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Zhendong Liu, Zhitao Liu, Yuanjun Zhang, Yan Li, Bo Liu, Kexiang Zhang

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<u>miR-24</u> represses metastasis of human osteosarcoma cells by targeting Ack1 via AKT/MMPs pathway

Zhendong Liu, Zhitao Liu, Yuanjun Zhang, Yan Li, Bo Liu, Kexiang Zhang * Address:

Department of Orthopedic Surgery, The Third Xiangya Hospital of Central South University, Changsha 410013, China

*Corresponding authors. E-mail: kexiangzhang216@163.com

Running Title: miR-24 represses osteosarcoma metastasis by targeting Ack1

Zhendong Liu zhendongliu2@163.com

Zhitao Liu zhitaoliu22@163.com

Yuanjun Zhang yuanjunzhang2@163.com

Yan Li yanli2214@163.com

Bo Liu boliu233@163.com

Kexiang Zhang kexiangzhang216@163.com

Abstract

The expression levels of the protein tyrosine kinase Ack1 has been reported to be dysregulated in various cancers and involve in oncogenesis and progression. However, the expression and role of Ack1 in osteosarcoma remains unknown. In this study, we found that Ack1 were evidently upregulated in human osteosarcoma tissues and cell lines. In addition, the clinical data showed that high expression level of Ack1 is closely associated with clinical stage and positive distant metastasis, and negatively correlated with overall survival. Then, bioinformatics prediction and luciferase reporter assay indicated Ack1 as a direct target of miR-24, and Ack1 could be downregulated by miR-24 at both the mRNA and protein expression levels. Moreover, Ack1 expression levels were inversely correlated with that of miR-24 in osteosarcoma tissues. Furthermore, functional assay showed that miR-24 significantly suppressed osteosarcoma progression partially mediated by inhibiting Ack1 expression. Finally, western bolt assay revealed that miR-24 regulate AKT/MMPs pathway via Ack1 in osteosarcoma cells. In conclusion, our study

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