

## PEER-REVIEW REPORT

**Name of journal:** *World Journal of Clinical Cases*

**Manuscript NO:** 72912

**Title:** Immunoabsorption therapy for Klinefelter syndrome with APS in a patient : A case report

**Provenance and peer review:** Unsolicited Manuscript; Externally peer reviewed

**Peer-review model:** Single blind

**Reviewer's code:** 00503257

**Position:** Editorial Board

**Academic degree:** MD, PhD

**Professional title:** Professor

**Reviewer's Country/Territory:** Japan

**Author's Country/Territory:** China

**Manuscript submission date:** 2021-12-24

**Reviewer chosen by:** Qi-Gu Yao (Online Science Editor)

**Reviewer accepted review:** 2022-01-14 08:07

**Reviewer performed review:** 2022-01-16 00:13

**Review time:** 1 Day and 16 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

<b>Peer-reviewer statements</b>	Peer-Review: [ <input checked="" type="radio"/> ] Anonymous [ <input type="radio"/> ] Onymous Conflicts-of-Interest: [ <input type="radio"/> ] Yes [ <input checked="" type="radio"/> ] No
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## SPECIFIC COMMENTS TO AUTHORS

This is an interesting case report. However, several concerns are arisen which should be addressed adequately. 1. It is unclear why Patients with Klinefelter syndrome associated with antiphospholipid syndrome showed refractory deep thrombosis. This issue should be clarified. 2. It is unclear why the authors choose immunoadsorption first in this patient. The rationale of this issue should be clarified. 3. This is only a case report with anecdotal clinical experience. Thus, the limitations of their experience should be stated clearly. 4. Also, more concise and logically writing style should be encouraged. The whole description should be much shortened.

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**Peer-review model:** Single blind

**Reviewer's code:** 00863327

**Position:** Peer Reviewer

**Academic degree:** MD, PhD

**Professional title:** Full Professor

**Reviewer's Country/Territory:** Taiwan

**Author's Country/Territory:** China

**Manuscript submission date:** 2021-12-24

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2022-01-27 14:58

**Reviewer performed review:** 2022-01-28 13:12

**Review time:** 22 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

<b>Peer-reviewer statements</b>	Peer-Review: [ <input checked="" type="checkbox"/> ] Anonymous [ <input type="checkbox"/> ] Onymous Conflicts-of-Interest: [ <input type="checkbox"/> ] Yes [ <input checked="" type="checkbox"/> ] No
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## SPECIFIC COMMENTS TO AUTHORS

In this interesting case report, the authors described a 31-year-old male Klinefelter syndrome with chromosome 47 XXY type, who suffered from deep vein thrombosis (DVT) of lower limbs. In addition to elevated serological levels of factor VIII, he had increased titers of antiphospholipid antibodies (aPL) with repeated measurements, leading to the diagnosis of antiphospholipid syndrome (APS). In addition to long-term anticoagulant therapy, he received immunoadsorption, immunosuppressants and intravenous immunoglobulin treatments without the recurrent thrombotic events during one-year follow up. The findings in the manuscript might be relevant to the clinical application. Nevertheless, there are some issues needed to be clarified as follows. 1. After the first thrombotic event, the reported patient was prescribed with daily use of direct-acting oral anticoagulant (DOAC) without the recurrent thrombosis for more than 2 years. His DVT recurred after quitting the use of anticoagulant, and there were no other rheumatological manifestations like skin rash and Raynaud's phenomenon. The International Congress on Antiphospholipid Antibodies (Lupus 2020;29:1571), recommends that DOACs should not be used for recurrent thrombosis while on standard-intensity vitamin K antagonists (VKA), and other treatment options such as increased VKA INR target range, standard treatment dose of low-molecular-weight-heparin (LMWH). and the addition of antiplatelet therapy. Furthermore, the reported patients is a victim of genetic disorder with the only prenotation of venous thrombosis, not a systemic rheumatic disease patient like systemic lupus erythematosus with miscellaneous autoimmune manifestations. Therefore, for this case, the authors should further discuss the possibility of other alternative regimens such

as switch from DOAC to warfarin therapy, and the rationale for using immunosuppressive agents. 2. In Figure 1, figure 1A pointed by the red arrow seemed to be the disappearance of pulmonary embolism without a filling defect, and figure 1B pointed by the red arrow seemed to be the presence of pulmonary embolism with a filling defect. 3. In Figure 3, both sides of the y-axis were labelled with anti-cardiolipin antibody IgG, and there were no demonstration of lupus anticoagulant levels.

## RE-REVIEW REPORT OF REVISED MANUSCRIPT

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**Professional title:** Professor

**Reviewer's Country/Territory:** Japan

**Author's Country/Territory:** China

**Manuscript submission date:** 2021-12-24

**Reviewer chosen by:** Xue-Li Chen (Quit 2022)

**Reviewer accepted review:** 2022-02-16 00:36

**Reviewer performed review:** 2022-02-16 00:40

**Review time:** 1 Hour

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Peer-reviewer</b>	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous



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statements

Conflicts-of-Interest: [ ] Yes [Y] No

#### **SPECIFIC COMMENTS TO AUTHORS**

The revised MS would be OK. I have no further concerns.