NITROLINOLEATE INFLUENCES ENDOTHELIAL CELL MIGRATION AND ANGIOGENESIS IN VITRO

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Background: Nitrolinoleate (LNO2), a nitroalkene derivative of linoleic acid that belongs to a new class of lipid mediators, has been demonstrated to donate nitric oxide (NO) and to posses a role in cell signaling. Since neovascularization has been suggested as a causative factor for atherosclerotic plaque growth and destabilization, and 'NO donors have been shown to regulate angiogenesis, the objective of the present study was to examine the effects of LNO₂ on angiogenesis in vitro. Methods: Angiogenesis was evaluated by measuring proliferation and cell migration of human umbilical endothelial cell (HUVEC) and endothelial cell sprouting by the rat aortic ring assay. Results: No effect of the LNO₂ on the endothelial cell proliferation (1-10 uM) was observed. However, HUVEC migration was significantly induced by LNO₂ (1-10 uM), whereas LNO₂ at 10 uM augmented endothelial cell sprouting from rat aortic rings. Interestingly, equimolar doses of its precursor, the linoleic acid, showed no effect on endothelial cell migration. Furthermore, the LNO₂ effects on endothelial cell migration was significantly prevented by a 'NO scavenger pretreatment [2-(4carboxyphenyl)-4,4,5,5-tetramethylimidazoline-1-oxyl-3-oxide (carboxy PTIO, 100 uM)]. **Conclusions**: Our findings indicate that LNO₂ might regulate endothelial cell migration by a *NO-dependent mechanism, and that this lipid mediator may have a major role in the neoangiogenesis process. Financial Support: FAPESP, CNPQ/INSTITUTO do MILÊNIO REDOXOMA