impedance images in relation to a reference. EIT images are usually represented on a pixel grid, where all image elements are the same size. Image reconstruction calculate images of change in tissue properties between a baseline measurement (reference) and the current frame, expressed as a percentage; these images are then represented by colour coding, which depends on the choice of the baseline frame (not unified between manufacturers). An EIT waveform is then defined as a sequence of impedance change values over time, represented in different ROI (or regions of interest), where image pixels are chosen to reflect regional changes associated with relevant effects (regions where ventilation related impedance changes occur).

Hence, obtaining functional EIT, we are able to visualize global and local ventilation activity over a period of time (selected by the user), picturing a map of ventilation distribution. Functional EIT produces EIT measures, defined as values derived from an EIT data acquisition. There are average global and regional measures (for example: regional or global ventilation delay or tidal variation), measures of spatial distribution of ventilation (for example, anterior/posterior ventilation ratio, center of ventilation, global inhomogeneity (GI) index] and examination-specific measures (e.g., regional respiratory

system compliance or regional intratidal gas distribution). However, we must point out an important drawback; only regions of the thorax that change their impedance over time will be represented by the EIT images.

Clinical applications in acute respiratory distress syndrome (ARDS)

ARDS is relatively frequent in the intensive care patient population. ARDS is characterized by diffuse alveolar damage, associated with an increase in alveolar and capillary permeability due to mechanisms of tension, stretching and shear between alveolar units, leading to accumulation of interstitial and alveolar edema. Release of inflammatory mediators causes bronchoconstriction, embolic phenomena, vasoconstriction of the pulmonary arterioles, and finally fibrosis of the lung parenchyma. This pathophysiologic phenomena leads, on one hand, to an increase in the overall weight of the lung (causing alveolar collapse in the dependent lung regions due to gravity) and on the other hand, triggers an inflammatory response that determines the presence of multiorgan failure (15-17).

Physicians have been looking for methods to optimize alveolar recruitment, maintaining an open lung and limiting pulmonary overdistension. Protocolled mechanical ventilation settings are not adapted to each patient's individual needs, and adjusting mechanical ventilation parameters is still the main bone of contention in clinical practice. We must highlight that ARDS has heterogeneous properties. Therefore, overall parameters reflecting the condition of the lung as a whole are insufficient to adequately target protective ventilation (18-21). All the systems that are being used for routine lung monitoring—graphic charts and ventilator data, radiological or gasometric tests—have limitations: either they provide information without regional specificity, or they do not provide progressive and continuous data over time.

In search of better ways to monitor pulmonary functions, EIT plays the lead. Lung EIT can be a diagnostic and a guiding tool for an adequate optimization of mechanical ventilation in critically ill patients, especially on ARDS, where mechanical ventilation remains a challenge.

Guidance on mechanical ventilation settings (22-39)

"Protective" ventilatory strategies have been designed to try to optimize mechanical ventilation and minimize its potential damages. These strategies are mainly based on the application of low tidal volumes and maneuvers designed to increase the functional residual capacity, trying to reduce cyclic alveolar collapse and overdistension of the lung.

As mentioned above, global parameters such as pressure-volume curves or measurement of the respiratory system compliance do not reliably translate what is actually happening in the lung, especially regarding to regional distribution of the tidal volume administrated. EIT images of the regional distribution of ventilation and changes in lung volume in real time highlight the relationship between the dependent and non-dependent lung regions. This dynamic evaluation makes EIT a helpful tool to optimize ventilator parameters on an individualized basis, as EIT may assist in defining mechanical ventilation settings, assess distribution of tidal volume and of end-expiratory lung volume (EELV) and contribute to titrate positive end-expiratory pressure (PEEP)/tidal volume combinations.

Studies performed in the last decade (23-32) have demonstrated that EIT-guided ventilation results in improved respiratory mechanics, improved gas exchange and reduced histologic evidence of ventilator-induced lung injury in an animal model. Careful titration of PEEP following maximal lung recruitment effectively reverses existing alveolar collapse and prevents further alveolar closure. EIT-based algorithms that estimate recruitable alveolar collapse, avoiding hyperdistension, have been one of the most interesting topics over the last years. Lung EIT has been used to assess and quantify the global and regional changes in lung impedance at the end of exhalation, quantifying gains (recruitment) and losses (overdistention or de-recruitment), granting a more realistic evaluation of different ventilator modes or recruitment maneuvers, and helping in the identification of responders and nonresponders to such maneuvers (Figure 1).

Costa et al. (33) demonstrated good correlation between EIT and CT estimates of lung collapse during decremental PEEP trials after a maximal lung recruitment maneuver. GI index has been used to describe the inhomogeneity of tidal volume distribution in a cross-sectional lung plane, restricting the analysis to one lung region. In 2010 Zhao et al. (35,36) demonstrated that by adding small changes of PEEP, the gradient of the GI value indicated the direction of beneficial PEEP alteration. This same parameter was used in Liu et al. (37), where regions with overdistension were defined as those that presented high air levels on end-expiration but no air coming in or out during tidal breathing.

Recently, Eronia et al. (38) studied the feasibility of personalized PEEP selection based on its efficacy in

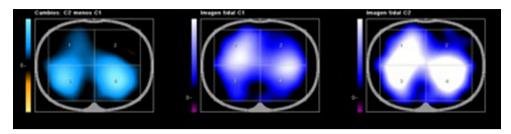


Figure 1 EIT images showing impedance change after recruitment maneuver in patient suffering ARDS. Impedance change defined by colour coding scheme: impedance increase represented as blue-white, whilst impedance decrease represented as purple, and zero as black with a uniform color region around zero. Left side image, reference or basal EIT image; middle image, PEEP 5–10 cmH₂O; right-side image, PEEP 10 cmH₂O. EIT, electrical impedance tomography; ARDS, acute respiratory distress syndrome; PEEP, positive end-expiratory pressure.

stabilizing the EELV increase induced by a recruitment maneuver, using EIT as a tool to monitor EELV changes (parallel to end-expiratory lung impedance changes). Their study protocol consisted in three consecutive steps, comparing PEEP titration using the ARDSnet guidelines and PEEP titration with EIT after applying a continuous positive airway pressure of 40 cmH₂O for 40 s, measuring end-expiratory lung impedance variation (ΔΕΕLΙ) thirty seconds and 10 minutes after the recruitment maneuver; if ΔEELI decreased more than 10% of ΔEELIstart a new recruitment maneuver was performed, and PEEP increased by 2 cmH₂O; and if ΔEELI decreased less than 10% of ΔEELIstart, PEEP was left unchanged. They demonstrated that bedside PEEP setting based on sustained recruitment following a recruitment maneuver as visualized by EIT was achievable in the majority of the patients, leading to application of higher PEEP values and allowing quantification of regional overdistension associated with PEEP increase (*Figures 2,3*).

EIT has also been used to set PEEP for ECMO treated severe ARDS patients, providing real-time monitoring during ultra-protective ventilation with very low tidal volume. Franchineau *et al.* (39) emphasize the importance of preventing cyclic end-expiratory collapse, associated with increased ECMO blood flow, which might unnecessarily prolong ECMO duration and enhance the risk of ECMO-related complications. In the study they evaluate EIT ability to monitor a decremental PEEP trial (20 to 0 cmH₂O) in a 5 cmH₂O steps; EIT-based best compromise PEEP (defined in this case as the lowest pressure able to limit EIT-assessed collapse to under 15% with the least overdistension) was around 10 to 15 cmH₂O, drawing attention to the broad variability in optimal PEEP observed in severe ARDS patients.

Lastly, high flow nasal cannula (HFNC) has been used in hypoxemic respiratory failure due to its physiological mechanisms (administration of high inspired oxygen fraction and creation of positive oropharyngeal airway pressure). HFNC under EIT monitoring has shown improved oxygenation by increasing both end-expiratory lung volume and tidal volume, regardless of body position suggesting an increase in functional residual capacity (40,41) (Figure 4).

So far, great progress has been made. However, even though EIT is becoming a more day-to-day tool regarding mechanical ventilation settings, we must remind that all parameters (severity of illness, hemodynamic tolerance, CT images, respiratory mechanics, ...) must be taken into account when titrating PEEP. According to (42) the multicenter, prospective cohort LUNG-SAFE (Large Observational Study to Understand the Global Impact of Severe Acute Respiratory Failure) study, 53% of ARDS patients were ventilated with a tidal volume of less than 7 mL/kg, and 82.6% of the patients received a PEEP of less than 12 cmH₂O, whilst prone positioning was used in 16.3% of patients with severe ARDS. Mechanical ventilation guidelines (43) suggest higher rather than lower levels of PEEP, taking into account that higher PEEP strategies are not associated with significant differences in barotrauma, new organ failure, ventilator-free days and mortality as compared with a lower PEEP strategy. ARDS still appears to be undertreated and associated with high mortality rate (in the LUNG-SAFE study, mortality as high as 46.1% was observed for those with severe ARDS). PEEP titration based on PaO₂/FiO₂ ratio may not be a correct approximation; a more complete and individually personalized understanding or ARDS lung mechanics and its interaction with the ventilation is needed (44). As

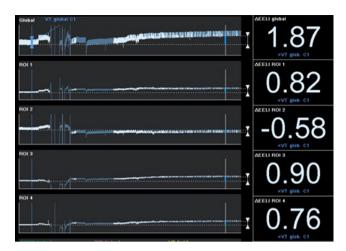


Figure 2 PEEP titration after recruitment maneuver, demonstrating end-expiratory lung impedance changes stabilization (parallel to end-expiratory lung volume increase) after recruitment. PEEP, positive end-expiratory pressure.

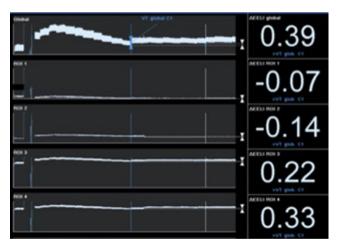


Figure 3 EIT waveforms from a patient suffering septic shock due to peritonitis. The patient was initially ventilated on a pressure-control mode (settings: PEEP 10 mmH₂O, FiO₂ 0.7, PaO₂/FiO₂ ratio 190). A recruitment maneuver was carried out, in a 4 cmH₂O steps every two minutes, until PEEP value reached 18 cmH₂O. A decremental PEEP trial was performed afterwards (using a 2 cmH₂O stepdown) until PEEP value reached 2 cmH₂O (driving pressure was kept on 12 cmH₂O). Tidal volume and oxygen saturation were monitored during incremental and decremental PEEP trials. Optimal PEEP was interpreted as the PEEP value with the best possible compliance value. After both trials, PEEP value was set on 6 cmH₂O; no derecruitment was observed, and driving pressure was decreased to 10 cmH₂O, witnessing a decrease in impedance change on anterior ROI and an increase on posterior ROI. EIT, electrical impedance tomography; PEEP, positive end-expiratory pressure.

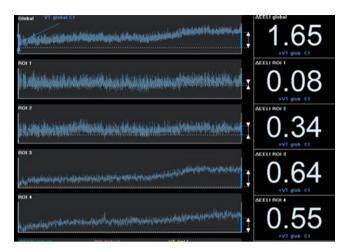


Figure 4 EIT waveforms (global impedance and regions-of-interest waveforms represented as four quadrants: anterior right (ROI 1), anterior left (ROI 2), posterior right (ROI 3), posterior left (ROI 4). Increase in end-expiratory lung impedance variation (ΔΕΕLΙ), and therefore increase of functional residual capacity, after application of high flow nasal cannula, with a gas flow of 60 L/min. EIT, electrical impedance tomography.

Gattinoni *et al.* (45) discuss, in order to set tidal volume and PEEP we must keep in mind hemodynamics and lung mechanics, setting ventilator parameters knowing lung recruitment capacity (avoiding atelectrauma) and keeping the balance with an adequate tidal volume (avoiding overstretching phenomena), optimizing lung size and homogeneity.

Spontaneous breathing trials (SBT) (46-50)

Spontaneous breathing during mechanical ventilation improves lung compliance; patients preserve their end expiratory lung volumes more easily, and avoid redistribution of ventilation to the ventral parts of the lung. Atelectrauma, induced by cyclic collapse and tidal recruitment, appears to be reduced. EIT can also play an important role in demonstrating the advantages of partial ventilatory support. Moreover, EIT is capable of visualizing and quantifying spatial ventilation distribution in real time (49), taking advantage of the GI index to predict in advance whether planned SBT would lead to clinical impairment.

Prone positioning (and other positions)

Current scientific evidence derived from clinical trials and

meta-analysis suggest that ventilation in the prone position should be part of the ventilatory strategy in patients with severe ARDS. EIT allows a close monitoring on changes in ventilation caused by these position changes, and contributes to assess the functional effect on the lung system in response to position changes, particularly when there is a highly unbalanced distribution of ventilation (51-53).

Clinical applications in other situations

Occasionally, abrupt decrease in regional lung ventilation may endanger the patient's life, requiring a rapid therapeutic action to avoid worse outcomes. EIT may guide towards the diagnosis of pneumothorax, atelectasis, malpositioning of the endotracheal tube or pleural effusion (54). Though EIT imaging is not pathognomonic, it provides additional information that, along with clinical findings, contribute to achieve a prompt diagnosis. Costa *et al.* (55) created algorithms for the detection of this pneumothorax using the EIT, demonstrating a sensitivity of 100% for its diagnosis, even in cases of small pneumothorax (under 20 mL).

EIT may be used as a supplementary tool along with conventional pulmonary function tests, as it enables the assessment of both spatial and temporal heterogeneity of reginal function over time or after an intervention (56,57). Furthermore, EIT has also been used to assess patients with cystic fibrosis, identifying regional airway obstruction, and monitoring regional ventilation during breathing aids or physiotherapy (58-60).

Transient lung depressurization during suctioning of secretions or bronchoscopic maneuvers may be evident through EIT monitoring (61,62). Wolf et al. (63) showed that lung volumes decreased after three suctioning maneuvers during clearance of secretions, whilst Lindgren et al. (64) demonstrated that bronchoscopy initially leads to localized auto-PEEP due to the reduced transverse area available for airflow, and that suctioning leads to a decrease in lung aeration and compliance even when a closed suctioning system was used. EIT may even be able to guide lung sampling with non-bronchoscopic bronchoalveolar lavage (especially when bronchoalveolar lavage through bronchoscopy is contraindicated) (65).

Assessment of pulmonary edema often requires invasive measurements, even though they are not always reliable. New technologies can help us in this conundrum, such as lung ultrasound or EIT. In an experimental animal model (66), a novel noninvasive technique using changes in thoracic EIT with lateral body rotation was able to estimate extravascular lung

water. This lung water/EIT ratio may be able to guide fluid management, while minimizing ventilator-induced lung injury.

Even though EIT has been extensively developed in lung monitoring, we must not forget to mention new lines of research, such as the assessment of cardiac function and lung perfusion (67-70). EIT-based perfusion monitoring may allow image of local perfusion and mapping of regional ventilation-perfusion ratios. Some of the approaches described to separate cardiac from respiratory impedance changes are by holding the breath, by use of modeling techniques that may separate both signals while breathing (heartbeat-related EIT signal variations) or by intravenous injection of hypertonic saline (which serves as contract agent due to its much higher conductivity). Nevertheless, clinical validation in this field is still pending.

Limitations and problems to solve

The advantages of pulmonary EIT have been reviewed. However, the constraints and limitations of this technique need to be addressed. Although it is a fairly simple technique to apply, it may pose a risk when placed in patients with spinal injury, thus it is considered a relative contraindication. The belt should not be placed over areas of damaged or inflamed skin. Poor quality EIT images could be obtained from morbid obese patients with body mass index above 50. Measurements of EIT are sensitive to body movements, hence their use in patients with uncontrolled body movements may not be reliable.

Regarding the interference of the EIT and pacemakers or defibrillators, the application of an alternating current could interfere in its functioning. Also, cardiac defibrillations should not be performed at the same time as EIT, since the energy needed for defibrillation can propagate through the electrode-belt and decrease its effectiveness.

EIT measurement encompasses cross-sectional slice of 5-10 cm of thorax, and we assume that other lung regions behave similarly. The clinical use and plausibility of EIT measurements depends then on proper belt position, proper impedance visualization, correct analysis and data interpretation (71).

The images provided translate lung tissue impedance changes, not absolute values. We will always need for a control measure (a reference) taken before or at the beginning of the morbid process to be able to determine or estimate the observed changes. Additionally, EIT captures only ventral-to-dorsal regional ventilation distribution; Bikker *et al.* (72) report different ventilation distribution between cranial and

caudal lung levels during decremental PEEP trials, concluding that no single optimal PEEP level exists for the whole lung.

Conclusions

Despite the possible limitations of the existing EIT systems, this device have meant a remarkable technological advance in the field of lung monitoring and mechanical ventilation adjustments. The final appraisal of the indications is especially promising in the light of the first results, although its implantation on a daily basis will be the result of the scientific community's acquired experience over the years.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

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