

# SODIUM CHLORIDE 3%

## NEWBORN USE ONLY

2017

<b>Alert</b>	Osmolarity: 1027 mOsm/L. <sup>1</sup> Sodium supplementation is not always appropriate and fluid restriction may be appropriate in the management of hyponatraemia. Treatment should always be tailored to the cause.
<b>Indication</b>	Treatment of hyponatraemia.
<b>Action</b>	Sodium is the major cation of extracellular fluid.
<b>Drug Type</b>	Sodium chloride 3% contains 30 g/L sodium chloride, equivalent to 0.5 mmol/mL of sodium.
<b>Trade Name</b>	Sodium chloride 3%
<b>Presentation</b>	Sodium chloride 3% – 1000 mL.
<b>Dosage/Interval</b>	<p><b><u>Severe hyponatraemia &lt; 120 mmol/L or symptomatic hyponatraemia</u></b></p> <p>IV: Give sodium chloride 3% at 0.5 mmol/kg/hour (1 mL/kg/hour) until symptoms abate or sodium <math>\geq</math> 120 mmol/L.*</p> <p>Then give sodium chloride 3% at 0.15 mmol/kg/hour (0.3 mL/kg/hour) for 48 hours or until desired sodium is achieved.</p> <p>[Therapeutic goal is to increase sodium by 7 mmol/L/day]</p> <p>*1 mL/kg sodium chloride 3% will raise serum sodium by approximately 1 mmol/L.<sup>2</sup></p> <p><b><u>IV supplementation</u></b> Start at 2–4 mmol/kg/day and increase as required.</p>
<b>Route</b>	IV
<b>Maximum Dose</b>	
<b>Preparation/Dilution</b>	<b>IV:</b> Sodium chloride 3% can be given undiluted.
<b>Administration</b>	<b>IV:</b> Sodium chloride 3% – Can be given undiluted as an infusion, preferably through large vein.
<b>Monitoring</b>	IV: Watch the local site for signs of extravasation. Monitor serum sodium as per clinical team's recommendation.
<b>Contraindications</b>	IV: No information.
<b>Precautions</b>	Impaired renal function, cardiac insufficiency, pre-existing oedema with sodium retention.
<b>Drug Interactions</b>	No information.
<b>Adverse Reactions</b>	Hypernatraemia, volume overload, congestive heart failure, respiratory distress. Hyperchloraemia, hypercalciuria. Disseminated intravascular coagulation (DIC) is associated with inadvertent injections of sodium chloride into blood vessels of the uterus or placenta due to hypernatraemic shock; not reported in infants. Osmotic demyelinating syndrome. Fever. IV site: Extravasation, phlebitis, venous thrombosis.
<b>Compatibility</b>	<b>IV Fluids:</b> Glucose 5%, glucose 10%, glucose 5% in sodium chloride 0.9%, glucose 5% in sodium chloride 0.45%, sodium chloride 0.9%, sodium chloride 0.45%.  Y site: No information.
<b>Incompatibility</b>	<b>IV Fluids:</b> Fat emulsion.  Y site: No information.  Amino acid solutions – No information.
<b>Stability</b>	
<b>Storage</b>	Store at room temperature, 20–25°C
<b>Special Comments</b>	Osmolarity of undiluted hypertonic sodium chloride is $>$ 1000 mOsm/L, posing the risk of extravasation for peripheral IV solutions. <sup>3,4</sup> Monitor for extravasation when infused peripherally at higher rates.

	<p>Total body water is traditionally calculated as weight x 0.6 in children. Greater total body water content in newborns should be considered and therefore should be calculated as weight x 0.75.<sup>2,5</sup></p>
<b>Evidence summary</b>	<p><u>IV correction for severe and/or symptomatic hyponatraemia</u></p> <p>The body of evidence to base recommendations in this clinical setting is extremely limited, particularly in neonatal populations. Recommendations are based on expert opinion, which have been extrapolated from adult consensus guidelines<sup>6,7</sup> and take into account specific neonatal safety concerns (see Safety below). In acute hyponatraemia, where the risk of sequelae is greater than that of osmotic demyelination, the correction should be rapid.<sup>8</sup></p> <p>Aim to increase sodium by 1–2 mmol/L per hour until symptoms abate or a safe level of sodium is achieved (<math>\geq 120</math> mmol/L).<sup>9</sup> Once the safe level is achieved, suggested subsequent goals are 6–8 mmol/L in 24 hours, 12–14 mmol/L in 48 hours and 14–16 mmol/L in 72 hours.<sup>10</sup> (LOE IV, GOR C)</p> <p><u>Sodium deficit calculation</u></p> <p style="text-align: center;">Deficit in mmol = (desired sodium – serum sodium) x total body water</p> <p>Total body water is traditionally calculated as weight x 0.6 in children. Greater total body water content in newborns should be considered and therefore should be calculated as weight x 0.75.<sup>2,5</sup> (LOE IV, GOR C)</p> <p><u>Oral supplementation</u></p> <p>A randomised, controlled trial of 4 mmol/kg/d of sodium versus placebo from DOL 7 to 35 in infants born 24–31 weeks (53 infants) showed higher sodium levels and increased weight gain in the intervention group.<sup>11</sup> A randomised, controlled trial of 4 mmol/kg/d of sodium versus placebo from DOL 4 to 14 in infants born at 29–34 weeks (20 infants) showed higher sodium levels and increased weight gain in the intervention group.<sup>12</sup> There are also three case-control studies that report similar findings with respect to sodium levels and growth in preterm infants supplemented with oral sodium.<sup>13–15</sup> A systematic review comparing higher versus lower sodium intake for preterm infants is in progress.<sup>16</sup> These findings support the use of oral sodium supplements to correct hyponatraemia and potentially improve growth. (LOE II, GOR B)</p> <p><u>Safety</u></p> <p>An historical case-control study identified 42/350 (12%) ELBW NICU admissions with an episode of hyponatraemia (Na &lt; 125 mmol/L [range 113–124]) that lasted &gt; 6 hours (median 1.5 days).<sup>17</sup> Rates of abnormal head ultrasound (IVH or PVL) and abnormal neurological examination were higher in the hyponatraemic group (p &lt; 0.03; p &lt; 0.001 respectively). Correction <math>\geq 0.5</math> mmol/L/h showed a trend toward higher rates of abnormal neurological examination. In paediatric and adult populations, multiple cohort studies and reviews have concluded that in patients with chronic hyponatraemia (<math>\geq 48</math> hours), neurologic sequelae due to osmotic demyelination are associated with more rapid rates of correction.<sup>7,9</sup></p> <p>In summary, rapid correction of hyponatraemia may be detrimental to neurological outcome during myelination of the newborn brain.<sup>17</sup> In adult populations, osmotic demyelination syndrome can usually be avoided by limiting correction of chronic hyponatremia to &lt; 10 to 12 mmol/L in 24 hours and to &lt; 18 mmol/L in 48 hours. These estimates should be regarded as approximate limits and not goals of therapy.<sup>7</sup> (LOE IV, GOR C)</p> <p><u>Osmolarity and Osmolar load</u></p> <p>A retrospective, matched-cohort study of 352 children <math>\leq 18</math> years evaluated the incidence of phlebitis or infiltration associated with peripheral administration of parenteral nutrition with an osmolarity &gt;1000 mOsm/L vs <math>\leq 1000</math> mOsm/L.<sup>18</sup> There were 151 neonates in the study. There were no differences between patients who did or did not develop adverse events in terms of age or weight. Administration of PPN with osmolarity &gt;1000 mOsm/L vs <math>\leq 1000</math> mOsm/L significantly increased infiltration (17% vs 7%; odds ratio [OR], 2.47; 95% confidence interval [CI], 1.24–4.94; p = 0.01) and the combined composite end point of phlebitis or infiltration (45% vs 34%; OR, 1.65; 95% CI, 1.07–2.54; p = 0.02). In multivariate analysis, osmolarity &gt;1000 mOsm/L was an</p>

	<p>independent risk factor for developing complications (OR, 1.67; 95% CI, 1.08–2.52; p = 0.02).<sup>18</sup> (LOE III, GOR C)</p> <p>A prospective, observational study in adults suggests that osmolar load (i.e. number of milliosmoles per hour, calculated as osmolarity x infusion rate) is a better predictor than osmolarity alone for phlebitis.<sup>19</sup> They found an osmolarity rate of 84–99 mOsm/hour was associated with a 4–27% rate of phlebitis. They did not report on other injuries such as extravasation. The infusion rates suggested in our formulary have low osmolar load and are considered to carry minimal risk of phlebitis (Consensus opinion).</p>
<p><b>References</b></p>	<ol style="list-style-type: none"> <li>1. Micromedex solutions. Accessed on 18 July 2017.</li> <li>2. Zieg J. Evaluation and management of hyponatraemia in children. <i>Acta Paediatr</i> 2014;103:1027-34.</li> <li>3. Dugan S, Le J, Jew RK. Maximum tolerated osmolarity for peripheral administration of parenteral nutrition in pediatric patients. <i>Journal of Parenteral and Enteral Nutrition</i>. 2014 Sep;38(7):847-51.</li> <li>4. Timmer JG, Schipper HG. Peripheral venous nutrition: the equal relevance of volume load and osmolarity in relation to phlebitis. <i>Clinical Nutrition</i>. 1991 Apr 1;10(2):71-5.</li> <li>5. Modi N, Bétrémieux P, Midgley J, Hartnoll G. Postnatal weight loss and contraction of the extracellular compartment is triggered by atrial natriuretic peptide. <i>Early Hum Dev</i> 2000;59:201-8.</li> <li>6. Spasovski G, Vanholder R, Allolio B, Annane D, Ball S, Bichet D, et al. Clinical practice guideline on diagnosis and treatment of hyponatraemia. <i>Eur J Endocrinol</i> 2014; 170: G1–47.</li> <li>7. Verbalis JG, Goldsmith SR, Greenberg A, Schrier RW, Sterns RH. Hyponatremia treatment guidelines 2007: expert panel recommendations. <i>Am J Med</i> 2007;120:S1-21.</li> <li>8. Marcialis MA, Dessi A, Pintus MC, Irmesi R, Fanos V. Neonatal hyponatremia: differential diagnosis and treatment. <i>J Matern Fetal Neonatal Med</i> 2011;24:75-9.</li> <li>9. Assadi F. Hyponatremia: a problem-solving approach to clinical cases. <i>J Nephrol</i>. 2012;25(4):473-80.</li> <li>10. Sterns RH, Nigwekar SU, Hix JK. The treatment of hyponatremia. <i>Semin Nephrol</i> 2009;29:282-99.</li> <li>11. Isemann B, Mueller EW, Narendran V, Akinbi H. Impact of Early Sodium Supplementation on Hyponatremia and Growth in Premature Infants: A Randomized Controlled Trial. <i>Jpen: Journal of Parenteral &amp; Enteral Nutrition</i> 2016;40:342-9.</li> <li>12. Vanpee M, Herin P, Broberger U, Aperia A. Sodium supplementation optimizes weight gain in preterm infants. <i>Acta Paediatr</i>. 1995;84:1312-1314.</li> <li>13. Sulyok E, Rascher W, Baranyai Z, Ertl T, Kerekes L. Influence of NaCl supplementation on vasopressin secretion and water excretion in premature infants. <i>Biology of the Neonate</i> 1993;64:201-8.</li> <li>14. Ayisi RK, Mbiti MJ, Musoke RN, Orinda DA. Sodium supplementation in very low birth weight infants fed on their own mothers milk, I: effects on sodium homeostasis. <i>East Afr Med J</i>. 1992;69:591-595.</li> <li>15. Al-Dahhan J, Haycock GB, Nichol B, Chantler C, Stimmler L. Sodium homeostasis in term and preterm neonates. <i>Arch Dis Child</i> 1984;59:945-950.</li> <li>16. Chan W, Chua MYK, Teo E, Osborn DA, Birch P. Higher versus lower sodium intake for preterm infants (Protocol). <i>Cochrane Database of Systematic Reviews</i> 2017: CD012642.</li> <li>17. Bhatti S, Tsirka A, Bigini-Quinn P, La Gamma EF. Does Hyponatremia Result in Pontine Myelinolysis and Neurological injury in Extremely Low Birth Weight (ELBW) Micropremies?† <i>Pediatric Research</i> 1997;41:140.</li> <li>18. Clark E, Giambra BK, Hingl J, Doellman D, Tofani B, Johnson N. Reducing risk of harm from extravasation: a 3-tiered evidence-based list of pediatric peripheral intravenous infusates. <i>Journal of Infusion Nursing</i> 2013;36:37-45.</li> <li>19. Pereira-da-Silva L, Henriques G, Videira-Amaral JM, Rodrigues R, Ribeiro L, Virella D. Osmolality of solutions, emulsions and drugs that may have a high osmolality: aspects of their use in neonatal care. <i>The Journal of Maternal-Fetal &amp; Neonatal Medicine</i> 2002;11:333-338.</li> </ol>

# SODIUM CHLORIDE 3%

## NEWBORN USE ONLY

2017

<b>Original version Date: 06/09/2017</b>	<b>Author: NMF Consensus Group</b>
<b>Current Version number: 1.0</b>	<b>Version Date: 06/09/2017</b>
<b>Risk Rating: Medium</b>	<b>Due for Review: 06/09/2020</b>
<b>Approved by: As per Local policy</b>	<b>Approval Date: As per Local policy</b>

### Authors Contribution

Original author/s	Chris Wake, Srinivas Bolisetty
Pharmacy Review	Ushma Trivedi
Expert review	
Evidence Review	Timothy Schindler
Final content and editing review	Ian Whyte
Facilitator/s	Srinivas Bolisetty