

## ***Comparison of SpO<sub>2</sub>/FiO<sub>2</sub> ratio, Oxygenation index, ventilator ratio and SpO<sub>2</sub>/PaCO<sub>2</sub> ratio, SpO<sub>2</sub>/PEEP ratio with PaO<sub>2</sub>/FiO<sub>2</sub> ratio in critically ill patients***

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### **ABSTRACT**

**Comparison of SpO<sub>2</sub>/FiO<sub>2</sub> ratio, Oxygenation index, ventilator ratio and SpO<sub>2</sub>/PaCO<sub>2</sub> ratio, SpO<sub>2</sub>/PEEP ratio with PaO<sub>2</sub>/FiO<sub>2</sub> ratio in critically ill patients.**

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Trend towards noninvasive, easy to use monitor was always a challenge. Numerous indices have been used to monitor the progress of patients on positive pressure ventilation. The present study compares different indices in a mixed larger intensive care unit (ICU) population. In a prospective observational study Arterial blood gases (ABG) analyses were obtained from 225 patients under mechanical ventilation in a polyvalent adult ICU. Values of ideal body weight (IBW), Body mass index (BMI), PAO<sub>2</sub>, PaO<sub>2</sub>/FiO<sub>2</sub> ratio (PFR), SpO<sub>2</sub>/FiO<sub>2</sub> ratio (SFR), SpO<sub>2</sub>/PEEP ratio (SPR), SpO<sub>2</sub>/PaCO<sub>2</sub> ratio (SPCR), Oxygenation index(OI) and Ventilatory ratio (VR) were calculated; and further correlation analysis was conducted. In Pressure control ventilation mode a relative strong relation between PFR and SFR and OI was found; yet further regression analysis implies that no direct replacement of PFR with SFR can be made without limitations, in clinical setting. In Volume control ventilation mode moderate relation was found between SFR and PFR. In the present study a moderate relation was found between SFR and

PFR. The results agree with previous published studies; the differences among them lie in the different design of each one of them. The authors believe that- given the fact that one still considers using broadly PFR as index of oxygenation- SFR can be used safely as a surrogate for PFR only for certain disease states. Larger series are needed in order to define those patients groups and these pathophysiological conditions.

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## INTRODUCTION

Measures of oxygenation are traditionally used to monitor the progress of patients on positive pressure ventilation. Numerous indices have been such as the arterial to alveolar O<sub>2</sub> difference, the intrapulmonary shunt fraction, the oxygen index and the PaO<sub>2</sub>/FiO<sub>2</sub> ratio, the oxygenation index, the ventilatory ratio, etc. Of these different indices the PaO<sub>2</sub>/FiO<sub>2</sub> ratio has been adopted for routine use because of its simplicity. This ratio is included in the Lung Injury Score<sup>1</sup>, the American–European Consensus Conference ARDS Definition<sup>2</sup> and the ARDS Berlin definition<sup>3</sup>. Yet, even this index has problems such as the need of an arterial line in situ, the PEEP dependence and the nonlinear behavior in relation with FiO<sub>2</sub> changes<sup>4</sup>.

Several authors have proposed different, often less invasive ways to monitor oxygenation or ventilation in patients in ALI. Such indices are SpO<sub>2</sub>/FiO<sub>2</sub> (SFr)<sup>5</sup>, the already popular in pediatric populations Oxygenation Index (OI) (which is defined as  $\text{FiO}_2 / (\text{Mean Airway Pressure} / \text{PaO}_2)$ <sup>6</sup> and the Ventilatory ratio (VR). The latter compares actual measurements and predicted values of minute ventilation and PaCO<sub>2</sub> and it is defined as  $(\text{MV}_{\text{measured}} \times \text{PaCO}_{2\text{ measured}}) / (\text{MV}_{\text{predicted}} \times \text{PaCO}_{2\text{ measured}})$ , where  $\text{MV}_{\text{predicted}}$  is taken 100 ml.kg<sup>-1</sup>.min<sup>-1</sup> based on predicted body weight and PaCO<sub>2measured</sub> is taken 37.5mmHg (5 kPa)<sup>7</sup>. Predicted Body Weight (kg) is calculated using the formula 50+0.91 (centimetres of

height–152.4) for males, and 45.5+0.91 (centimetres of height–152.4) for females (ARDS Network calculator)<sup>8-9</sup>.

A previous smaller study identified a strong relation between PFr, SFr and OI but not VR not only in ALI patients; suggesting that these markers may be used interchangeably as bedside indices of oxygenation in critically ill patients<sup>10</sup>. The aim of the present study is to compare the aforementioned indices along with the SpO<sub>2</sub>/PaCO<sub>2</sub> ratio (SPCr) and SpO<sub>2</sub>/PEEP ratio (SPr) in a mixed larger ICU population.

## MATERIAL AND METHODS

During an 11 month period a prospective observational study was carried out in a polyvalent adult ICU. Since this was an observational study and arterial blood samples were taken as part of their standard therapy, there was no need for an approval of regional ethics committee. Arterial blood gases (ABG) analyses were obtained from 225 patients under mechanical ventilation: Pressure or Volume Control. Two independent measurements were taken from each patient under the same mode of ventilation. Demographic data (age, sex, body weight and BSA), APACHE score II, diagnosis, mode of ventilation and values of PaO<sub>2</sub>, PaCO<sub>2</sub>, FiO<sub>2</sub>, mean airway pressure-Pm, peak airway pressure- Pp, PEEP, Compliance (C), expiratory Resistance (R<sub>e</sub>), minute ventilation (MV), respiratory frequency (RR), tidal vol-

ume (Vt) and hemoglobin concentration (Hb) were recorded. Exclusion criteria were: age<18yr old, presence of chronic pulmonary disease (obstructive or restrictive), mechanical ventilation duration<24h, unstable clinical status 3h before and 3h after the time of measurement.

Values of ideal body weight (IBW), Body mass index (BMI), PAO<sub>2</sub>, PaO<sub>2</sub>/FiO<sub>2</sub> ratio (PFr), SpO<sub>2</sub>/FiO<sub>2</sub> ratio (SFr), SpO<sub>2</sub>/PEEP ratio (SPr), SpO<sub>2</sub>/PaCO<sub>2</sub> ratio (SPCr), Oxygenation index (OI) and Ventilatory ratio (VR) were then calculated using MS Office Excel 2007.

Further statistical analysis (SPSS v.19 - IBM SPSS Inc., USA) included comparison of the two modes of ventilation, descriptive analysis and Shapiro-Wilk W test for normality of the data of every parameter. Relationships between the different indices were also explored. In case of a strong correlation (>0.80) and when applicable (assumption of normality), a regression analysis was performed for certain indices. Finally, residual analysis control of the prediction equation was conducted.

## RESULTS

The mean age of all patients included in the study was 61.2 years old and the mean APACHE score was 15.4±1.8. Mean characteristics of the two groups are shown in table 1.

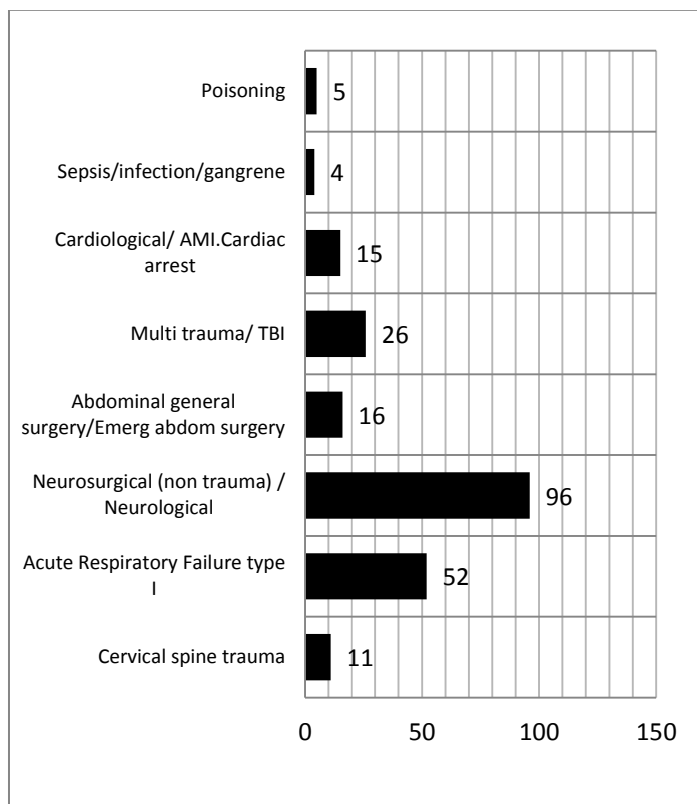
**Table 1.** Characteristics of the two groups

Mode	APACHE II *	Age	Height *	Weight *	Hb *	P peak *	RR*	Compliance static*	P mean
PCV	15.9 ± 2.3	62.9 ± 15.3	176.1 ± 10.5	83.2 ± 12.7	27. 6± 4.6	29.4 ± 4.2	12.9 4 ± 4.2	47.7 ± 16.8	15.3 ± 2.4
VCV	14.4 ± 1.7	57.9 ± 18	172.8 ± 11.0	81.1 ± 15.2	27 ± 5.2	27.6 ± 4.3	11.8 ± 2.8	49 ± 11.8	13.7 ± 1.9

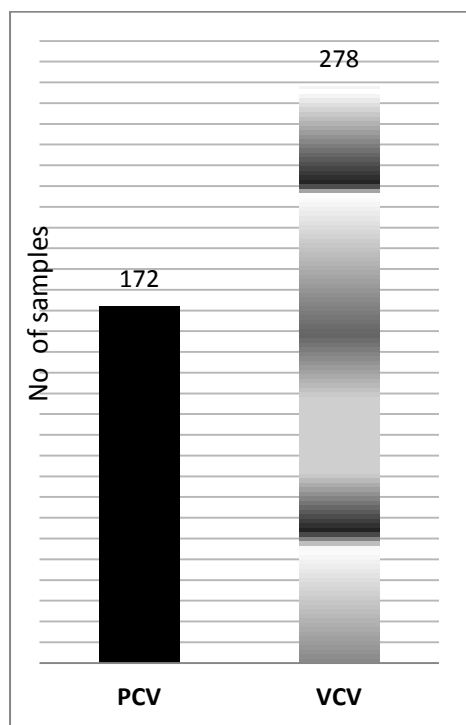
Values are mean ± SD, \*p>0.05

The distribution of the patients according to diagnosis and the distribution of the ABG samples based on the mode of mechanical ventilation are displayed in Figure 1.

**Figure 1.** A) Distribution of patients based on diagnosis B) Distribution of ABG samples based on mode of ventilation.



A



**Table 2.** Descriptive statistics of selected parameters (rounded to two decimals).

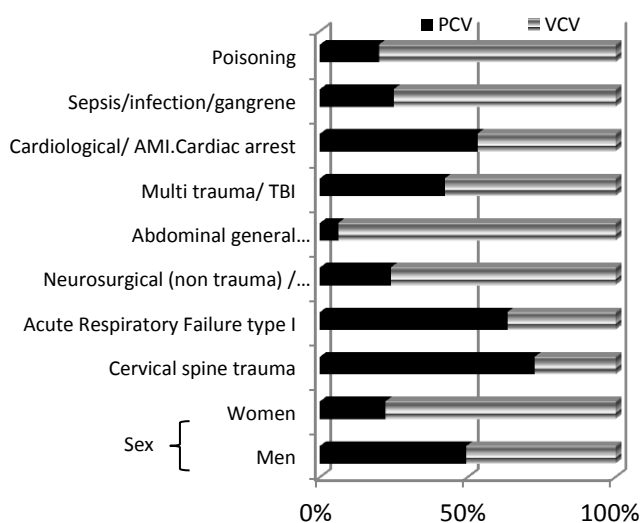
	PEEP		SpO <sub>2</sub> /FiO <sub>2</sub> % ratio		PaO <sub>2</sub> /FiO <sub>2</sub> % ratio		Oxygenation Index	
	PCV	VCV	PCV	VCV	PCV	VCV	PCV	VCV
<b>Mode</b>								
<b>Mean</b>	9.65	6.9	168.1	211	212.5	310	9.33	4.28
<b>SEM</b>	0.21	0.13	5.95	3.99	10.56	10.86	0.61	0.17
<b>Median</b>	10	7	163.3	240	200.4	291	7.78	4.05
<b>SD</b>	1.9	1.53	55.19	46.56	97.98	126.1	5.72	2.01

SEM: standard error of mean, SD: standard deviation.

**B**

Distribution of patients based on sex, diagnosis and mode of ventilation is shown in Figure 2. Descriptive statistics of the seven parameters of interest are displayed in table 2.

**Figure 2.** Distribution of patients by diagnosis, sex and mode of ventilation.



	Ventilatory Ratio		SpO <sub>2</sub> /PEEP ratio		SpO <sub>2</sub> /PaCO <sub>2</sub> ratio	
	PCV	VCV	PCV	VCV	PCV	VCV
<b>Mode</b>						
<b>Mean</b>	1.62	1.35	10.67	15.02	2.38	2.62
<b>SEM</b>	0.057	0.030	0.27	0.28	0.05	0.03
<b>Median</b>	1.64	1.30	9.58	14.14	2.37	2.60
<b>SD</b>	0.53	0.35	2.55	3.35	0.49	0.37

SEM: standard error of mean, SD: standard deviation

Correlations between the parameters of interest are displayed in table 3. Coefficients (Spearman or Pearson) are chosen according to normality test results. Different relations are noticed for different modes of ventilation

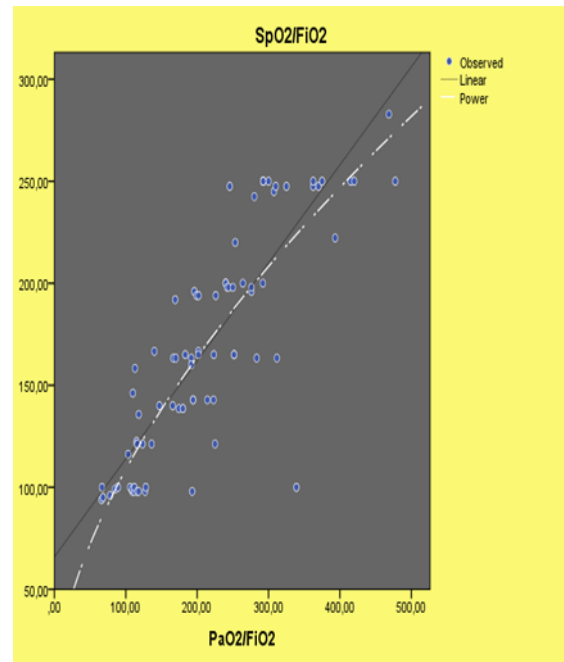
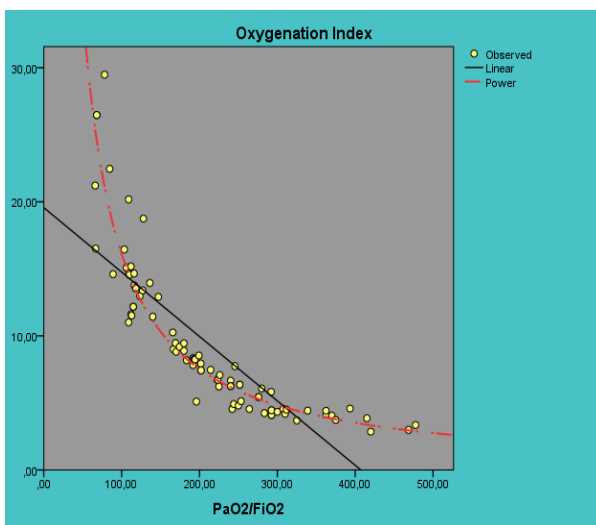
**Table 3.** Significant ( $p < 0.05$  two tailed) and moderate/strong (Spearman  $\rho > 0.6$ ) correlations between the selected parameters.

VCV	PEEP	SFr	SPCr	PFr	SPr	OI	VR
PEEP		W	W,NS	W	-0.97§	W	W,NS
SFr	W		W	0.616	W	-0.627	W,NS
SPCr	W,NS	W		W	W	W	W
PFr	W	0.63	W		W	-0.742	W,NS
SPr	-0.97§	W	W	W		W	W
OI	W	W	W	-0.742	W		W,NS
VR	W,NS	W,NS	W	W,NS	W,NS	W,NS	

NS:non significant, W: weak relation (rounded to nearest 2<sup>nd</sup> decimal), SFr: SpO<sub>2</sub>/FiO<sub>2</sub> % ratio, PFr: PaO<sub>2</sub>/FiO<sub>2</sub> % ratio, SPCr: SpO<sub>2</sub>/PaCO<sub>2</sub> ratio, OI: Oxygenation Index, VR:ventilatory ratio. §:Pearson r (further regression analysis conducted).

Further regression analysis (when applicable) revealed that in PCV mode only 72.7% of SFr change and 92.8% of OI change can be explained by PFr change; yet, in VCV mode no regression analysis could be conducted (figure 3).

**Figure 3.** Regression analysis in PCV mode.



$$OI = 2515.1 \times PFr^{-1.096}, R^2=0.928 (p < 0.05)$$

$$SFr = 7,12 \times PFr^{0.592}, R^2=0.736 (p < 0.05)$$

## DISCUSSION

The ideas of using SpO<sub>2</sub> instead of PaO<sub>2</sub> and the possibility of replacing an invasive index (PFr) with a non-invasive index (SFr) are not new. The larger study comes from Rice et al. (2613 measurements), who report that SFr of 235 and 315 correlate ( $r=0.89, p < 0.001$ ) with PFr of 200 and 300 ( $SFr = 64 + 0.84x (PFr)$ ), respectively, for diagnosing and following up adult patients with ALI and ARDS<sup>5</sup>. In children, Pandharipande et al. found that the total and respiratory SOFA scores obtained with imputed SF values correlate ( $\rho \sim 0,85, p < 0.001$ ) with the corresponding SOFA score using PF ratios. Both the derived and original respiratory SOFA scores similarly predict outcomes<sup>11</sup>. Their patients were either undergoing

general anesthesia or were part of the ARMA study<sup>8</sup>, they did not incorporate covariates such as hemoglobin, age, comorbid illnesses, smoking history, body mass index and positioning of patient (supine versus prone).

In our study, general characteristics of the 2 groups were comparable (both BMI and hemoglobin), with the exception of age and Pmean. All patients were supine; yet, we also did not incorporate comorbid illness and smoking history. The difference of Pmean is affected by the different values of PEEP for each group (table 2). The inhomogeneity of the mechanical ventilation mode distribution among various diagnoses is dominant. Yet, as we did not conduct any diagnosis - based analysis, we did not take this kind of differences in account. In PCV mode a relative strong relation between PFr and SFr and OI was found; yet further regression analysis implies that no direct replacement of PFr with SFr can be made without limitations, in clinical setting. In VCV mode moderate relation was found between SFr and PFr, though no regression analysis was conducted. The above findings may reflect the different characteristics between the 2 groups (diagnosis, number of samples). No correlation was found between the other indices.

Hence; at first sight, it appears that the correlation between SFr and PFr is difficult to be applied in everyday clinical setting. Why? We

suggest that the reason is the relation SpO<sub>2</sub> to PaO<sub>2</sub>.

Numerous mathematical models that have been proposed to describe the standard and nonstandard HbO<sub>2</sub> “equilibrium” dissociation curves since the pioneering work of Hill and Adair<sup>12-13</sup>. From the relatively “simple” Severinghaus equation to the empirical equations of Kelman and Siggaard-Andersen; Thomas modification and, the more complex, mathematical computational models of Easton, Buerk-Bbridges and Nickalls, the aim is to reliably calculate SpO<sub>2</sub> from PaO<sub>2</sub> and *vice versa*- i.e. to replace PFr with SFr for a given FiO<sub>2</sub><sup>13-20</sup>. Yet, data of these models come mainly from subjects (human and animals) with “normal” physiology. In addition, each model has its drawbacks, the detailed reference of which is beyond the scope of this article. Large series of arterial blood samples from Gøthgen et al and Sommers and al. only confirm the complexity of the problem<sup>20-21</sup>.

Finally, another issue is the overall clinical relevance of the PFr. Since PFr has a lot of limitations: PEEP dependence, nonlinear behavior of PFr in dependence of FiO<sub>2</sub>, etc<sup>3-4</sup>, are we trying in vain to replace a poor index of oxygenation with another maybe less reliable? The authors of this study agree with former reports that “accuracy should be balanced with feasibility”<sup>23</sup>. We do need a mathematical model complex enough to reliably reflect gas



exchange in lung disease state, but simple enough to use; or more than one model that can be used under certain conditions.

However, seek for a noninvasive index in place of an invasive or seek for the conditions that this replace could be applied are not vain. Till the time of the writing of this article (Dec 2014) PFr is one of the most used and popular index; that why we should continue the research in order to find the conditions that they'll allow us to use a noninvasive index (SFr) instead.

## CONCLUSION

In the present study a moderate relation was found between SFr and PFr. The results agree with previous published studies; the differences among them lie in the different design of each one of them. The authors believe that given the fact that one still considers using broadly PFr as index of oxygenation- SFr can be used safely as a surrogate for PFr only for certain disease states. Larger series are needed in order to define those patients groups and these pathophysiological conditions.

## REFERENCES

1. Murray J, Matthay M, Luce J, et al. An expanded definition of the adult respiratory distress syndrome. *Am Rev Respir Dis* 1998; 138:720-3.
2. Bernard GR, Artigas A, Brigham KL, et al. The American-European Consensus

Conference on ARDS. Definitions, mechanisms, relevant outcomes and clinical trial coordination. *Am J Respir Crit Care Med* 1994; 149:818-24.

3. The ARDS Definition Task Force. Acute Respiratory Distress Syndrome: The Berlin Definition. *JAMA* 2012;307:2526-33.
4. Aboab J, Luis B, Jonshon B, et al. Relation between PaO<sub>2</sub>/FiO<sub>2</sub> ratio and FiO<sub>2</sub>: a mathematical description. *Intensive Care Med* 2006; 32:1494-7.
5. Rice TW, Wheeler AP, Bernard GR, et al. Comparison of SpO<sub>2</sub>/FiO<sub>2</sub> ratio to PaO<sub>2</sub>/FiO<sub>2</sub> ratio in patients with acute lung injury or ARDS. *Chest* 2007; 132:410-7.
6. Ortiz RM, Cilley RE, Bartlett RH. Extracorporeal membrane oxygenation in pediatric respiratory failure. *Pediatr Clin North Am*. 1987; 34:39-46.
7. Sinha M, Fauvel N, Singh S, et al. Ventilatory ratio: a simple bedside measure of ventilation. *Br J Anaesth* 2009;105:692-7.
8. Ventilation with lower tidal volumes as compared with traditional tidal volumes for acute lung injury and the acute respiratory distress syndrome. The ARDS Network. *N Engl J Med* 2000; 342:1301-8.
9. Charron C, Repesse X, Bouferrache K, et al. PaCO<sub>2</sub> and alveolar dead space are

- more relevant than PaO<sub>2</sub>/FiO<sub>2</sub> ration in monitoring the response of prone positioning in ARDS patients: a physiological study. *Crit. Care* 2011; 15:R175.
10. Aslanidis T, Myrou A, Chytas E, et al. Relation between PaO<sub>2</sub>/FiO<sub>2</sub>ratio. SpO<sub>2</sub>/FiO<sub>2</sub> ratio. Oxygenation Index and Ventilation ratio in critically ill patients *Crit Care* 2013; 17(Sup.2):34.
  11. Pandharipande P, Shintani A, Hangerman H., et al. Derivation and validation of SpO<sub>2</sub>/FiO<sub>2</sub> ratio to impute for PaO<sub>2</sub>/FiO<sub>2</sub> ratio in the respiratory component of the Sequential Organ Failure Assessment (SOFA) Score. *Crit. Care* 2009; 37(4):1317-21.
  12. Adair G. S. The hemoglobin system. VI. The oxygen dissociation curve of hemoglobin. *J. Biol. Chem.* 1925; 63:529-45.
  13. Hill A. V. The possible effects of the aggregation of the molecules of haemoglobin on its dissociation curves. *J. Physiol.* 1910; 40:iv-vii.
  14. Kelman GR. Digital computer subroutine for the conversion of oxygen tension into saturation. *Journal of Applied Physiology* 1966; 21: 1375.
  15. Severinghaus JW. Simple, accurate equations for human blood O<sub>2</sub> dissociation computations. *Journal of Applied Physiology* 1979; 46: 599-602.
  16. Buerk, D, Bridges E.W. A simplified algorithm for computing the variation in oxyhemoglobin saturation with pH, PCO<sub>2</sub>, T and DPG. *Chem. Eng. Commun.* 47:113-124, 1986.
  17. Easton D. M. Oxyhemoglobin dissociation curve as expoexponential paradigm of asymmetric sigmoid function. *J. Theor. Biol.* 1979; 76:335-49.
  18. Dash RK, Bassingthwaigte A. Erratum to: Blood HbO<sub>2</sub> and HbCO<sub>2</sub> Dissociation Curve saturation Varied O<sub>2</sub>, CO<sub>2</sub>, pH, 2,3-DPG and Temperature Levels. *Annals of Biomedical Engineering*, 2010; 38 :1683-1701.
  19. Nickalls RWD. Inverse solutions of the Severinghaus and Thomas equations which allow PaO<sub>2</sub> to be delivered directly from SaO<sub>2</sub>. 2011; [www.nickalls.org/dick/papers/anes/severinghaus.pdf](http://www.nickalls.org/dick/papers/anes/severinghaus.pdf)
  20. Gøthgen IH, Siggaard-Andersen O, Kokholm G. Variations in the hemoglobin-oxygen dissociation curve in 10079 arterial blood samples. *Scand J Clin Lab Invest* 1990; 203:87-90.
  21. Summers J, Jose R, Durcan J, et al. Validation of the severinghaus oxygen dissociation curve in critically ill adult patients. 22nd ESCIM annual congress. *Intensive Care Med* 2009; 35 (1):S49



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22. Wagner PD. Assessment of gas exchange in lung disease: balancing accuracy against feasibility. Crit. Care 2007, 11:182.
23. El-Khatib MF, Gammaledine GW. Clinical relevance of PaO<sub>2</sub>/FiO<sub>2</sub> ratio. Crit. Care 2008; 12:407.
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**Author Disclosures:**

Authors Aslanidis Th, Myrou A, Tsirona Ch, Kontos A, Giannakou-Peftoulidou M have no conflicts of interest or financial ties to disclose.

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