

REVIEWER'S REPORT

Manuscript No.: IJAR-57090

Title: Cyclic ACTH-Independent Cushing Syndrome in an Adolescent Revealing Multinodular Adrenal Hyperplasia: A Case Report.

Recommendation:

Accept after minor revision.....

Rating	Excel.	Good	Fair	Poor
Originality		✓		
Techn. Quality		✓		
Clarity	✓			
Significance		✓		

Reviewer's ID: JPR-198

Detailed Reviewer's Report

This is a clinically valuable and well-presented case report of a rare endocrine condition: cyclic ACTH-independent Cushing syndrome due to multinodular adrenal hyperplasia in an adolescent. The diagnostic journey is logically described, the clinical details are sufficient, and the discussion highlights important learning points. However, several minor to moderate deficiencies need correction before publication. No major scientific flaws are present.

Major Comments

Add a structured abstract before the introduction with the following headings:

- **Background** (1–2 sentences on rarity of condition)
- **Case Presentation** (key clinical, biochemical, imaging, and histopathological findings)
- **Conclusion** (main takeaway message)

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Issues to address:

Issue	Recommendation
No patient consent statement	Add a sentence after case presentation: <i>"Written informed consent was obtained from the patient's parents for publication of this case report and accompanying images."</i>
Ferriman-Gallwey score	Mentioned but no numerical values provided. If available, add baseline and follow-up scores. If not, remove or state "qualitative improvement."
Ketoconazole rationale	Why ketoconazole over metyrapone or other steroidogenesis inhibitors? Add a brief justification.
Histopathology – nodule sizes and mitotic index	If available, add size range of nodules and Ki-67 index. If not available, add a limitation sentence: <i>"Quantitative morphometric and immunohistochemical analysis was not available."</i>
In discussion: Negative Liddle test	The manuscript states it "argued against PPNAD." This is correct but note that Liddle test has low sensitivity/specificity. Add a sentence: <i>"However, the Liddle test is no longer considered definitive, and genetic testing for PRKAR1A mutations would be required to definitively exclude PPNAD."</i>
Genetic testing	Given consanguineous parents and young onset, genetic testing (e.g., ARMC5, PRKAR1A, GNAS) is not mandatory for a case report but should be mentioned as a limitation. Add: <i>"Genetic testing was not performed due to limited availability, representing a study limitation."</i>
Unilateral surgery in bilateral disease	Your explanation is adequate (left gland was major functional source, proven by postoperative resolution). However, clarify: was right adrenal function assessed postoperatively? Add: <i>"The right adrenal micronodules have not required intervention to date, and the patient remains on hydrocortisone replacement."</i>