

「分泌性タンパク質 p53PAD7 と Hippo シグナル経路による増殖抑制機構の解明」

“Secreting protein p53PAD7 inhibits cell proliferation via the Hippo signaling pathway”

○滝川 雅大、渡邊 祐三、岡部 篤史、金田 篤志、石川 冬木、定家 真人、大木 理恵子

The *p53PAD7* gene is frequently methylated in a part of cancers. The *p53PAD7* gene is known as a target of p53 that suppresses cell proliferation, however, the molecular function of p53PAD7 is totally unknown. Here, we found that p53PAD7 is secreted to the cell culture medium and that apoptosis is induced in the presence of p53PAD7 protein. We hypothesized that p53PAD7 acts as a ligand that is received by specific receptors. To identify the candidate receptors, we performed immunoprecipitation and mass spectrometric analysis together with a heterobifunctional crosslinker, and successfully identified protocadherin FAT1 and FAT4 as p53PAD7 receptors. Human FAT1 and FAT4 are homologs of *Drosophila melanogaster* Fat, which is a receptor of the Hippo signaling pathway that regulates cell proliferation. Consistently, we observed activation of the Hippo signaling pathway when cells were treated with purified p53PAD7. Taken together, our results suggest that cell proliferation is regulated by p53PAD7 via the Hippo signaling pathway.

OE5-1-4 **Secreting protein p53PAD7 inhibits cell proliferation via the Hippo signaling pathway**

Masahiro Takikawa^{1,2}, Atsushi Okabe⁴, Atsushi Kaneda⁴, Fuyuki Ishikawa³, Mahito Sadaie¹, Rieko Ohki² (¹Dept. Applied Biological Sci., Tokyo Univ. of Sci., ²Lab. of Fundamental Oncol., Natl. Cancer Ctr. Res. Inst., ³Grad. Sch. of Biostudies, Kyoto Univ., ⁴Dept. Mol. Oncology, Grad. Sch. of Med., Chiba Univ.)

分泌性タンパク質 p53PAD7 と Hippo シグナル経路による増殖抑制機構の解明

滝川 雅大^{1,2}、岡部 篤史⁴、金田 篤志⁴、石川 冬木³、定家 真人¹、大木 理恵子² (¹東京理科大・理工・応用生物科学科、²国立がん研セ・研・基礎腫瘍学ユニット、³京都大・院生命、⁴千葉大・院医・分子腫瘍学)