66-year-old woman with a pituitary mass

MEDICINE

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HPI

- 66 yo Taiwanese F with hx of seizure disorder was seen in NSG clinic after recent MRI revealed a pituitary mass (presenting symptoms-decreased vision, headaches)
- Has a hx of seizure disorder for 7 years and has been on carbamazepine
- Oct. 2011, her neurologist checked her MRI and discovered a suprasellar mass, ~1.5 in. diameter encroaching the optic chiasm

HPI continued...

- baseline poor vision in left eye-hx of epiretinal membrane surgery in left eye in 2007
- Best corrected acuity is 20/30 in rt. eye and 20/200 in lt. eye
- Automated visual fields per Optho revealed "a right temporal hemi-defect with mild nasal changes and left eye is diffusely abnormal"
- Ophtho recommended surgery

HPI continued...

- Seen by Dr. Lesniak in NSG clinic in 1/2012 who recommended a craniotomy to allow optimal decompression of the optic chiasm
- Ordered labs to evaluate pituitary function

Prolactin- 7.69 (4.8-23.3 ng/ml) FSH- 43.8 miU/ml LH- 24.3 miU/ml ACTH- 63 pg/ml (<52 pg/ml) IGF1- 172 (69-200 ng/ml) TSH- 1.56 (0.3-4 mcU/ml) Ft4- 0.81 (0.9-1.7 ng/dl)

HPI continued...

- Underwent lt. craniotomy on 1/31
- Immediately after surgery, patient complains of blindness in both eyes
- STAT head CT shows "blood products within resection cavity"
- Emergently taken back to the OR 1/31 for re-exploration, removal of blood contents and to free optic nerve

HPI continued....

- Ophtho consulted and vision continued to improve during hospital course
- Started on dexamethasone 6 mg q6h after surgery (tapered down by ~1 mg a day over 5 days until tapered off)
- Patient scheduled to leave for rehab on Monday 2/6, Endo consulted for evaluation

Past Medical History

- Seizure disorder (dx 7 years ago)
- Diet controlled diabetes
- Hypertension
- Newly diagnosed pituitary mass
- Hx of cataract surgery in both eyes
- Hx of epiretinal membrane surgery in left eye in 2007 with poor vision in left eye at baseline

Home meds:

Carbamazepine 200 mg TID Diovan 80 mg daily

Hospital Meds:

Brimondine eye drops Carbamazepine 200 mg TID Valsartan 80 mg po daily Novolog 1-5 unit sliding scale

Social Hx, Family Hx, ROS

- Social Hx
- Married and lives with her husband, who is a retired Ob-gyn in NW Indiana.
- -Has 2 grown daughters.
- Originally from Taiwan
- Not working
- Denies smoking, etoh, drugs
- <u>Family Hx-</u> no hx of pituitary tumors, both her parents died of "natural causes." Both daughters are healthy.
- ROS-
- -Denies any weight changes
- -vision has been poor for years
- -infrequent headaches.
- -denies galactorrhea
- -no changes in ring size
- -denies muscle weakness

Physical Exam

□ VS: 97° F, 118/59, RR 18, 98% on RA, 5'2", 139 lbs

Gen: NAD

Head: s/p craniotomy, stitches in place

Eyes: ptosis in R>L, right cranial nerve III palsy documented per

ophtho

Neck: no thyromegaly

Heart: +S1/S2, no murmurs

Lungs: CTA b/l

Abdomen: soft, ntnd, +bs

Extremities: no c/c/e

Neuro: Alert and oriented x 3, no focal deficits, muscle

strength normal, right cranial nerve III palsy, other cranial nerves

intact

Labs

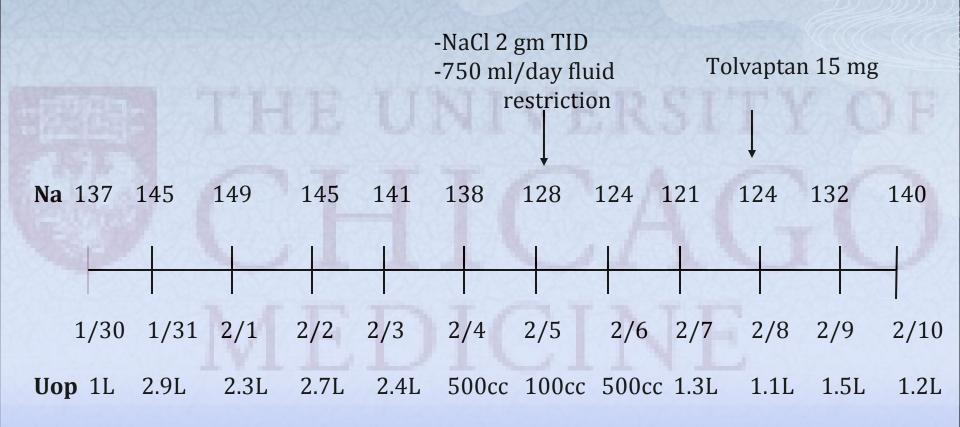
□ <u>138 | 104 | 22 /</u>120 4.1 | 22 | 0.4 \

A1c-6.5

MEDICINE

12.1 \ <u>10.7 /</u>311 / 33 \

Serum sodium and Urine output



Additional labs from 2/5

- Plasma Osm-263 (289-308 mOsm/kg)
- Urine Osm- 776 (500-800 mOsm/kg)
- Urine Na- 159 mEq/L

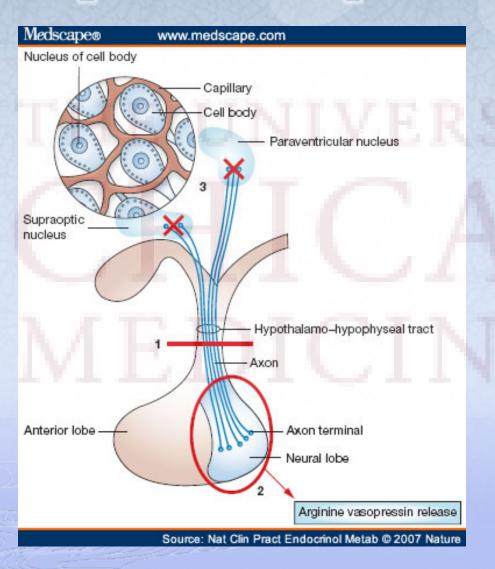
SIADH

- □ Plasma osm <270 mosmol/kg H₂0
- Inappropriate urine concentration
 - Uosm > $100 \text{ mosmol/kg H}_2\text{O}$
- Euvolemic
- Elevated urine Na >40 mmol/L

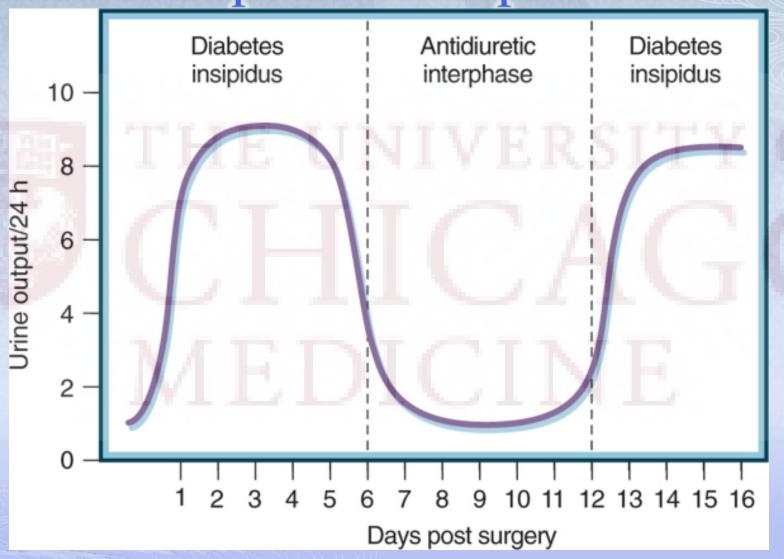
SIADH and pituitary surgery

- Hyponatremia is a late common complication of pituitary surgery, occurring in 21-35% of cases
- Major cause is inappropriate ADH release from injured pituitary gland
- Fall in serum sodium is most severe on the 6-7th post-operative day

Triphasic Response

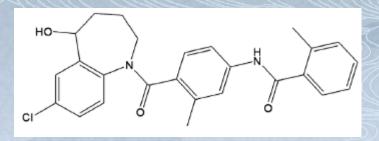


Triphasic Response



Williams Textbook of Endocrinology, 12th ed.

Tolvaptan



 An arginine vasopressin (AVP) receptor antagonist with affinity for AVP receptor subtypes V2 and V1a.

 Antagonism of the V2 receptor promotes excretion of free water, resulting in net fluid loss, increased urine output, decreased urine osmolality, and subsequent restoration of serum Na levels.

Tolvaptan

- Tolvaptan was approved by the U.S. Food and Drug Administration on May 19, 2009.
- Should only be used in a hospital setting where serum
 Na can be monitored closely
- Too rapid correction of hyponatremia (e.g., ≥ 12 mEq/L/24 hours) can cause central pontine myelinolysis resulting in dysarthria, mutism, dysphagia, lethargy, spastic quadriparesis, seizures, coma and death.
- Patient should be allowed to drink when thirsty while on the medication

Carbamazepine and SIADH

- Acts by stimulating release of ADH and potentiates its action by promoting reabsorption of water
- Has been used as an alternative treatment for DI
- Development of SIADH has been reported in the literature since 1973
- Risk of carbamazepine-induced hyponatremia has been associated with increased age (>30 years)
- Patients with additional risk factors (ie. Pituitary surgery) could be at increased risk of developing SIADH

Hospital Course

- Patient's sodium stabilized after receiving 1 dose of tolvaptan
- Her vision also improved.
- Sent to rehab with salt tablets and fluid restriction
- Serum sodium to be monitored daily while at rehab and NaCl tabs to be tapered
- Outpatient Endo f/u and Ophtho f/u.

Take Home Points

- Typical triphasic response of urine volume follows sectioning of the pituitary stalk induced by surgery.
- The 1st phase of diabetes insipidus occurs immediately postoperatively and continues to day 6.
- The second phase of antidiuresis occurs from day 7-day
 12.
- The third phase is the reocurrence of diabetes insipidus on day 13.
- Carbamazepine can increase sensitivity to ADH.
- Tolvaptan is an effective treatment of post-operative SIADH.

References

- Sherlock M, Thompson CJ. The syndrome of inappropriate antidiuretic hormone: current and future management options. Eur J Endocrinol. 2010 Jun;162 Suppl 1:S13-8. Epub 2010 Feb 17. Review. PubMed PMID: 20164215.
- Gold PW, Robertson GL, Ballenger JC, Kaye W, Chen J, Rubinow DR, Goodwin FK, Post RM. Carbamazepine diminishes the sensitivity of the plasma arginine vasopressin response to osmotic stimulation. J Clin Endocrinol Metab. 1983;57(5):952.
- Kamiyama T, Iseki K, Kawazoe N, Takishita S, Fukiyama K. Carbamazepine-induced hyponatremia in a patient with partial central diabetes insipidus. Nephron. 1993;64(1):142.
- Sachdeo RC, Wasserstein A, Mesenbrink PJ, D'Souza J. Effects of oxcarbazepine on sodium concentration and water handling. Ann Neurol. 2002;51(5):613.
- Palm C, Pistrosch F, Herbrig K, Gross P. Vasopressin antagonists as aquaretic agents for the treatment of hyponatremia. Am J Med. 2006 Jul;119(7 Suppl 1):S87-92. Review. PubMed PMID: 16843091.
- Vaghasiya RP, Devita MV, Michelis MF. Serum and urine responses to the aquaretic agent tolvaptan in hospitalized hyponatremic patients. Int Urol Nephrol. 2011 May 24. [Epub ahead of print] PubMed PMID: 21607553.