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Journal Name:	British Journal of Medicine and Medical Research
Manuscript Number:	2014_BJMMR_13820
Title of the Manuscript:	Dose-dependent Modulation of Lipid Parameters and Inflammatory Biomarkers by $\delta\textsc{-}Tocotrienol$ in Hypercholesterolemic Subjects
Type of the Article	Original Research Article

General guideline for Peer Review process:

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

To know the complete guideline for Peer Review process, reviewers are requested to visit this link:

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PART 1: Review Comments

Compulsory REVISION 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)
Minor REVISION comments	The authors evaluated the dose-responsive effects of tocopherol-free δ-tocotrienol on serum lipid parameters, and several inflammatory biomarkers (TNF-α, IL-4, IL-6, IL-8) including circulating microRNAs (miRNAs) in hypercholesterolemic subjects for 30-weeks. The results show that low doses of δ-tocotrienol administered for 4 weeks are effective in lowering lipid parameters and also are observed to downregulate potentially pathological several inflammatory processes. Conversely, higher doses of this agent upregulates pro-inflammatory activity. The report includes novel information that adds value to ongoing efforts to develop new anti-inflammatory agents from natural sources. The study design is adequate and the scientific background is solidly grounded.	
	Minor comments: - Use of the English language in the present manuscript is somewhat awkward. The authors occasionally use terms and	

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phraseology that are not in common use – and may thus confuse readers. Some grammatical errors were also noted. Thus, the paper would benefit by review and "tweaking" of the language use by a native speaker of English.

- The Introduction contains useful information – however it should be expanded somewhat to encompass a more comprehensive description of the topic area. Some additional references are also appropriate. This Reviewer feels that it is particularly important for the authors to provide background hypercholesterolaemia and its adverse impact on cardiac parameters. Here, this reviewer suggests citing the following article published in International Journal of Molecular Sciences 2013 17;14(9):19086-108.
- This reviewer thinks in addition to including more detail in the Introduction, the Discussion would benefit by elaborating on some of the points covered in this section of the manuscript. Specifically, it would be informative to readers if the authors would include additional information on some potential therapeutic approaches previously

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published by other investigators. Here, this reviewer suggests citing the following articles: (1) *Current Pharmaceutical Design* 2013;19(39):6896-905; and (2) *Molecular and Cellular Biochemistry* 2011 Feb;348(1-2):205.

- This reviewer believes that the format used to display the figures in the present manuscript is not optimal for reader clarity and requirements of the journal. It is recommended that a simpler version for graphic display be chosen.
- The authors should elaborate and focus on human (and/or veterinary) relevance of the findings.
- Reference format must be uniform according to standards of the journal. It is suggested that some citing program (e.g. EndNote) be used to accomplish this task.
- The authors should elaborate (i.e. add detail) to the sections describing miRNA, and cytokines (which will likely be accomplished incidental to extending the Introduction and Discussion).



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Optional/General comments	

Reviewer Details:

Name:	Anonymous
Department, University & Country	Hungary

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