血管内皮細胞におけるMTA1発現は血管新生阻害の標 的分子となりうる

Ishikawa, Mizuho Division of Pathological Biochemistry, Faculty of Medicine, University of Tottori

Osaki, Mitsuhiko Division of Pathological Biochemistry, Faculty of Medicine, University of Tottori

Yamagishi, Makoto DCBMS, Graduate School of Frontier Sciences, The University of Tokyo

Onuma, Kunishige Division of Pathological Biochemistry, Faculty of Medicine, University of Tottori

他

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MTA1 expressed in endothelial cells is a candidate target molecule for inhibiting angiogenesis

石川瑞穂¹, 尾崎充彦^{1,2}, 山岸誠³, 小沼邦重¹, 井藤久雄^{1,4}, 岡田太^{1,2}, 遠藤英也⁵

- 鳥取大・医・病態生化学
 Div. of Pathol. Biochem., Fac. of Med., Tottori Univ.
 2 鳥取大・染色体工学センター
- Ctr. Chromo. Engineering., Tottori Univ.
- ³ 東大・院新領域創成科学・メディカルゲノム専攻 DCBMS, Grad. Sch. Front. Sci., The Univ. of Tokyo.
- 4 井野口病院

Inokuchi Medical Center

⁵ 東大・医科研・分子発癌分野 Dept. Cancer. Biol., Inst. Med. Sci., The Univ. of Tokyo.

Metastasis-associated proteins, such as S100A4 and MTA1, have been studied for over two decades, but correlation between them is not understood. A recent report suggesting that silencing of S100A4 in endothelial cells suppresses in vitro tube formation and in vivo tumor angiogenesis, motivated us to examine MTA1 from the same perspective. In this study, we showed that the suppression of MTA1 in endothelial cells by murine MTA1 siRNA (mMTA1 siRNA) inhibited tube formation in vitro. We found that mMTA1 siRNA has an anti-angiogenic activity in vivo targeting endothelial cells using directed in vivo angiogenesis assay (DIVAA). Moreover, we found mMTA1 siRNA inhibited tumor angiogenesis in a xenograft model. Further, we revealed that inhibition of angiogenesis by mMTA1 siRNA mediates downregulation of S100A4 followed by promoting phosphorylation of non-muscle myosin IIA (NMIIA). These data suggested that silencing of MTA1 in endothelial cells could be used as a new strategy to regress tumors, by inhibiting tumor angiogenesis.