Stress Index Can Be Accurately and Reliably Assessed by Visually Inspecting Ventilator Waveforms

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Data Supplement

Contents

- Figure E1 to E3: Off-line dynamic airway pressure-time (Paw-t) curve profile analysis and stress index (SI) classification by dedicated software
- 2. Figure E4 Example of unqualified Paw-t curve
- Table E1 Off-line software determined stress index and classification in the first 50 assessments
- Table E2 Accuracy of visual stress index classification among different reference stress index classifications and among different ventilator brands
- 5. Table E3 Contingency table for diagnostic accuracy analysis
- 6. Training manual for visual classification of stress index



Figure E1 Identification of the constant flow segment

ICU-Lab 2.5 Software Package (KleisTEK, Bari, Italy) was used in off-line dynamic airway pressure-time (Paw-t) curve profile analysis. The software identified the beginning and the end of recorded breath by a threshold value in the flow tracing (0.2 L/sec). The constant flow segment was identified as the fluctuation was within \pm 3% of the steady value. To eliminate the influence of on-flow and off-flow transients, the segment was further narrowed for 50 ms after the beginning and before the end of the constant flow portion. If a constant segment could not be found or the length of selected segment was less than one-third of the entire inspiratory phase, data were discarded but documented.





After confirming the qualified constant flow segment, Paw-t curve within the corresponding segment was automatically fitted by the equation:

 $Paw = a \times time^b + c$

The R values were calculated automatically and if lower than 0.95, the data were discarded but documented. In the equation, a represents the slope of Paw-t curve as time equal to 1 s; c is the airway pressure as time of 0 s; and b is a dimensionless parameter named as SI, which reflects the shape of the Paw-t curve and is related to respiratory system elastance (Ers). Assuming that the airway resistive and viscoelastic properties remain invariant, the change of slope along the Paw-t curve is mostly contributed to the change of Ers during tidal inspiration.

Reference:

Ranieri VM, Zhang H, Mascia L, Aubin M, Lin CY, Mullen JB, et al. Pressure-time curve predicts minimally injurious ventilatory strategy in an isolated rat lung model. Anesthesiology 2000;93(5):1320-1328.



Figure E3 Paw-t curve shape classification according to the SI value

Paw-t curve shape can be classified into three categories:

- an SI between 0.9 and 1.1 corresponding to a linear shape of qualified Paw-t curve segment that describes an unchanged Ers with time;
- 2) an SI < 0.9 corresponding to an upward concavity shape that represents a decreasing Ers with time
- an SI > 1.1 corresponding to a downward concavity shape that represents an increasing Ers with time.

Reference:

Terragni P, Bussone G, Mascia L. Dynamic airway pressure-time curve profile (Stress Index): a

systematic review. Minerva Anestesiol 2016;82(1):58-68.



Figure E4 Example of unqualified Paw-t curve

Panel a is the example of unqualified Paw-t curve analyzed by the software. The unstable flow at the end inspiratory caused that the length of constant flow segment (0.32s) was less than one-third of the inspiratory phase (1.02s). Simultaneously, the Paw during the unstable flow increased obviously. Panel b shows the same unqualified Paw-t curve shown on the ventilator screen. We did not find the unstable flow during on-site visual inspection. At the bedsides, an abrupt elevation in Paw-t waveform might suggest an unstable inspiratory flow pattern.

SI value	No. of assessments	No1	No2	No3	No4	No5	No6	No7	No8	No9	No10
	single breath	0.81	0.90	0.95	0.98	1.04	0.86	0.79	0.92	0.95	1.11
	2-min mean value	0.76	0.90	0.94	1.07	1.03	0.86	0.87	0.91	0.99	1.12
classification	single breath	downward concavity	linear	linear	linear	linear	downward concavity	downward concavity	linear	linear	upward concavity
	2-min mean value	downward concavity	linear	linear	linear	linear	downward concavity	downward concavity	linear	linear	upward concavity
	No. of assessments	No11	No12	No13	No14	No15	No16	No17	No18	No19	No20
SI value	single breath	0.97	1.01	1.03	0.92	0.78	0.91	1.00	0.95	1.18	0.94
	2-min mean value	1.08	1.01	1.03	0.92	0.78	0.90	0.91	0.91	1.16	0.91
classification	single breath	linear	lincor	lineer	lineer	downword concevity	lincor	lineer	lineer	unword concernity	lineer
	2 min mean value	linear	linear	linear	linear	downward concavity	linear	linear	linear	upward concavity	linear
	2-min mean value	linear	mica	inical	lineai	downward concavity	inical	linear	inical	upward concavity	inicai
SI value	No. of assessments	No21	No22	No23	No24	No25	No26	No27	No28	No29	No30
	single breath	0.89	0.93	0.95	0.96	0.81	0.97	0.94	0.87	1.07	1.02
	2-min mean value	0.87	0.93	0.99	0.94	0.79	0.98	0.94	0.84	1.05	1.05
classification	single breath	downward concavity	linear	linear	linear	downward concavity	linear	linear	downward concavity	linear	linear
clussification	2-min mean value	downward concavity	linear	linear	linear	downward concavity	linear	linear	downward concavity	linear	linear
	No. of assessments	No31	No32	No33	No34	No35	No36	No37	No38	No39	No40
SI value	single breath	0.99	1.04	0.85	0.88	0.88	1.05	1.05	0.87	0.88	0.98
	2-min mean value	1.05	1.06	0.81	0.88	0.88	1.09	1.03	0.89	0.89	0.95
classification	single breath	linear	linear	downward concavity	downward concavity	downward concavity	linear	linear	downward concavity	downward concavity	linear
	2-min mean value	linear	linear	downward concavity	downward concavity	downward concavity	linear	linear	downward concavity	downward concavity	linear
						-				-	
	No. of occorrecto	No.41	Na 42	N-42	No.44	No.45	No.46	No.47	NI-49	No.40	No 50
SI value	ino. Of assessments	11041	0.04	0.02	0.96	0.97	1.12	11047	1 11	0.07	1.21
	Single breath	1.10	0.94	0.93	0.80	0.82	1.15	0.90	1.11	0.97	1.21
	2-min mean value	1.14	0.92	0.93	0.88	0.88	1.14	0.91	1.15	0.91	1.28
classification	single breath	upward concavity	linear	linear	downward concavity	downward concavity	upward concavity	linear	upward concavity	linear	upward concavity
	2-min mean value	upward concavity	linear	linear	downward concavity	downward concavity	upward concavity	linear	upward concavity	linear	upward concavity

Table E1 Off-Line Software Determined Stress Index and Classification in the First 50 Assessments

In the first 50 off-line analyses, the stress index (SI) from the single breath and the 2-min mean value before the single breath were both measured in the off-line analysis. The SI was classification into three types: an SI between 0.9 and 1.1 corresponding to a linear shape; an SI < 0.9 corresponding to a downward concavity shape; an SI > 1.1 corresponding to a upward concavity shape. An identical SI classification was found in each assessment.

stress mach clussifications and among arreferer ventilator oranas						
		Accuracy (95% confidential interval)				
Reference stress index	Linear	94% (89%-98%)				
	Downward concavity	91% (83%-99%)				
	Upward concavity	95% (79%-100%)				
	p	0.20				
Ventilator brands	Servo-I	91% (85%-98%)				
	AVEA	92% (88%-97%)				
	PB 840	93% (86%-100%)				
	p	.0.68				

Table E2 Accuracy of visual stress index classification among different referencestress index classifications and among different ventilator brands

		Reference SI classification	Total	
		Downward concavity	Linear	
	Downward concavity	50	2	52
	Linear	5	123	128
	Total	55	125	180
		Upward concavity	Linear	
uo	Upward concavity	19	6	25
sificati	Linear	1	119	120
Visual SI class	Total	20	125	145
		Combined downward and upward concavity	bined downward Linear pward concavity	
	Combined downward and upward concavity	69	8	77
	Linear	6	117	123
	Total	75	125	200

 Table E3 Contingency table for diagnostic accuracy analysis

Training Manual for Visual Classification of Stress Index by Inspecting the Ventilator Screen Waveforms

Version 2017.02.23

Contents

- Fundamental principle of airway pressure-time (Paw-t) curve profile analysis and its clinical use;
- Standard method for off-line stress index (SI) fitting by ICU-Lab software and classification;
- 3) Method for visual SI classification by inspecting the flowtime and Paw-t waveforms on the ventilator screen.

1) Airway pressure-time curve profile analysis and stress index

Lung-protective ventilation, mainly composed of small tidal volume, low plateau and driving pressures and adequate positive end-expiratory pressure (PEEP), has been recommended in mechanically ventilated patients (E1). Evidences suggest that the assessment of elastic properties of the respiratory system is necessary to identify the proper settings of the ventilator and minimize ventilator-induced lung injury (E2).

Since first introduced in 2000 (E3), analysis of dynamic airway pressure-time (Paw-t) curve profile has been investigated to differentiate tidal recruitment and hyperinflation in animal experiments (E3-E8) and clinical studies (E9-E12). The shape of Paw-t curve within the constant inspiratory flow section is related to the change of respiratory system elastance (Ers). A linear, downward and upward concavity shape indicates the Ers remaining constant, decreasing or increasing during tidal inspiration, respectively (E13, E14). Fitting of the Paw-t curve also yields a dimensionless coefficient, i.e. stress index (SI), quantitatively describing the shape of the curve. This method is considered as a non-invasive and real-time respiratory mechanics monitoring at the bedside.

Experiments in pig model with acute lung injury showed that an SI < 0.9 indicated tidal recruitment and an SI > 1.1 indicated tidal hyperinflation (E6, E7). Histological and inflammatory evidences also revealed that markers of injurious ventilation were minimized when ventilator was set to maintain SI range of 0.9 to 1.1 (E3-E5). In patients with acute respiratory distress syndrome, titrating PEEP to obtain SI between 0.9 and 1.1 attenuated alveolar hyperinflation (E9) and improved recruitment volume (E10), lung elastance and oxygenation (E9).

2) Airway pressure-time curve analysis by ICU-Lab software

For off-line Paw-t curve analysis, flow and airway pressure data are collected. A heated Fleisch pneumotachograph (VitalographInc, Lenexa, KS, USA) is placed between the Y-piece of the ventilator circuit and the endotracheal tube to measure the flow. A KT 100D-2 pressure transducer (KleisTEK, Bari, Italy) is connected proximally to the endotracheal tube to measure the airway pressure. Pressure transducer and Fleisch pneumotachograph are connected to an ICU-Lab Pressure Box by 80 cm rigid tube lines. The pressure transducer is calibrated by a water column, and the pneumotachograph by a 1-L calibration syringe (SN: 554-2266, Hans Rudolph, Inc. Shawnee, KS, USA). Signals are continuously displayed and saved (ICU-Lab 2.5 Software Package, KleisTEK, Bari, Italy) on a laptop computer for further analysis, at a sample rate of 200 Hz.

Pressure transducer and Fleisch pneumotachograph





Downloaded data are analyzed by ICU-Lab.



During off-line analysis, the software identifies the beginning and the end of recorded breath by a threshold value in the flow tracing (0.2 L/sec). The constant flow segment is identified as the fluctuation is within \pm 3% of the steady value. To eliminate the influence of on-flow and off-flow transients, the segment is further narrowed for 50 ms after the beginning and before the end of the constant flow portion. If a constant segment cannot be found or the length of selected segment is less than one-third of the entire inspiratory phase, data are discarded.

After confirming the qualified constant flow segment, Paw-t curve within the corresponding segment is automatically fitted by the equation:

```
Airway pressure=a \times time^b + c
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The *R* values will be calculated automatically and if lower than 0.95, the data are discarded. In the equation, *a* represents the slope of Paw-t curve as time equal to 1 s; *c* is the airway pressure as time of 0 s; and *b* is a dimensionless parameter named as SI, which reflects the shape of the Paw-t curve and is related to Ers (E3, E6). Assuming that the airway resistive and viscoelastic properties

remain invariant, the change of slope along the Paw-t curve is mostly contributed to the change of Ers during tidal inspiration.



Paw-t curve shape can be classified into three categories (E11):

- an SI between 0.9 and 1.1 corresponding to a linear shape of qualified Paw-t curve segment that describes an unchanged Ers with time;
- an SI < 0.9 corresponding to a downward concavity shape that represents a decreasing Ers with time
- an SI > 1.1 corresponding to an upward concavity shape that represents an increasing Ers with time.



3) Visual classification of stress index by inspecting the flow-time and Paw-t waveforms on the ventilator screen

The middle point on the constant inspiratory flow is identified (red dot). Then the corresponding point on the Paw-t waveform is confirmed (red circle). Thereafter, a tangent line (red line) passing through the point on the Paw-t waveform is marked using a ruler. The relationship of this tangent line and Paw-t waveform is visually inspected.





The relationship of the tangent line marked by ruler and Paw-t waveform can be classified into three types:

1) the Paw-t waveform almost coincident with the tangent line indicates an SI value between 0.9 and 1.1, and is judged as linear shape (**A**); 2) the two side of Paw-t waveform both deviating downward from the tangent line indicates an SI < 0.9 and is judged as downward concavity shape (**B**); or 3) the two side of Paw-t waveform both deviating upward from the tangent line indicates SI > 1.1 and is judged as upward concavity shape (**C**).

Training tests

For the ongoing slides, please use the method for visual inspection and discriminate the shape of Paw-t waveform shown on the ventilator screen as linear, downward or upward concavity shape.

Please use the full screen function in your computer during the estimation. You can use the ruler to measure the computer screen.

- Sex: M
- Age: 56
- Diagnosis: ARF
- Ventilator settings
 - RR:20
 - VT (ml): 500
 - PEEP (cmH2O): 15
 - FiO₂: 80%



- Sex: M
- Age: 65
- Diagnosis: ARF
- Ventilator settings
 - RR:15
 - VT (ml): 450
 - PEEP (cmH2O): 12
 - FiO₂: 50%



- Sex: F
- Age: 60
- Diagnosis: Glioma
- Ventilator settings
 - RR:20
 - VT (ml): 500
 - PEEP (cmH2O): 8
 - FiO₂: 40%



- Sex: M
- Age: 50
- Diagnosis: Postoperative
- Ventilator settings
 - RR:25
 - VT (ml): 600
 - PEEP (cmH2O): 5
 - FiO₂: 30%



- Sex: M
- Age: 51
- Diagnosis: TBI
- Ventilator settings
 - RR:20
 - VT (ml): 500
 - PEEP (cmH2O): 8
 - FiO₂: 50%



- Sex: M
- Age: 58
- Diagnosis: ARF
- Ventilator settings
 - RR:20
 - VT (ml): 540
 - PEEP (cmH2O): 18
 - FiO₂: 50%



- Sex: F
- Age: 48
- Diagnosis: SAH
- Ventilator settings
 - RR:20
 - VT (ml): 500
 - PEEP (cmH2O): 12
 - FiO₂: 40%



- Sex: M
- Age: 38
- Diagnosis: Glioma
- Ventilator settings
 - RR:20
 - VT (ml): 350
 - PEEP (cmH2O): 5
 - FiO₂: 40%


- Sex: F
- Age: 60
- Diagnosis: TBI
- Ventilator settings
 - RR:20
 - VT (ml): 350
 - PEEP (cmH2O): 6
 - FiO₂: 50%



- Sex: M
- Age: 28
- Diagnosis: TBI
- Ventilator settings
 - RR:25
 - VT (ml): 450
 - PEEP (cmH2O): 10
 - FiO₂: 40%



- Sex: M
- Age: 42
- Diagnosis: Postoperative
- Ventilator settings
 - RR:20
 - VT (ml): 500
 - PEEP (cmH2O): 5
 - FiO₂: 40%



- Sex: F
- Age: 72
- Diagnosis: SAH
- Ventilator settings
 - **- RR**:18
 - VT (ml): 500
 - PEEP (cmH2O): 6
 - FiO₂: 40%



- Sex: F
- Age: 32
- Diagnosis: ARF
- Ventilator settings
 - RR:25
 - VT (ml): 520
 - PEEP (cmH2O): 8
 - FiO₂: 40%



- Sex: M
- Age: 54
- Diagnosis: TBI
- Ventilator settings
 - **- RR**:18
 - VT (ml): 540
 - PEEP (cmH2O): 15
 - FiO₂: 50%



- Sex: M
- Age: 63
- Diagnosis: SAH
- Ventilator settings
 - **- RR**:18
 - VT (ml): 580
 - PEEP (cmH2O): 9
 - FiO₂: 30%



- Sex: F
- Age: 50
- Diagnosis: TBI
- Ventilator settings
 - RR:20
 - VT (ml): 470
 - PEEP (cmH2O): 10
 - FiO₂: 40%









- Sex: M
- Age: 53
- Diagnosis: Glioma
- Ventilator settings
 - **- RR**:18
 - VT (ml): 560
 - PEEP (cmH2O): 5
 - FiO₂: 30%



- Sex: M
- Age: 58
- Diagnosis: TBI
- Ventilator settings
 - **–** RR:16
 - VT (ml): 520
 - PEEP (cmH2O): 8
 - FiO₂: 40%







- Sex: M
- Age: 40
- Diagnosis: postoperative
- Ventilator settings
 - **–** RR:16
 - VT (ml): 560
 - PEEP (cmH2O): 6
 - FiO₂: 30%



- Sex: F
- Age: 65
- Diagnosis: SAH
- Ventilator settings
 - RR:25
 - VT (ml): 400
 - PEEP (cmH2O): 6
 - FiO₂: 40%



ICU-Lab pictures for each patients

Patient NO1 SI=1.01 linear shape



Patient NO2 SI=0.96 linear shape



Patient NO3 SI=1.00 linear shape



Patient NO4 SI=0.98 linear shape



Patient NO5 SI=0.94 linear shape



Patient NO6 SI=1.19 upward concavity shape



Patient NO7 SI=1.20 upward concavity shape



Patient NO8 SI=0.82 downward concavity shape



Patient NO9 SI=0.82 downward concavity shape



Patient NO10 SI=0.98 linear shape



Patient NO11 SI=1.05 linear shape


Patient NO12 SI=0.96 linear shape



Patient NO13 SI=0.91 linear shape



Patient NO14 SI=1.02 linear shape



Patient NO15 SI=1.00 linear shape



Patient NO16 SI=1.04 linear shape



Patient NO17 SI=1.04 linear shape



Patient NO18 SI=0.96 linear shape



Patient NO19 SI=0.97 linear shape



Patient NO20 SI=0.83 downward concavity shape



	Linear shape	Downward concavity shape	Upward concavity shape
Numbers	15 (75)	3 (15)	2 (10)
SI (range)	0.91-1.05	0.82-0.83	1.19-1.20

The characters of 20 assessments

Data are presented as number of patients (%)

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