

Overcoming mental health care barriers for postpartum depression in developing countries: Zuranolone as a game changerAqsa Eeman¹, Muhammad Afaq Khan², Hamza Ashraf³, Haider Ashfaq⁴

Dear Madam, Postpartum depression (PPD) is defined as the depressive episode occurring during pregnancy or within the first four weeks after childbirth—although symptoms may arise much later. PPD has been under-researched and frequently under-diagnosed, especially due to lack of awareness, limited access to mental health care, and insufficient training for healthcare providers in recognizing and diagnosing PPD. According to literature from lower-middle-income countries, the PPD is estimated to affect about one in five women.¹

This condition encompasses a wide spectrum of physical and psychological symptoms, including depressed mood, sleep and appetite disturbances, loss of concentration, psychomotor symptoms and suicidal thoughts. The impact extends beyond the mother's health, as affected women are less likely to breastfeed, attend well-child visits, or complete infant immunisations. Depressed females have a lower likelihood of breastfeeding, attending well-child visits, completing infant immunizations, use of safety devices and the essential caution for the infants' posture during its sleep and active hours also gets neglected. Hence, the severity of the mother's depression is directly linked to its impact on the child's health.

Although Brexanolone is currently the first line drug in treating PPD, it has many limitations. One of the major problems in prescribing brexanolone is the need for the patient to be admitted to the hospital for careful monitoring. Moreover, its intravenous (IV) administration is impractical for many patients. The cost of its treatment, ranging somewhere between \$15,000 to \$34,000 per vial, makes it inaccessible to the masses.²

On the other hand, Zuranolone, a neuroactive steroid (NAS), is a positive allosteric modulator for both synaptic and extra synaptic GABAARs, which has been proven to be

an alternative treatment choice for treating PPD. It is the second NAS and first oral medication to receive approval from the FDA for adults suffering from PPD.³ It has a higher efficacy than brexanolone as it can be easily administered once daily and is proven to be more GABA A receptor selective with better oral bioavailability.⁴

A phase 3, double-blind, randomized placebo-controlled trial demonstrated that zuranolone, when taken once daily for two weeks, led to significant improvements in the depressive symptoms.⁵ Zuranolone was well tolerated and demonstrated safety, with only mild side effects such as nausea, dizziness, and sedation. These factors not only make zuranolone a superior treatment option for PPD but also position it as a potential game changer for families affected by maternal depression, particularly in lower-middle-income countries.

In conclusion, lower-middle income countries like Pakistan face significant challenges in raising awareness and providing treatment for mental health conditions such as PPD. The lack of awareness and limited healthcare infrastructure further complicate the situation. Introducing effective treatments like Zuranolone, which offers a more practical and accessible solution, can greatly improve the well-being of mothers experiencing postpartum depression. Enhancing maternal mental health benefits not only immediate families but also contributes to broader societal well-being.

Disclaimer: None.

Conflict of interest: None.

Funding disclosure: None.

DOI: <https://doi.org/10.47391/JPM.22362>

References

1. Wang Z, Liu J, Shuai H, Cai Z, Fu X, Liu Y, et al. Mapping global prevalence of depression among postpartum women. *Transl Psychiatry* 2021;11:543. doi: 10.1038/s41398-021-01663-6.
2. Cornett EM, Rando L, Labbé AM, Perkins W, Kaye AM, Kaye AD, et al. Brexanolone to treat postpartum depression in adult women. *Psychopharmacol Bull* 2021;51:115–30.
3. Parikh SV, Aaronson ST, Mathew SJ, Alva G, DeBattista C, Kanes S, et al. Efficacy and safety of zuranolone co-initiated with an

¹Final Year MBBS Student, Rehman Medical College, Peshawar, Pakistan; ²Final Year MBBS Student, Jinnah Medical College, Peshawar, Pakistan; ^{3,4}4th Year MBBS Student, Allama Iqbal Medical College, Lahore, Pakistan.

Correspondence: Hamza Ashraf. e-mail: a93.hamza@gmail.com

ORCID ID: 0009-0007-9121-6091

Submission completed: 16-10-2024 **1st Revision received:** 09-01-2025

Acceptance: 01-02-2025 **2nd Revision received:** 31-01-2025

- antidepressant in adults with major depressive disorder: results from the phase 3 CORAL study. *Neuropsychopharmacology* 2024;49:467–75. doi: 10.1038/s41386-023-01751-9.
4. Althaus AL, Ackley MA, Belfort GM, Gee SM, Dai J, Nguyen DP, et al. Preclinical characterization of zuranolone (SAGE-217), a selective neuroactive steroid GABAA receptor positive allosteric modulator. *Neuropharmacology* 2020;181:108333. doi: 10.1016/j.neuropharm.2020.108333.
 5. Deligiannidis KM, Meltzer-Brody S, Gunduz-Bruce H, Doherty J, Jonas J, Li S, et al. Effect of zuranolone vs placebo in postpartum depression: a randomized clinical trial. *JAMA Psychiatry* 2021;78:951–9. doi: 10.1001/jamapsychiatry.2021.1559.

Author Contribution:

AE: Writing-original draft, writing, review and editing.

MAK: Writing-original draft, writing, review, editing, project validation and supervision.

HA: Writing, review, editing and project validation.

HA: Writing, review and editing.