## Role of Sodium Ion Exchange in Plasma Membrane in Cardiac Hypertrophy

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## Summary

To test whether Na+/H+ (sodium/hydrogen) exchange plays an important role in neurohumoral signal transduction in cardiac hypertrophy, we studied the effects of Na+/H+ exchange inhibitors on ion exchange and protein synthesis in rat cardiomyocytes stimulated with neurohumoral factors. Norepinephrine enhanced Na+/H+ exchange through α<sub>1</sub>-adrenergic receptor, leading to intracellular alkalinization and increase in Ca<sup>2+</sup> concentration. Na<sup>+</sup>/H<sup>+</sup> exchange inhibitors, i. e. amilolide and ethyl-isopropyl-amilolide (EIPA), inhibited norepinephrineinduced protein synthesis. Angiotensin II increased intracellular pH and Ca<sup>2+</sup> through Na<sup>+</sup>/H<sup>+</sup> exchange activated by protein kinase C, and enhanced protein synthesis associated with expression of the protooncogenes (c-fos, c-myc). Amilolide also attenuated the angiotensin II-induced protein synthesis. Furthermore, arginine-vasopressin increased intracellular pH and Ca2+ through V1-receptor, which was attenuated by EIPA. Thus, we conclude that Na+/H+ exchange is involved in signal transduction through  $\alpha_1$ -adrenergic, angiotensin II and  $V_1$ receptor in rat cardiomyocyte growth.