Thrombolysis in Massive Pulmonary Embolism Based on the Volumetric Capnography

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This is the first report of a patient submitted to chemical thrombolysis due to massive pulmonary embolism (PE) during the postoperative period of neurosurgery, in whom due to the lack of adequate clinical conditions, no imaging assessment was performed. Clinical, gasometric and capnographic data allowed the decision to perform the thrombolysis with safety. The P(a-et)CO₂ gradient decreased from 46.4 mmHg to 11.8 mmHg (normal < 5 mmHg) and the end-tidal alveolar dead space fraction decreased from 0.85 to 0.37 (normal < 0.15) from the pre-thrombolysis period to the 7th day post-thrombolysis. We conclude that the volumetric capnography (VC) was useful in the patient’s diagnosis and clinical follow-up.

Introduction

Pulmonary embolism (PE) is a frequent disorder, of which diagnostic confirmation is difficult to achieve. National studies have shown that approximately 3% to 5% of the necropsies disclose the presence of emboli in the pulmonary vessels and, in 68% of these cases, it is the cause of death. It is estimated that in 75% of the cases, the diagnosis is not achieved¹ and the standard imaging assessments are not always available, mainly in secondary hospitals. In contrast, the volumetric capnography (VC) is a non-invasive bedside test, which is available even at smaller hospitals. The VC can help diagnose patients with a diagnostic suspicion of PE²-⁴.

This report presents the case of a patient who, in spite of anticoagulant therapy, developed sudden alteration in the hemodynamic parameters presumably due to massive PE. Considering the rapid deterioration of the hemodynamic state, although the diagnosis of PE was not confirmed by imaging assessment, the patient was submitted to chemical thrombolysis. The VC was performed before the thrombolysis and for seven subsequent days.

Key words
Pulmonary embolism; capnography; thrombolytic therapy.

Case Report

A 22-year-old female patient, admitted at the Intensive Care Unit (ICU) of a tertiary hospital developed difficult-to-control diabetes insipidus after the excision of a frontal brain tumor (astrocytoma), together with septic conditions and mild hemodynamic alterations. On the 24th postoperative day, the patient was going through the mechanical ventilation (MV) weaning process, with the following parameters: FiO₂ = 0.30; spontaneous breathing (30 rpm), PEEP = 5 cmH₂O; pHa = 7.50; PaO₂ = 62.2 mmHg; PaCO₂ = 27.6 mmHg; HCO₃⁻ = 21.8 mmol/l; BE = -0.1 mmol/l; SatO₂ = 94.8%; PaCO₂/FiO₂ ratio = 207.

The patient suddenly started to present respiratory difficulty, hypoxemia and hemodynamic worsening. With the diagnostic hypothesis of massive PE, a transthoracic echocardiogram was performed, which disclosed moderate dilatation of the right chambers, moderate tricuspid regurgitation and pulmonary artery systolic pressure = 50 mmHg. The echocardiogram (ECHO) showed the presence of S, Q₁, T₂ pattern.

After this episode, the MV was adjusted to: FiO₂ = 1; SIMV (12/38 rpm); PEEP = 8 cmH₂O and TV = 500 ml; the subsequent arterial gasometry showed: pH = 7.26; PaO₂ = 44.5 mmHg; PaCO₂ = 54.6 mmHg; HCO₃⁻ = 23.7 mmol/l; BE = -2.7 mmol/l; SatO₂ = 71.7%; PaO₂/FiO₂ ratio = 44.5. Dobutamine was started, aiming at the hemodynamic function improvement. Three hours later, while still on mechanical ventilation, the patient presented the following parameters: FiO₂ = 1; SIMV (14/27 rpm); PEEP = 10 cmH₂O; TV = 500 ml, arterial gasometry showed: pH = 7.22; PaO₂ = 50.7 mmHg; PaCO₂ = 54.7 mmHg; HCO₃⁻ = 21.3 mmol/l; BE = -6.2 mmol/l; SatO₂ = 82.3%; PaCO₂/FiO₂ ratio = 50.7.

Through the VC, the end-tidal CO₂ pressure (PetCO₂) was determined (CO₂SMO PLUS 8100 Dixtal/Novametrix®) which, associated to the arterial gasometry (Radiometer ABL 700™), allowed the calculation of different derived indices, such as end-tidal alveolar dead space fraction (AVDSL), the late dead space fraction (fDSL), the CO₂ arterial-alveolar gradient [Pa(et)CO₂]; and the slope of phase III of the CO₂ spirogram (Slp III).

Considering the patient’s rapid deterioration, imminent risk of death and VC values indicative of increase in the alveolar dead space - compatible with PE²-⁷ - the medical team, after the family had given the informed consent, chose to institute the thrombolytic treatment with alteplase (IV 100 mg/2 hours).

Discussion

A recent systematic literature review⁸ concluded the
bleeding is the most frequent complication of thrombolysis, of which risk is around 6-20%, with intracranial bleeding being the most feared complication. Thus, the risk of bleeding is what defines the contraindications for thrombolysis. These relative contraindications are recent active bleeding or intracranial disease (less than 6 weeks), trauma, visceral biopsy, gastrointestinal bleeding, coagulation disorders, kidney or liver failure, puncture of vessel not susceptible to compression and pericarditis. As the patient presented severe hemodynamic instability HR = 168 bpm, MAP = 52 mmHg, SpO\textsubscript{2} = 83% and FiO\textsubscript{2} = 1), she did not present the adequate clinical conditions for transportation to another institution in order to undergo image assessment (helicoidal angiotomography). However, the patient was assessed by VC, which disclosed values suggestive of PE\textsuperscript{2-7}.

The capnography estimates the value of the alveolar dead space\textsuperscript{2-7}. The association between the VC results with those of the arterial gasometry allows the calculation of several indices, from which variables the extension of the alveolar dead space can be inferred and, consequently, the presence and extension of the occluded areas of the pulmonary artery system\textsuperscript{2-7}.

The following capnographic parameters were determined:
1. PetCO\textsubscript{2};
2. Slp III;
3. P(a-et)CO\textsubscript{2} (normal value < 5 mmHg);
4. AVDSf, calculated by the formula: PaCO\textsubscript{2} - PetCO\textsubscript{2} / PaCO\textsubscript{2}, where PetCO\textsubscript{2} is end-tidal CO\textsubscript{2} \textsuperscript{7} (normal value < 0.15);
5. fDlate, obtained from the extrapolation of the current expired volume at 15% of the estimated total pulmonary capacity (TPC): fDlate = PaCO\textsubscript{2} - Pet(15\% TPC) / PaCO\textsubscript{2} \textsuperscript{7} (normal value < 0.12).

During an embolic event, the ventilation/perfusion ratio imbalance is accentuated and, as a consequence, the variables that express the alveolar dead space also present alterations\textsuperscript{2-7}. The higher the calculated value, the higher the degree of vascular network obstruction, and thus, the more extensive the alveolar dead space is, considering a correlation between the extension of the area without perfusion and the obtained value\textsuperscript{2-7}. The studied variables tended to normalization after the thrombolytic treatment, indicating the occurrence of vessel rechanneling.

The capnographic variables presented the following behavior:
- PetCO\textsubscript{2} (reference value: 36.7 ± 3.7 mmHg\textsuperscript{7}): 8.2 (pre-thrombolysis); 13.1 (after 24h); 17.7 (after 48h); 16.4 (after 72h); 19.9 (after 96h); 19.0 (after 120h); 20.3 (after 144h); 20.4 mmHg (after 168h);
- Slp III (reference value: 7.5 ± 2.4 mmHg/l\textsuperscript{7}): 0.26 (pre-thrombolysis); 0.77 (after 24h); 3.96 (after 48h); 8.77 (after 72h); 7.62 (after 96h); 5.07 (after 120h); 6.6 (after 144h); 8.47mmHg/l (after 168h);
- fDlate\textsuperscript{7}: 0.85 (pre-thrombolysis); 0.68 (after 24h); 0.54 (after 48h); 0.55 (after 72h); 0.39 (after 96h); 0.38 (after 120h); 0.41 (after 144h); 0.31 (after 168h).

To illustrate that (Figures 1 and 2), two capnographic variables muras [P(a-et)CO\textsubscript{2} and AVDSf] are demonstrated, which, when associated to arterial gasometry can be easily obtained through any capnographer available in the market and performed at the bedside and at any hospital.

In this dramatic case report, one can observe that, when strictly applied, VC can be useful as a diagnostic tool and in the therapeutic follow-up of patients that cannot be submitted to imaging assessment. The VC also showed to be a useful tool in decision-making at the initial phase of PE assessment and O CV mresolution, as the progressive
pulmonary reperfusion resulted in a decrease of the alveolar dead space and thus, a tendency towards the normalization of the capnographic variables.

One can speculate that the increase in the dead space was due to the presence of PEEP. However, Blanch et al. showed that there is no alteration in the dead space when the PEEP is at the levels observed in the present study. In this case, all VC measurements were carried out with a PEEP of 5 cmH$_2$O.

After the thrombolytic therapy and the stabilization of the hemodynamic picture, a head computed tomography was carried out, which did not disclose bleeding. The patient was discharged from the ICU and subsequently, from the hospital.

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References


