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PRIMARY CARE:

Michelle P B Guppy, Sharon M Mickan, and Chris B Del Mar

"Drink plenty of fluids": a systematic review of evidence for this recommendation in acute respiratory infections

BMJ 2004; 328: 499-500 [Full text]

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▼ SHOULD WE RESTRICT FLUIDS IN RESPIRATORY INFECTIONS—NO PROOF YET.

Sunit Singh (22 May 2004)

Fluids and URTI's

27 February 2004

Peter J Collignon,
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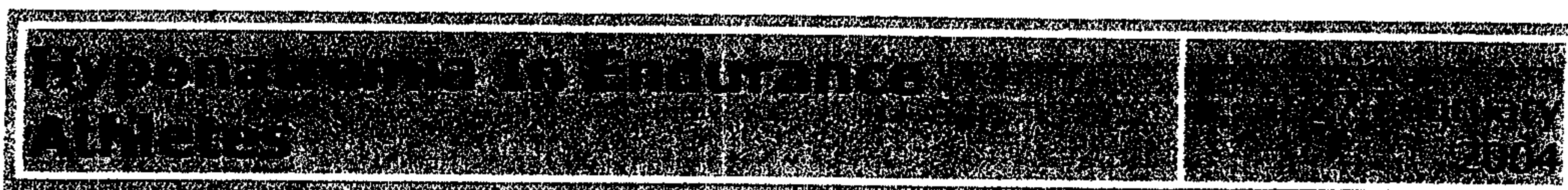
Send response to
journal:
Re: Fluids and URTI's

I worry that the authors of the recent article "Drink plenty of fluids": a systematic review of evidence for this recommendation in acute respiratory infections" have left readers with recommendations and implications that are not supported by the data they have reviewed. Upper respiratory tract infections

(URTI's) and episodes of bronchitis are very common and presumably outnumber episodes of pneumonia by a factor of more than 100. The only data they give to show excess fluid may potentially be harmful in acute respiratory infections is from studies with moderate to severe pneumonia. While they give a number of theoretical reasons why antidiuretic hormone may be increased in respiratory infections, most of those mechanisms would not be relevant in conditions where pneumonia was not present. Their article would have been better entitled "Drink plenty of fluids": a systematic review of evidence for this recommendation in moderate to severe pneumonia".

The title of their article has implication for all respiratory infections including URIs I however do not believe the data they have reviewed should be used to extrapolate for conditions other than moderate to severe pneumonia (and with these latter cases one would hope close medical or hospital supervision was taking place and so hyponatraemia could be avoided). We need to ensure that we do not leave the community with the implication that this advice applies to the much more common URIs. As they state in the first paragraph of their paper it appears self-evident that there are many benefits in keeping patients with less serious respiratory tract infections well hydrated. They have presented no data to show that this "common sense" approach should not continue to be the case.

Competing interests: None declared



Bill D. Misner Ph.D.,
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Send response to
journal:
[Re: Hyponatremia In
Endurance Athletes](#)

Each year during the heat early spring and summer, my e-mail is flooded with reports from endurance athletes who drink too much fluid, take too few electrolytes, or consume too much carbohydrates for energy during a hyperthermic prolonged endurance event.

Proportionate to the increase in temperature and humidity above 60 degrees F. and 60% respectively in relationship to the athlete's exposure to training-induced adaptations are the reports of dilutional hyponatremia. The endurance athlete requires approximately 10-14 days training in exposure to similar heat and humidity as imposed during an event that lasts 3-6 hours in length at an exercise pace of 75-85% VO2 Maximum Heart Rate. If 10-14 days of adaptation to heat and humidity is accomplished, the athlete is also advised to consume from 24-28 fluid ounces liquid with 0.3-0.7 g sodium and no more than 280 calories from carbohydrates each hour divided

dose to prevent dilutional hyponatremia.

If the adaptation training exposure is not completed, the fluids, electrolytes, and carbohydrate caloric replenishment formula may lead to severe dehydration, tempting the athlete to consume too much fluids above 1 liter per hour, leading to dilutional hyponatremia. It is observed that the athletes who consume in excess of 1 liter fluid per hour in prolonged endurance events are most likely to experience the symptomatic malaise observed in 1st-stage hyponatremia.

Competing interests: None declared



Yiu Wing Cheung,
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I am not convinced that drinking fluid is harmful to all patients with respiratory infection (RTI).

Send response to
journal:
Re: Plenty of Fluid and
Respiratory Infection

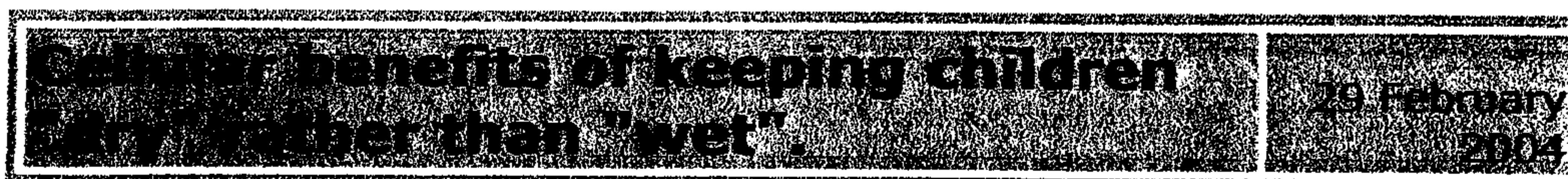
It is not correct to think that increased antidiuretic hormone (ADH) secretion in lower respiratory tract infection (LRTI) is a common phenomenon. Hyponatraemia is a well known complication of LRTI but it is not common. Furthermore, it is difficult to prove that the incidence or the death of the LRTI related hyponatraemia cases were associated with increased water intake or not.

I believed that hyponatraemia could still occur even with normal fluid intake. Fluid restriction may not be helpful to all LRTI. It can be even harmful if we restricted fluid too much to the extent of causing dehydration.

I did not think that the data the authors found could be extrapolate to patients with upper respiratory infection (URTI) because there was no definite evidence.

In Hong Kong, the newspaper had quoted the study and said "plenty of fluid in flu can be harmful"! I worried the community might have misconception about the finding of the study.

Competing interests: None declared



Richard G Fiddian-
Green,
None
None

Guppy et al report that they had found "two prospective prevalence studies [which had] reported hyponatraemia at rates of 31% and 45% for children with moderate to severe pneumonia. None of these

Send response to
journal:

Re: Cellular benefits of
keeping children "dry"
rather than "wet".

children showed clinical signs of dehydration. Symptoms associated with hyponatraemia were not reported, but four children with a serum sodium below 125 mmol/l died during one study" (1).

The fall in tissue pH that may develop either regionally or systemically in acute illnesses and severe exercise, because of unreversed ATP hydrolysis, is accompanied by a rise in ionised $[Ca^{++}]$ and may cause intracellular calcium overload which if excessive is toxic to cells. This may be accompanied by a rise in intracellular $[Na^{+}]$ because the sodium pump is inhibited by an extracellular acidosis because protons are buffered intracellularly in exchange for potassium. The hyponatraemia might be compounded by activation of the $Na^{+}-Ca^{2+}$ exchanger (2). Activation of pH regulatory mechanisms, including Na^{+}/H^{+} exchange, may also compound the severity of the hyponatraemia to a degree limited by its stimulation of Na^{+}/K^{+} -ATPase activity (3).

These changes could account for the hyponatraemia seen in the children with respiratory infections, and its association with poor outcomes. They might also account for the antidiuretic hormone secretion that increases in proportion with the extent of lung parenchymal involvement. In fetal sheep normocapnic hypoxia releases vasopressin and hypercapnia/acidemia augments the response to hypoxia (4).

Fluid intake, particularly if excessive, may compound the severity of these changes in patients with acute respiratory infections in the same manner that it does in marathon runners (5). The fluid intake could have an adverse effect by causing oedema and increasing the diffusional distances for the transport of oxygen, nutrients and metabolic by-products between capillaries and cells. They might also make matters worse by decreasing the a-v pressure gradient in capillaries by increasing venous outflow pressure. Some degree of dehydration may have the reverse effects and increase capillary flow rates as observed in the electronic discussion of Noakes' paper. Keeping patients "dry" rather than "wet" improves outcome patients in haemorrhagic shock.

Shock, defined as the presence of an impairment of tissue energetics evident from the presence of a gastric intramucosal acidosis, was undoubtedly present in those children with respiratory infections who died. In which case the translocation of endotoxin, and its release of cytokines, may have been another compounding factor as proposed in marathon runners.

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Competing interests: None declared

Ronald Eccles,
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Martez Jawad, Medical
Director

Send response to
journal:
Re: No harm in fluids
for colds and flu

Editor- The article by Guppy, Mickan and Del Mar as a systematic review of evidence for the effects of fluid intake on the course of acute respiratory infections has been interpreted in the press as a warning against fluid intake whilst suffering from acute upper respiratory tract infections such as common cold or flu. Headlines in newspapers and the inter-net warn that the 'age old advice on fluids for colds' is now disputed by research out of Australia.

The authors have mischievously taken an old folk-lore on fluids and colds and presented evidence on severe lower respiratory infections in infants to make a case that intake of fluids may be harmful. The press and public have got the wrong message because of the confusion in the article. The saying "Drink plenty of fluids" is generally accepted to refer to common colds and is not normally associated with acutely ill and hospitalised infants. By linking these two together the authors have created a scare story, but they have not addressed the main issue of the folk-lore. There are no controlled clinical trials in the literature to support any beneficial effect of maintaining fluid intake whilst suffering from an acute upper respiratory tract infection, but neither is there any evidence that indicates that this remedy is harmful in any way when applied as intended to colds and flu.

Competing interests: RE and MJ conduct sponsored

RF - FAMILY

clinical trials for the pharmaceutical industry on common cold medicines at the Common Cold Centre.

Drink plenty of fluids in acute respiratory infections 16 March 2004

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Stevens Rob

Send response to
journal:

Re: 'Drink plenty of
water' may be fatal

We read with interest Guppy et al's recent paper [1] as we have concerns that routinely advising patients to increase their fluid intake, especially in the elderly and after some procedures, can have serious consequences. Post operative urological patients are often routinely advised to drink plenty of fluids to keep a urinary catheter flushed. It is not unusual for these patients to drink an extra 5 litres of water per day in addition to their normal fluid intake. A recent case has highlighted to us that older patients don't always handle this extra volume well, which is consistent with age related renal changes [2]. Polydipsia can lead to severe hyponatraemia with serious consequences, including death, even in apparently healthy individuals with normal renal function [3 - 5].

References

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Competing interests: None declared

Drink plenty of fluids in acute respiratory infections 22 March 2004

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We read with interest this report by Guppy et al. However, their final comment that "we should be cautious about universally recommending increased

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Send response to
journal:
Re: 'Drink plenty of
fluids' in acute
respiratory infections

fluids to patients, especially those with infections of the lower respiratory tract" should perhaps be further qualified.

Firstly, the studies used to support their comment relate mainly to paediatric patients where the observed risk of hyponatraemia was >30%. It cannot be assumed that the same applies in adult patients. Indeed, Fine et al in cohorts of 14,199 and 38,039 adult patients with community acquired pneumonia (CAP) reported hyponatraemia ($\text{Na} < 130 \text{ mmol/l}$) in only 7.7% and 6.5% respectively.[1]

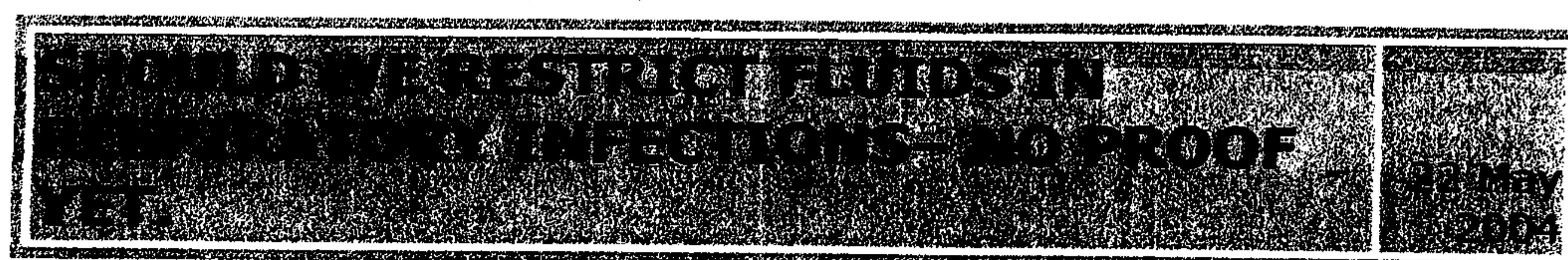
Secondly, as hyponatraemia is a recognised poor prognostic factor in CAP [1](this is indicated by the authors as well), most patients with hyponatraemia would have disease severe enough to warrant hospital referral. This is reflected in the extremely low incidence of hyponatraemia in adults with CAP treated in the community -0.7% of 944 out-patients compared to 6.1% of 1343 in-patients in the large prospective Pneumonia PORT cohort study.[1]

Therefore, we believe the risks to adults with CAP treated in the community from hyponatraemia are not as high as suggested by the authors. Patients with severe CAP need hospital referral. In these patients, hyponatraemia will be detected from 'routine blood tests' performed to assess disease severity. A powerful marker of prognosis is a raised urea which occurs in >20% of hospitalised adults with CAP [1], mostly as a result of dehydration. In these patients, fluid replacement is essential.

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Competing interests: None declared



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Send response to
journal:

The article "Drink plenty of fluids " a systematic review of evidence for this recommendation in acute respiratory infections" is misleading. The authors also have not reviewed the studies available in adults and animal models of pneumonias. The authors observe "giving increased fluids to patient with respiratory infection may cause harm", and comment that "fluid restriction may be appropriate management." They

Re: SHOULD WE
RESTRICT FLUIDS IN
RESPIRATORY
INFECTIONS- NO
PROOF YET.

base their recommendation on frequent occurrence of hyponatraemia in lower respiratory infections and higher risk of death in those with severe form of hyponatremia. Implicit in the above recommendation is the assumption that hyponatraemia caused deaths. With the current evidence this assumption is not tenable; a cause and effect relationship can not be assigned between hyponatraemia and death. It is equally possible that hyponatraemia was a marker of severe illness. In all the four patients who died, hyponatraemia was present at admission, and had nothing to do with fluids that they received.¹ These patients had more severe disease, a point that was emphasized in the original paper.¹ Indeed, hyponatraemia occurs more frequently in all kinds of seriously ill patients irrespective of the primary diagnosis.² Hyponatraemia in patients with pneumonia and bronchiolitis has been attributed to impaired water excretion³⁻⁵. However, fluid restriction across the board in all the patients with respiratory infection may not be justified as about two thirds of patient with pneumonia or bronchiolitis do not have hyponatremia Why should they be given restricted fluids? Moreover, in all the patients hyponatraemia may not be because of water retention. An important mechanism of hyponatraemia in critically ill patients is translocation of sodium from extracellular to intracellular space, and leakage of intracellular solutes, so called 'sick cell syndrome. In a recent study in critically ill adults an accumulation of non-diffusible osmotically active solutes in plasma, measured as increased 'osmolar gap', was observed in more than 50% of hyponatraemic patients.⁶ In experimental model of sepsis Hannon and Boston showed significant intracellular shift of sodium and suggested that hyponatraemia and hypo-osmolality in sepsis was caused by intracellular shift of sodium and dilution of extracellular space as a result of water retention.⁷ We found a significant increase in RBC sodium and impaired sodium-potassium pump coinciding with hyponatraemia in septicemic children.⁸ In another study we found that intracellular (RBC) sodium was significantly increased in hyponatraemic patients with pneumonia while it was normal in hyponatraemic patients having acute diarrhoeal illness (unpublished data). None of these arguments is directed towards advocating liberal fluid therapy in lower respiratory infections. Plasma volume expansion resulted in increased extravascular lung water in a canine model of pneumococcal lobar pneumonia⁹ and increased the extent of pneumonia in a canine model of acute Pseudomonas pneumonia.¹⁰ If one is inclined to use fluid restriction, perhaps it should be confined to hyponatremic patients and that too after correcting hypoxemia. A study published in 1970 had shown that an expansion of plasma volume in acute phase of

pneumonia in elderly patients improved cardiac output and decreased arteriovenous oxygen difference.¹¹ However, one cannot be categorical on either side. One should take a balanced approach till such time that randomised controlled studies answer the question.

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Competing interests: None declared

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