Tetsuya Nakamoto, M.D., Ph.D.

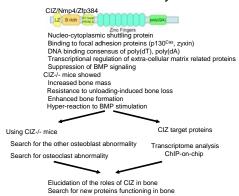
Department of Medicine and Rheumatology COE Research Associate Professor

Main Research Projects

- 1. Roles of CIZ in bone
- 2. Transcriptional targets of CIZ

Abstract

The nucleocytoplasmic shuttling protein CIZ regulates the transcription of extracellular matrix-related genes. In CIZ-deficient mice, increased bone mass and resistance to unloading-induced bone loss were observed. These phenotypes were caused by enhanced bone formation by osteoblasts. I expect that the other osteoblast functions



are also enhanced in CIZ-deficient mice. At the same time, I search for the targets of CIZ protein, using transcriptome analysis and ChIP assay. Based on these results, I am looking for new genes that are involved in bone formation.

Papers

- 1 . Hino, K., Nakamoto, T., Nifuji, A., Morinobu, M., Yamamoto, H., Ezura, Y., Noda, M. Deficiency of CIZ, a nucleocytoplasmic shuttling protein, prevents unloading-induced bone loss through the enhancement of osteoblastic bone formation in vivo, Bone 40:852-860,2007
- **2.** Morinobu, M., Nakamoto, T., Hino, K., Tsuji, K., Shen, Z.J., Nakashima, K., Nifuji, A., Yamamoto, H., Hirai, H., Noda, M. The nucleocytoplasmic shuttling protein CIZ reduces adult bone mass by inhibiting bone morphogenetic protein-induced bone formation. J Exp Med 201:961-970, 2005
- 3. Nakamoto, T., Shiratsuchi, A., Oda, H., Inoue, K., Matsumura, T., Ichikawa, M., Saito, T., Seo, S., Maki, K., Asai, T., Suzuki, T., Hangaishi, A., Yamagata, T., Aizawa, S., Noda, M., Nakanishi, Y., Hirai, H. (2004). Impaired spermatogenesis and male fertility defects in CIZ/Nmp4-disrupted mice. *Genes to Cells* 9, 575-589.
- 4. Shen, Z. J., Nakamoto, T., Tsuji, K., Nifuji, A., Miyazono, K., Komori, T., Hirai, H. and Noda, M. (2002). Negative Regulation of Bone Morphogenetic Protein/Smad Signaling by Cas-interacting Zinc Finger Protein in Osteoblasts. J Biol Chem 277, 29840-29846.

5. Nakamoto, T., Yamagata, T., Sakai, R., Ogawa, S., Honda, H., Ueno, H., Hirano, N., Yazaki, Y. and Hirai, H. (2000). CIZ, a zinc finger protein that interacts with p130(cas) and activates the expression of matrix metalloproteinases. Mol Cell Biol 20, 1649-58.