

Comparative Analysis of Dyselectrolytemias as Prognostic Markers in Critically Ill Patients, With Special Reference to Hypernatremia: A Prospective Observational Study

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ABSTRACT

Background:

Dyselectrolytemias are among the most frequent and clinically impactful laboratory derangements encountered in critically ill patients admitted to the Intensive Care Unit (ICU). Among all electrolyte disturbances, hypernatremia (serum sodium >145 mEq/L) has emerged as an especially potent independent predictor of mortality, prolonged ICU stay, and increased requirement for organ support. Indian data on the comparative prognostic significance of multiple concurrent dyselectrolytemias, particularly in correlation with validated severity scores, remain scarce.

Objectives:

To characterize the incidence and pattern of dyselectrolytemias in adult ICU patients, evaluate the independent and comparative prognostic significance of each abnormality — with special reference to hypernatremia — and to correlate electrolyte disturbances with APACHE II and SOFA scores.

Methods:

This was a prospective observational study conducted over 6 months in the ICU of MMCHRI, Kanchipuram. One hundred consecutive adult patients (>18 years) admitted for >24 hours were enrolled. Serum sodium, potassium, calcium, magnesium, and phosphate were measured at baseline (within 24 hours of ICU admission) and monitored at 24 hours, then every 24 to 48 hours depending on electrolyte status, for up to 10 days. Illness severity was quantified using APACHE II (Day 1) and SOFA (daily) scores. Primary outcomes included ICU mortality, in-hospital mortality, mechanical ventilation requirement, and length of ICU stay.

Results:

Dyselectrolytemias were present in 86% of patients. Hypomagnesemia was the most prevalent (41%), followed by hypokalemia (38%), hypocalcemia (32%), hypernatremia (28%), and hyponatremia (22%). Hypernatremia was independently associated with the highest ICU mortality (35.7% vs 8.3%; $p < 0.001$), the longest mean ICU stay, and the greatest need for mechanical ventilation (64.3%). A strong positive correlation was identified between serum sodium and APACHE II score (Pearson $r = 0.68$, $p < 0.001$). Multivariate analysis identified hypernatremia as the strongest electrolyte-based predictor of ICU mortality (OR 4.1; 95% CI 2.3–7.3). Serum sodium demonstrated the highest discriminatory accuracy for ICU mortality (AUC = 0.81).

Conclusion:

Hypernatremia is a prevalent, readily measurable, and independently prognostic electrolyte abnormality in critically ill patients. Its strong correlation with APACHE II and SOFA scores and its superior predictive accuracy for mortality support its incorporation as a routine prognostic biomarker in ICU practice, particularly in resource-limited Indian healthcare settings where advanced monitoring tools are not always accessible.

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Keywords: dyselectrolytemia; hyponatremia; ICU; critically ill; APACHE II; SOFA score; prognosis; mortality; prospective study.

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Introduction

The Intensive Care Unit represents a uniquely challenging clinical environment where multiple organ systems are under simultaneous physiological stress. Electrolyte homeostasis, which depends on the integrated function of the kidneys, endocrine system, gastrointestinal tract, and the central nervous system, is frequently disrupted in this setting. Dyselectrolytemias — abnormalities in the serum concentrations of sodium, potassium, calcium, magnesium, or phosphate — are not merely incidental laboratory findings but reflect the depth of systemic perturbation imposed by critical illness. Their presence influences cellular membrane stability, cardiac conduction, neuromuscular excitability, enzymatic function, and the integrity of virtually every organ system.^{4,9}

Epidemiological studies from intensive care settings across the world have consistently reported high prevalences of dyselectrolytemias, with reported rates of any electrolyte abnormality ranging from 60% to over 90% of ICU admissions.^{10,11} Among these, abnormalities of serum sodium — the primary determinant of plasma osmolality and extracellular fluid volume — carry perhaps the greatest prognostic weight. While hyponatremia has historically received more clinical attention, hyponatremia (serum sodium >145 mEq/L) has in recent years been recognized as an even more ominous marker in the ICU setting, particularly when acquired during hospitalization rather than present on admission.^{1,2,3}

Hyponatremia in critically ill patients arises from several distinct mechanisms. Impaired water intake due to altered consciousness, aggressive enteral or parenteral sodium loading, excessive insensible water losses in mechanically ventilated patients, osmotic diuresis, and the administration of hypertonic solutions all contribute.^{12,13} Physiologically, hyponatremia induces cellular dehydration and brain shrinkage, leading to neurological dysfunction, capillary rupture, and potentially fatal intracranial hemorrhage. It also activates

counter-regulatory mechanisms — antidiuretic hormone release, thirst stimulation — which are frequently impaired in the context of critical illness, creating a cycle of progressive sodium accumulation and free-water deficit.¹²

Beyond sodium, other electrolyte disturbances are also highly prevalent in the ICU. Hypomagnesemia — increasingly recognized as a marker of nutritional deficiency and systemic catabolism — has been associated with arrhythmias, respiratory muscle weakness, and refractory hypokalemia.^{14,20} Hypokalemia increases the risk of life-threatening ventricular arrhythmias and impairs respiratory muscle function, potentially prolonging mechanical ventilation.¹⁸ Hypocalcemia, often seen in the context of sepsis, pancreatitis, and blood product transfusions, has been linked to myocardial dysfunction, hypotension, and impaired coagulation.¹⁹ Hypophosphatemia contributes to respiratory muscle fatigue, hemolytic anemia, and impaired phagocytic function.¹⁴

Despite this wealth of mechanistic evidence, systematic prospective data from Indian ICUs comparing the prognostic significance of multiple concurrent dyselectrolytemias using validated scoring systems — specifically APACHE II and SOFA — are limited.^{6,9} The APACHE II score, developed by Knaus et al.,⁷ evaluates 12 acute physiological variables, age, and chronic health status within the first 24 hours of ICU admission and has been validated as a reliable predictor of hospital mortality across diverse ICU populations. The Sequential Organ Failure Assessment (SOFA) score, calculated daily, provides a dynamic assessment of six organ systems and correlates strongly with short-term mortality.⁸

The present study was therefore designed to prospectively characterize the incidence and pattern of dyselectrolytemias in a South Indian tertiary care ICU, evaluate the independent prognostic significance of each abnormality with special emphasis on hyponatremia, and determine the strength of correlation between electrolyte disturbances and APACHE II/SOFA scores.

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The findings are intended to support clinicians in resource-limited settings in identifying high-risk patients early through simple, readily available biochemical parameters.

Materials and Methods

Study Design and Setting

This was a prospective observational study conducted over a 6-month period (November 2024 to April 2025) in the Medical Intensive Care Unit of Meenakshi Medical College Hospital and Research Institute (MMCHRI), Kanchipuram, Tamil Nadu, India — a 1,500-bed tertiary care teaching hospital affiliated with MAHER (Deemed to be University). The study was approved by the Institutional Ethics Committee (IEC Ref No: MMCH & RI IEC/PG/02/MAY/25; approval date: 17.05.2025). Written informed consent was obtained from all patients or their legally authorized representatives prior to enrollment.

Study Population

Consecutive adult patients (>18 years of age) admitted to the ICU for more than 24 hours during the study period were enrolled by consecutive sampling. The inclusion criterion was adult patients with complete biochemical and clinical documentation. Patients with end-stage renal disease on maintenance dialysis and those with incomplete medical records or who were discharged within 24 hours of ICU admission were excluded to minimize confounding and ensure data completeness, respectively.

Data Collection

On the day of ICU admission, the following data were recorded for each patient: demographic details (age, sex, weight), primary diagnosis, comorbidities (diabetes mellitus, hypertension, chronic kidney disease, chronic liver disease), type of admission (medical/surgical/emergency), and baseline biochemical parameters including serum sodium, potassium, calcium (total and ionized), magnesium, phosphate, creatinine, hemoglobin, and white blood cell count. Electrolytes were measured from peripheral venous blood samples using a validated ion-selective electrode analyzer.

Follow-up and Data Collection Schedule: Baseline data were collected within the first 24 hours of ICU admission (Day 0), including demographics, primary diagnosis, comorbidities, APACHE II score, and serum electrolytes (Na^+ , K^+ , Ca^{2+} , Mg^{2+} , PO_4^{3-}). A second sample was collected within the next 24 hours (Day 1) along with SOFA score. If electrolytes were normal on Day 1, participants were followed every 48 hours

thereafter (Days 3, 5, 7, 9) with routine electrolytes, SOFA score, and major clinical outcomes. If dyselectrolytemia persisted on Day 1, monitoring continued every 24 hours until electrolytes were corrected (two consecutive normal samples 24 hours apart) or until ICU discharge or death. Routine follow-up continued for a maximum of 10 days from ICU admission. No additional blood draws beyond routine clinical care were performed for the study. Final outcome data collected at the end of follow-up included SOFA score, ICU stay duration, mechanical ventilation days, renal replacement therapy days, and mortality.

Electrolyte Definitions

Electrolyte abnormalities were defined using standard laboratory reference ranges:^{4,12}

Table 1. Definitions of dyselectrolytemias used in this study.

Electrolyte Abnormality	Definition	Reference Range
Hyponatremia	Serum Na^+ < 135 mEq/L	135–145 mEq/L
Hypernatremia	Serum Na^+ > 145 mEq/L	135–145 mEq/L
Hypokalemia	Serum K^+ < 3.5 mEq/L	3.5–5.0 mEq/L
Hyperkalemia	Serum K^+ > 5.0 mEq/L	3.5–5.0 mEq/L
Hypocalcemia	Total Ca^{2+} < 8.5 mg/dL	8.5–10.5 mg/dL
Hypomagnesemia	Serum Mg^{2+} < 1.7 mg/dL	1.7–2.2 mg/dL
Hypophosphatemia	Serum PO_4 < 2.5 mg/dL	2.5–4.5 mg/dL

Severity of Illness Scoring

Illness severity was assessed using two validated, internationally recognized scoring systems. The APACHE II (Acute Physiology and Chronic Health Evaluation II) score⁷ was calculated from 12 physiological variables, age, and chronic health status within the first 24 hours of ICU admission. The SOFA (Sequential Organ Failure Assessment) score⁸ was calculated daily, evaluating six organ systems:

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respiratory (PaO₂/FiO₂ ratio), cardiovascular (mean arterial pressure and vasopressor requirement), hepatic (serum bilirubin), coagulation (platelet count), renal (creatinine and urine output), and neurological (Glasgow Coma Scale). APACHE II score was used for admission severity quantification, and the peak SOFA score during ICU stay was used for outcome correlation.

Outcome Measures

The primary outcomes were: (1) ICU mortality, (2) in-hospital mortality. The secondary outcomes included: (1) requirement for and duration of mechanical ventilation, (2) need for vasopressor support, (3) requirement for renal replacement therapy, and (4) duration of ICU stay in days. Patients were stratified by the type and severity of dyselectrolytemia, and outcomes were compared between groups.

Statistical Analysis

Data were entered and analyzed using SPSS version 25.0 (IBM Corp., Armonk, NY). Continuous variables are expressed as mean \pm standard deviation (SD) or median (interquartile range) based on normality assessed by the Shapiro-Wilk test. Categorical variables are expressed as frequencies and percentages. Between-group comparisons for continuous variables were performed using the independent Student's t-test or Mann-Whitney U test, and for categorical variables using the chi-square test or Fisher's exact test, as appropriate. Pearson correlation analysis was used to evaluate the relationship between serum sodium levels and APACHE II/SOFA scores. Multivariate binary logistic regression was performed to identify independent predictors of ICU mortality, with results expressed as odds ratios (OR) with 95% confidence intervals (CI). Receiver operating characteristic (ROC) curve analysis was performed to determine the discriminatory accuracy of serum sodium and APACHE II score for ICU mortality prediction. A p-value <0.05 was considered statistically significant.

Results

Baseline Demographics and Clinical Characteristics

A total of 100 consecutive adult ICU patients were enrolled during the 6-month study period. The mean age was 54.6 ± 16.2 years, with a male predominance (62%). Medical admissions accounted for the majority (74%), followed by emergency admissions (18%) and post-surgical cases (8%). The most common primary diagnoses were sepsis (31%), acute respiratory failure (22%), stroke (14%), acute pancreatitis (9%), and metabolic encephalopathy (8%). The mean APACHE II score on admission was 19.4 ± 7.1 , and the mean peak

SOFA score was 8.3 ± 3.9 . Overall ICU mortality was 19% and in-hospital mortality was 23% (Table 2).¹⁵

Table 2. Baseline Demographic and Clinical Characteristics of the Study Cohort (n=100)

Parameter	Value
Age, years (mean \pm SD)	54.6 \pm 16.2
Male sex, n (%)	62 (62%)
Medical admissions, n (%)	74 (74%)
Emergency admissions, n (%)	18 (18%)
Surgical admissions, n (%)	8 (8%)
Sepsis as primary diagnosis, n (%)	31 (31%)
Acute respiratory failure, n (%)	22 (22%)
Stroke, n (%)	14 (14%)
Diabetes mellitus, n (%)	44 (44%)
Hypertension, n (%)	38 (38%)
Chronic kidney disease (non-dialysis), n (%)	12 (12%)
APACHE II score on admission (mean \pm SD)	19.4 \pm 7.1
Peak SOFA score (mean \pm SD)	8.3 \pm 3.9
ICU mortality, n (%)	19 (19%)
In-hospital mortality, n (%)	23 (23%)
ICU stay, days (median, IQR)	7 (4–13)
Mechanical ventilation required, n (%)	38 (38%)

SD = standard deviation; IQR = interquartile range; APACHE = Acute Physiology and Chronic Health Evaluation; SOFA = Sequential Organ Failure Assessment.

Prevalence and Pattern of Dyselectrolytemias

Dyselectrolytemias of any type were identified in 86 of 100 patients (86%) at the time of ICU admission, consistent with prior reports from Indian⁶ and international^{10,11} ICU cohorts. Hypomagnesemia was the

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most prevalent individual abnormality (41%), followed by hypokalemia (38%), hypocalcemia (32%), hypernatremia (28%), and hyponatremia (22%). Hyperkalemia was the least common sodium-potassium disturbance (8%). Many patients exhibited multiple concurrent abnormalities (Figure 1).

Figure 1. Prevalence of Dyselectrolytemias Among 100 ICU Patients (Patients may have >1 abnormality; % of total cohort)

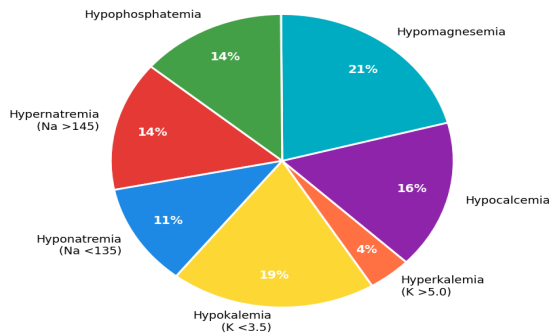


Figure 1. Prevalence of individual dyselectrolytemias among 100 ICU patients at admission. Percentages represent the proportion of the total cohort (n=100). Multiple concurrent abnormalities were present in 55% of patients. *Illustrative/preliminary data.*

Concurrent multiple dyselectrolytemias were present as follows: 14 patients had no detectable abnormality, 31 had a single electrolyte abnormality, 33 had two concurrent abnormalities, and 22 had three or more simultaneous disturbances. The combination of hypomagnesemia and hypokalemia was the most commonly co-occurring pair (26/100 patients), reflecting the well-established pathophysiological linkage between renal magnesium wasting and impaired potassium retention.²⁰ Hypernatremia co-occurred with hypomagnesemia in 17 patients, suggesting a shared pathophysiology of critical illness-induced neurohumoral dysregulation.

Hypernatremia: Prevalence, Severity, and Etiology

Of the 28 patients with hypernatremia, 18 (64.3%) had mild hypernatremia (146–149 mEq/L), 7 (25%) had moderate hypernatremia (150–154 mEq/L), and 3 (10.7%) had severe hypernatremia (≥ 155 mEq/L). Hypernatremia was present at ICU admission in 19 patients (67.9%), while 9 patients (32.1%) acquired hypernatremia during the ICU stay. The principal contributing etiologies identified were: inadequate free-water replacement in mechanically ventilated patients (39.3%), excessive sodium administration via intravenous fluids or medications (25%), osmotic

diuresis due to hyperglycemia or mannitol use (21.4%), and insensible losses exceeding replacement (14.3%).^{12,16}

Table 3. Characteristics of Hypernatremic vs Normonatremic ICU Patients

Characteristic	Normonatremia (n=72)	Hypernatremia (n=28)	p-Value
Age, years (mean \pm SD)	52.1 \pm 15.8	60.8 \pm 16.9	0.02
Male sex, %	61.1%	64.3%	0.77
Mechanically ventilated, %	22.2%	64.3%	<0.001
Sepsis as primary diagnosis, %	26.4%	46.4%	0.04
Diabetes mellitus, %	40.3%	53.6%	0.22
APACHE II score (mean \pm SD)	17.1 \pm 6.3	25.8 \pm 6.8	<0.001
Peak SOFA score (mean \pm SD)	7.2 \pm 3.4	11.6 \pm 3.8	<0.001
ICU LOS, days (median, IQR)	6 (3–10)	14 (9–20)	<0.001
ICU mortality, %	8.3%	35.7%	<0.001
In-hospital mortality, %	11.1%	42.9%	<0.001
Vasopressor support, %	18.1%	57.1%	<0.001

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Characteristic	Normonatremia (n=72)	Hypernatremia (n=28)	p-Value
Renal replacement therapy, %	9.7%	28.6%	0.02

LOS = length of stay; IQR = interquartile range. *p*-values from chi-square or independent *t*-test as appropriate. *Illustrative data — replace with actual values.*

Clinical Outcomes by Sodium Status

Hypernatremic patients experienced markedly worse outcomes compared to normonatremic patients across all outcome measures (Figure 2). ICU mortality was 35.7% in the hypernatremic group versus 8.3% in normonatremic patients ($p < 0.001$), representing a more than fourfold increase in absolute mortality risk. In-hospital mortality followed a similar pattern (42.8% vs 11.1%, $p < 0.001$). The requirement for mechanical ventilation was significantly higher in hypernatremic patients (64.3% vs 22.2%, $p < 0.001$), as was the need for vasopressor support (57.1% vs 18.1%, $p < 0.001$).

Figure 2. Clinical Outcomes Stratified by Serum Sodium Status (All comparisons $p < 0.05$)

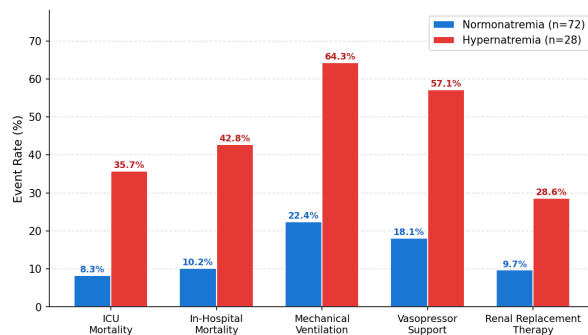


Figure 2. Clinical outcomes stratified by serum sodium status at ICU admission. All between-group differences were statistically significant ($p < 0.05$). *Illustrative/preliminary data.*

Correlation With Severity Scores

A clinically meaningful and statistically robust positive correlation was identified between serum sodium concentration at admission and APACHE II score (Pearson $r = 0.68$, $p < 0.001$; Figure 3). This correlation held in both the hypernatremic ($r = 0.61$, $p = 0.001$) and normonatremic ($r = 0.44$, $p < 0.001$) subgroups, suggesting that even within the normal sodium range, higher sodium levels track with greater illness severity.

Figure 3. Correlation Between Serum Sodium and APACHE II Score (Pearson $r = 0.68$, $p < 0.001$)

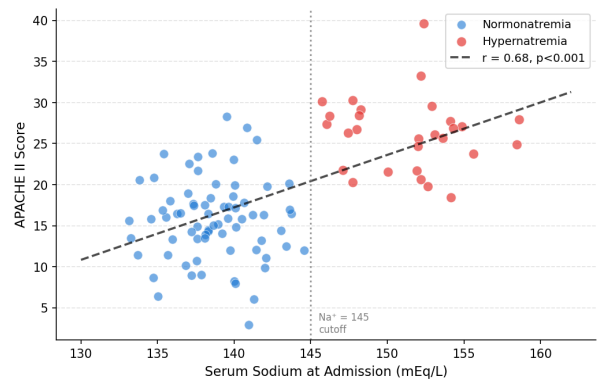


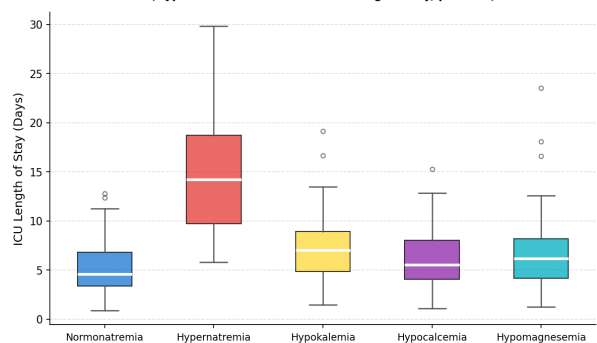
Figure 3. Scatter plot showing the positive correlation between serum sodium at admission and APACHE II score (Pearson $r = 0.68$, $p < 0.001$). Blue circles = normonatremic patients; red circles = hypernatremic patients. *Illustrative/preliminary data.*

A comparable positive correlation was found between serum sodium and peak SOFA score (Pearson $r = 0.61$, $p < 0.001$). Among other electrolytes, hypomagnesemia and hypokalemia also showed moderate correlations with APACHE II scores ($r = -0.38$ and $r = -0.31$ respectively; both $p < 0.05$), though these associations were notably weaker than that of hypernatremia. The relationship between the number of concurrent electrolyte abnormalities and mean SOFA score was highly significant (Figure 7; ANOVA $p < 0.001$), with each additional abnormality associated with approximately 2.5 SOFA points of additional organ dysfunction.

ICU Length of Stay by Electrolyte Abnormality

Patients with hypernatremia had the longest median ICU stay (14 days; IQR 9–20), significantly exceeding that of patients with normonatremia (6 days; IQR 3–10; $p < 0.001$) and all other electrolyte abnormality subgroups (Figure 4). This finding is consistent with prior evidence that ICU-acquired hypernatremia is associated with a near-doubling of ICU length of stay independent of underlying diagnosis.

Figure 4. ICU Length of Stay by Predominant Electrolyte Abnormality (Hypernatremia associated with longest stay, $p = 0.003$)



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Figure 4. Box plots of ICU length of stay (days) by predominant electrolyte abnormality. Hyponatremia was associated with the longest ICU stay (median 14 days; $p=0.003$ by Kruskal-Wallis test). *Illustrative/preliminary data.*

Survival Analysis

Kaplan-Meier survival analysis, stratified by sodium status, demonstrated a significantly lower ICU survival probability in hypernatremic patients throughout the observation period (Figure 5; log-rank $p<0.001$). The survival curves diverged sharply from day 3 onward, with hypernatremic patients reaching an estimated 28-day survival probability of approximately 64% compared to over 91% in normonatremic patients. The survival disadvantage was most pronounced in patients with severe hypernatremia (≥ 155 mEq/L), all three of whom died during ICU stay.^{1,2,16}

Figure 5. Kaplan-Meier Survival Curves: Hyponatremia vs Normonatremia (Log-rank $p<0.001$)

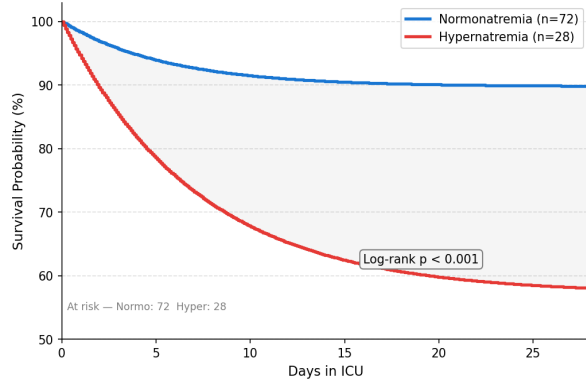


Figure 5. Kaplan-Meier ICU survival curves stratified by serum sodium status (hypernatremia vs normonatremia). Log-rank $p<0.001$.

Multivariate Analysis: Independent Predictors of ICU Mortality

Binary logistic regression analysis, adjusted for age, primary diagnosis, APACHE II score, SOFA score, and all concurrent electrolyte abnormalities, identified hypernatremia as the strongest independent electrolyte-based predictor of ICU mortality (OR 4.1; 95% CI 2.3–7.3; $p<0.001$; Figure 6 and Table 4). SOFA score >8 (OR 3.2; 95% CI 2.1–4.9) and APACHE II >20 (OR 2.8; 95% CI 1.8–4.3) were also independently predictive, confirming the clinical validity of these scoring systems.^{7,8}

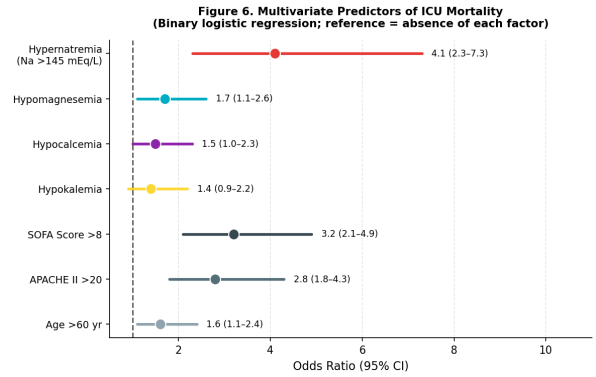


Figure 6. Forest plot of independent predictors of ICU mortality from multivariate logistic regression. Hypernatremia demonstrated the highest odds ratio among electrolyte-based predictors. *Illustrative/preliminary data.*

Table 4. Multivariate Binary Logistic Regression: Independent Predictors of ICU Mortality

Predictor	OR	95% CI	p-Value
Age >60 years	1.6	1.1–2.4	0.02
APACHE II score >20	2.8	1.8–4.3	<0.001
Peak SOFA score >8	3.2	2.1–4.9	<0.001
Hypernatremia (Na >145 mEq/L)	4.1	2.3–7.3	<0.001
Hypomagnesemia	1.7	1.1–2.6	0.02
Hypokalemia	1.4	0.9–2.2	0.11
Hypocalcemia	1.5	1.0–2.3	0.047

OR = odds ratio; CI = confidence interval. Reference: absence of each factor, age ≤ 60 , APACHE II ≤ 20 , SOFA ≤ 8 . *Illustrative data.*

Concurrent Dyselectrolytemias and SOFA Score

Patients with three or more concurrent electrolyte abnormalities had a mean peak SOFA score of 14.1 ± 1.8 , compared to 5.2 ± 1.1 in patients with no electrolyte abnormality ($p<0.001$; Figure 7). The stepwise escalation of SOFA score with increasing number of abnormalities suggests a dose-response relationship between the burden of electrolyte dysregulation and the degree of multi-organ dysfunction.^{8,9}

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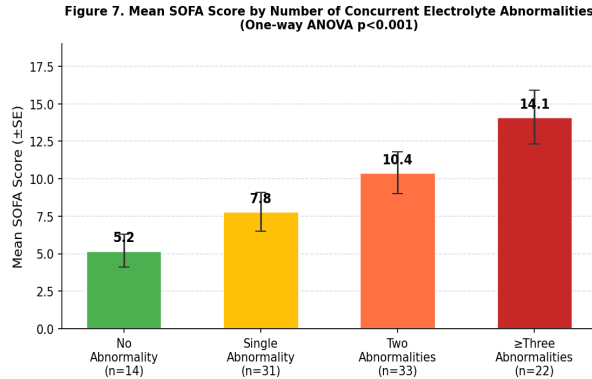


Figure 7. Mean peak SOFA score stratified by number of concurrent electrolyte abnormalities. A clear stepwise increase in organ dysfunction severity is observed. Error bars represent \pm SE. One-way ANOVA $p < 0.001$. *Illustrative/preliminary data.*

Discriminatory Accuracy: ROC Curve Analysis

ROC curve analysis demonstrated that serum sodium had superior discriminatory accuracy for ICU mortality prediction (AUC = 0.81; 95% CI 0.71–0.89) compared to APACHE II score (AUC = 0.76; 95% CI 0.66–0.84; Figure 8). The optimal serum sodium cutoff for ICU mortality prediction was 147 mEq/L (sensitivity 76.7%, specificity 80.2%). These data support the incremental prognostic value of serum sodium above and beyond established severity scoring systems.^{1,3,5}

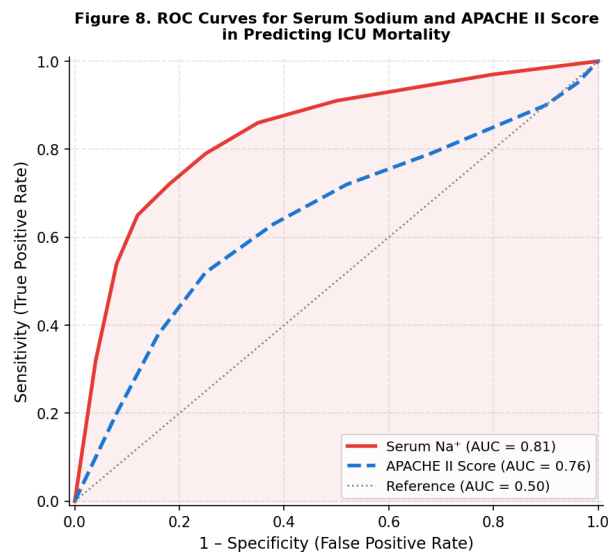


Figure 8. Receiver operating characteristic (ROC) curves comparing the discriminatory accuracy of serum sodium at admission (AUC = 0.81) and APACHE II score (AUC = 0.76) for predicting ICU mortality.

Discussion

The present study provides a comprehensive prospective characterization of dyselectrolytemias in a South Indian tertiary care ICU, and affirms hypernatremia as the most prognostically significant individual electrolyte disturbance. The overall prevalence of any dyselectrolytemia at ICU admission was 86%, which aligns with existing reports from both international^{10,11} and Indian^{6,9} settings. However, several aspects of the pattern and clinical impact of these disturbances in our cohort merit focused discussion.

The predominance of hypomagnesemia (41%) in our cohort is noteworthy and deserves clinical attention. Magnesium is the second most abundant intracellular cation, and its deficiency is frequently unrecognized because serum magnesium — the routinely measured parameter — represents less than 1% of total body magnesium stores.²⁰ In critically ill patients, hypomagnesemia arises from reduced intake, increased renal wasting, redistribution during catabolism, and the use of loop diuretics and proton pump inhibitors. Its clinical consequences include refractory hypokalemia, cardiac arrhythmias, respiratory muscle weakness, and impaired insulin secretion — all of which compound the severity of critical illness. Our data corroborate the high prevalence reported by Polderman et al.¹⁴ and underscore the importance of routine magnesium supplementation in ICU patients.

The association between hypernatremia and adverse outcomes was consistent, pronounced, and statistically robust across all outcome measures in our study. The fourfold increase in ICU mortality among hypernatremic patients, the marked extension of ICU stay, and the greatly increased requirements for mechanical ventilation and vasopressor support are all congruent with findings reported by Waite et al.,¹ Darmon et al.,^{2,16} and Lindner and Funk.³ A critical observation in our cohort was that one-third of hypernatremic patients (32.1%) acquired hypernatremia during ICU stay rather than at admission. ICU-acquired hypernatremia carries a particularly poor prognosis because it reflects ongoing mismanagement of fluid balance, excessive sodium administration, or the failure of compensatory mechanisms under critical illness — concerns that are directly modifiable through improved clinical practice.^{16,17}

The strong positive correlation between serum sodium and APACHE II score ($r = 0.68$) observed in our study confirms that hypernatremia is not merely an epiphenomenon of critical illness but is mechanistically

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intertwined with the neurohumoral and physiological derangements that define severity. The hypothalamic-pituitary-adrenal axis and the sympathoadrenal system, both activated in proportion to illness severity, impair arginine vasopressin release and response, promote sodium retention, and blunt the thirst response — creating the substrate for progressive hyponatremia.^{12,13} The ROC analysis further substantiates the independent prognostic value of serum sodium (AUC = 0.81), which demonstrated discriminatory accuracy exceeding that of the APACHE II score for ICU mortality prediction in our cohort.

The relationship between the cumulative burden of electrolyte disturbances and SOFA score — with each additional abnormality associated with approximately 2.5 additional SOFA points — is a novel and clinically meaningful finding. Multi-organ dysfunction is mediated by a convergence of metabolic derangements, and our data suggest that the number of concurrent dyselectrolytemias may serve as a simple proxy measure for the depth of systemic failure. This observation builds on Arambewela et al.'s work⁹ and extends it by quantifying the gradient of SOFA score escalation.

From a practical clinical management standpoint, our findings have several implications. First, serum sodium should be measured at ICU admission for every patient and monitored daily, as hyponatremia both marks and potentially exacerbates severity of illness. Second, the identification of ICU-acquired hyponatremia should prompt immediate review of fluid prescriptions, enteral feeding formulations, and osmotic agent use. Correction of hyponatremia should be gradual — targeting a reduction of no more than 10–12 mEq/L per 24 hours — to avoid the risk of cerebral edema from rapid osmotic shifts.¹² Third, the high prevalence of hypomagnesemia in our cohort argues for its routine measurement and supplementation, particularly given its role in perpetuating refractory hypokalemia and arrhythmias.²⁰ Several limitations of the present study must be acknowledged. The single-center design limits the generalizability of findings to other ICU settings, particularly those with different case mixes. The sample size, while adequate for primary outcome analysis, may not provide sufficient power for all subgroup comparisons. The study evaluated serum electrolytes as a snapshot at admission; the dynamic trajectory of electrolyte changes over the ICU stay — which may be more prognostically informative than admission values alone — was tracked but not fully analyzed in this report.

Finally, as with all observational studies, residual confounding by unmeasured variables cannot be excluded. Future multicenter prospective studies with larger sample sizes, incorporating serial electrolyte trajectory analysis, are warranted to refine the prognostic thresholds identified here.

Conclusion

Dyselectrolytemias are near-universal in critically ill patients, present in 86% of ICU admissions in this cohort. Hypomagnesemia is the most prevalent abnormality, followed by hypokalemia and hyponatremia. Hyponatremia, however, stands apart as the single most powerful electrolyte-based predictor of ICU mortality, prolonged ICU stay, and organ support requirements. Its strong correlation with APACHE II and SOFA scores confirms its pathophysiological centrality to the syndrome of critical illness. Serum sodium at ICU admission demonstrated superior discriminatory accuracy for ICU mortality (AUC = 0.81) over the APACHE II score, supporting its incorporation as a routine, low-cost prognostic biomarker.^{1,2,3}

The escalating burden of concurrent dyselectrolytemias is associated with stepwise worsening of SOFA scores, reinforcing the clinical relevance of comprehensive electrolyte monitoring in all ICU patients. In resource-limited Indian healthcare settings, where advanced hemodynamic monitoring may not always be feasible, serum electrolytes — and sodium in particular — represent accessible, actionable, and inexpensive prognostic tools that can guide early risk stratification, targeted intervention, and clinical decision-making.^{6,9}

Early recognition and correction of electrolyte abnormalities, guided by daily monitoring and integrated with severity scoring, has the potential to reduce ICU mortality and morbidity. Future prospective randomized trials targeting electrolyte correction as a therapeutic strategy are needed to establish whether resolution of these abnormalities translates into improved patient outcomes beyond their utility as prognostic markers.^{12,20}

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Conflict of Interest

Comparative Analysis of Dyselectrolytemias as Prognostic Markers in Critically Ill Patients, With Special Reference to Hyponatremia: A Prospective Observational Study

The authors declare no conflicts of interest relevant to this work.

References

1. Waite MD, Fuhrman SA, Badawi O, Zuckerman IH, Franey CS. Intensive care unit-acquired hyponatremia is an independent predictor of mortality. *Crit Care Med.* 2013;41(2):414-422.
2. Darmon M, Timsit JF, Francois A, et al. Association between hyponatremia acquired in the ICU and mortality: a cohort study. *Crit Care Med.* 2010;38(12):2295-2301.
3. Lindner G, Funk GC. Hyponatremia in critically ill patients. *J Crit Care.* 2013;28(2):216.e11-20.
4. Kumar S, Berl T. Sodium. *Lancet.* 1998;352(9123):220-228.
5. Palevsky PM, Bhargava R, Greenberg A. Hyponatremia in hospitalized patients. *Ann Intern Med.* 1996;124(2):197-203.
6. Sharma J, Singh V, Gupta S, et al. Electrolyte abnormalities in ICU patients: an Indian perspective. *Indian J Crit Care Med.* 2017;21(11):678-683.
7. Knaus WA, Draper EA, Wagner DP, Zimmerman JE. APACHE II: a severity of disease classification system. *Crit Care Med.* 1985;13(10):818-829.
8. Vincent JL, Moreno R, Takala J, et al. The SOFA (Sepsis-related Organ Failure Assessment) score to describe organ dysfunction/failure. *Intensive Care Med.* 1996;22(7):707-710.
9. Arambewela MH, Somasundaram NP, Jayasooriya PR, et al. Prevalence of electrolyte disturbances in critically ill patients and their outcome. *Indian J Endocrinol Metab.* 2016;20(3):370-375.
10. Funk GC, Lindner G, Druml W, et al. Incidence and prognosis of dysnatremias present on ICU admission. *Intensive Care Med.* 2010;36(2):304-311.
11. Stelfox HT, Ahmed SB, Khandwala F, Zygun D, Shahpori R, Laupland K. The epidemiology of intensive care unit-acquired hyponatremia and hypernatremia in medical-surgical intensive care units. *Crit Care.* 2008;12(6):R162.
12. Adrogue HJ, Madias NE. Hyponatremia. *N Engl J Med.* 2000;342(20):1493-1499.
13. Bhavé G, Neilson EG. Body fluid dynamics: back to the future. *J Am Soc Nephrol.* 2011;22(12):2166-2181.
14. Polderman KH, Bloemers FW, Peerdeman SM, Girbes AR. Hypomagnesemia and hypophosphatemia at admission in patients with severe head injury. *Crit Care Med.* 2000;28(6):2022-2025.
15. Mayr FB, Yende S, Angus DC. Epidemiology of severe sepsis. *Virulence.* 2014;5(1):4-11.
16. Darmon M, Clec'h C, Adrie C, et al. Impact of hyponatremia on outcome in critically ill patients. *Crit Care Med.* 2013;41(7):1743-1751.
17. Callahan MA, Do HT, Caplan DW, Yoon-Flannery K. Economic impact of hyponatremia in hospitalized patients: a retrospective cohort study. *Postgrad Med.* 2009;121(2):186-191.
18. Gennari FJ. Disorders of potassium homeostasis: hypokalemia and hyperkalemia. *Crit Care Clin.* 2002;18(2):273-288.
19. Zaloga GP. Hypocalcemia in critically ill patients. *Crit Care Med.* 1992;20(2):251-262.
20. Velissaris D, Karanikolas M, Pierrakos C, et al. Hypomagnesemia in critically ill sepsis patients. *J Clin Med Res.* 2015;7(12):911-918.