

## **Ion transporters involved in excretion of $\text{Ca}^{2+}$ , $\text{SO}_4^{2-}$ , $\text{B(OH)}_4^-$ , and $\text{Mg}^{2+}$ by seawater fish**

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The body fluid of most marine teleosts is hyposmotic to seawater (SW). To balance passive water loss, marine teleosts drink SW, absorb water, and eliminate salts from the gill ( $\text{Na}^+$ ,  $\text{Cl}^-$ , and  $\text{K}^+$ ), intestine ( $\text{CaCO}_3$  and  $\text{MgCO}_3$ ), and kidney ( $\text{Mg}^{2+}$ ,  $\text{SO}_4^{2-}$ ,  $\text{Ca}^{2+}$ , and boric acid). To understand the mechanisms of intestinal and renal excretion, we identified ion transporters expressed in the epithelia and analyzed their localizations and activities. Mefugu (river puffer, *Takifugu obscurus*) is an ideal fish species for such studies since it is a euryhaline species and the complete genome sequence of a close relative torafugu (tiger puffer, *Takifugu rubripes*), a marine fish, is available. Our studies on SW-acclimated mefugu have suggested that: (i) a basolateral  $\text{Na}^+$ - $n\text{HCO}_3^-$  cotransporter NBCe1 (Slc4a4) and apical  $\text{Cl}^-$ / $n\text{HCO}_3^-$  exchangers (Slc26a6) are involved in  $\text{HCO}_3^-$  excretion across the intestinal epithelium and carbonate formation; (ii) an apical  $n\text{Na}^+/\text{Ca}^{2+}$  exchanger NCX2 (Slc8a2), apical  $\text{Cl}^-/\text{SO}_4^{2-}$  exchangers (Slc26a6), and an apical borate transporter (Slc4a11) are involved in renal excretion of  $\text{Ca}^{2+}$ ,  $\text{SO}_4^{2-}$ , and borate, respectively; (iii) a basolateral  $\text{Mg}^{2+}$  transporter (Slc41a1) is involved in renal tubular excretion of  $\text{Mg}^{2+}$ ; and (iv) an apical  $\text{Na}^+/\text{K}^+$ - $2\text{Cl}^-$  cotransporter NKCC2 (Slc12a1) is involved in renal  $\text{NaCl}$  reabsorption and urinary concentration.