Case report

Diaphragm pacing failure secondary to deteriorated chest wall mechanics: When a good diaphragm does not suffice to take a good breath in

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A B S T R A C T

Diaphragm pacing allows certain quadriplegic patients to be weaned from mechanical ventilation. Pacing failure can result from device dysfunction, neurotransmission failure, or degraded lung mechanics (such as atelectasis). We report two cases where progressive pacing failure was attributed to deteriorated chest wall mechanics. The first patient suffered from cervical spinal cord injury at age 45, was implanted with a phrenic stimulator (intrathoracic), successfully weaned from ventilation, and permanently paced for 7 years. Pacing effectiveness then slowly declined, finally attributed to rib cage stiffening due to ankylosing spondylitis. The second patient became quadriplegic after meningitis at age 15, was implanted with a phrenic stimulator (intradiaphragmatic) and weaned. After a year hypoventilation developed without obvious cause. In relationship with complex endocrine disorders, the patient had gained 31 kg. Pacing failure was attributed to excessive mechanical inspiratory load. Rib cage mechanics abnormalities should be listed among causes of diaphragm pacing failure and it should be kept in mind that a “good diaphragm” is not sufficient to produce a “good inspiration”.

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Introduction

Diaphragm pacing allows patients suffering from high cervical spinal cord lesions who are quadriplegic and ventilator-dependent to be separated from their mechanical ventilator. Diaphragm pacing has clinical benefits and reduces health costs [1–5]. All of the currently available diaphragm pacing techniques rely on a similar principle: electrodes are implanted in the vicinity of the phrenic nerve (either intrathoracically during a video-assisted mini-thoracotomy [6] -hook wire electrodes-) and connected to an external stimulator that provides a modular stimulation current. After a post-implantation reconditioning period aimed at correcting diaphragmatic disuse atrophy [7], successful diaphragm pacing generally results in stable tidal volumes that are obtained with stable stimulation intensities [6]. Deteriorated ventilation can result from device dysfunction, neuromuscular issues including neurotransmission failure, or degraded respiratory mechanics (e.g. airway obstruction, atelectasis, etc.). These events generally translate in rapid onset clinical, spirometric and blood gases abnormalities. We report two observations of quadriplegic patients in whom a previously fully efficient diaphragm pacing slowly lost efficacy, resulting in full time or part time return to mechanical ventilation, without any of the usual mechanisms of diaphragm pacing failure being identified. In both cases, major changes in the mechanical properties of the chest wall appeared as plausible explanations of diaphragm pacing failure.
One year later, tidal volumes had fallen to less than 250 mL and the fracture, mainly via the intrathoracic approach. Approximately 50 patients have been implanted with phrenic stimulators, mostly via the intrathoracic approach.

Case #1
This patient suffered from a high cervical spinal injury (C1/C2 fracture, traffic accident) resulting in flaccid quadriplegia and full ventilator dependency in May 2000, aged 45. Diagnostic phrenic nerve stimulation demonstrated bilateral diaphragm responses, allowing diaphragm pacing. In May 2001, the patient was implanted with intrathoracic phrenic stimulators (Atrostim™, Atrotech, Finland). After 4 weeks of diaphragm reconditioning, complete ventilator weaning was obtained, the patient rapidly using diaphragm pacing 24 h a day and then doing so for several years. Decreased pacing efficiency was first noted in 2007, with the need to increase stimulation intensities to maintain tidal volume at the level needed to keep alveolar ventilation adequate (650–700 mL). The situation slowly deteriorated over the following years, with the appearance of dyspnoea in the sitting position, decreased tidal volumes, and partial failure by total loss of ventilatory autonomy leading to nocturnal, and then permanent ventilator dependency in 2012 (tidal volume 200–250 mL). Phrenic nerve function was verified several times, always with normal phrenic nerve conduction times and ample, and symmetrical diaphragm action potentials. A progressive decrease in tracheal pressure and abdominal movements following phrenic nerve stimulation was noted. During this course, device replacement, aggressive correction of malnutrition and aggressive urologic management of chronic urinary tract infection (both liable to be responsible of muscle weakness and/or neuromuscular transmission impairment) were conducted without modifying the course of diaphragm pacing deterioration. In May 2011, high-resolution chest computed tomography revealed widespread vertebral fusion suggestive of ankylosing spondylitis (Fig. 1). This diagnostic had indeed been established prior to the accident having led to quadriplegia (HLA B-27 positive), the patient reporting the occasional use of non-steroidal anti-inflammatory drugs to control intermittent back pain before he became quadriplegic. It was concluded that the disease had evolved over years, the patient not perceiving pain because of the spinal lesion, and diaphragm pacing failure was attributed to thoracic ankylosis preventing diaphragm contractions to adequately inflate the rib cage. The patient died in May 2012 of severe pneumonia.

Case #2
At age 15, this patient suffered from meningococcal meningitis complicated by acute cerebral oedema leading to compressive spinal cord ischaemia. Magnetic resonance imaging did not show spinal cord lesions below C3. Diagnostic phrenic nerve stimulation demonstrated bilateral diaphragm response, allowing diaphragm pacing. In September 2009, the patient was implanted with intradiaphragmatic phrenic stimulators (NeurRXDPS™, Synapse Biomedical, Oberlin, OH, USA). After nine months (July 2010), diaphragm pacing was considered efficient, producing tidal volumes of 500 mL on average and achieving normocapnia (PaCO₂ 39 mmHg). One year later, tidal volumes had fallen to less than 250 mL and the patient was hypercapnic (PaCO₂ 51 mmHg) in spite of maximal stimulation intensities (25 mA). Phrenic stimulation induced-thoracic expansion had become almost undetectable. No device dysfunction was identified. Phrenic nerve conduction was normal, as was tracheal twitch pressure in response to bilateral phrenic stimulation. Over the same time frame, complex hormonal disorders led to massive weight gain (from 67 to 104 kg, body mass index 22.6–35.1 kg/m²), with an ulcerotic diagnostic of Klinefelter syndrome. Diaphragm pacing failure was attributed to excessive chest wall and abdominal weight preventing diaphragm contractions to adequately inflate the rib cage.

Discussion
These two observations point at alterations of chest wall compliance as a mechanism of diaphragm pacing failure that has seemingly not been described before. In both cases, extensive workup failed to find any technical dysfunction of the stimulators, and any patent abnormality of the phrenic nerves. In the first case, malnutrition of chronic infection could have been responsible for diaphragmatic dysfunction, but their aggressive correction did not improve the situation. Chronic diaphragm fatigue could also be discussed, but the patient had been paced continuously between 2000 and 2007 without any reduction in ventilation. In the hypothesis of diaphragm fatigue, improvement should have been observed when the patient was returned to nocturnal mechanical ventilation.

In ankylosing spondylitis, vertebral fusion (Fig. 1), responsible for the typical "bamboo spine" aspect, is associated with sternoclavicular and sternocostal ankylosis (present in our patient), and at times with costovertebral joints abnormalities [8]. These abnormalities are naturally bound to deteriorate the transformation of diaphragm strength into rib cage movement. In addition, the coordinated action of the diaphragm and abdominal muscles is an important determinant of the compensation of rib cage stiffness in ankylosing spondylitis [9]. In our patient, the lack of any abdominal muscle activity probably aggravated the impact of ankylosis on tidal volume production. Massive obesity is associated with a restrictive ventilatory defect and decreased rib cage compliance [10]. This corresponds to an elastic load which, to be overcome, requires the respiratory muscles to increase the work that they develop [11] in response to an augmented neural drive [12]. Although this has apparently not been specifically studied, the conjunction of obesity and respiratory muscle weakness is bound to aggravate the deleterious effects of rib cage mechanical deterioration on the production of ventilation. In spite of diaphragm pacing, our second patient represents an extreme case of respiratory weakness, because of the absence of co-contraction of thoracic muscles with the diaphragm and of the absence of any possibility to compensate the inspiratory load by active expiration. Diaphragm pacing also prevent the extension of diaphragm activity over expiration that has been described as a feature of respiratory load compensation in obesity [13].

Conclusion
In conclusion, our observations are practical reminders that the inspiratory action of the diaphragm does not only depend on diaphragm contractile properties, but also of a diaphragm “contractile environment” that includes diaphragm geometry, abdominal compliance [14,15], and rib cage compliance. It is also a reminder that breathing is intimately dependent on, and interferes with, spinal cord and costovertebral joints mechanics [16]. That a “good diaphragm” is not sufficient to produce a “good inspiration” must be kept in mind when managing patients with diaphragm pacing.
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Patients consent

The patients admitted to the department of respiratory and critical care of the Pitié-Salpêtrière hospital are systematically informed that their medical data can be used, anonymously, for research and publication purposes and are given the option to agree or refuse. The two patients described in this case report agreed to this use.

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