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The actin cytoskeleton and macrophages: Function and regulation of the WASp/WAVE family member, WAVE2, in macrophages

Wassim Aboukheir

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Cell migration is a fundamental process required for normal immune system function,embryonic development,and tissue repair. Nevertheless,this process also contributes to the pathogenesis of several diseases,such as chronic inflammatory disease and tumor cell invasion. The actin cytoskeleton stands central in the process of cell migration. CSF-1 is an important physiological chemoattractant for macrophages. The mechanisms by which CSF-1 affects actin cytoskeleton and induces macrophage migration are not fully understood.In particular,very little is known regarding the role of WAVE2 protein in response to CSF-1. We have identified for the first time WAVE2 as a pivotal factor required for CSF-1-induced cell migration in macrophages.We have also suggested a model for the regulation of WAVE2 activity in vivo in which IRSp53 plays a central role in mediating the interaction between Rac1 and WAVE2.Dissecting the fundamental mechanisms underlying WAVE2-dependent CSF-1-induced cell migration is important for understanding macrophage-mediated diseases,like tumor metastasis, and therefore holds the promise of newly effective therapeutic approaches.

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