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How is acute hypophosphataemia treated in adults?

Prepared by UK Medicines Information (<u>UKMi</u>) pharmacists for NHS healthcare professionals Before using this Q&A, read the disclaimer at <u>https://www.sps.nhs.uk/articles/about-ukmi-medicines-gas/</u> Date prepared: 20/07/2017

Background

There are no national guidelines for the treatment of acute hypophosphataemia and practice varies widely across hospital Trusts. Following a thorough search of the literature this guidance has been prepared and adopted in Leeds Teaching Hospitals NHS Trust (LTHT). The use of phosphate for other indications such as re-feeding syndrome or use in the critical care setting has not been considered.

Reference ranges for serum phosphate vary between laboratories. For the purpose of this document, the reference range used for phosphate is 0.8 - 1.5 mmol/L.

Phosphate deficiency can be caused by (1-4)

- redistribution of phosphate into cells (e.g. respiratory alkalosis, drug therapy (insulin, catecholamines))
- increased urinary excretion (e.g. metabolic or respiratory acidosis, hyperparathyroidism)
- decreased intestinal absorption (e.g. antacid abuse, vitamin D deficiency, chronic diarrhoea)

Symptomatic hypophosphataemia is usually observed when plasma phosphate falls below 0.3 mmol/L (1,3). Symptoms may include (1-4):

- myopathy, rhabdomyolysis, weakness
- respiratory failure
- arrhythmias, cardiomyopathy
- · irritability, confusion, hallucinations, somnolence, convulsions, coma

Answer

At LTHT phosphate replacement is prescribed for patients with severe hypophosphataemia (serum phosphate concentration < 0.3mmol/L). For patients with moderate hypophosphataemia (serum phosphate concentration 0.3 - 0.6 mmol/L), phosphate replacement can be considered if the patient is symptomatic or following consideration of the clinical risks and benefits (6). Although this document offers guidance, the dose of phosphate to correct hypophosphataemia should be determined on an individual patient basis.

Phosphate is renally excreted and should be used with caution in patients with severe renal impairment (5). Phosphate should be used with caution in patients who have low serum calcium concentrations as these may decrease further when phosphate is replaced (3-5).

Oral phosphate replacement

- In the UK a licensed oral phosphate preparation is Phosphate Sandoz[®] effervescent tablets (7). Each tablet contains phosphate 16.1mmol, sodium 20.4mmol and potassium 3.1mmol (8). A common dose is 1 2 tablets three times a day (6). The dose should be reviewed daily and adjusted according to phosphate levels (6).
- Oral phosphate supplements should not be taken with aluminium, calcium or magnesium salts as these will bind phosphate and reduce its absorption (5).





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Parenteral phosphate replacement

- Intravenous therapy is indicated if the patient has severe hypophosphataemia or is symptomatic. Intravenous therapy may also be considered for patients who are unlikely to absorb oral agents (6).
- Phosphates Polyfusor[®] may be a suitable intravenous product. Each 500ml Phosphates Polyfusor[®] contains phosphate 50mmol, potassium 9.5mmol and sodium 81mmol (4). Alternative intravenous phosphate preparations are available, however their use may be limited to intensive care wards due to their high potassium content (6).
- Doses for intravenous phosphate vary in the literature and suggested regimens have included 0.2 0.5mmol/kg/day up to a maximum of 50mmol (4,5,8).
- Table 3 gives some suggested doses of Phosphates Polyfusor[®] based on weight for patients with normal renal function. Reduced doses may be necessary in patients with impaired renal function (4).

Serum	Weight 40 - 60kg		Weight 61 - 80kg		Weight 81 - 120kg	
phosphate concentration	Amount of phosphate	Volume of polyfusor	Amount of phosphate	Volume of polyfusor	Amount of phosphate	Volume of polyfusor
< 0.3mmol/L	25 mmol	250 mL	35 mmol	350 mL	50 mmol	500 mL
0.3 - 0.6 mmol/L (if oral route not suitable)	10 mmol	100 mL	15 mmol	150 mL	20 mmol	200 mL

Table 3: Suggested doses of Phosphates Polyfusor[®] adapted from Taylor et al. (9).

Summary

- There is no national guidance on the treatment of hypophosphataemia and practice varies widely across hospital Trusts. The guidance in this document reflects practice at Leeds Teaching Hospitals NHS Trust.
- Phosphate replacement should be prescribed for patients with severe hypophosphataemia (serum phosphate concentration < 0.3 mmol/L). For patients with moderate hypophosphataemia (serum phosphate concentration 0.3 0.6 mmol/L), phosphate replacement should be considered if the patient is symptomatic or following a consideration of the clinical risks and benefits.
- In moderate hypophosphataemia where the patient is asymptomatic, oral phosphate therapy should be considered if dietary modification is unsuitable. A dose of Phosphate Sandoz[®] effervescent tablets for hypophosphataemia is 1-2 tablets three times daily. The dose should be reviewed daily and adjusted according to phosphate levels.
- In severe hypophosphataemia, in symptomatic patients and when the oral route is not appropriate, intravenous phosphate therapy may be considered. Doses for intravenous phosphate vary in the literature and suggested regimens have included 0.2-0.5mmol/kg/day





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up to a maximum of 50mmol (see Table 3) however local practices may vary. Phosphates Polyfusor[®] is a commonly used product for this indication.

- The required dose from a Phosphates Polyfusor[®] is usually given over 12 24 hours but can be given over 6 12 hours.
- Phosphate is renally cleared. Phosphate (especially via the intravenous route) should be used with caution in patients with renal impairment.
- Phosphates Polyfusors[®] should be administered with caution to patients with cardiac failure, peripheral or pulmonary oedema, impaired renal function or conditions predisposing to hyperkalaemia due to the potassium and sodium content of Phosphates Polyfusors[®].
- Patients with hypocalcaemia should have their calcium corrected before replacing phosphate to prevent further <u>hypocalcaemia</u>.

Limitations

This Q&A is designed for adult patients only. This guidance is not suitable for chronic hypophosphataemia, patients with complex medical problems, or those with renal impairment or refeeding syndrome. The dose of phosphate to correct hypophosphataemia should be determined on an individual patient basis. There are no national guidelines for the treatment of hypophosphataemia, and practice varies widely across hospital Trusts.

References

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Search strategy

Original:

- Embase [terms used: hypophosphataemia (exp), hypophosphataemia/dt (drug therapy), limited to Publication Types Review]
- Medline [terms used: hypophosphataemia (exp), hypophosphataemia/dt (drug therapy), limited to Review Articles]
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- Embase [terms used: hypophosphataemia (exp), hypophosphataemia/dt (drug therapy), limited to Publication Types Review. Publication Year 2015-2017]
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