



SDI Review Form 1.6

Journal Name:	International Journal of Biochemistry Research & Review
Manuscript Number:	Ms_IJBCRR_23824
Title of the Manuscript:	Adipokines, Receptor activator of nuclear factor-κB ligand, And Midkine: An Intricate biomarkers Network Involved In Pathogenesis And Activity Of Rheumatoid Arthritis in Egyptians
Type of the Article	

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This journal's peer review policy states that **NO** manuscript should be rejected only on the basis of '**lack of Novelty**', provided the manuscript is scientifically robust and technically sound.

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PART 1: Review Comments

	Reviewer's comment	Author's comment (if agreed with reviewer, correct the manuscript and highlight that part in the manuscript. It is mandatory that authors should write his/her feedback here)
<u>Compulsory</u> REVISION comments	In this study serum concentration of TNF-alpha, adiponectin, visfatin and RANKL as well as the expression of midkine in peripheral blood cells was measured in 40 newly diagnosed patients with rheumatoid arthritis and 20 matched control subjects. It is demonstrated that these cyto- and adipokines were elevated in RA patients, were higher in active vs. inactive RA subgroup, and were positively correlated with DAS-28. The study is well-performed and the manuscript is in general well-written. However, there are also some important concerns: 1) The results are not entirely innovative since at least some of these factors have been previously measured in RA. 2) Adiponectin is mentioned as the proinflammatory adipokine in several places in the manuscript. Although adiponectin is proinflammatory in some systems, it reduced inflammation in the majority of studies. Thus, it should not be classified as "proinflammatory". 3) It is suggested that adipokines of interest could be useful in diagnosis and/or monitoring of RA. This is the over-interpretation of results. Diagnostic value of adipokines was not compared to traditional criteria in this study. In addition, as all patients were with recently diagnosed RA, nothing about monitoring could be concluded. 4) What kind of adiponectin (total or HMW) was measured? 5) Some details about ELISA kits should be provided such as detection limit, intra- and interassay coefficients of variation. 6) Correlations between adipo- and cytokines measured in this study should be analysed. In addition, some multivariate analysis would be interesting, for	



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	example to establish which of adipo- and cytokines of interest are the strongest and independent predictors of RA severity.	
Minor REVISION comments	1) It should be specified in the Abstract that MDK mRNA was measured in peripheral blood cells. 2) Page 1: TNF-alpha is classified as the adipokine which is not strictly correct. Although TNF-alpha is produced by adipose tissues, there are other sources of it such as inflammatory cells. Thus, especially if measured only in serum, TNF-alpha should not be classified as the adipokine because the source of its elevation is not clear.	
Optional/General comments		

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