

ORIGINAL ARTICLE

MiR-22 alleviates the proliferation and metastasis of melanoma by targeting FASN

Jun Qiu^{1*}, Yanhua Yi^{2*}, Yong Miao¹, Zhiqi Hu¹

¹Department of Plastic and Aesthetic Surgery, Nanfang Hospital of Southern Medical University, Guangzhou, China. ²Department of Aesthetic Surgery, Affiliated Hospital of Guilin Medical University, Guilin, China

*Jun Qiu and Yanhua Yi contributed equally to this work

Summary

Purpose: To explore the role of MicroRNA-22 (miR-22) in the development of malignant melanoma and the underlying mechanism.

Methods: Potential miRNAs binding FASN (fatty acid synthetase) were predicted by bioinformatics analysis and miR-22 was selected. Their binding relationship was confirmed by dual-luciferase reporter assay. MiR-22 and FASN levels in 40 clinical samples of melanoma were detected. Expression correlation between miR-22 and FASN was assessed by Pearson correlation test. To uncover the role of miR-22 in regulating cell phenotypes of malignant melanoma, M21 and A375 cells were transfected with miRNA-NC, miR-22 mimics or miR-22 mimics+FASN-OE, respectively. Proliferative and metastatic abilities in each group were determined by cell counting kit-8 (CCK-8), 5-Ethynyl-2'-deoxyuridine (EdU) and Transwell assay, respectively.

Results: MiR-22 was the target gene binding the oncogene FASN. Downregulated miR-22 and upregulated FASN were observed in melanoma tissues, showing a negative correlation between them. Overexpression of miR-22 inhibited proliferative, migratory and invasive abilities in M21 and A375 cells. Notably, overexpression of FASN abolished the inhibitory effects of miR-22 on proliferative and metastatic abilities in melanoma.

Conclusions: MiR-22 is lowly expressed in the malignant melanoma samples. Overexpression of miR-22 inhibits proliferative and metastatic abilities in melanoma by targeting FASN and negatively regulating its level. MiR-22 may be a promising therapeutic target of melanoma.

Key words: malignant melanoma, MiR-22, FASN, proliferation, metastasis

Introduction

Melanoma is caused by the malignant deterioration of melanocytes distributed in the stroma, most of which are formed by the canceration of normal moles and pigmented plaques [1]. Although the incidence of malignant melanoma accounts for only 1% of skin tumors, its mortality is extremely high [2]. Melanoma rapidly progresses, leading to local or distant metastasis in a short period. Therefore, clarifying potential mechanisms of prolifera-

tion and metastasis in melanoma contributes to improve its prognosis.

MicroRNAs (miRNAs/miRs) are a kind of short, single-stranded non-coding RNAs with a length of about 22nt. They mainly regulate target gene expressions at the post-transcriptional level. In recent years, miRNAs have been considered to be closely related to tumorigenesis and tumor progression [3]. Increasing evidences have indicated that dysregu-