

Research Article

Prone Positioning Improves Oxygenation and Outcome of Trauma Patients with Severe Acute Respiratory Distress Syndrome (ARDS)

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Abstract

Introduction: The acute respiratory distress syndrome (ARDS) was proposed as an acute onset respiratory failure with PaO_2/FiO_2 ratio <200 (regardless of positive end-expiratory pressure (PEEP) level), bilateral infiltrates on chest X-ray and pulmonary artery occlusion pressure below 18 mmHg. The main therapeutic management of ARDS is based on the concept of protective lung strategy ventilation. Some authors reported about increase in oxygenation in the prone position (PP) in both direct types of ARDS; however, the clinical significance of it still remains questionable. We review and analyze potential clinical benefits of using the prone position in trauma critically ill patients with severe ARDS.

Methods: This is an observational, retrospective study provided in university teaching hospital from January 2006 and June 2012. In this study, we retrospectively examined clinical data of 33 trauma and 56 non-trauma critically ill adult patients suffering from severe ARDS and managed by application of prone positioning (PP) during General Intensive Care Unit (GICU) stay.

Results: We found no difference in demographic data and in the rate of complications between both study groups after application of PP. The trauma patients with severe acute lung injury demonstrated remarkable improvement in lung compliance (from 18.72 ± 9.52 to 29.285 ± 6.26 , p<0.05,) and significant decreases in peak inspiratory pressure (PIP) in the prone position (from 33.48 ± 9.52 cm H2O to 21.06 ± 7.06 cm H2O, p<0.05) The mortality rate was lower in trauma group patients treated by PP (p<0.005) than in non-trauma ICU population.

Conclusions: Our study showed clinical benefit by application of PP in the treatment of severe ARDS in post-trauma critically ill patients.

Keywords: Prone position; Multiple trauma; Severe ARDS

Introduction

Acute Respiratory Distress Syndrome (ARDS) was described in 1967 in context of acute dyspnea, severe hypoxemia, decreased lung compliance and bilateral diffuse lung infiltrates on chest radiograph [1]. The first definition of ARDS was proposed in 1994 as an acute onset respiratory failure with PaO2/FiO2 ratio <200 (regardless of PEEP level), bilateral infiltrates on chest X-ray and pulmonary artery occlusion pressure below 18 mmHg [2,3]. Since 2012, ARDS has been defined as an arterial oxygen partial pressure (PaO₂) to inspiratory oxygen fraction (FiO₂) ratio less than 200 and PEEP or CPAP>5 cm H₂O according to the American European Consensus Conference (AECC) [4]. The therapeutic management of ARDS has also changed over the last two decades. The cornerstone of therapeutic strategy in ARDS began from a new concept of ventilatory support - "protective lung strategy" - which includes mechanical ventilation with small tidal volumes, low plateau pressure and application of relatively high levels of positive end-expiratory pressure (PEEP) [5-7]. In recent years, inhalation of Nitric Oxide (NO), prone position (PP), high frequency ventilation and ECMO devices have successfully demonstrated their ability to improve oxygenation in critically ill ARDS patients [8,9].

The effectiveness of PP in treatment of severe ARDS patients remains questionable. An increase in oxygenation in the prone position has been demonstrated in both direct (lung contusion, aspiration) and indirect (sepsis, SIRS, burns, pancreatitis) pathophysiological types of ARDS [10]. Previously published data [11,12] demonstrated no significant survival benefit in patients with ARDS or in subgroups of patients with moderate and severe hypoxemia. However, recently published data [13] shows a significant improvement in survival from early application of prone positioning in severe (PO₂\FiO, ratio less

J Trauma Treat ISSN: 2167-1222 JTM, an open access journal than 150) ARDS in a mixed general ICU population. Moreover, use of the prone position was demonstrated to provide clinical benefit especially in direct acute lung injury (post-traumatic lung contusion -ARDS/acute lung injury- ALI) [14]. In this paper, we review and analyze potential clinical benefits of using the prone position in trauma versus non-trauma types of severe ARDS in critically ill patients admitted to the intensive care unit at Soroka Medical Center.

Patients and Methods

The study is observational and retrospective. The Human Research and Ethics Committee at Soroka Medical Center in Beer-Sheva, Israel approved this study. We collected clinical data from all cases of critically ill patients suffering from ARDS admitted to the General Intensive Care Unit (GICU) in Soroka Medical Center between January 1999 and June 2005. Soroka Medical Center is a tertiary care facility with 1100 inpatient beds, including 12 beds in the General Intensive Care Unit (GICU). All clinical data was extracted from Patients' Register Database, General Intensive Care Unit, Soroka Medical Center during

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those six years.

All adult critically ill patients suffering from severe ARDS who were managed by application of Prone Positioning (PP) during GICU admission were included in the present study. According to the admission diagnosis, all severe ARDS patients treated with PP were retrospectively allocated to trauma and non-trauma study groups.

The ARDS criteria and prone positioning protocol

The criteria of severe ARDS include diffuse patchy infiltrates on chest radiograph, PaO_2/FiO_2 ratio less than 100 and (PEEP) of >10 cm H₂O [4]. All severe ARDS patients were mechanically ventilated on volume or pressure control modes. Prone positioning was applied to critically ill patients with severe ARDS ($PaO_2/FiO_2 < 100$, $FiO_2 = 1.0$). Patients were placed back in the supine position when the PaO_2/FiO_2 ratio had improved to more than 200 and/or FiO₂ more than 0.6.

Variables, measures, primary and secondary outcome

The demographic data, cause for admission, APACHE-II score, patients' co-morbidities, and length of ICU stay, clinical data of oxygenation/ventilation parameters (FiO_2 , PaO_2/FiO_2 , $PaCO_2$, PEEP level, lung compliance) and in-ICU mortality were collected. The primary outcome endpoint of the present study is ICU mortality rate. Lung mechanics and oxygenation parameters are the secondary outcomes.

Statistical Analysis

For categorical variables, proportions were compared using Fisher's Exact Test or Chi Square, as appropriate. Continuous variables were analyzed with Student's *t*-test or the Wilcoxon Rank Sum Test, depending on the validity of the normality assumption (Pearson's chi-squared test). A two-tailed p-value of <0.05 was considered to be significant. All analysis was performed using SPSS version 17 (SPSS, Chicago, IL).

Results

In total, two hundred and ten (210) critically ill patients with ARDS were hospitalized in the General Intensive Care Unit over the six years period (Table 1). Of those 210, 89 patients fulfilled the criteria for severe ARDS ($PaO_2/FiO_2 < 100$). Those patients were ventilated by FiO_2 1.0 and treated by placement in the prone position. Those patients were included in the present study.

Thirty-three 33 patients had admission diagnosis of multiple trauma (Trauma group) and fifty-six patients 56 with admission diagnosis other than trauma were included in Non-trauma group (see an explanation in Table 1).

There was no significant difference in demographic data, APACHE score and probability of death index between trauma and non-trauma population. Patients in both study groups were hemodynamically stable before (Group 1, 81.6 ± 23.68 mmHg vs Group 2, 82.78 ± 19.29 mmHg) and after (Group 1, 87.06 ± 17.35 mmHg vs Group 2, 87.82 ± 17.85 mmHg) application of PP. The overall complication rate was similar in both study groups (10%, *p* NS, Table 1). The mortality rate was significantly lower in trauma patients after application of PP than in the non-trauma ICU population (p<0.005, Table 1).

The response of oxygenation with the use of prone positioning was similar in both study groups [increase in PaO_2/FiO_2 ratio >200 from 81.626 ± 18.376 (Group 1) and 76.444 ± 17.398 (Group 2) after prone position application, p NS, (Table 2). The PEEP levels were

unremarkably changed after PP in trauma (from 9.33 ± 3.763 to 7.655 ± 2.175 mmHg) and non-trauma (from 9.196 ± 2.932 to 8.765 ± 2.905 mmHg, p NS, (Table 2) persons. However, trauma patients with severe acute lung injury demonstrated remarkable improvement in lung compliance (from 18.72 ± 9.52 to 29.285 ± 6.26 , p<0.05,) and significant decreases in peak inspiratory pressure (PIP) in the prone position (from 33.48 ± 9.52 cm H₂O to 21.06 ± 7.06 cm H₂O, p<0.05) (Table 2). Finally, lung compliance was significantly higher in trauma group patients after PP treatment (29.285 ± 6.26 vs 20.428 ± 6.56) than in non-trauma group (Table 2).

PIP after PP treatment declined significantly also in trauma group (21.06 \pm 7.06 vs. 27.127 \pm 8.11 cm H₂O, p<0.05, Table 2). Total time in the prone position was higher in trauma group patients (54 \pm 59.94 h vs. 34.196 \pm 43.056 h, p<0.005, Table 2). Time on mechanical ventilation and total length of ICU stay after PP management were significantly less in trauma group population than non-trauma persons (p<0.05, Table 2). Injury severity score was similar between survived and non-survived trauma patients (p NS, Table 2).

Discussion

Severe ARDS caused by direct (pneumonia, aspiration, traumatic pulmonary contusion, inhalational injury etc.) or indirect (sepsis, severe SIRS etc.) etiologic factors are characterized pathophysiologically by widespread lung and endothelial injuries [15,16]. Post-traumatic severe ARDS is primarily related to alveolar and capillary damage by intense direct kinetic force on the thorax as opposed to a systemic inflammatory reaction (SIRS)/sepsis-induced ARDS which develops as a result of increased cytokine blood levels leading to alveolar edema, increased alveolar-capillary permeability, apoptosis and, eventually, cell necrosis [17,18].

Thus, the "pulmonary contusion" in trauma patients with ARDS is a self-limited insult contrary to the systemic inflammatory reaction, which continues progressively, causing on-going indirect lung injury until the severe SIRS or sepsis is successfully managed by the clinicians. In both direct and indirect types of pathophysiological mechanisms of ARDS, the prone positioning improves oxygenation and respiratory mechanics [19,20]. Prone positioning provides more homogeneous

	Trauma***	Non-trauma	Dualua
	(n=33)	(n=56)#	P value
Age (mean ± SD)	39.87 ± 14.66	52.35 ± 19.22	NS
Gender (male/female)	12/11 (33)	30/26 (56)	NS
APACHE II score 1 (units)*	22.38 ± 6.52	23.92 ± 7.96	NS
APACHE II score 2 (units)**	24.6 ± 5.77	27 ± 6.94	NS
Complications rate (overall, n) ^{##} 3/33 6/56 NS	3/33	6/56	NS
Mortality rate (%)	10/33 (22.7%)	34/56 (51.1%)	0.005
ISS (for trauma patients)	Survivors	Nonsurvivors	P value
iss (ior trauma patients)	294 + 971	24 086 + 9 69	NS

*APACHE II score 1was calculated on admission to ICU; APACHE II score 2 was calculated on day when application of Prone Position (PP) has been initiated.

**P value has been considered to be statistically significant if less than 0.05; NSnon significant.

***Multiple trauma injuries included: head injury, blunt chest and abdomen trauma, spine and extremity fractures. ISS: Injury Severity Score. #Non-trauma group patients included several diagnoses on admission to ICU: sepsis/septic shock, acute pancreatitis, pneumonia, blood products transfusion for non-trauma patients, massive intracerebral hemorrhage.

^{##}Complications on PP were included pressure sores on chest and abdomen surface (totally 9 patients for both study groups, 10%).

Table 1: Demographic data (mean \pm SD, %) and clinical outcome endpoints of patients on prone positioning.

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	Trauma (n=33)	Non-trauma (n=33)	P value**
PaO ₂ /FiO ₂ before PP (torr)	81.626 ± 18.376	76.444 ± 17.398	NS
PaO ₂ /FiO ₂ after PP (torr)	200.556 ± 45.822	200.241 ± 44.08	NS
FiO ₂ before PP (%)	87.57 ± 18.376	87.41 ± 17.398	NS
FiO ₂ after PP (%)	51.034 ± 15.199	52.127 ± 18.1702	NS
PEEP (cmH ₂ O) before PP	9.33 ± 3.763	9.196 ± 2.932	NS
PEEP (cmH ₂ O) after PP	7.655 ± 2.175	8.765 ± 2.905	NS
Compliance (ml/cmH ₂ O) Before PP	18.72 ± 9.52	19.33 ± 8.48	NS
Compliance (ml/cmH ₂ O) After PP	29.285 ± 6.26	20.428 ± 6.56	<0.05
PIP (cmH ₂ O) before PP	33.48 ± 9.52	31.839 ± 8.48	NS
PIP (cmH ₂ O) after PP	21.06 ± 7.06	27.127 ± 8.11	<0.05
Prone positioning characteristics			
Time on PP (hours) on PP *** (h)	54 ± 59.94	34.196 ± 43.056	<0.005
Time to improve oxygenation	8.76 ± 11.245	8.67 ± 11.6	NS
Ventilatory support after PP	8.98 ± 13.1	14.9 ± 16.2	<0.05
Length of ICU stay after PP (days)	10.67 ± 13.155	18.06 ± 18.75	<0.05

**P value has been considered to be statistically significant less than 0.05

***Time of improvement oxygenation (PaO,/FiO, ratio >200 or/and FiO,<0.6) since initiation of prone position

ICU- intensive care unit, PP- prone positioning, PEEP- positive end-expiratory pressure

Table 2: Clinical data of pulmonary parameters (mean ± SD, %) of critically ill patients in the prone position

distribution of transpulmonary pressure and a subsequent decrease in alveolar inflation pressure over the non-dependent (dorsal) and dependent (ventral) lung zones [21,22]. The perfusion in dorsal (dependent) regions is also improved with the use of prone positioning, which may be explained by the effects of gravity. Lung mechanics have been shown to improve after repositioning patients from prone to supine position and such observation is related to increased respiratory system compliance and decreased inspiratory pressure [23,24].

There is some clinical data which shows remarkable improvement in survival of mixed critically ill population after application of PP [13]. However, none of the clinical studies we reviewed showed an increase in survival in homogenic (for example, direct pathophysiological mechanism) critically ill patients with severe ARDS.

In the present study, a remarkable improvement of lung mechanics and survival were showed in trauma patients. Most of them had direct chest injury (pulmonary contusion). It might be a good explanation of consistent potential benefit of prone positioning application after initial self-limited lung injury. In our present study, we argue that the prone positioning application failed to achieve positive constant physiological effects because of continuous indirect lung injury in the non-trauma group.

Despite the potential complications and risks (inability to perform CPR, damage to peripheral nerves and eyes, elevation of intra-abdominal pressure, etc), use of the prone position has been demonstrated to be a safe and beneficial tool, especially in multiple trauma patients [25]. Hale et al. [26] demonstrated safe application of prone positioning in adult burn patients with severe ARDS. D'Ignazio et al. [27] reviewed two cases of severe blunt trauma patients with cervical spine and pelvis involvement who developed ARDS and were safely managed by prone positioning. Moreover, Kenn et al. [28] effectively treated a 34-week pregnant patient with blunt chest trauma and subsequent ARDS on prone positioning during 8 hours. Previously published data by Davis et al showed a decrease in mortality rate, fewer days on the ventilator and shorter length of hospital stay after using original prone kinetic therapy protocol (4 hours cycle per day use of kinetic therapy bed) in ARDS trauma patients Our data correlates well with that data and demonstrates a remarkable decrease in mortality rate in post-trauma critically ill patients. In contrast to the aforementioned study, we left our patients in the prone position until target physiological parameters ($PaO_2/FiO_2>200$, $FiO_2<0.6$) had been reached.

Application of PP might be complicated by loss of chest tubes, arterial and venous access catheters, and endotracheal or tracheostomy tubes in critically ill patients and, rarely, by development of peripheral nerve injuries, skin necrosis, or damage to the eyes [29-31]. The ICU physicians are unable to perform cardiopulmonary resuscitation in the event of cardiac arrest on PP [31]. In our study, the overall complications rate in trauma and non-trauma group was 10% (Table 1), which correlates well with previously published data [25].

Our study has a number of limitations. The present study is retrospective and observational. We did not use a standard protocol. Thus, trauma group patients remained on prone position significantly longer than non-trauma persons. The patients' clinical data were collected during 1999-2005. We also planned to observe survived patients for a long-term outcome (during the next 5 years). However, we were not able to get a consistent outcome data from the patients' records [32].

Conclusions

In conclusion, our study showed clinical benefit by application of PP in the treatment of severe ARDS in post-trauma critically ill patients compared to non-trauma counterparts. We propose that a large multicenter prospective study would help to clarify the precise physiological mechanisms in which prone positioning application is advantageous in post-trauma critically ill patients.

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