



# THE PRIMARY CARE COMPANION FOR CNS DISORDERS

## **Supplementary Material**

**Article Title:** Central and Extrapontine Myelinolysis in the Setting of Hyperglycemia

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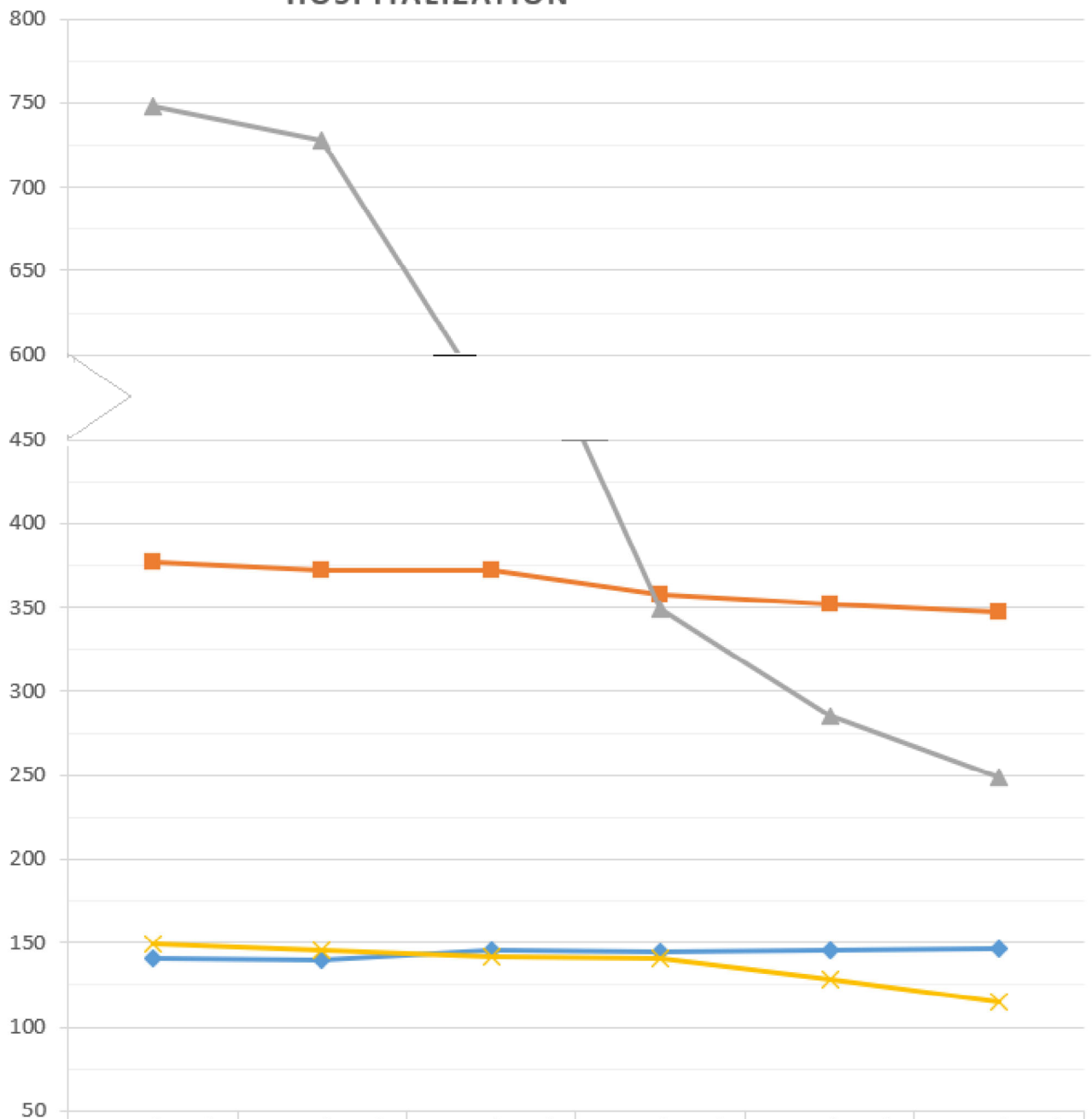
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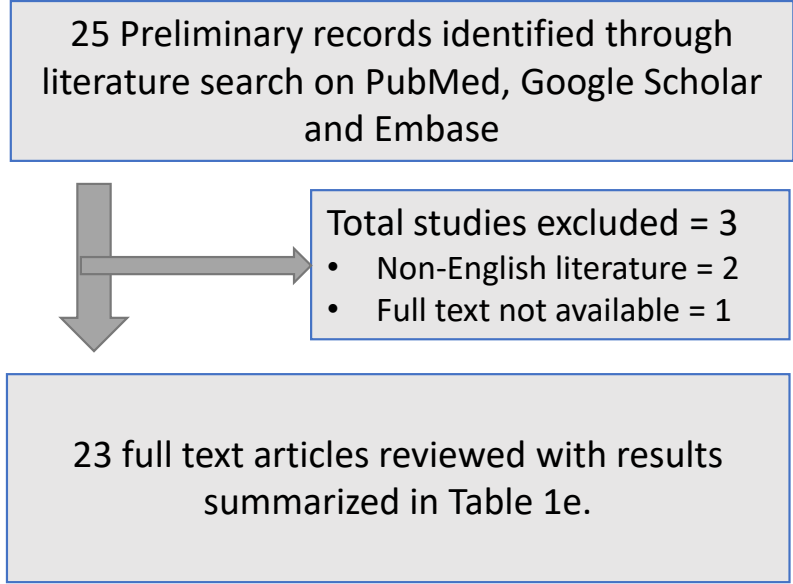
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## Supplementary Figure 1

### LABORATORY TRENDS DURING INITIAL 24 HOURS OF HOSPITALIZATION



	Day 0 (2100)	Day 1 (0200)	Day 1 (0430)	Day 1 (0730)	Day 1 (1300)	Day 1 (2030)
Sodium (mEq/L)	141	140	145	144	145	146
Osmolarity (mOsm/Kg)	377	372	372	358	352	347
Glucose (mg/dl)	748	728	569	349	285	249
BUN (mg/dl)	149	145	142	141	128	115



**Supplementary Figure 2: Flow Diagram of Literature Screening**

Supplementary Table 1: Literature Summary of Cases With Osmotic Demyelination Syndrome Associated With Hyperglycemia

Case Report	Age (years)	Sex	Glucose (mg/dl)	Osmolality (milli osmoles/Kg)	Na (mEq /L)	BUN (mg/dl)	Hb A1c %	K (mEq /L)	Presentation	Outcomes
<b>ODS cases attributed to hyperglycemic hyperosmolar state</b>										
Our Case	61	M	748	374	141	145	10.9	4.7	Found unconscious followed by persistent dysarthria and quadriparesis.	Discharge to inpatient rehab at 1 month. Improved mobility and swallowing but required total care.
Hirosawa et al (e1)	55	M	1011	324	126	43.8	17.8	3.2	AMS x 3 days and dysmetria.	Return to baseline over several weeks
Saini et al (6)	45	F	491	307	132	4	18	4.3	Ataxia, R UE pronator drift x 2 weeks	Gradual improvement in gait by second week
Pliquett et al (e2)	55	M	524	296	133	-	17.6	-	Dysmetria and Dysarthria x 5 days. Liver cirrhosis	Discharged to outpatient rehab Able to walk by 3 months
Hegazy et al (4)	43	F	828	-	181	11	-	4.6	AMS. Brisk plantar response developed during admission	Complete recovery by week 4
McComb et al (e3)	54	F	954	-	169	-	-	-	Obtunded	Deceased at 21 days
Mao et al (e4)	55	M	685	318	134	-	17.5	4.3	R focal seizures which evolved into EPC, R hemiplegia.	Regain of function by 1 m
Guerrero et al (5)	25	M	> 700	-	-	-	-	-	L hemiparesis developed as AG closed	-
Rodríguez-Velver et al (e5)	47	F	838	320	133	21	10.1	4.6	AMS and GTC. Worsening weakness on 24 hours	Return to baseline by 6 months
Kusumoto et al (e6)	87	F	1000	459	179	-	10.8	5.1	Fever, involuntary trunk and UE movements followed by coma	Return of spontaneous speech reported on 1 year followup
Gouveia et al (e7)	38	M	1225	412	154	38	-	5.4	AMS x 5 days, h/o chronic alcoholism	Remained poorly arousable, transferred to inpatient rehab
Yoshikawa et al (e8)*	84	F	465	308	113	168	-	6.3	AMS, worsened on HD 8.	Died in a few weeks
Bline et al (e9)*	14	F	> 600	-	130	64	13.8	2.8	Obtunded and emesis x 4days. Decline in mental status on HD 6.	Return to baseline by week 6
Kim et al (e10)	61	M	627	324	133	43.9	18.1	3.4	Dysarthria, dysphagia, dysmetria x 10 days. H/O cirrhosis	Gradually regained swallowing and mobility over weeks.
Kote et al (e11)	37	F	482	327.66	140	30.5	8.8	3.35	Dysarthria and dysmetria x 10-14 days. Exam decline after 6 hours of initiating treatment	Deceased at 15 days
Kumar et al (e12)	62	M	542	316	135	38	10.6	3.8	Dysphagia, dysarthria, and ataxia x 10-14 days	Improved dysphagia, dysarthria and walk independently upon discharge.
Lee et al (e13)	36	M	823	336	145	-	-	-	Dysphagia, dysarthria, and ataxia Chronic alcoholism	Dysphagia and ataxia resolved by 1 month, Dysarthria persisted at 4-month follow up.
Sharma et al (e14)	20	F	402	318	142	-	14.2	4.2	Dysarthria and generalized weakness x 15 days	Return to baseline by 30 days.
Yoong et al (e15)	53	M	594.6	340	135	-	14	4.6	Frequent falls and dysarthria x 2 months	Near complete recovery reported
Ramineni et al (e16)	50	M	546	318	136	66	13	3.6	Dysarthria, ataxia and generalized weakness x 10 days	Mild dysarthria at 1 month. Independent in all ADLs
Talluri et al (e17)	45	M	178	317	140	95	-	3.9	Intermittent ataxia, dysarthria and pseudobulbar affect	Return to baseline at 8 weeks
<b>Cases attributed to treatment of hyperglycemic hyperosmolar state</b>										
O'Malley et al (e18)	49	F	1910	399	134	23.3	-	2.2	Drowsy. No focal deficits. Flaccid quadriparesis noted on day 9 when weaned off sedation	Inpatient rehab, near complete recovery at 6 months
Burns et al (e19)	93	M	524	343	137	48	-	4.6	AMS and emesis. Ataxia developed 48 hours after admission	Improved gait at 1 month
Hsieh et al (e20)	29	M	646	-	138	-	-	2	AMS x 3 days. Declined 40 hours after admission	Remained vent dependent x 6 weeks, discharged to rehab.

\*Unclear if acute treatment played any role in development of ODS  
Abbreviations: AMS = Altered Mental Status; UE = Upper extremities; R = Right, L = Left  
H/O = History of; GTC = Generalized tonic clonic seizure; HD = Hospital day

**Supplementary Table 2: Literature Search Terms Used for the Review of Osmotic Demyelination Syndrome Associated With Hyperglycemia**

- Osmotic Demyelination Syndrome
- Osmotic Pontine Myelinolysis
- Extra pontine Myelinolysis
- Hyperglycemia
- Hyperosmolar Hyperglycemia
- Diabetic Ketoacidosis

## SUPPLEMENTARY REFERENCES

- e1. Hirosawa T. Osmotic demyelination syndrome due to hyperosmolar hyperglycemia. *Blood*. 2018;275:295.
- e2. Pliquett RU, Noll A, Ibe R, Katz A, Ackmann C, Schreiber A, et al. Hyperglycemia-related central pontine demyelination after a binge-eating attack in a patient with type-2 diabetes: a case report. *BMC endocrine disorders*. 2018;18(1):18.
- e3. McComb RD, Pfeiffer R, Casey J, Wolcott G, Till D. Lateral pontine and extrapontine myelinolysis associated with hypernatremia and hyperglycemia. *Clinical neuropathology*. 1989;8(6):284.
- e4. Mao S, Liu Z, Ding M. Central pontine myelinolysis in a patient with epilepsy partialis continua and hyperglycaemic hyperosmolar state. *Annals of Clinical Biochemistry*. 2011;48(1):79-82.
- e5. Rodríguez-Velver KV, Soto-García AJ, Zapata-Rivera MA, Montes-Villarreal J, Villarreal-Pérez JZ, Rodríguez-Gutiérrez R. Osmotic demyelination syndrome as the initial manifestation of a hyperosmolar hyperglycemic state. *Case reports in neurological medicine*. 2014;2014.
- e6. Kusumoto K, Koriyama N, Kojima N, Ikeda M, Nishio Y. Central pontine myelinolysis during treatment of hyperglycemic hyperosmolar syndrome: a case report. *Clinical Diabetes and Endocrinology*. 2020;6(1):23.
- e7. Gouveia CF, Chowdhury TA. Managing hyperglycaemic emergencies: an illustrative case and review of recent British guidelines. *Clin Med (Lond)*. 2013;13(2):160-2.
- e8. Yoshikawa R, Sako A, Kitagawa H, Hamasaki H, Okubo T, Hiraishi C, et al. The Development of Central Pontine Myelinolysis in a Type 1 Diabetic Patient With Anti-Insulin Antibody and Similar Clinical Manifestations to Autoimmune Polyglandular Syndrome 2012.
- e9. Bline K, Singh D, Poeppelman R, Lo W, O'Brien N. Extrapontine Myelinolysis and Microhemorrhages: Rare Finding in Pediatric Diabetic Ketoacidosis. *Pediatric Neurology*. 2018;89:68-70.
- e10. Kim JH, Park NH, Park JY, Kim S-J. Serial Magnetic Resonance Imaging Findings in Hyperglycemia-Related Osmotic Demyelination Syndrome: A Case Report. *Journal of the Korean Society of Radiology*. 2020;81(1):243-7.
- e11. Kote SS, Khandelwal A, Pathak DG, Nath R. An unusual case of osmotic demyelination syndrome without electrolyte changes in a patient with diabetes. *Journal of Neuroanaesthesiology and Critical Care*. 2016;3(02):145-8.
- e12. Kumar M, Nithyananthan P, Balaji T, Kannan R, Rajendran K. A case of hyperglycemia induced osmotic demyelination syndrome. *International Journal of Research in Pharmaceutical Sciences*. 2018;9(4):1262-4.
- e13. Lee S-P, See T-T, Kuo K-H. Central pontine myelinolysis in a chronic alcoholic patient with hyperglycemic hyperosmotic state. *Acta Neurol Taiwan*. 2013;22(3):142-3.
- e14. Sharma C, Kumawat BL, Panchal M, Shah M. Osmotic demyelination syndrome in type 1 diabetes in the absence of dyselectrolytaemia: an overlooked complication? *Case Reports*. 2017;2017.
- e15. Yoong CL, Lo F, Chew D. Sugars, Slips and Slurs: A Diabetic Patient with Hyperosmolar Hyperglycaemic State and Osmotic Demyelination Syndrome.

- e16. Ramineni K, Reddy K, Prusthi BK, Jakkani R. Pontine myelinolysis as the presenting complication of type 2 diabetes mellitus. *Indian journal of endocrinology and metabolism*. 2018;22(3):434-5.
- e17. Talluri S, Charumathi R, Khan M, Kissell K. Atypical presentation of central pontine myelinolysis in hyperglycemia. *Endocrinology, Diabetes & Metabolism Case Reports*. 2017;2017(1).
- e18. O'Malley G, Moran C, Draman MS, King T, Smith D, Thompson CJ, et al. Central pontine myelinolysis complicating treatment of the hyperglycaemic hyperosmolar state. *Annals of clinical biochemistry*. 2008;45(4):440-3.
- e19. Burns JD, Kosa SC, Wijdicks EF. Central pontine myelinolysis in a patient with hyperosmolar hyperglycemia and consistently normal serum sodium. *Neurocritical care*. 2009;11(2):251-4.
- e20. Hsieh H-C, Wu S-H, Chiu C-C, Ko K-C. Excessive sodium bicarbonate infusion may result in osmotic demyelination syndrome during treatment of diabetic ketoacidosis: a case report. *Diabetes Therapy*. 2019;10(2):765-71.