THE EFFECT OF POSITIVE PRESSURE VENTILATION ON RENAL FUNCTION IN CRITICALLY ILL PATIENTS; EFFECT ON RENNIN- ALDOSTERON SYSTEM

Thesis
Submitted for Partial Fulfillment of MD Degree
In
Critical Care

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﴿ قَالُواْ سُبْحَنَكَ لَا عِلْمَ لَنَا إِلَّا مَا عَلَمْ لَنَا إِلَّا مَا عَلَمْ تَنَا إِلَّا مَا عَلَمْ تَنَا إِلَّا مَا عَلَمْ تَنَا أَلِكُ أَنْتَ الْعَلِيمُ الْعَكِيمُ ﴾ عَلَمْتَنَا إِنّكَ أَنْتَ الْعَلِيمُ الْعَكِيمُ ﴾ عَلَمْتَنَا إِنَّكَ أَنْتَ الْعَلِيمُ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَظَمِينَ الْعَلَمْ الْعَظَمِينَ الْعَلَمْ الْعَظَمِينَ الْعَلَمْ الْعَلَمُ الْعَلَمْ الْعَلَمْ الْعَلَمْ الْعَلَمُ الْعُلِيمُ الْعُلَمُ الْمُ الْعَلَمُ الْمُعَلِمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ اللّهُ الْعَلَمُ الْعَلَمُ اللّهُ الْعَلَمُ اللّهُ اللّهُ الْعَلَمُ الْعُلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلْمُ الْعَلَمُ الْعُلِمُ الْعَلَمُ الْعِلْمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعِلْمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعَلَمُ الْعِلْمُ الْعِلْمُ الْعِلْمُ الْعَلِمُ الْعَلَمُ الْعِلْمُ الْعَلَمُ الْعِلْمُ الْعَلِمُ الْعَلَمُ الْعِلْمُ الْعُلِمُ الْعُلِمُ الْعُلِمُ الْعُلْمُ الْعُلِمُ الْعُلِمُ الْعُلِمُ اللّهُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلِمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلِمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلْمُ الْعُلِمُ الْعُلْمُ الْعُلِمُ الْعُلْمُ الْعُلِمُ الْعُلْمُ الْعُلْمُ الْعُلِمُ الْعُلِمُ الْعُلْمُ الْعُلِم

سورة البقرة آية (٣٢)

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ABSTRACT

Mechanical ventilation is associated with numerous organ-system complications. **The Aim of the study** was to investigate the effect of positive pressure ventilation on renin—aldosterone system and to investigate their prognostic value in relation to ICU course and outcome. **Results**: Mechanical ventilation caused elevation of renin and aldosterone levels especially with higher levels of PEEP.Renin and aldosterone levels in mechanically ventilated patients were significantly correlated with kidney function test (urea and Creatinine), length of mechanical ventilation, need for hemodynamic support, APACHE IV score and mortality.

Keywords: mechanical ventilation, lung –kidney crosstalk, reninaldosterone system

List of Abbreviations:

ADH		Antidiuretic hormone
ALT		Alanine aminotransferase
APACH IV		Acute Physiology and Chronic Health evaluation
ANP		Atrial naturitic peptide
ARDS		Adult Respiratory Distress Syndrome
ARF		Acute renal failure
AST		Aspartate aminotransferase
ВРМ		Beat per minute
COPD		Chronic obstructive pulmonary disease
CPAP		Continuous positive airway pressure
CVP		Central venous pressure
DBP		Diastolic blood pressure
DM		Diabetic
DVT	:	Deep venous thrombosis
ELIZA		enzyme – linked immunosorbent assay
ET		endothelin
FRC		functional residual capacity
GFR	:	Glomerular filtration rate
GI		Gastrointestinal
HDS		Hemodynamic support
HR		Heart rate
HTN		Hypertension
ICU		intensive care unit
IHD		Ischemic heart disease

IPPV intermittent positive pressure ventilation LOMV Length of mechanical ventilation LOS Length of ICU stay LV left ventricle LVEDV Left ventricular end-diastolic volume MODS Multiple organ dysfunction syndrome MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume ZEEP Zero PEEP	IL-6	Interleukin- 6
LOS Length of ICU stay LV left ventricle LVEDV Left ventricular end-diastolic volume MODS Multiple organ dysfunction syndrome MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-\alpha tumor necrosis factor-\alpha UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	IPPV	intermittent positive pressure ventilation
LV left ventricle LVEDV Left ventricular end-diastolic volume MODS Multiple organ dysfunction syndrome MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-\alpha tumor necrosis factor-\alpha UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	LOMV	Length of mechanical ventilation
LVEDV Left ventricular end-diastolic volume MODS Multiple organ dysfunction syndrome MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	LOS	Length of ICU stay
MODS Multiple organ dysfunction syndrome MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	LV	left ventricle
MOF Multiple organ faliure MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	LVEDV	Left ventricular end-diastolic volume
MV Mechanical ventilation NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	MODS	Multiple organ dysfunction syndrome
NO Nitric oxide PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	MOF	Multiple organ faliure
PEEP Positive end expiratory pressure PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	MV	Mechanical ventilation
PGE Prostaglandins PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	NO	Nitric oxide
PPV Positive pressure ventilation PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	PEEP	Positive end expiratory pressure
PVR pulmonary vascular resistance RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	PGE	Prostaglandins
RBF Renal blood flow ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	PPV	Positive pressure ventilation
ROC Receiver operator characteristic RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	PVR	pulmonary vascular resistance
RR Respiratory rate RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	RBF	Renal blood flow
RV right ventricle RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	ROC	Receiver operator characteristic
RVEDV right ventricular end-diastolic volume SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	RR	Respiratory rate
SBP Systolic blood pressure SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	RV	right ventricle
SD Standard deviation TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	RVEDV	right ventricular end-diastolic volume
TNF-α tumor necrosis factor-α UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	SBP	Systolic blood pressure
UOP Urine output VAP Ventilator associated pneumonia Vt Tidal volume	SD	Standard deviation
VAP Ventilator associated pneumonia Vt Tidal volume	TNF-α	tumor necrosis factor-α
Vt Tidal volume	UOP	Urine output
	VAP	Ventilator associated pneumonia
ZEEP Zero PEEP	Vt	Tidal volume
	ZEEP	Zero PEEP

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INTRODUCTION

Mechanical ventilation is a standard component of intensive care unit management of critically ill patients and is widely used for respiratory support and improves oxygenation in critically ill patients. Mechanical ventilation is often a life-saving intervention, but carries many potential complications. There are numerous mechanisms underlying the development of complications in the ventilated patient. Complications may result from the endotracheal tube or tracheotomy tube (or with noninvasive ventilation, the mask), or from the effects of positive-pressure ventilation. (1)

The ventilatory strategy required to maintain adequate gas exchange may exacerbate, or even initiate, significant lung injury and inflammation. Mechanical ventilation can produce lung injury that is functionally and histologically indistinguishable from that seen in ARDS. Mechanical ventilation has also been shown to have significant effects on lung levels of inflammatory cells and soluble mediators. These inflammatory mediators may get released from the alveolar space into the general circulation and play a critical role in the pathophysiology of systemic inflammatory response, multiple system organ failure and shock. (2)

Mechanical ventilation may also affect distal organ function via effects on cardiac output, as well as the level of oxygenation and the distribution of blood to the various organ systems. Positive intrathoracic pressure impairs venous return, decreases ventricular distensibility, and causes decreased ventricular filling. The decreased venous return leads to a decrease in right ventricular preload, which through sustained pressure changes in the cardiopulmonary vasculature leads to a sustained, decreased left ventricular afterload. Ultimately, decreased left ventricular preload decreases left ventricular afterload. These changes reduce cardiac output because although left ventricular afterload is reduced, the decreased left ventricular filling has a greater effect on cardiac output. Positive end-expiratory pressure (PEEP) may reduce cardiac output by causing a further increase in intrathoracic pressures. (3)

Because the kidneys receive 20% to 25% of cardiac output, any decrease in cardiac output caused by PEEP affects RBF. RBF is primarily affected by PEEP because of sympathetic activation related to increased plasma renin activity. Although total RBF is relatively unchanged, blood flow is redistributed from the cortical to the juxta medullary nephrons. This redistribution would be associated with decreased urine output, decreased Creatinine clearance, and an increased fractional resorption of sodium. (4)

PEEP further affects the hormonal and sympathetic pathways. The effect is due to an increase in sympathetic tone, which is caused by increased plasma renin activity and decreases GFR because of decreased blood flow. PEEP has a transient effect on aortic blood pressure, and this effect reflexively activates the sympathetic nervous system through aortic and (sino) carotid baroreceptors. Changing renal function then slowly affects intravascular volume.