Dysnatremia v2.2: Table of Contents

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Postop Neurosurgery At Risk for Diabetes Insipidus

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Background

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Dysnatremia v2.2: Postop Neurosurgery At Risk for Hyponatremia

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Monitoring

Orders

- Serum sodium and serum osmolality qam x 3 days
- Daily weight
- Strict intake and output
- If no void over 8 hours, bladder scan or ask patient to void

Call Contact Provider for

- Sodium <135
- Intake and output positive > 40 ml/kg over 8 hours
- Urine output <0.5 ml/kg/hr or no void over 8 hours
- Urine output >4 ml/kg/hour x 2 voids in a row
- Mental status change (particularly decreased responsiveness)

Inclusion Criteria

 Patients with procedure at high risk for low sodium

Exclusion Criteria

Age <1 year

Procedures at High Risk for Low Sodium

- Craniotomy
 - Craniosynostosis repair/ cranial vault expansion/frontal orbital advancement
 - Hemispherectomy/lobectomy
 - · Placement of Grid and strip
 - Tumor resection/biopsy
- Endoscopic 3rd ventriculostomy (ETV)
- Insertion of lumbar drain
- Laser ablation
- · Subgaleal shunt insertion
- VP shunt externalization/external ventricular drain

Diagnostic Criteria for SIADH

- Serum sodium <135 mEq/L AND
- Serum osm <275 mOsm/kg AND
- · Euvolemic or volume overloaded

Orders

 Check serum sodium, serum osm, urine osm, urine specific gravity, urine sodium, BUN, creatinine

Consider more frequent neuro checks

Serum sodium?

Sodium >135 Continue monitoring

Sodium <135 (if not patient's normal baseline)

Serum

Osm?

Serum Osm ≥275

No, volume down (see triangle)

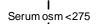
Pseudohyponatremia?

Serum sodium <135 and serum osm ≥275

Consider pseudohyponatremia

Check glucose and lipids

Off Pathway



Euvolemic?

Yes, euvolemic/volume overload

SIADH

Cerebral Salt Wasting?

Serum sodium <135, serum osm <275, Intravascular depletion

- · Consider cerebral salt wasting
- Run normal saline fluids at maintenance rate, or increase fluid rate if already running NS fluids

4 hours

Monitoring

 4-6 hours after change in therapy, recheck serum sodium, serum osm, urine osm, urine specific gravity

Next steps individualized to patient

Treatment options

- Restrict fluids, take into account volume status, IV medication volume, and medications that cause low sodium
- Do not add sodium (not a problem of sodium deficiency)
- Do not give NS bolus to raise serum sodium
- If sodium is 120 or less → go to PICU, consider hypertonic saline
- If sodium does not improve, consider alternative diagnoses

Assessment

• Continue to assess volume status, heart rate, etc

Signs of intravascular depletion: tachycardia, prolonged capillary refill, decreased weight, BUN elevated



4 hours

Dysnatremia v2.2: Periop Neurosurgery At Risk for Diabetes Insipidus

Approval & Citation

Summary of Version Changes

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Inclusion Criteria

 Postop brain tumor for known craniopharyngioma or transsphenoidal hypophysectomy or any other transcranial surgery involving pituitary or hypothalamus

Exclusion Criteria

Age <1 year

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How Dysnatremia Occurs

Diagnostic Criteria for Diabetes Insipidus (DI)

All are present:

- Serum sodium > 145 mEq/L
- Serum osmolality >300 mOsm/kg
- Urine osmolality <300 mOsm/kg
- Urine output > 4 ml/kg/hr

Neurosurgery Clinic

- · Assess for history of DI
- Is patient awakening at night to urinate?
- Labs
 - Serum sodium
 - Serum osmolality
 - Urine osmolality
 - Intake and output

PASS Clinic

 Confirm DI assessment has been done and is current

Endocrine Assessment

- If urgent, contact Endocrine consult provider on call
- If not urgent, schedule Endocrine clinic visit
- Known DI→use Preop Known DI page

Operating Room

If high risk for DI

• Insert temporary central IJ line

High risk for DI Criteria

- Cranopharyngioma or
- Perisellar mass ≥ 2cm and age <10 years

All patients

Foley catheter

Suspicious 5 4 1

or has Diagnosis

of DI

- Treat diabetes insipidus if criteria met
 - A higher sodium level may be appropriate if mannitol was given
 - Start vasopressin infusion at 0.5 milliUnit/kg/hr. Increase 0.5 milliUnit/kg/hr every 10-15 minutes to urine output 0.5 to 2 ml/kg/hr.
 - NS IV 1/3 maintenance
 - Anesthesiologist checks sodium and manages fluids to avoid hyponatremia
 - Do not replace urine output if on vasopressin infusion
 - Replace operative losses as required





Dysnatremia v2.2: Postop Neurosurgery At Risk for Diabetes Insipidus

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From Periop

Monitoring (PICU)

- Serum Sodium q4h
- · Strict intake and output
- · Daily weight
- Call for urine output >4 ml/kg/hr or serum sodium >145, and:
 - Obtain serum sodium, serum osm, urine osm, urine sodium
 - Consider ordering vasopressin for DI to bedside
 - If urine output >6 ml/kg/hr, may start vasopressin infusion while waiting for lab results after discussing with PICU attending

Inclusion Criteria

 Postop brain tumor for known craniopharyngioma or transsphenoidal hypophysectomy or any other transcranial surgery involving pituitary or hypothalamus

Exclusion Criteria

- Age <1 year
- Acute neurologic injury
 - Acute or chronic kidney injury

Phase 1

(PICU)

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Diagnostic Criteria for Diabetes Insipidus (DI)

All are present:

- Serum sodium > 145 mEq/L
- Serum osmolality >300 mOsm/kg
- Urine osmolality <300 mOsm/kg
- Urine output > 4 ml/kg/hr

High risk for DI

- Craniopharyngioma or
- Perisellar mass ≥2 cm and age <10 years

-DI Present-

Vasopressin IV

- Start vasopressin infusion at 0.5 milliUnit/kg/hr
- Increase 0.5 milliUnit/kg/hr every 10-15 minutes until urine output <2 ml/kg/hr
- If NPO, use NS IV 1/3 maintenance rate
- Be judicious in giving additional fluids, use only for clear intravascular depletion
- Serum sodium in 2 hours, then q4h
- Call for
 - Serum sodium <140
 - Urine output <0.5 or >2 ml/kg/hr x 2 hours in a row
 - Discuss with PICU attending whether to change infusion rate or stop IV fluids
- Consult Endocrine
- If tolerating PO
 - · Allow PO intake to thirst
 - Discontinue IV fluids
 - Discontinue vasopressin starting at 0800 on postop day 2

Urine output > 4 mL/kg/hr and urine specific gravity < 1.010

*Criteria for next desmopressin dose

- 1. Urine output > 4 mL/kg/hr AND
- 2. At least 4 hours from last desmopressin dose **AND**
- 3. Urine specific gravity < 1.010 OR serum sodium rising over the last two checks

Phase 2 (PICU or

acute care)

Desmopressin PRN

- First dose desmopressin typically 50-100 mcg
- Serum sodium q4h, urine specific gravity each void (max 1/hour; q4h if foley in place)
- When *criteria for next desmopressin dose met, call Endocrine to discuss timing and size of dose

Endocrine determines it is appropriate to schedule desmopressin

Acute Care (Floor) Criteria

- Tolerating PO
- If patient has NO intact thirst drive, establish fluid intake plan
- Manageable sodium monitoring
- Has tolerated oral desmopressin for 24 hours

Desmopressin Scheduled

- Desmopressin BID (sometimes TID)
- Serum sodium BID 2 hours prior to scheduled dose
- Hold dose for sodium <135 and call Endocrine
- When planning for discharge, call Endocrine to discuss plan for home fluid balance and sodium monitoring



Phase 3

Dysnatremia v2.2: Periop Known Diabetes Insipidus

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Inclusion Criteria

 Known diabetes insipidus on scheduled desmopressin at home

Exclusion Criteria

- Age <1 year
- Acute or chronic kidney injury

PASS Clinic

- Assess patient
 - What is desmopressin dose/frequency
 - How much patient is urinating (how well is DI controlled)
 - Any special feeding needs, oral intake
- · Include in PASS clinic visit summary
 - Most recent sodium
 - · Procedure details
 - Date/time (first case preferred)
 - · Length of case
 - Planned disposition
 - · Patient instructions
 - · Take usual desmopressin dose before procedure
 - Clears until 2 hours before procedure
- Labs
 - Serum sodium
 - If telehealth visit, contact endocrinologist to determine if serum sodium is required prior to day of procedure

Contact Endocrinologist via In Basket to develop plan for DI management

Default Plan of Care

- Schedule as first case in the morning
- Give usual desmopressin morning dose
- Include length of case and disposition (home, acute care, critical care) and admitting service
- If long case, plan for desmopressin during case (suggested IV desmopressin dose)
- Anesthesiologist
 - Run IV fluids at 2/3 maintenance plus intraoperative losses, unless otherwise specified (ie g-tube feeding plan)
 - · Do not replace urine output
 - Check sodium and manage fluids to avoid hyponatremia.
 If diagnostic imaging is reason for sedation, and patient is at clinical baseline, serum sodium is not required.



Dysnatremia v2.2: **ED or Acute Care Known Diabetes Insipidus**

Approval & Citation

Summary of Version Changes

Explanation of Evidence Ratings

ED

Assessment

- Is patient tolerating oral intake (can keep down liquids after ondansetron)?
- Any decreased input or increased losses (vomiting, diarrhea, dehydration)?
- Any mental status change?
- Any history of adrenal insufficiency?
- · Assess volume status and perfusion
- What is time and dose of last desmopressin?
- · What is time of last void?
- Chart and calculate I/O
- Check serum sodium (ePOC)
- · Discuss with Endocrinology
 - What is target sodium (typically 140-150)
 - If history of adrenal insufficiency, consider stress dose hydrocortisone

Management

- Strict intake and output
- What should happen before patient leaves ED?
 - Give desmopressin if necessary
 - Start IV fluids if necessary
 - · Enter acute care orders

Inclusion Criteria

- Known neurogenic diabetes insipidus on scheduled desmopressin at home
- Concern for acute illness

Exclusion Criteria

- Age <1 year
 - Acute or chronic kidney

Free Water Deficit Equation

Free water deficit = $60\% \times weight$,

See calculator (for SCH Only)

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PICU Transfer Criteria

- Unable to take desmopressin by mouth
- Abnormal mental status
- Seizure during this illness with high or low sodium
- Severe intravascular depletion

Be cautious giving fluid bolus; may cause rapid drop in sodium or rapid increase in urine output.

kg x (1 - target Na/current Na)

Discuss with Endocrinology

Plan for Initial Management (

Sodium <135

0-5

above

target

sodium

Do not give desmopressin or IV fluids without Endocrinology approval **Tolerating Enteral Intake**

· Give desmopressin if due per home schedule

• Enteral intake water to thirst (if no intact thirst: per home feeding plan)

Not Tolerating Enteral Intake

• Give desmopressin if due per home schedule

Consult Endocrine

• D5½NS at 2/3 maintenance rate (may not need overnight fluids)

5-10 above target

sodium

Tolerating Enteral Intake

- Give desmopressin if due per home schedule
- Enteral intake water to thirst (if no intact thirst: per home feeding plan)
- Consider goal enteral water intake over next 6 hours

Not Tolerating Enteral Intake

- Give desmopressin if due per home schedule
- D5½NS to replace water deficit over 24 hours
- Replace urine output > 2 ml/kg/hour over the next 2 hours
- Serum sodium q4h
- If sodium not decreasing, add insensibles and replace full urine output

11-20 above target sodium

Tolerating Enteral Intake

- Give desmopressin if due per home schedule, or urine output > 4 ml/kg/ hour and > 4 hours from the last dose
- Enteral Intake water with calculated max sodium decrease serum sodium 10 mmol/L in 24 hours
- Serum sodium q4h

Not Tolerating Enteral Intake

- Give desmopressin if due per home schedule, or urine output > 4 ml/kg/hour and > 4 hours from the last dose
- D5½NS to decrease serum sodium 10 mmol/L in 24 hours
- Replace urine output > 2ml/kg/hour over the next 2 hours
- Serum sodium q4h
- If sodium not decreasing, add insensibles and replace full urine output

Not Tolerating Enteral Intake or No Intact Thirst

Transfer to PICU for vasopressin infusion



Patients with a chronically elevated sodium are at risk for cerebral edema if their sodium is corrected too quickly (>0.5 mEq/hr)

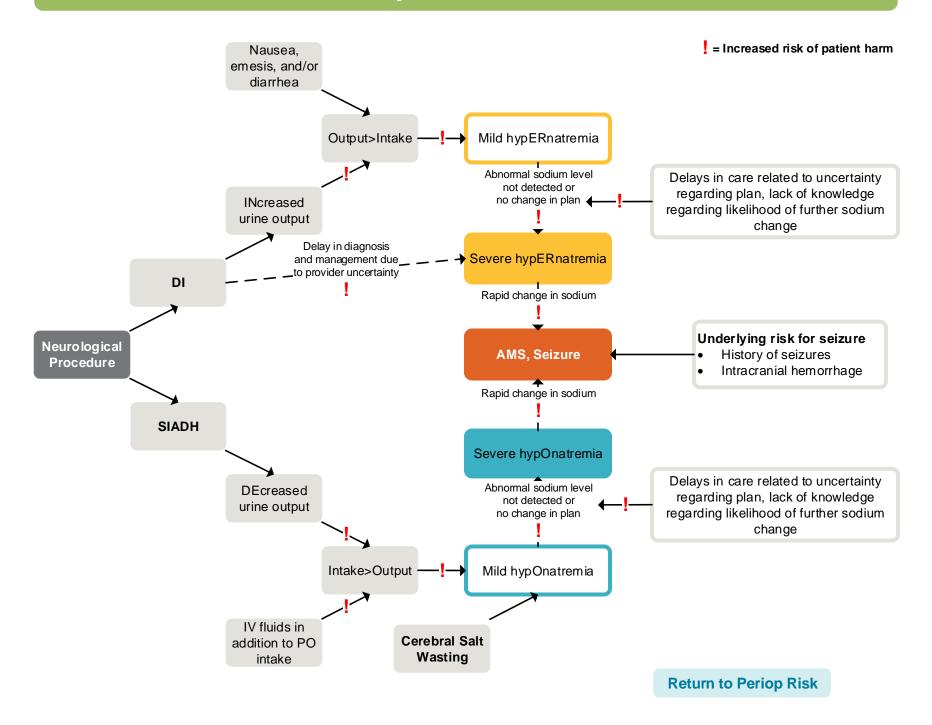
≥21 above target sodium

Tolerating Enteral Intake

- If abnormal mental status, go to PICU
- Give enteral water to decrease sodium 10 mmol/L in 24 hours
- Serum sodium q4h
- Consider PICU admission



How Dysnatremia Occurs



Periop Neurosurgery at risk for DI: "Assess for History of DI"

- Diabetes insipidus (DI) is a condition characterized by polyuria, and typically, corresponding polydipsia
- Neurogenic DI is due to a deficiency of anti-diuretic hormone (ADH)
 - ADH is produced in the pituitary gland
 - Deficiency can occur due to growth of a mass near the pituitary, due to surgery near the pituitary gland, or due to any other traumatic injury to the pituitary gland
- Untreated patients with DI produce large volumes of dilute urine, day and night, and many of them are often thirsty as well
- Untreated patients with DI often have a serum osmolality > 300 mOsm/kg, urine osmolality < 300 mOsm/Kg, and they may have an elevated serum sodium.



Periop Known Diabetes Insipidus: Suggested IV desmopressin dose

- While conversion from oral desmopressin to IV desmopressin is inexact for any individual patient, literature supports a range of 1/100 – 1/800 for a dose conversion
- A rule of thumb: an IV dose of 1/500th the oral dose provides anti-diuresis for a period of time without overtreatment



Phase 2: Desmopressin PRN

- Patients with a new diagnosis of DI after neurosurgery are known to have one of three outcomes over the next 10-14 days:
 - 1. Their DI is transient and it self-resolves
 - 2. They develop syndrome of inappropriate antidiuretic hormone (SIADH), which causes low urine output and a declining serum sodium
 - 3. They have permanent DI and will require lifelong treatment
- Giving a dose of desmopressin to a patient who has outcome #1 or #2 could place them at risk for a large decline in their serum sodium and dangerous hyponatremia
- The PRN desmopressin phase is designed to follow urine output and serum sodium closely so that desmopressin is only given when it is safe to do so



Phase 3: Scheduled Desmopressin

- Once a patient has been established to have permanent DI, they can have their desmopressin scheduled, rather than given PRN
- The goal of this phase is to establish safe dosing of desmopressin in preparation for discharge home
- When planning for discharge, it is important to determine lab follow-up and whether the family will monitor intake and output at home



"Call Fors" for patients at risk for hyponatremia

- Patients recovering from neurosurgery who have a large positive imbalance of intake and output or low overall urine output are at high risk to develop hyponatremia
- The "call fors" and labs that they trigger are designed to detect hyponatremia and to intervene before it becomes severe (sodium < 125)
- It is also important to consider that, although rare, some patients may develop DI which will be detected due to polyuria (urine output >4mL/kg/hr)



SIADH vs Cerebral Salt-Wasting (CSW)

- SIADH and Cerebral Salt-Wasting can be difficult to distinguish diagnostically, and they require different methods of treatment
- CSW patients should have evidence of volume depletion, because their sodium level is low due to inappropriate loss of sodium in the urine:
 - Tachycardia
 - Delayed capillary refill
 - Decrease in weight
 - Other signs of dehydration appropriate for the age (i.e. sunken fontanelle)
- Patients with SIADH will have normal or increased volume status, because their sodium level is low due to excess water retention



ED Assessment: History

- After establishing the ABCs (airway, breathing, circulation), the next most important determination for a patient with DI is whether they are due for a desmopressin dose
- Almost all patients with DI are on scheduled dosing of desmopressin and parents or the patient themselves should be able to report when their last dose was taken, and whether their last dose is wearing off (resulting in excessive urine output, i.e. "breakthrough urination")
- Assessing the presence of vomiting, diarrhea, mental status change, or fever will help determine whether the patient may need to be admitted



ED Assessment: Measure and chart intake and output

- For a patient with DI, measurement of their intake and output is just as important as other classic vital signs
- Unlike other pediatric patients, urine output is not a good assessment of hydration status for patients with DI
 - Instead, their urine output is almost entirely dependent on the amount of desmopressin in their system
- A urine output of > 4 mL/kg/hr is almost always an indication that they require a dose of desmopressin urgently
- A diaper scale is located in the dirty utility room in the ED. Zero the scale with a dry diaper first.



Known DI Management

- Unlike most other pediatric patients, those that have DI may have a difficult time tolerating the relatively large water and sodium load that comes with a normal saline (NS) bolus
- If they have recently taken a dose of desmopressin, the NS bolus may cause a rapid drop in their sodium
- If they are due for a dose of desmopressin, the NS bolus may cause a marked increase in their urine output, and their sodium may rise suddenly
- For these reasons, NS boluses should be used cautiously in patients with DI, and only when they have clear evidence of intravascular volume depletion causing poor perfusion



Known DI: Management

- Chronic hypernatremia (present at least 24 hours) should be corrected slowly to prevent rapid shifts in body water causing cerebral edema
- Retrospective studies have shown that for those with chronic hypernatremia, a maximum change in the sodium of 10-12 mEq/L per 24 hours is the fastest safe rate of correction
- For this reason, a change in sodium of more than 0.5 mEq/L/hr (i.e. > 2 point decrease in 4 hours) in a patient with chronic hypernatremia should prompt a discussion regarding a change in the fluid management



Dysnatremia Approval & Citation

Approved by the CSW Dysnatremia team for October 29, 2018 go live

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Retrieval Website: http://www.seattlechildrens.org/pdf/dvsnatremia-pathwav.pdf

Please cite as:

Seattle Children's Hospital, Werny D, Dervan L, Fenstermacher S, Hauptman J, Hrachovec J, Richardson N, Symons J, Turner A, Valdivia H, Zapata L, 2018 October. Dysnatremia Pathway.

Available from: http://www.seattlechildrens.org/pdf/dvsnatremia-pathway.pdf



Evidence Ratings

This pathway was developed through local consensus based on published evidence and expert opinion as part of Clinical Standard Work at Seattle Children's. Pathway teams include representatives from Medical, Subspecialty, and/or Surgical Services, Nursing, Pharmacy, Clinical Effectiveness, and other services as appropriate.

When possible, we used the GRADE method of rating evidence quality. Evidence is first assessed as to whether it is from randomized trial or cohort studies. The rating is then adjusted in the following manner (from: Guyatt G et al. J Clin Epidemiol. 2011;4:383-94.):

Quality ratings are downgraded if studies:

- Have serious limitations
- Have inconsistent results.
- If evidence does not directly address clinical questions
- If estimates are imprecise OR
- If it is felt that there is substantial publication bias

Quality ratings are *upgraded* if it is felt that:

- The effect size is large
- If studies are designed in a way that confounding would likely underreport the magnitude of the effect OR
- If a dose-response gradient is evident

Guideline – Recommendation is from a published guideline that used methodology deemed acceptable by the team.

Expert Opinion – Our expert opinion is based on available evidence that does not meet GRADE criteria (for example, case-control studies).

Quality of Evidence:

○○○○ High quality

○○○ Moderate quality

○○○○ Low quality

©OOO Very low quality

Guideline

Expert Opinion

Summary of Version Changes

- Version 1.0 (10/29/2018): Go live.
- Version 2.0 (3/18/2019): Updated perioperative workflow.
- Version 2.1 (10/22/2020): Aligned verbiage to correspond with Epic.
- Version 2.2 (5/11/2021): Clarified serum sodium requirements for PASS clinic and definition for high risk for diabetes insipidus post neurosurgery. Added navigation buttons.

Medical Disclaimer

Medicine is an ever-changing science. As new research and clinical experience broaden our knowledge, changes in treatment and drug therapy are required.

The authors have checked with sources believed to be reliable in their efforts to provide information that is complete and generally in accord with the standards accepted at the time of publication.

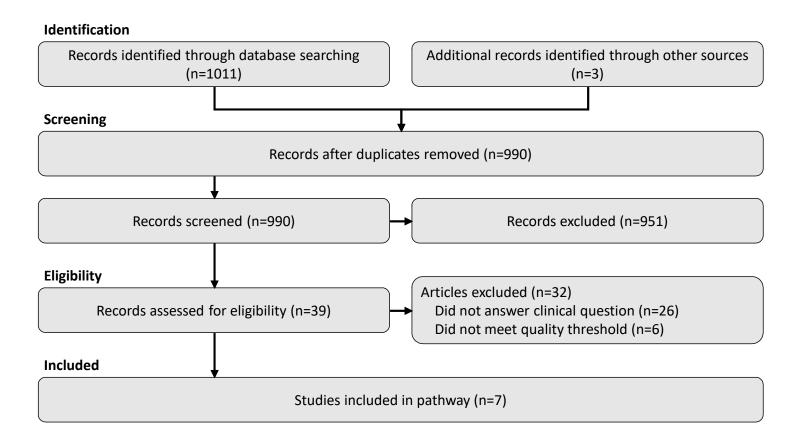
However, in view of the possibility of human error or changes in medical sciences, neither the authors nor Seattle Children's Healthcare System nor any other party who has been involved in the preparation or publication of this work warrants that the information contained herein is in every respect accurate or complete, and they are not responsible for any errors or omissions or for the results obtained from the use of such information.

Readers should confirm the information contained herein with other sources and are encouraged to consult with their health care provider before making any health care decision.

Bibliography

Search Methods

Literature searches were conducted in two phases and executed by Jackie Morton, Medical Librarian. The initial search in November 2017, targeted synthesized literature on the topics of dysnatremia, diabetes insipidus and sodium blood levels. The search was executed in Ovid Medline, Cochrane Database of Systematic Reviews, Embase, National Guideline Clearinghouse and TRIP. The search was limited to items published in English, from 2007 to date. The second search, in January 2018, was conducted in Medline and Embase to retrieve both synthesized and primary studies. The topics searched were diabetes insipidus and neurosurgical procedures, hyponatremia and Syndrome of Inappropriate Antidiuretic Hormone Secretion (SIADH). The search was limited to items published in English, from 1998 to date. The team added one citation containing primary research cited by a recent protocol. Results were exported to RefWorks for system de-duplication, then to Excel for the screening process.



Flow diagram adapted from Moher D et al. BMJ 2009;339:bmj.b2535

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Bibliography

- Coulthard MG, Long DA, Ullman AJ, Ware RS. A randomised controlled trial of hartmann's solution versus half normal saline in postoperative paediatric spinal instrumentation and craniotomy patients. Arch Dis Child. 2012;97(6):491-496. Accessed 20120521; 1/18/2018 3:18:49 PM. https://dx.doi.org/10.1136/archdischild-2011-300221.
- de Sola-Morales O, Riera M. Urea for management of the syndrome of inappropriate secretion of ADH: A systematic review. Endocrinol nutr. 2014;61(9):486-492. Accessed 20141202; 1/18/2018 3:15:27 PM. https://dx.doi.org/10.1016/j.endonu.2014.04.006.
- Matarazzo P, Tuli G, Verna F, et al. Management of sodium metabolism derangements in children treated for hypothalamic-hypophyseal tumors. J Pediatr Biochem. 2010;1(4):289-296. Accessed 1/18/2018 3:34:06 PM. 10.1055/s-0036-1586385.
- Mukherjee KK, Dutta P, Singh A, et al. Choice of fluid therapy in patients of craniopharyngioma in the perioperative period: A hospital-based preliminary study. Surg Neurol Intl. 2014;5(Supplement). Accessed 1/18/2018 3:34:06 PM. 10.4103/2152-7806.136399.
- Pratheesh R, Swallow DMA, Joseph M, et al. Evaluation of a protocol-based treatment strategy for postoperative diabetes insipidus in craniopharyngioma. Neurol India. 2015;63(5):712-717. Accessed 11/10/2017 12:54:53 PM. 10.4103/0028-3886.166533.
- Saito M, Kikuchi Y, Kawarai Lefor A, Hoshina M. High-dose nebulized budesonide is effective for mild asthma exacerbations in children under 3 years of age. Eur Ann Allergy Clin Immunol. 2017;49(1):22-27.
- Wise-Faberowski L, Soriano SG, Ferrari L, et al. Perioperative management of diabetes insipidus in children [corrected]. J Neurosurg Anesthesiol. 2004;16(1):14-19. Accessed 20031216; 1/18/2018 3:18:49 PM.