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From here onwards, the paper should be written with line spacing of 1.5.

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The abstract should not exceed 250 words and should be structured in: Objective, Methods, Results, Conclusions.

**Key words**

Provide up to 5 key words using Mesh terms for indexing purposes.

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Text should not exceed 5000 words from the Introduction to the Conflict of Interest Statement, excluding the title page, abstract and references.

**Introduction**

The introduction should put the focus of the manuscript into a broader context, and keep in mind readers who are not experts in the field. The introduction should conclude with a brief statement of the overall aim of the study.

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**Materials and methods**

This section should provide enough details to allow full replication of the study by suitably skilled investigators.

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Include a review of the key literature. If there are relevant controversies or disagreements in the field, they should be mentioned so that a non-expert reader can delve into these issues further. The discussion should consider the major conclusions of the work along with some explanation and/or speculation on their significance. How do the conclusions affect the existing assumptions and models in the field? How can future research build on these observations? What are the key experiments that remain? The discussion should be concise with solid arguments.

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If a study has not been submitted to an ethics committee prior to commencing, retrospective ethics approval usually cannot be obtained and it may not be possible to consider the manuscript for peer review. How to proceed in such cases is at the Editor(s)’ discretion.

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**Acknowledgements**

Within this section, list those contributors who have not met the authorship criteria. Authors should describe all funding sources (e.g. full name of funding organizations, grant numbers).

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References should be limited to the most essential and relevant, published allegedly in the last decade, identified in the text with consecutive numerals (with numbers in superscript) and listed at the end of the manuscript in the order in which they are cited (use the numbering function of your word processor – do not number by hand).

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