



CORRELATION OF BASELINE SERUM ALBUMIN LEVELS WITH PROGNOSIS OF GUILLAIN-BARRÉ SYNDROME

Neurology

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ABSTRACT

INTRODUCTION: Guillain-Barre syndrome (GBS) is a polyradiculoneuropathy characterized by a rapidly progressive bilateral paresis of the limbs. There is a limited number of studies in the literature which have explored the correlation between prognosis of GBS patients and serum albumin levels. So the present study was carried out.

MATERIAL AND METHODS: Prospective Longitudinal study was conducted among 60 patients admitted in the Neurology Department of MDM Hospital, Dr.SN Medical College, Jodhpur over a period of two years from January 2017 to December 2018. Serum albumin levels were determined. Patients were assigned into two groups. One with low albumin level (GROUP 1) and other with normal albumin level (GROUP 2). And the groups were compared based on Hughes's disability scores.

RESULTS: The albumin levels were negatively correlated with the Hughes' scores (admission/discharge). After treatment mean of the disability score in group 1 was significantly higher ($p < 0.05$) as compared to group 2. The difference of means of disability score {end line - baseline} was significantly higher in group 1 (0.09) as compared to group 2 (-0.92).

CONCLUSION: This study determined albumin level as an independent factor for assessing the prognosis of GBS.

KEYWORDS

Guillain-Barré Syndrome; polyradiculoneuropathy; Serum albumin levels; Prognosis.

INTRODUCTION:

Guillain-Barré syndrome (GBS) is a disorder of the peripheral nerves typified by rapidly ascending weakness evolving usually over days to one to four weeks.

Type of GBS are acute inflammatory demyelinating polyradiculoneuropathy (AIDP), acute motor axonal neuropathy (AMAN) and acute motor sensory axonal neuropathy (AMSAN).

The reported mortality of GBS in whole population ranged from 0.89 to 1.89 cases (median 1.11) per 100,000 people¹

Prognosis can be determined with serum albumin, glucose, cortisol and sodium levels as well as neutrophil / lymphocyte (N/L) ratio which is reported to be an inflammatory biomarker²⁷.

No prognostic biomarkers are available for GBS. Determining the albumin level at initial stage has advantage as its level may change with disease progression and procedure i.e. plasmapheresis, related fluctuations. Very few studies have been carried out in the past to show serum albumin level as the prognostic marker of GBS. Thus the current study was carried out with the above aim.

MATERIAL AND METHODS

A hospital based Prospective Longitudinal study was conducted among 60 cases of Guillain-Barre syndrome admitted in the Neurology Department of MDM Hospital, Dr.SN Medical College, Jodhpur over a period of two years i.e. January 2017 to December 2018.

Demographics, age, sex, clinical features, electrophysiology subtype, and treatment-related outcomes were assessed. A diagnosis was based on the criteria of the Brighton Collaboration GBS Working Group⁸.

Each patient was evaluated according to Hughes et al.'s disability score at the time of hospital admission and discharge⁹.

According to HDS, stage 1-3 was considered to represent good prognosis, and stage 3 or higher was considered to represent poor prognosis.

All the patients underwent physical and neurological examinations, complete blood count, electrolyte levels liver and kidney function tests, and lipid profile.

At our hospital, a serum albumin range of 3.4-5.4 gm/dL is considered normal.

To find out the prognostic value of serum albumin level in case of GBS the patients were divided into two groups. GROUP 1 with patients having S.albumin level < 3.4 and GROUP 2 S.albumin levels ≥ 3.4

The disorder was grouped into AIDP, AMAN, and AMSAN subgroups. The patients were categorized into the demyelinating form (AIDP) and axonal form (AMAN and AMSAN)

Patients criteria

Inclusion criteria

- Both sexes are included.
- Age: any age.
- All patients met diagnosis of GBS

Exclusion criteria

- severe organ failure
- Thyroid dysfunction
- Adrenal dysfunction
- Drug induced polyneuropathy.

Statistical analyses

All statistical analyses were performed using the SPSS 22.0 software package and Instat graphpad software. Descriptive statistics included mean \pm SD, number, and percentage. Student's *t*-test was used to compare continuous variables. A *p*-value of < 0.05 was considered statistically significant for all comparisons and correlations.

RESULT

Sixty patients were enrolled in this study. Of all the patients with GBS, 38 were men (63.33%), and 22 were women (36.66%). The mean age of the patient group was 34.57 ± 15.72 . Two patients died. Intravenous immunoglobulin was administered to all the patients except those having Hughes's disability score of 2.

The baseline mean serum albumin levels at the time of admission were 3.57 ± 0.29 and 18.33% of the patients ($n = 11$) had hypoalbuminemia.

The albumin levels were negatively correlated with the Hughes' scores (admission/discharge).

Fourty four of the patients had AIDP, 9 had AMAN, and 7 had

AMSAN. The Table 1 shows the comparisons of the demographic features and laboratory findings among the subgroups.

No significant relationship was found between S. uric acid, disability score before and after treatment. However, the mean albumin levels were significantly higher in axonal subtype as compared to AIDP (Table 1).

Table1: Comparisons of age, serum markers and disability score between AIDP and axonal subgroups

	AIDP (n=)	Axonal	p value
Age	3.11±13.9	38.56±19.9	0.24
Serum Albumin levels	3.51±0.3	3.73±0.21	0.009
Serum Uric acid	4.53±0.98	4.89±0.63	0.17
CSF protein	81.22±62.76	120.32±90.31	0.06
Disability score before	3.18±0.54	3.38±0.5	0.22
Disability score after	2.55±0.87	2.38±0.5	0.97

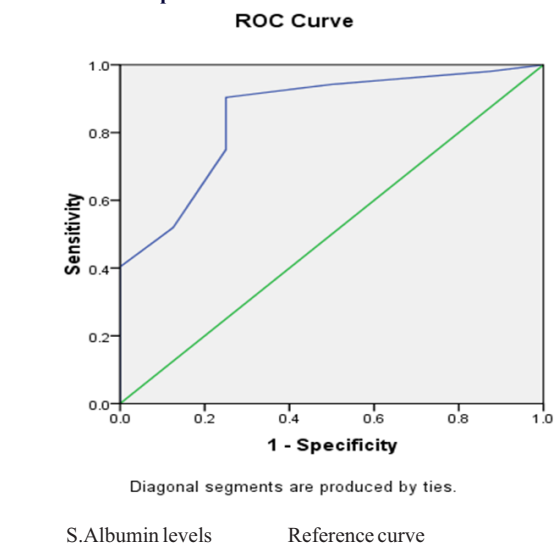
After treatment mean of the disability score in group 1 was significantly higher ($p < 0.05$) as compared to group 2. The difference of means of disability score {end line - baseline} was significantly higher in group 1 (0.09) as compared to group 2 (-0.92). (Table 2). This showed the negative correlation of S. albumin level and Hughes' disability score.

Table2: Comparison of means of Hughes' disability score in relation to S. albumin levels

Disability score	GROUP 1 (S. albumin < 3.4 gm/dl)	GROUP 2 (S. albumin ≥ 3.4 gm/dl)	p value
Baseline	3.18±0.75	3.24±0.48	0.73
Endline (After treatment)	3.27±1.10	2.33±0.59	0.0002
Score difference	0.09±1.04	-0.92±0.40	<0.0001

A cut-off S. albumin level of 3.35 predicted GBS, with 90% sensitivity and 75% specificity (ROC area under the curve [AUC] of 0.852, 95% CI, 0.712–0.992, $p < 0.001$). Fig: 1

Figure 1: Receiver Operating characteristic Curve (ROC) analysis of S. albumin for prediction of GBS



DISCUSSION

Our study demonstrated that there was a negative correlation between albumin levels and Hughes' scores (admission/discharge). With normal albumin level the disability score was less as compared to high Hughes' disability score with patients having low albumin level.

Albumin is a late-reacting negative acute-phase protein¹⁰.

In a study it was determined that a low serum albumin level is a strong marker of poor outcome in the setting of acute illness^{11,12}. GBS is also an acute illness and needs a prognostic marker so we tried to find out albumin as an independent model for clinical outcome in GBS. The

present study demonstrated that hypoalbuminemia is common in patients with GBS and there is a negative correlation between albumin levels and GBS disability. At time of admission there was no significant difference in the means of disability score. The results of our study are comparable with an international study, Willem-Jan R. et al¹³ that established an association of low albumin level with poor outcome in Intravenous Immunoglobulins treated GBS.

The results of our study are again corroborated by the outcomes of studies done by Ozdemir HH et al¹⁴, Naglaa Mohamed El-Khayat et al¹⁵, Ozlem Ethemoglu et al¹⁶, Yitao Zhang et al¹⁷ and Ghulam Shabbir et al¹⁸.

A study, done by Vincent JL et al¹⁹, Sapjiazko MJ et al²⁰ and Mamary AJ et al²¹ focusing on ICU and critically ill patients identified serum albumin as a biomarker for survival and the need for mechanical ventilation.

In this study, the most common Nerve Conduction Study pattern was demyelinating type, and this was comparable to the previous studies by Tavee and coauthors²². The infectious event is described to appear in 40-70% of patients²³⁻²⁵. In our series up to 43.5% of cases have had the infectious event, and respiratory infection was the most frequent one among all these infections.

In a nutshell, decreased albumin levels at the time of presentation of patient are associated with worse outcomes according to our study. However even now larger prospective studies are needed to support the findings of the present study.

REFERENCES

- Sejvar JJ, Baughman AL, Wise M, Morgan OW. Population incidence of Guillain-Barré syndrome: a systematic review and meta-analysis. *Neuroepidemiology*. 2011;36(2):123.
- Strauss J, Aboab J, Rottmann M, et al. Plasma cortisol levels in guillain-barré syndrome. *Crit Care Med* 2009;37:2436-40.
- Fokkink W-JR, Walgaard C, Kuitwaard K, et al. Association of albumin levels with outcome in intravenous immunoglobulin-treated guillain-barré syndrome. *JAMA Neurol* 2017;74:189-96.
- Huang Y, Ying Z, Quan W, et al. The clinical significance of neutrophil-to-lymphocyte ratio and monocyte-to-lymphocyte ratio in Guillain-Barré syndrome. *Int J Neurosci* 2018;128:729-35.
- Wang Y, Li G, Yang S, et al. Fasting glucose levels correlate with disease severity of Guillain-Barré syndrome. *PLoS one* 2015;10:e014575.
- Sipilä JO, Kauko T, Soilu-Hänninen M. Admission sodium level and prognosis in adult Guillain-Barré syndrome. *Int J Neurosci* 2017;127:344-9.
- Sahin S, Cinar N, Karsidag S. Are cerebrospinal fluid protein levels and plasma neutrophil/lymphocyte ratio associated with prognosis of Guillain Barré syndrome? *Neurol Int* 2017;9:7032.
- Sejvar JJ, Kohl KS, Gidudu J, Amato A, Bakshi N, Baxter R et al. Brighton Collaboration GBS Working Group. Guillain-Barré syndrome and Fisher syndrome: case definitions and guidelines for collection, analysis, and presentation of immunization safety data. *Vaccine*. 2011;29(3):599-612.
- Hughes RA, Newsom-Davis JM, Perkin GD, Pierce JM. Controlled trial prednisolone in acute polyneuropathy. *Lancet*. 1978;2(8093):750-3.
- Tsirpanlis G, Bagos P, Ioannou D, Bleta A, Marinou I, Lagouranis A et al. Serum albumin: a late-reacting negative acute-phase protein in clinically evident inflammation in dialysis patients. *Nephrol Dial Transplant*. 2005;20(3):658-9.
- Herrmann FR, Safran C, Levkoff SE, Minaker KL. Serum albumin level on admission as a predictor of death, length of stay, and readmission. *Arch Intern Med*. 1992;152(1):125-130.
- Vincent JL, Dubois MJ, Navickis RJ, Wilkes MM. Hypoalbuminemia in acute illness: is there a rationale for intervention? a meta-analysis of cohort studies and controlled trials. *Ann Surg*. 2003;237(3):319-334.
- Fokkink WR, Walgaard C, Kuitwaard K, Tio-Gillen AP, van Doorn PA, Jacobs BC. Association of Albumin Levels With Outcomes in intravenous Immunoglobulin-Treated Guillain-Barré Syndrome. *JAMA Neurol*. 2017; (2): 189-196.
- Ozdemir HH. Analysis of the albumin level, neutrophil-lymphocyte ratio, and platelet-lymphocyte ratio in Guillain-Barré syndrome. *Arq Neuropsiquiatr*. 2016;74(9):718-722.
- Naglaa Mohamed El-Khayat, Maha Ali Nada*, Heba Hamed El-Sayed, May Ahmad Nasr. Factors associated with prognosis of Guillain-Barré syndrome. *Clin Psychol Cog Sci*. 2018; 2(1):29-31.
- Ozlem Ethemoglu, Mustafa Calik. Effect of serum inflammatory markers on the prognosis of adult and pediatric patients with Guillain-Barré syndrome. *Neuropsychiatric Disease and Treatment* 2018;4:255-260.
- Yitao Zhang, Yanyin Zhao and Yi Wang. Prognostic factors of Guillain-Barré syndrome: a 111-case retrospective review. *Chinese Neurosurgical Journal*. 2018; 4:2-9.
- Ghulam Shabbir, Sumaira Fazal Nabi, Danyal Ahmed. Association of serum albumin levels and guillain barre syndrome (gbs) outcome. *Pakistan journal of neurological sciences* 2018; 13 (2):1-6.
- Vincent JL. Relevance of albumin in modern critical care medicine. *Best Pract Res Clin Anaesthesiol*. 2009;23(2):183-191.
- Sapjiazko MJ, Brant R, Sandham D, Berthiaume Y. Nonrespiratory predictor of mechanical ventilation dependency in intensive care unit patients. *Crit Care Med*. 1996;24(4):601-607.
- Mamary AJ, Kondapaneni S, Vance GB, Gaughan JP, Martin UJ, Criner GJ. Survival in patients receiving prolonged ventilation: factors that influence outcome. *Clin Med Insights Circ Respir Pulm Med*. 2011;5:17-26.
- Tavee JO, Polston D, Zhou L, et al. Sural sensory nerve action potential, epidermal nerve fiber density, and quantitative sudomotor axon reflex in the healthy elderly. *Muscle Nerve* 2014; 49(4):564-569.
- van Doorn PA, Ruts L, Jacobs BC. Clinical features, pathogenesis, and treatment of Guillain-Barré syndrome. *Lancet Neurol*. 2008;7(10):939-50.
- Vucic S, Kiernan MC, Cornblath DR. Guillain-Barré syndrome: an update. *J Clin Neurosci*. 2009;16(6):733-41.

25. Telleria-Diaz A, Calzada-Sierra DJ. Guillain Barre syndrome. *Rev Neurol*. 2002;34(10):966-76.
26. Aladro-Benito Y, Conde-Sendin MA, Munoz-Fernandez C, Perez-Correa S, Alemany-Rodriguez MJ, Fiuza-Perez MD, et al. Guillain-Barre syndrome in the northern area of Gran Canaria and the island of Lanzarote. *Rev Neurol*. 2002;35(8):705-10.
27. A prospective study on the incidence and prognosis of Guillain-Barre syndrome in Emilia-Romagna region, Italy (1992-1993). Emilia-Romagna Study Group on Clinical and Epidemiological Problems in Neurology. *Neurology*. 1997;48(1):214-21.
28. Lyu RK, Tang LM, Cheng SY, Hsu WC, Chen ST. Guillain-Barre syndrome in Taiwan: a clinical study of 167 patients. *J Neurol Neurosurg Psychiatry*. 1997;63(4):494-500.
29. Cuadrado JI, de Pedro-Cuesta J, Ara JR, Cemillan CA, Diaz M, Duarte J, et al. Guillain-Barre syndrome in Spain, 1985-1997: epidemiological and public health views. *Eur Neurol*. 2001;46(2):83-91.