

## Supporting Text

**Methodology.** In Table 2, the prerace total body water ( $TBW_1$ , column C) was calculated as 0.6 times prerace weight (i.e., column B); change TBW ( $\Delta TBW$ ) during the race (column D) was calculated as the difference between fluid loss and fluid intake until normalization of serum  $[Na^+]$  or until the subject was discharged from hospital on the grounds of clinical recovery;  $Na^+$  balance during the race (column E) was calculated as the difference between  $Na^+$  intake and  $Na^+$  loss during recovery; measured post race serum  $[Na^+]$  (column F) was the value reported in the published studies; predicted postrace serum  $[Na^+]$  (column G) was calculated according to the equation of Nguyen and Kurtz (1)

$$[Na^+]_{2p} = \frac{([Na^+]_{1p} + 23.8)TBW_1 + 1.03\Delta E}{TBW_1 + \Delta TBW} - 23.8$$

in which  $[Na^+]_{1p}$  = the prerace serum  $Na^+$  at the start of the balance study;  $TBW_1$  is as calculated as for column C;  $\Delta E$  is net  $Na^+$  gain or loss from intake and output records during recovery; and  $\Delta TBW$  is net fluid loss from intake and output records during recovery; the difference between the measured (column F) and the predicted (column G) postrace serum  $[Na^+]$  is given in column H; the amount of osmotically-active  $Na^+$  that was inactivated (negative values) or conversely the amount of osmotically-inactive sodium activated to  $Na^+$  (positive values) to explain the discrepancies in column H is listed in column I. This calculation is made using the equation of Nguyen and Kurtz (1) for a  $\Delta TBW = 0$ . For  $\Delta TBW = 0$ , the Nguyen–Kurtz equation simplifies to

$$\Delta E = \frac{TBW_1}{1.03} \times \Delta[Na^+]_p$$

[2]

Column J provides the reference for the origin of these data.

Table 3 provides the calculations for the same 18 subjects to calculate the amount of sodium that was either osmotically-activated or inactivated during the recovery period of the study after the race and the total Na<sup>+</sup> balance for the entire duration of the study. The athletes are split on the basis of those athletes who showed an activation of osmotically-inactive, exchangeable sodium during the recovery period (top group) and those who inactivated osmotically-active Na<sup>+</sup> during recovery (bottom group). Columns B and C repeat the body weights and measured postrace serum [Na<sup>+</sup>] reported in Table 2 and which for this calculation is [Na<sup>+</sup>]<sub>1p</sub>. Column D lists the measured serum [Na<sup>+</sup>] at the end of the study period; Column E is the predicted postrecovery serum [Na<sup>+</sup>] based on the Nguyen-Kurtz equation (1) in which [Na<sup>+</sup>]<sub>1p</sub> = postrace serum [Na<sup>+</sup>] at the start of the balance study;  $TBW_1 = 0.6 \times \text{PreRace Wt} + \text{net water loss during hospitalization}$ ;  $\Delta TBW = \text{net water loss during the study}$ ; and  $\Delta E = \text{net sodium gain during the balance study}$ . Column F lists the predicted minus the observed postrecovery serum [Na<sup>+</sup>]; and Column G records the calculated amount of sodium that was osmotically-activated (positive values) or osmotically inactivated (negative values) during recovery to explain these discrepancies. As in Table 2 (column I), this calculation is made for a  $\Delta TBW = 0$ , using the simplified Nguyen-Kurtz equation already described; Column H lists the amount of Na<sup>+</sup> infused or ingested during recovery; whereas Column I records the calculated total Na<sup>+</sup> balance for the entire prerace to postrecovery period.

**Appendix: Estimate of the Size of the Osmotically Inactive Exchangeable Sodium Stores.** In Edelman *et al.*'s empirical equation (3), plasma water sodium concentration ([Na<sup>+</sup>]) was found to be primarily a linear function of the exchangeable Na (Na<sub>e</sub>) plus the exchangeable potassium (K<sub>e</sub>) divided by the TBW so that  $[Na^+] \propto (Na_e + K_e)/TBW$ . The equation also had a non-zero intercept which Edelman *et al.* thought to be a *variable* measure of exchangeable, but osmotically inactive sodium plus potassium. Nguyen and Kurtz (1) showed the y-intercept to be the sum of 4 terms, one of which includes the sum of osmotically inactive Na<sub>e</sub> and K<sub>e</sub>. Thus, in Edelman *et al.*'s hypothesis the term for osmotically inactive sodium plus potassium would equal the y-intercept value, with the other three terms summing to zero.

Exchangeable sodium ( $Na_e$ ) consists of both osmotically active ( $Na_{e,act}$ ) and osmotically inactive ( $Na_{e,inact}$ ) sodium (1). Exchangeable sodium is  $\approx 70\%$  of total body sodium. Since total body sodium is  $\approx 60$  meq/kg body weight, exchangeable sodium is about 42 meq/kg (4).

Because  $Na_e = Na_{e,act} + Na_{e,inact}$ ,

$$Na_{e,inact} = Na_e - Na_{e,act}$$

All  $Na_{e,act}$  is in body water. In intracellular fluid, its concentration is 14 meq/liter, and in extracellular fluid, 140 meq/liter (5).

Consider a 70-kg person. Total exchangeable sodium is  $70 \text{ kg} \times 42 \text{ meq/kg} = 2,940$  meq. Assume that TBW (in liters) is 42 liters, distributed as two-thirds or 28 liters of ICF, and 1/3 or 14 liters of ECF.

Osmotically active sodium can be estimated thus:

$$\text{ICF: } 28 \text{ liters} \times 14 \text{ meq/liter} = 392 \text{ meq}$$

$$\text{ECF: } 14 \text{ liters} \times 140 \text{ meq/liter} = 1960 \text{ meq}$$

$$\text{TOTAL } Na_{e,act}: 392 + 1960 = 2352 \text{ meq.}$$

Thus,  $Na_{e,inact}$  is:

$$2940 - 2352 = \approx 600 \text{ meq.}$$

With respect to body weight:  $600 \text{ meq}/70 \text{ kg} = 8.6 \text{ meq/kg}$ .

With respect to body water:  $600 \text{ meq}/42 \text{ liters} = 14 \text{ meq/liter body water}$ .

This number should be viewed as an estimated average only. It likely varies interindividually (3) and it has been demonstrated to vary significantly intraindividually over time (6–8).

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