

Dear Sir,

Thank you for your e-mail and the valuable comments/suggestions of the reviewers.

Please find enclosed our manuscript entitled "Antiviral Drugs for Treatment of Human Monkeypox: A Systematic Review/Meta-analysis: Antivirals for Monkeypox" modified according to the valuable recommendations of the reviewers; Modifications in RED.

Here we answer all inquiries/suggestions of the reviewers:

## **FIRST ROUND**

### **Reviewer A:**

5. Ethical aspects. Does the manuscript have a paragraph on ethical aspects, where it mentions approval by the ethics committee, informed consent, and strict compliance with research ethics? **No**

**Answer: Systematic reviews and Meta-analysis are exempted from Ethical Committees Approvals.**

9. References. Quality of bibliographic references and if they are in accordance with the Vancouver format. Poor

**Answer: All references now follow Vancouver style.**

10. Redaction. Is the manuscript correctly written? Does it contain any spelling or grammar mistakes? Needs some language corrections.

**Answer: Done.**

11. Contributions. What are the main weaknesses of the manuscript and how the author can do to improve it

**Answer:**

**Small number of included clinical trials that investigate monkeypox treatment. limited number of patients.**

**This systematic review includes only systemic antivirals not topical one.**

**Till now, no antiviral founded to be FDA approved for monkeypox treatment.**

## **JUSTIFICATION**

Before the aim, lucid and concise justification for the study should be provided.

**Answer: Done.**

## **INTRODUCTION**

The introduction can be improved by the following:

Succinct information on Epidemiology of the disease should be included.

**Answer: Done**

Antiviral drugs previously tested for the treatment of human monkeypox like trifluridine should be included in the introduction aspect.

**Answer: This study includes only systemic antiviral, topical one was excluded.**

**This was reported in the limitations of study.**

Introduction can be made more robust by capturing antiviral drugs that offer cross protection to Monkey-pox disease.

**Answer: Done.**

## **RESULTS**

What does N in Table 2 Represent? if it's just serial numbers, then the numbering should be corrected.

Answer: Done.

Language use in Table 2 should be checked, for instance, for study 14 "Eligible and consented participants will be randomized 1:1 to receive " also for study 15, "In this study, 530 eligible participants will be randomized" , likewise study 16 "The study will provide evidence on the efficacy and safety"

Answer: Done.

DISCUSSION: Authors should enhance the discussion by providing: More detailed information on the pharmacodynamics/pharmacokinetics data.

Answer: Done.

REFERENCING: inconsistent referencing Style

Answer: All references now follow Vancouver style.

CONCLUSION: The conclusion is too terse.

Answer: Modified.

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**Reviewer B:**

Summary: Presents the general idea of the topic, objectives, research methods, results and conclusions, written in an objective and concise manner; and are found according to the maximum number of words per section. Poor

Answer: Done

Redaction: Is the manuscript correctly written? Does it contain any spelling or grammar mistakes? Requires a thorough assessment of grammar and spelling.

Answer: Done

Contributions: What are the main weaknesses of the manuscript and how the author can do to improve it: Manuscript is filled of flaws author need to rearrange the manuscript in correct order.

Answer: Done

**SECOND ROUND**

Reviewer A:

**REVIEWERS'S COMMENTS**

**ABSTRACT**

Inconsistency in writing monkeypox. Do not capitalize the name of the disease.

Answer: Done.

**Capitalise the names all drugs, e.g. Tecovirimat, Cidofovir, Brincidofovir.**

Answer: Done.

**KEYWORDS**

What is the relevance of October 6 University to your studies? It should be deleted.

Answer: Done. October 6 University Deleted.

Monkeypox which is the focus of the manuscript is conspicuously omitted as part of the keywords.

Answer: Done. Monkeypox Added.

## **INTRODUCTION**

**Abbreviations like Rab9 GTPase, TIP47 should be written in full first time, before subsequent abbreviations in the write-up.**

Answer: Done.

**Justification for the study is still not stated.**

Answer: Done.

Till date, there is limited consensus on the optimal treatment strategy for human monkeypox.