

Renal Regulation of Acid Base Balance

by

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Abstract

A mathematical model has been developed to describe the reabsorption of bicarbonate in the kidney. The model is parameterized to simulate the reabsorption in the rat proximal convoluted tubule. The "total CO₂" reabsorption rate (including HCO₃⁻ and H₂CO₃ as well as CO₂) is predicted as a function of the following physiological parameters: transcellular potential difference, bulk radial velocity, intracellular pH, cell membrane permeabilities, and CO₂ hydration reaction rate in the cells and at the brush border.

The model is based on a scheme in which all protons generated within the cells, by a combination of uncatalyzed and enzyme-catalyzed hydration of CO₂, are secreted into the lumen to titrate HCO₃⁻, forming CO₂ and H₂CO₃. The results for total inhibition of the responsible enzyme (carbonic anhydrase) are consistent with the most recent experimental findings. For the fully catalyzed reaction in the cells and at the brush border the difference of HCO₃⁻ out of the cells is rate limiting for the intracellular generation of protons. The results show that the HCO₃⁻ generated within the cells diffuses out at both the brush border and basolateral membranes. There is net diffusion of HCO₃⁻ from the cells to the lumen (secretion) whether the intracellular reaction is catalyzed or uncatalyzed.

With normal enzyme activity diffusion of CO₂ from lumen to cells accounts for almost all of the net reabsorption of CO₂. With enzyme inhibition, however, H₂CO₃ accounts for most of the remaining total CO₂ reabsorption.

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