

Characterization Of *RECK* Gene Expression And Its Alternative Isoforms In Melanoma Cells

Jacomasso, T¹; Barbosa, FAL¹; Lima, MT²; Sogayar, MC²; Winnischofer, SMB¹

¹Setor de Ciências Biológicas, Universidade Federal do Paraná, Paraná, Brazil.

²Instituto de Química, Universidade de São Paulo, São Paulo, Brazil.

RECK encodes a membrane-anchored glycoprotein that suppresses both invasion and metastasis by negatively regulating matrix metalloproteinases, namely: MMP-2, MMP-9 and MT1-MMP. Previous studies showed a positive correlation between *RECK* expression in tumor samples and better prognosis for patients with lung, pancreatic and colorectal cancers. In the present study, we characterized the mRNA expression levels of *MMPs* (particularly: *MMP-2*, *MMP-9* and *MT1-MMP*) and their inhibitors: *TIMPs* (1-3) and *RECK* (canonical form and alternative isoforms, namely: *RECK B*, *RECK C*, *RECK D* and *RECK J*) in a panel of ten human melanoma cell lines that represent the different degrees of malignancy (radial growth phase - RGP, vertical growth phase - VGP and metastatic phase). The expression profiles of these genes were investigated through quantitative real time RT-PCR assays. Preliminary results indicate that mRNA levels of the *RECK* alternative isoforms in comparison to *RECK* canonical form tend to be higher in RGP/VGP cell lines than in metastatic cells. Specifically, *RECK C* isoform, that has an extra exon between exons 2 and 3 of the canonical form, is significantly increased in the RGP/VGP cells in comparison to metastatic cells ($p < 0.01$). We also found a positive correlation between the expression of *RECK C* isoform and *TIMP-3*. Our results suggest that *RECK C* isoform might have an important role in melanoma model, distinct to the *RECK* canonical form. Better understanding of *RECK* gene regulation may contribute to uncover the mechanisms of tumor progression and to develop new strategies for cancer therapy.

Keywords: *RECK* gene, Matrix metalloproteinase, Melanoma

Supported by: FAPESP, CNPq, CAPES and FINEP.