# The Timed Inspiratory Effort: A Promising Index of Mechanical Ventilation Weaning for Patients With Neurologic or Neuromuscular Diseases

Leonardo Cordeiro de Souza PhD PT, Fernando Silva Guimarães PhD PT, and Jocemir Ronaldo Lugon PhD MD

BACKGROUND: The aim of this study was to evaluate the performance of the recently described timed inspiratory effort (TIE) index in comparison with 4 other previously reported indices as to the weaning outcome in patients with neurologic or neuromuscular disorders. METHODS: This observational prospective study included subjects undergoing weaning from mechanical ventilation. The performance of the indices was evaluated by calculation of the area under the receiver operating characteristic curves. The areas under the curve were compared using the Hanley and McNeil method. P values < .05 were considered significant. RESULTS: Seventy-two subjects (57  $\pm$  20 y old) were selected for the study. Forty-three subjects were weaned, and 21 died during the study period. The mean duration of mechanical ventilation was  $22.3 \pm 19.4$  d. The areas under the curve of 5 weaning predictors (TIE index, integrative weaning index, noninvasive tension-time index, maximum inspiratory pressure, and breathing frequency/tidal volume index) were significantly higher than those of the other indices. The TIE index had the largest area under the curve  $(0.96 \pm 0.02)$  in comparison with the integrative weaning index (0.82  $\pm$  0.05, P = .009), noninvasive tension-time index (0.80  $\pm$  0.05, P = .001), maximum inspiratory pressure  $(0.77 \pm 0.06, P = .001)$ , and breathing frequency/tidal volume index ( $0.72 \pm 0.06$ , P = .001). CONCLUSIONS: In patients with neurologic or neuromuscular impairment, the TIE index had a better performance than the best weaning indices used in clinical practice. Key words: mechanical ventilation; weaning; maximum inspiratory pressure; respiratory muscle; critical care. [Respir Care 2015;60(2):231–238. © 2015 Daedalus Enterprises]

## Introduction

Previous studies have reported that ventilator weaning and extubation failures in subjects with neurologic and neuromuscular diseases are related to severity of the injury.<sup>1,2</sup> Moreover, lower Glasgow coma scores on admission and/or lower pulmonary capacity and inadequate toilet have been associated with increased duration of ventilation.<sup>1</sup>

Weaning decisions based solely on expert clinical judgment are not always correct.<sup>3</sup> Previous reports suggest that clinical judgment based on traditional respiratory parameters and the level of neuromuscular impairment<sup>4-7</sup> may prevent unnecessary delay of withdrawal from mechanical ventilation and mitigate complications such as diaphragm atrophy and weakness.<sup>8,9</sup>

A number of procedures to verify patients' readiness for weaning have been proposed.<sup>10</sup> Currently, the 3 weaning readiness assessments should include the following steps: measurement of weaning predictor indices, unassisted breathing trial (ie, low-pressure support or T-tube trial), and extubation trial.<sup>11</sup>

Dr de Souza is affiliated with the Medical Science Post-graduation Program, and Dr Lugon is affiliated with the Nephrology Division, Department of Medicine, Fluminense Federal University, Niterói, Rio de Janeiro, Brazil. Dr Guimarães is affiliated with the Physiotherapy School, Federal University of Rio de Janeiro, Rio de Janeiro, Brazil.

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Correspondence: Leonardo Cordeiro de Souza PhD PT, Icaraí Hospital, Rua Marquês de Paraná 233, 5° Andar, CEP 24030-215, Centro, Niterói, Rio de Janieiro, Brazil. E-mail: leonardo.uti@gmail.com.

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Spontaneous ventilation is dependent on the drive from the central nervous system, the capacity of the respiratory muscles, and the load imposed on them. Factors most likely to be predictive of extubation outcome are those assessing the effectiveness of breathing efforts and/or the load of the respiratory muscles.<sup>12</sup> In fact, univariate indices that examine a single aspect of physiologic function often have poor predictive power probably because they do not fully reflect all the pathophysiologic processes affecting weaning outcome.<sup>13-15</sup> Accordingly, predictions that use multivariate indices that integrate a number of physiologic functions are more accurate.<sup>13-17</sup>

On reviewing the literature for ventilator weaning, 5 indices demonstrated enough accuracy for predicting the weaning outcome: frequency/tidal volume  $(f/V_T)$ ,<sup>10,13</sup> maximum inspiratory pressure with unidirectional valve  $(P_{Imax(UV)})$ ,<sup>10,13,16,18,19</sup> integrative weaning index (IWI),<sup>16</sup> tension-time index of inspiratory muscles  $(TT_{mus})$ ,<sup>14,17</sup> and timed inspiratory effort (TIE).<sup>19</sup>

Unfortunately, conventional weaning predictive indices have less accuracy in patients with neurologic or neuromuscular diseases.<sup>20</sup> We hypothesized that the recently reported TIE index, which combines the ability to generate an inspiratory pressure with the time needed to achieve P<sub>Imax</sub>, could perform better in such a group of patients. Thus, the objective of this study was to compare the predictive performance of the TIE index with the 4 best previously described weaning indices in patients with neurologic or neuromuscular diseases.

## Methods

## Subjects and Settings

This study was conducted in the ICU of a general hospital. The ICU is a closed 33-bed unit attended by 4 physicians certified and trained in critical care, two 24-h respiratory physiotherapists, 2 senior nurses, and nurse assistants (at a ratio of one for 3 patients).

The study was approved by the ethics committee of the Fluminense Federal University (number 259/09). Informed consent was obtained from each subject, whenever possible, or from the subject's next-of-kin. We utilized the same database used in a previous report,<sup>19</sup> complementing the analysis focusing on subjects with neurologic or neuro-muscular diseases.

All mechanically ventilated subjects who were clinically stable and judged ready to undergo a weaning trial by their primary physicians and respiratory physiotherapists were enrolled in this prospective observational study. Inclusion criteria included: 18 y or older, a neurologic or neurosurgical condition, and mechanical ventilation for > 24 h. Subjects were accepted for enrollment if they met the American Thoracic Society/European Respiratory So-

## QUICK LOOK

### Current knowledge

Weaning and extubation failures in patients with neurologic and neuromuscular disorders are associated with increased disease severity. The timed inspiratory effort (TIE) index combines the ability to generate an inspiratory pressure with the time needed to achieve the maximum inspiratory pressure ( $P_{Imax}$ ). The role of the TIE index in predicting weaning success in neurologic and neuromuscular disease has not been studied.

## What this paper contributes to our knowledge

In a small group of subjects with neurologic disease or neuromuscular impairment, TIE was superior to the breathing frequency/tidal volume ratio,  $P_{Imax}$ , and integrative weaning index in predicting weaning success. Large trials are required before adoption of this predictive index.

ciety 2007 guidelines to start a weaning trial.<sup>10</sup> These criteria were as follows: resolution of the acute phase of the disease for which the subject was intubated, adequate cough reflex or absence of excessive tracheobronchial secretions, stable cardiovascular status (heart rate of  $\leq 120$  beats/min and systolic arterial blood pressure of 90–160 mm Hg, with no or minimal use of vasopressors), stable metabolic status, adequate oxygenation (arterial oxygen saturation [S<sub>aO2</sub>] of > 90% on F<sub>IO2</sub>  $\leq 0.4$  or P<sub>aO2</sub>/F<sub>IO2</sub>  $\geq 200$  mm Hg with PEEP  $\leq 8$  cm H<sub>2</sub>O), adequate ventilatory pattern (breathing frequency of  $\leq 35$  breaths/min) with pressure support of  $\leq 16$  cm H<sub>2</sub>O, no significant respiratory acidosis (pH > 7.30), and adequate mentation or a Glasgow coma score of > 10 (only for subjects with an endotracheal tube).

Exclusion criteria were as follows: tracheal stenosis, intracranial pressure of > 20 mm Hg, sedation, overt cardiac failure or hemodynamic instability, and signs of systemic infection/re-infection during the weaning process. The Taema eXtend ventilator (Air Liquide, Gentilly, France) was used.

#### **Study Procedures**

Discontinuation from mechanical ventilation was attempted when the physician in charge judged that the subject was ready to be weaned according to the aforementioned criteria. Before testing, all subjects were under pressure support ventilation (ranging from 5 to 16 cm  $H_2O$ ), with PEEP varying from 5 to 8 cm  $H_2O$ . The subjects were not sedated (they were all between +1 and -1 on the Richmond Agitation-Sedation Scale<sup>21</sup>) and were allowed to breathe spontaneously for 10 s before performing the test.<sup>22-23</sup> They were in a semirecumbent position (45°), with their airways suctioned, and were pre-oxygenated for 2 min with an F<sub>IO<sub>2</sub></sub> of 1.0.<sup>22-23</sup> The endotracheal tube cuff was hyperinflated to prevent air leakage during measurements.<sup>22-23</sup>

First, the variables that could be registered under mechanical ventilation were obtained. Second, spirometry was performed, and the subject was returned to mechanical ventilation. After stabilization, subjects were connected to a vacuometer and a unidirectional valve. After measurements of all indices, subjects were to left to recover for 5-10 min and underwent a 30-min spontaneous breathing trial with a T-piece at a  $F_{IO_2}$  of 0.4. All subjects were continuously monitored by pulse oximetry and electrocardiography, with continuous respiratory physiotherapist attention. The following 5 weaning indices were evaluated:  $P_{Imax(UV)}$ , f/V<sub>T</sub>, IWI, TT<sub>mus</sub>, and TIE.

The f/V<sub>T</sub> index was measured with a bedside spirometer (Wright MK-20, Ferraris Medical, Hertford, United Kingdom) attached to an artificial airway for 1 min. The spontaneous V<sub>T</sub> was calculated by dividing the minute ventilation by the breathing frequency, and f/V<sub>T</sub> was calculated by dividing the breathing frequency by V<sub>T</sub> (average of 60 s, in L).<sup>13</sup>

The IWI uses the static compliance of the respiratory system (C<sub>RS</sub>), S<sub>aO<sub>2</sub></sub>, and f/V<sub>T</sub>: IWI = static C<sub>RS</sub> × S<sub>aO<sub>2</sub></sub>/(f/V<sub>T</sub>). The static C<sub>RS</sub> was calculated as described by Aboussouan et al<sup>24</sup> using the volume and pressure readout on the ventilator display. An inspiratory hold of 2 s was used to record the plateau pressure (P<sub>plat</sub>), and the static C<sub>RS</sub> was calculated by dividing the V<sub>T</sub> by the difference between the inspiratory P<sub>plat</sub> and PEEP: static C<sub>RS</sub> = V<sub>T</sub>/P<sub>plat</sub> – PEEP).<sup>16,19</sup>

 $TT_{mus}$  is obtained noninvasively and assesses the inspiratory muscle function by the following equation:  $TT_{mus} = (P_I/P_{Imax(UV)}) \times (T_I/T_{tot})$ , where  $P_I$  is inspiratory pressure estimated as  $5 \times P_{0.1} \times T_I$ ,<sup>25,26</sup>  $T_I$  is inspiratory time, and  $T_{tot}$  is the total respiratory cycle time obtained from the mechanical ventilation parameters during the testing period.<sup>14,17,25,26</sup>

The TIE index was recorded using the unidirectional valve method and calculated as the ratio between  $P_{Imax(UV)}$  and the corresponding time to achieve it in 60 s of observation.<sup>19</sup>  $P_{Imax(UV)}$  and respiratory drive ( $P_{0,1}$ ) were obtained using a digital vacuometer (MVD 300, GlobalMed, Porto Alegre, Rio Grande do Sul, Brazil). This device has a range of 300 cm H<sub>2</sub>O with a 1 cm H<sub>2</sub>O interval and is designed to perform measurements every 100 ms. Measurements were digitally recorded, allowing easy access to the parameter values whenever needed.<sup>18,19,23</sup> For this study,  $P_{Imax(UV)}$  was taken as the peak value in the first 20 s of observation, and  $P_{0,1}$  as the mean of the first 3 respiratory

cycles.<sup>18,19,23</sup> Subjects were not coached for the maneuver.<sup>18,19,22,23</sup>

The decision to return to mechanical ventilation was made by a respiratory physiotherapist and the physician in charge (both were blinded to the results of the indices evaluated) based on the signs of poor tolerance described below.<sup>8,10,16,27,28</sup> Weaning was considered successful if spontaneous breathing was sustained for > 48 h after withdrawal from mechanical ventilation (through the tracheostomy tube for tracheostomized subjects and after extubation for those who were orally intubated).<sup>10,16,19,23,27,28</sup>

The trial was stopped if at least one of the following intolerance criteria was present:  $S_{aO_2} < 90\%$  or  $P_{aO_2} < 60$  mm Hg with  $F_{IO_2} > 0.4$ ;  $P_{aCO_2} > 50$  mm Hg or increased by  $\geq 8$  mm Hg; arterial pH < 7.33 or decreased by 0.07 or more; breathing frequency > 35 breaths/min or increased by 50% for 5 min or longer; heart rate > 140 beats/min or a sustained increase or decrease of > 20%; mean blood pressure > 130 or < 70 mm Hg; or presence of agitation, diaphoresis, disorientation, or depressed mental status.<sup>10,16,19,23,27-29</sup> Subjects who demonstrated one of these signs during the spontaneous breathing trial or within 48 h after discontinuation from mechanical ventilation were considered to have failed weaning and were returned to ventilatory support.<sup>8,10,16,27,29</sup> Any death occurring within 48 h of weaning was also computed as a failure.

## **Statistical Analysis**

Continuous variables are expressed as mean  $\pm$  SD, and categorical variables as frequencies. Comparisons between groups were performed by the Mann-Whitney test or chisquare test as appropriate. The end point of the study was successful weaning. The performance of the indices to predict weaning outcome was evaluated by calculation of sensitivity, specificity, predictive values, and likelihood ratios. The areas under the receiver operating characteristic (ROC) curves, which are built integrating sensitivity and specificity, were also obtained; thresholds for the 5 weaning indices were calculated using the Youden method.<sup>30</sup> Areas under the curves were compared using the method of Hanley and McNeil.<sup>30</sup> P < .05 was considered significant.

Statistical analysis was performed employing SPSS 18.0 (SPSS, Chicago, Illinois). Comparison between ROC curves was accomplished using MedCalc 11.4.2.0. (Mariakerke Software, Mariakerke, Belgium).

### Results

From the 990 patients admitted to the ICU of the hospital between July 2011 and July 2012, 67 were eligible for the study (Fig. 1). Subjects who were successfully



Fig. 1. Flow of subjects in the study indicating stratification of the groups according to weaning outcome and exclusions.

weaned were in group 1, and those who failed the weaning trial were in group 2.

All subjects in groups 1 and 2 used orotracheal or tracheostomy tubes with internal diameters of 7.5–9.0 mm. Twenty-one subjects (31%) died before discharge, including 6 among the successfully weaned subjects.

The subjects' characteristics and the differences between the groups are provided in Table 1. The conditions precipitating ICU admissions were brain trauma (22 cases, 68% subjects with tracheostomies), stroke (27 cases, 66% subjects with tracheostomies), acute myopathy (19 cases, 89% subjects with tracheostomies), and cervical medullary injury (4 cases, 100% subjects with tracheostomies). Of the subjects with stroke, 23 had cerebral ischemia, 2 had intracerebral hemorrhage, and 2 had subarachnoid hemorrhage. Of the acute myopathy subjects, 6 had critical illness polyneuropathy, 3 had Guillain-Barré syndrome, 2 had polymyositis, one had cerebral palsy (as a sequel of meningitis), one had porphyria, one had inclusion body myositis, one had Parkinson disease, and 2 had myasthenia gravis.

Among the monitored parameters during the test, an increase in breathing frequency was the most common reason for returning subjects to mechanical ventilation. The mean values of the measured parameters used to calculate the weaning predictor indices as well as the indices themselves in both groups are provided in Table 2. Quality indicators and the cutoff values calculated from the ROC curves of the 5 weaning indices (TIE, IWI,  $TT_{mus}$ ,

 $P_{Imax(UV)}$ ), and  $f/V_T$ ) are provided in Table 3. The IWI exhibited a slightly higher sensitivity than the remaining ones, but the TIE index showed a better performance in every other quality indicator. The area under the curve of the 5 indices and the statistics between them are provided in Figure 2. The largest area under the curve was found for the TIE index (0.96 ± 0.02), followed by IWI (0.82 ± 0.05), TT<sub>mus</sub> (0.80 ± 0.05), P<sub>Imax(UV)</sub> (0.77 ± 0.06, *P* = .001), and  $f/V_T$  (0.72 ± 0.06). The values for the TIE index were statistically higher than those for every other index. A post hoc analysis was performed to calculate the area under the ROC curve for each index, separating subjects who had tracheostomies (Table 4). Again, the highest areas under the presence of a tracheostomy.

#### Discussion

The use of predictive weaning indices is one of the most studied phenomena in intensive care.11 However, their performance in patients with neuromuscular diseases has been found to be poor.<sup>12,16,19</sup> In this study, we evaluated the performance of 5 selected weaning indices in subjects with critical neurologic or neuromuscular diseases. In our sample, the weaning failure rate (39%) was greater than that in the literature, which is  $\sim 30\%$ .<sup>10</sup> This finding was not unexpected considering that we dealt with subjects with neurologic or neuromuscular diseases, who are more likely to have complications. Accordingly, our subjects had a prolonged duration of mechanical ventilation (22 d), a high tracheotomy rate (75%), a high death rate in the ICU (31%), and a high APACHE II score (17.9). Subjects who failed the weaning trial had longer ventilation times, higher tracheotomy rates, greater ICU mortality, and less brain trauma.

The mean of every tested respiratory parameter was statistically different between successful and failed weaning subjects (see Table 2). Cutoff points of the weaning indices provided in Table 3 were obtained by ROC curve analysis using the Youden index. The TIE index and  $P_{Imax(UV)}$  maintained the previously published cutoff values of 1 and 30,<sup>19</sup> whereas the other weaning indices did not.<sup>13,16,17</sup>

The previously mentioned specificities of the study group may in part account for these differences. Numbers regarding sensitivity, negative predictive values, and likelihood ratios of negative tests were comparable among the various weaning indices. In contrast, specificity, positive predicted values, and likelihood ratios of positive tests were markedly superior for the TIE index compared with the other weaning tests.

To better explore the relative performance of the weaning tests, ROC curves for each tested index were generated (see Fig. 2). The area under the curve of the TIE index was

#### Table 1. General Characteristics of Subjects

		Wea	Weaning		
	All Subjects	Success	Failure	P	
Subjects, $n$ (%)	72	43 (60)	29 (40)	Group 2 vs group 1	
Males/females, n	48/24	30/13	18/11	.43	
Blacks/whites, n	47/25	31/12	16/13	.10	
Age (mean $\pm$ SD), y	$57.3 \pm 20$	$56.7 \pm 18.9$	$58.3 \pm 22.4$	.39	
Mechanical ventilation (mean $\pm$ SD), d*	$22.3 \pm 19.4$	$19.9 \pm 20.6$	$26.5 \pm 17.1$	.02	
OTT/Trach, n	18/54	15/28	3/26	.04	
Death in ICU, $n$ (%)	21 (31)	6 (9)	15 (22)	.08	
APACHE II score (mean $\pm$ SD)	$17.9 \pm 6.4$	$17.1 \pm 5.5$	$19.3 \pm 7.5$	.21	
Conditions precipitating ICU admission, n (%)					
Brain trauma	22 (31)	17 (39)	5 (17)	.02	
Stroke	27 (39)	17 (40)	10 (35)	.25	
Acute myopathy	19 (25)	7 (16)	12 (41)	.36	
Cervical medullary injury	4 (5)	2 (5)	2 (7)	.62	
* Until the first weaning trial OTT = Subjects with ≤ 14 d of mechanical ventilation and orotrache	al tube				

Trach = Subjects with > 14 days of mechanical ventilation and tracheostomized

APACHE II = Acute Physiology and Chronic Health Evaluation II

Table 2.	Respiratory	Parameters	According	to Weaning	Outcome
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Parameter	Success, Mean $\pm$ SD (n = 43)	Failure, Mean $\pm$ SD (n = 29)	<i>P</i> *
P <sub>0.1</sub>	$3.02 \pm 0.94$	$2.60 \pm 0.69$	.03
P <sub>Imax(UV)</sub>	$39.26 \pm 21.45$	$22.93 \pm 11.28$	< .001
TIE index	$1.55\pm0.80$	$0.70\pm0.20$	< .001
f/V <sub>T</sub>	$75.32 \pm 29.44$	$94.00 \pm 22.97$	< .001
IWI	$59.34 \pm 22.08$	$37.23 \pm 18.51$	< .001
TT <sub>mus</sub>	$0.18\pm0.09$	$0.39\pm0.22$	< .001
$T_I/T_{tot}$	$0.43\pm0.17$	$0.55\pm0.18$	< .001
P <sub>I</sub>	$14.33 \pm 5.30$	$13.84 \pm 4.16$	.82
$P_I/P_{Imax(UV)}$	$0.41\pm0.10$	$0.69\pm0.29$	< .001
TI	$0.95\pm0.16$	$1.07\pm0.21$	.01
f	$26.40 \pm 4.71$	$30 \pm 5.83$	< .001
V <sub>T</sub>	$0.36\pm0.07$	$0.33\pm0.07$	.03

\* Nonparametric Mann-Whitney test

 $P_{0.1} =$  respiratory drive

P<sub>Imax(UV)</sub> = maximum inspiratory pressure with unidirectional valve

- $f/V_T$  ratio = breathing frequency/tidal volume ratio
- IWI = integrative weaning index
- TT<sub>mus</sub> = tension-time index of inspiratory muscles
- $T_I/T_{tot}$  = inspiratory time/total cycle time ratio
- P<sub>I</sub> = inspiratory pressure

significantly higher than those of the other tested weaning indices. The finding was confirmed when subjects with or without a tracheostomy were analyzed separately.

Historically, the consensus of the American Thoracic Society/European Respiratory Society is that the indices  $f/V_T$  and  $P_{Imax}$  are the most frequently used to indicate if patients are ready to undergo a weaning process.<sup>10</sup> Recent

studies have shed light on the clinical utility of new weaning indices: the  $TT_{mus}$  exhibited a surprisingly performance with an area under the curve of 1.0 in children with neurologic disorders<sup>17</sup>; the IWI showed an area under the curve of 0.96 in adult patients without neurologic diseases<sup>16</sup>; and the TIE index revealed an area under the curve of 0.90 in a general population of adults in the ICU. The results of the last 3 indices are superior to the better results reported for  $f/V_T$  (area under the curve = 0.89) and  $P_{Imax(UV)}$  (area under the curve = 0.80) in this setting.<sup>13,19</sup> In our study, the TIE index was found to have the best performance in predicting weaning outcome.

The TIE index critically evaluates how fast a patient can generate  $P_{Imax}$ , which is dependent on central stimuli and muscle power.<sup>19</sup> Our hypothesis was that TIE index reflects neuromuscular performance and can be particularly useful in identifying ready-to-wean patients in the class of subjects in our study. Our results allow us to suggest the TIE index as an eligible predictor of weaning in those patients for whom the performance of other indices such as  $f/V_T$  and  $P_{0.1}$  is recognized to be poor.<sup>20,27,31</sup> Indeed, the well-known low accuracy of other indices was perhaps the reason why this class of patients was used as an exclusion criterion in a previous study.<sup>16</sup>

Because this is the first investigation on the reliability of the TIE index for subjects with neurologic or neuromuscular diseases, the external validity of these results is limited by the small sample size. Therefore, the clinical application of our findings should await further studies with larger samples. It should also be noted that our study population represents a heterogeneous collection comprising

TIE = timed inspiratory effort

Index	Cutoff	Sensitivity	Specificity	Positive Predictive Value	Negative Predictive Value	Positive Likelihood Ratio	Negative Likelihood Ratio
TIE	> 1.00	0.86	0.96	97.3	80.6	22.4	0.14
IWI	> 37	0.90	0.69	83.7	79.2	3.26	0.17
TT <sub>mus</sub>	$\leq 0.27$	0.83	0.73	82.9	73.1	3.08	0.23
f/V <sub>T</sub>	≤ 93	0.85	0.61	77.8	72.7	2.22	0.24
P <sub>Imax(UV)</sub>	> 30	0.60	0.82	0.84	0.62	3.51	0.48

Table 3. Quality Indicators of Indexes Used to Predict Weaning Outcome

Cutoff values corresponds to the maximal Youden index.

TIE = timed inspiratory effort

IWI = integrative weaning index

 $TT_{mus}$  = tension-time index of inspiratory muscles  $f/V_T$  = frequency/tidal volume ratio

 $P_{Imax(\mathrm{UV})}$  = maximum inspiratory pressure with unidirectional valve



	f/V <sub>T</sub>	P <sub>Imax(UV)</sub>	TTmus	IWI
TIE	.0001	.0001	.001	.009
IWI	.02	.78	.75	
TTmus	.26	.54		
P <sub>Imax(UV)</sub>	.57			

Fig. 2. Area under the receiver operating characteristic curves of the 4 predictors of successful weaning in this study: timed inspiratory effort (TIE), integrative weaning index (IWI), tension-time index of inspiratory muscles ( $TT_{mus}$ ), maximum inspiratory pressure with unidirectional valve ( $P_{Imax(UV)}$ ), and frequency/tidal volume ratio ( $f/V_T$ ). *P* values for each comparison are provided.

Table 4.	Performance of Weaning Indexes as Evaluated by Areas
	Under the Receiver Operating Characteristic Curves in
	Subjects With Orotracheal Tubes or Tracheostomies

Index	Orotracheal Tube, Mean $\pm$ SD (n = 18)	Tracheostomy. Mean $\pm$ SD (n = 54)
TIE	$1.00 \pm 0.09$	$0.95 \pm 0.03$
TT <sub>mus</sub>	$0.91 \pm 0.09$	$0.75\pm0.06$
P <sub>Imax(UV)</sub>	$0.84 \pm 0.09$	$0.75\pm0.06$
IWI	$0.75 \pm 0.12$	$0.84\pm0.05$
$f/V_T$	$0.71\pm0.12$	$0.72\pm0.07$

TIE = timed inspiratory effort

TT<sub>mus</sub> = tension-time index of inspiratory muscles

 $P_{Imax(UV)} = maximum \ inspiratory \ pressure \ with \ unidirectional \ valve$ 

IWI = integrative weaning index f/V<sub>T</sub> ratio = frequency/tidal volume ratio

brain trauma, stroke, and neuromuscular diseases. Heterogeneous sedation levels due to residual effects of drugs could have a role in this regard, but the impact was attenuated by the regular use of the Richmond Agitation-Sedation Scale.<sup>21</sup> We think that it would be desirable to assess the performance of these indices in each one of these specific groups in the future. Despite these limitations, our results are encouraging and suggest that the TIE index can be a useful tool in the clinical management of patients whose weaning readiness is difficult to predict.

#### Conclusions

In summary, the TIE index was found to be useful in predicting successful weaning in subjects with critical neurologic diseases, and it performed better than the other weaning indices tested in this setting. We hope that regular utilization of this index might attenuate the burden of a wrong decision for extubation or unnecessary prolongation of mechanical ventilation in such patients.

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