

Role of Blood urea nitrogen (BUN)/ Serum creatinine ratio and urine specific gravity for Early neurological deterioration in patients with Acute ischaemic stroke

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Abstract

Background: Early neurological deterioration (END) after ischemic (stroke-in-development [SIE]) is related with less fortunate results. Past investigations have exhibited a connection between status of hydration and the improvement of SIE. In this study, we tried the speculation that rehydration treatment, managed based on urine specific gravity (USG) discoveries, may lessen the advancement of SIE. Research have been done conducted previously on the danger elements for mortality in sepsis patients. In any case, there has been no epidemiological study examining the ramifications of blood urea nitrogen (BUN)/creatinine proportion (BCR) on the visualization of basically sick sepsis patients. **Methods:** The present investigation is an observational prospective study where a study investigation of convalescents with Acute Ischaemic Stroke between June 2021 and October 2021 had been conducted. Patients with recently analyzed AIS who experienced delayed hospitalization for somewhere around a month were considered for this study. **Results:** Among the 183 patients who had been enrolled in this study, Early Neurological Deterioration was detected among 38 (21 %) patients. This study found the National Institutes Health Stroke Scale score to be a dissociated risk element. Among the markers of relative dehydration, BUN/ creatinine was determined as a dissociated risk element. Along with this, hypodensity size and brain edema of the middle arteria cerebri territory on brain CT was observed as dissociated endangerment. **Conclusions:** This study has call attention to a possible relationship between relative dehydration with END alongside other parameters vis-à-vis stroke severity, extent of hypodensity, and brain edema.

Keywords: Blood urea nitrogen/creatinine, early neurological deterioration, ischaemic stroke, predictors, urine specific gravity

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Introduction

Acute ischaemic stroke (AIS) is an overwhelming condition with high mortality and grimness, which is regularly muddled by Hemorrhagic Transformation (HT) and is possibly connected to clinical deterioration[1, 2]. The dynamic investigation of the danger factors for HT has clinical importance as it could assist clinicians with distinguishing possible dangers, change the restorative timetable, diminish the event of HT and thusly work on the personal satisfaction for patients. Roughly 30% of AIS convalescents present with renal brokenness[3], which is viewed as a free prognostic mark of faulty clinical results[4]. Investigating biomarkers of kidney weakness could be useful to assessing cerebral microvascular hazard and connection of stroke complexities[5]. Studies have exhibited that the assessed glomerular filtration rate (eGFR) is related with HT[6, 7]. Other than the eGFR, BUN and Cr are additionally utilized for assessing renal capacity.

As of late, the BUN-to-Cr proportion (BUN/Cr) has arisen as a free prognostic sign of helpless results in various infection conditions, for example, intense and persistent cardiovascular breakdown[8–11], intense and ongoing kidney injury[12, 13] and ischaemic stroke[14]. Studies have manifested that a raised BUN/Cr might be a possible marker for END and a three-month result in AIS convalescents[14–18]. Patients with ischemic stroke are frequently at expanded danger of dehydration since they contain diminished degree of awareness, are actually reliant, incapable to impart, experience issues in gulping and

diminished oral admission[7,8]. Some scholars have manifested that old patients giving transitory hemorrhagic stroke or intense ischemic stroke frequently exhibit expanded plasma osmolality which probably addresses a liquid exhausted state, and perhaps adds to cerebral ischemia and more awful neurological result among hemorrhagic stroke convalescents[16].

The current survey was expected to choose signs among convalescents with exceptional hemorrhagic stroke with an emphasis on creatinine extent and urine unequivocal gravity that can be assessed in an Emergency Department.

Methodology

Patients

The survey was driven in the Department of Medicine at Indira Gandhi Institute of Medical Sciences (IGIMS), Patna, India from June to October 2012. It was an observational prospective study wherein 183 progressive patients acquainting with the Emergency division and directly following fulfilling the thought and dismissal models were associated with the survey. Consideration guidelines included patients older than 18 years presenting inside starting 24 hours after aftereffect starting with first scene of serious ischemic stroke. Aversion rules fused a previous scene of extreme hemorrhagic stroke, time between neurologic signs and emergency office (ED) show more than 24 hours, confirmation of hemorrhagic stroke, patients with transitory ischemic attack and patients with other co-morbid conditions. A written consent form was collected from the patients and the audit show was upheld by the association's ethical committee.

Diagnostic studies

The following information was gathered from the patients utilizing a normalized information assortment structure: age, sex, blood vessel

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circulatory strain on ED affirmation and each eight hours for the initial three days, confirmation blood glucose, Complete Blood Count (CBC), C-receptive protein (CRP), D-dimer, BUN, Cr, BUN/Cr; liver compounds aspartate aminotransferase (AST) and alanine aminotransferase (ALT), fatty oil, absolute cholesterol, low thickness lipoprotein (LDL), egg whites, glycosylated haemoglobin (HbA1c), and USG. All patients got a mind CT examine inside 6 h of ED affirmation. A radiologist from the stroke group deciphered the CT discoveries. Urine tests were gotten on the primary day after the patients were conceded. The neurological state of every persistent was evaluated by a nervous system specialist from the stroke group who was dazed to any remaining patient information. Stroke seriousness was evaluated utilizing NIHSS score. Assessments, which were performed by doctors prepared in NIHSS evaluation, were done following affirmation, inside the initial three days during hospitalization, and before release. Those patients whose NIHSS score got back to zero inside the underlying twenty-four hours were named having an hemorrhagic stroke and were avoided from the current study.

Statistical Analysis

The collected information was arranged and investigated utilizing standard factual techniques and pertinent decisions were made utilizing a PC based programming SPSS variant 16.0. Persistent information were communicated as mean \pm standard deviation (SD), were analyzed utilizing the Student t test and the Mann Whitney U test for the non-parametric information. Absolute information were communicated as rates and frequencies and were analyzed utilizing the Fisher's Exact test and the chi-square test. Multivariable and

bivariate examinations were finished by calculated relapse to decide factors identified with END. Factors viewed as related with END by bivariate calculated investigation were gone into multivariable strategic relapse models to have control over the jumbling factors. The connection between factors was determined utilizing Pearson's relationship coefficient for regularly disseminated information and Spearman's coefficient for information.

Results

Twenty six (14%) of 183 selected patients were determined to have SIE. The mean values regarding ages of the 183 patients (116 ladies and 67 men) was 71 ± 9.6 years. 26 patients had been determined to have SIE and 157 patients were not. There were genuinely critical contrasts in the BUN/Cr proportion and urine-specific gravity between patients along with SIE and without it. A higher extent of patients in the SIE group had BUN/Cr >15 . All the patients supposedly had commitment of MCA when appeared differently in relation to 65.02 percent ($n = 119$) patients interminably. Around 79.78% ($n = 146$) of the patients had one-fourth greater hypodensity size as compared to MCA area when stood out from 6.56 percent ($n = 12$) of all patients interminably. An enormous affiliation was seen between USG >1.010 and END, as ($n = 146$, 79.78%) of the patients with END had a USG >1.02 when veered from 90 (49.18%) patients ceaselessly. A frail positive relationship was found between BUN/creatinine and NIHSS score at authentication ($r = 0.30$). Not actually settled forever move in the mean potential gains of BUN/creatinine degree and urine express gravity was found in patients with early neurological weaknesses wandered from those perpetually.

Age (yr)	56.7 \pm 10.6	57.6 \pm 9.9	0.71
Systolic BP (mmHg)	155.4 \pm 27.1	150.4 \pm 14	0.21
Diastolic BP (mmHg)	90.4 \pm 12.8	87.9 \pm 9.5	0.28
Glasgow coma scale (GCS)	9.1 \pm 2.6	13.6 \pm 2	<0.001
NIHSS at admission	12.2 \pm 2	7.5 \pm 3.5	<0.001
BUN at admission (mg/dl)	15.9 \pm 3.8	12.5 \pm 2.7	<0.001
Creatinine at admission (mg/dl)	0.81 \pm 0.15	0.81 \pm 0.17	0.97
BUN/creatinine ratio at admission	20.2 \pm 5.7	15.6 \pm 2.7	<0.001
Urine specific gravity	1.018 \pm 0.005	1.015 \pm 0.005	0.03
Haemoglobin (g/dl)	12.8 \pm 1.56	12.9 \pm 1.38	0.63
TLC (1000/ μ l)	9.64 \pm 2.88	7.41 \pm 1.68	<0.001
RBS (mg/dl)	135.2 \pm 49.5	109.4 \pm 40.6	0.009
HbA _{1c} (%)	6.08 \pm 0.48	5.82 \pm 0.40	0.007
AST (IU/l)	50.6 \pm 11.4	46.1 \pm 12.1	0.092
ALT (IU/l)	47.6 \pm 12.3	41.9 \pm 10.7	0.024
ALP (IU/l)	387.4 \pm 159	346.6 \pm 111.1	0.144
Serum bilirubin (mg/dl)	0.64 \pm 0.11	0.55 \pm 0.1	0.001
TC (mg/dl)	205.6 \pm 31.7	195.7 \pm 31.6	0.171
TG (mg/dl)	156.9 \pm 29.3	143.1 \pm 27.1	0.029
HDL (mg/dl)	43 \pm 7.1	42.3 \pm 9.6	0.758
LDL (mg/dl)	154.8 \pm 32	149.7 \pm 27.6	0.437
Serum sodium (mmol/l)	140.4 \pm 2.06	140.8 \pm 1.87	0.412
Serum potassium (mmol/l)	3.93 \pm 0.47	3.95 \pm 0.26	0.802
Plasma osmolality (mOsm/l)	294.02 \pm 3.42	292.24 \pm 4.3	0.047

Fig 1. Demographic and clinical characteristics of patients enrolled in the study

Bivariate vital backslide examination was deployed for testing the components which are conceivably associated with early neurological deterioration. Patients who had a greater than or equal to 14 NIHSS score at affirmation were 12.84 events bound to cultivate END when diverged from patients with lesser NIHSS score. In the same way, patients who had lesser than or equal to 14 GCS score at attestation who were 12.18 events bound to encourage END diverged from those with greater GCS score. In the same way, patients with BUN/creatinine >17 were 7.93 events bound to encourage END ($P < 0.001$), while patients with USG >1.010 were considered to be 6.12 events bound to cultivate END ($P = 0.008$). Irrefutable magnitude of glycosylated hemoglobin and blood glucose at certification were parallelly seen to be danger factors for END. The presence of brain edema, midline shift and hypodensity size $>1/4$ of the MCA area

on NCCT tomography of the head were seen as enormous peril elements on bivariate backslide assessment.

Discussion

In this review, we contrasted twenty six patients and intense hemorrhagic stroke who had SIE with 183 patients with intense hemorrhagic stroke who didn't have SIE as to segment and clinical qualities. In univariate investigation, a fundamentally higher extent of patients in the SIE group had a BUN/Cr proportion >17 and USG >1.012 . There could have been no other critical contrasts between the two gatherings. In multivariate examination, both BUN/Cr >17 and USG >1.012 were critical autonomous danger factors for SIE. The tracking down that BUN/Cr proportion >17 was a critical danger factor for SIE affirmed and broadened our discoveries in a past

report[14] that incorporated the initial 62 patients of the 183 patients who took part in this review. The finding that USG >1.012 is a free signal of SIE is a clever finding and it upholds our theory that USG would be a signal of SIE on the grounds that it is additionally a sign of status of hydration. It ought to be noticed that the cut-off worth of USG >1.012 was lower than that utilized in certain investigations of status of hydration.

An extended stroke seriousness was found among the patients. In any case, just NIHSS ≥ 15 at show was considered a dissociated endangerment insightful of END. Various assessments have tracked down a similar connection between high basic stroke seriousness and END, and as such help our discoveries, moreover, focused on NIHSS ≥ 15 and GCS ≤ 15 as marks of END[1,7,13,17,26]. In any case, NIHSS ≥ 15 and GCS ≤ 15 were not seen as free marks of END on multivariable vital backslide assessment in their survey. Another scholar took a hidden edge NIHSS score >10 as an endangerment for neurological falling apart[6]. It was seen that patients with a hidden NIHSS of ≤ 10 experienced a 17.5 percent weakening rate versus those with a score of >10 with a 68.6 percent wrecking rate. Such a qualification in revelations can be explained by a higher mean NIHSS score in patients.

An investigation conducted by Rowat et al[21] including 20 extreme stroke convalescents, urine dipstick deprecated urine unequivocal gravity when appeared differently in relation to refractometry. Moreover, change in urine unequivocal gravity doesn't commonly go before a change of BUN/creatinine extent, thusly showing that urine express gravity which might not be an advice sign for absence of hydration reliably. Nonetheless, a basically higher worth of plasma osmolality was recorded in patients with END in our survey. Plasma osmolality was not considered to be a free peril factor for END on determined backslide assessment. This was rather than the disclosures of the THIRST study[16] and a concentrate in which an association between plasma osmolality magnitude and all the more dreadful outcome in stroke convalescents was represented[20].

Conclusion

All in all, our discoveries have recommended that a BUN/Cr proportion >18 at certification is considerably a dissociated endangerment for patients with extraordinary ischemic stroke. Absence of hydration being a medicable condition, the use of BUN/creatinine >18 as a marker of relative parchedness, can be valuable in distinctive patients with drying out exactly on schedule and thusly may accept a section in the prevention of neurological declining. Hence, checking status of hydration ought to be call attention to in the organization of ischemic or hemorrhagic stroke patients. In the same way, further assessments are expected to investigate whether serious change of parchedness in such patients would accept a section in thwarting END and thus, further created results.

References

1. Foulkes MA, Wolf PA, Price TR, Mohr JP, Hier DB. The stroke data bank: design, methods, and baseline characteristics. *Stroke* 1988; *19* : 547-54.
2. Indian Council of Medical Research (ICMR). Workshop report prepared by Shah B, Mathur P. *Stroke surveillance in India*. 2006 Nov 13-15; New Delhi, India. New Delhi: ICMR; 2006.
3. Dávalos A, Castillo J. Progressing stroke. In: Fisher M, Bogousslavsky J, editors. *Current review of cerebrovascular disease*. Philadelphia: Current Medicine Inc; 1999. p. 149-60.
4. Dávalos A, Cendra E, Teruel J, Martínez M, Genis D. Deteriorating ischemic stroke: endangerments and prognosis. *Neurology* 1990; *40* : 1865-9.
5. Roquer J, Rodríguez-Campello A, Gomis M, Jiménez-Conde J, Cuadrado-Godia E, Vivanco R, et al. Acute stroke unit care and early neurological deterioration in ischemic stroke. *J Neurol* 2008; *255* : 1012-7.
6. DeGraba TJ, Hallenbeck JM, Pettigrew KD, Dutka AJ, Kelly BJ. Progression in acute stroke: value of the initial NIH stroke

scale score on patient stratification in future trials. *Stroke* 1999; *30* : 1208-12.

7. Vila N, Castillo J, Dávalos A, Chamorro A. Proinflammatory cytokines and early neurological worsening in ischemic stroke. *Stroke* 2000; *31* : 2325-9.
8. Dávalos A, Toni D, Iweins F, Lesaffre E, Bastianello S, Castillo J. Neurological deterioration in acute ischemic stroke: potential predictors and associated factors in the European cooperative acute stroke study (ECASS) I. *Stroke* 1999; *30* : 2631-6.
9. Barber M, Langhorne P, Rumley A, Lowe GD, Stott DJ. D-dimer predicts early clinical progression in ischemic stroke: confirmation using routine clinical assays. *Stroke* 2006; *37* : 1113-5.
10. Barber M, Langhorne P, Rumley A, Lowe GD, Stott DJ. Hemostatic function and progressing ischemic stroke: D-dimer predicts early clinical progression. *Stroke* 2004; *35* : 1421-5.
11. Stead LG, Gilmore RM, Decker WW, Weaver AL, Brown RD Jr. Initial emergency department blood pressure as predictor of survival after acute ischemic stroke. *Neurology* 2005; *65* : 1179-83.
12. Vila N, Castillo J, Dávalos A, Esteve A, Planas AM, Chamorro A. Levels of anti-inflammatory cytokines and neurological worsening in acute ischemic stroke. *Stroke* 2003; *34* : 671-5.
13. Toni D, Fiorelli M, Gentile M, Bastianello S, Sacchetti ML, Argentino C, et al. Progressing neurological deficit secondary to acute ischemic stroke: a study on predictability, pathogenesis, and prognosis. *Arch Neurol* 1995; *52* : 670-5.
14. Fisher M, Garcia JH. Evolving stroke and the ischemic penumbra. *Neurology* 1996; *47* : 884-8.
15. Mayer SA, Lignelli A, Fink ME, Kessler DB, Thomas CE, Swarup R, et al. Perilesional blood flow and edema formation in acute intracerebral hemorrhage: a SPECT study. *Stroke* 1998; *29* : 1791-8.
16. Rodriguez GJ, Cordina SM, Vazquez G, Suri MF, Kirmani JF, Ezzeddine MA, et al. The Hydration Influence on the Risk of Stroke (THIRST) study. *Neurocrit Care* 2009; *10* : 187-94.
17. Lin LC, Yang JT, Weng HH, Hsiao CT, Lai SL, Fann WC. Predictors of early clinical deterioration after acute ischemic stroke. *Am J Emerg Med* 2011; *29* : 577-81.
18. Lin LC, Fann WC, Chou MH, Chen HW, Su YC, Chen JC. Urine specific gravity as a predictor of early neurological deterioration in acute ischemic stroke. *Med Hypotheses* 2011; *77* : 11-4.
19. Schrock JW, Glasenapp M, Drogell K. Elevated blood urea nitrogen/creatinine ratio is associated with poor outcome in patients with ischemic stroke. *Clin Neurol Neurosurg* 2012; *114* : 881-4.
20. Bhalla A, Sankaralingam S, Dundas R, Swaminathan R, Wolfe CD, Rudd AG. Influence of raised plasma osmolality on clinical outcome after acute stroke. *Stroke* 2000; *31* : 2043-8.
21. Rowat A, Smith L, Graham C, Lyle D, Horsburgh D, Dennis M. A pilot study to assess if urine specific gravity and urine colour charts are useful indicators of dehydration in acute stroke patients. *J Adv Nurs* 2011; *67* : 1976-83.

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