

Neuropsychiatric manifestations of Posterior Reversible Encephalopathy Syndrome (PRES) after a severe obstetric complication

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Resumen

Introducción: Posterior Reversible Encephalopathy Syndrome (PRES) is conventionally defined as a clinical and imaging syndrome marked by symptoms such as headaches, encephalopathy, visual disturbances, and seizures. Although psychiatric symptoms might be present.

Clinical case: A 29-year-old female patient hospitalized after a hypovolemic cardiac arrest due to ectopic pregnancy rupture, previously healthy, after a two-week hospitalization started with suicidal ideas, visual hallucinations, headache, and seizures. She presented two episodes of tonic-clonic generalized seizures, depressive symptoms, and hypertension. A brain CT scan showing parieto-occipital edema compatible with PRES findings. After the diagnosis of PRES, antiepileptic medication, antihypertensive therapy, and second-generation antipsychotic drugs were started. After stabilization, there was clinical improvement, and following 40 days she was discharged with no neurological deficit.

Discussion: Patients with PRES have distinct symptoms like seizures, headaches, and visual disturbances, often mimicking other conditions, highlighting the importance of its accurate diagnosis for timely intervention. Moreover, the psychiatry manifestations also should be considered as a part of the clinical features.

Conclusions: Awareness of factors such as hypertension, pregnancy-related complications, and psychiatric manifestations associated with PRES is crucial for personalized and appropriate management to reduce potential risks of this disease.

Keywords: behavioral symptoms; brain edema; case report; reversible posterior leukoencephalopathy syndrome; seizure; neuroimaging.

Manifestaciones neurológicas y psiquiátricas del PRES después de una complicación obstétrica grave

Introducción: el síndrome de encefalopatía posterior reversible (PRES, según sus siglas en inglés) se define convencionalmente como un síndrome clínico y de imagen caracterizado por síntomas como dolores de cabeza, encefalopatía, alteraciones visuales y convulsiones, aunque los síntomas psiquiátricos también pueden presentarse.

Caso clínico: una paciente de 29 años, de sexo femenino, hospitalizada después de un paro cardíaco hipovolémico debido a la ruptura de un embarazo ectópico, previamente sana, tras dos semanas de hospitalización presentó ideas suicidas, alucinaciones visuales, dolor de cabeza y crisis epiléptica. La paciente presentó dos episodios de crisis tónico-clónicas generalizadas, alucinaciones, agitación psicomotora e hipertensión arterial. La tomografía computarizada de cráneo evidenció la presencia de edema vasogénico en la región parietooccipital del cerebro, hallazgos compatibles con PRES. Después del diagnóstico de PRES, se inició el tratamiento con medicamentos anticonvulsivantes, terapia antihipertensiva y medicamentos antipsicóticos de segunda generación. Tras la estabilización, se observó una mejora clínica y, después de 40 días, fue dada de alta sin déficits neurológicos.

Discusión: las crisis epilépticas, las cefaleas y los trastornos visuales suelen presentarse dentro de un extenso grupo de entidades patológicas, no siendo exclusivas en pacientes con PRES. Además, las manifestaciones psiquiátricas deben considerarse como parte relevante de las características clínicas que caracterizan esta enfermedad.

Conclusiones: es fundamental tener en cuenta factores como la hipertensión, las complicaciones relacionadas con el embarazo y las manifestaciones psiquiátricas como entidades comúnmente asociadas al PRES, y lograr así una gestión personalizada, con el fin de reducir los posibles riesgos de esta enfermedad.

Palabras clave: conducta y mecanismos de conducta, edema cerebral, informe de caso, síndrome de encefalopatía posterior reversible, crisis epiléptica, neuroimaging.

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Historia del artículo/Article info

Recibido/Received: November 23, 2023

Evaluado/Revised: April 2, 2024

Aceptado/Accepted: June 5, 2024

Publicado/Published online: July 18, 2024

Citación/Citation: Gómez Jordan S, Polo Trujillo M, Cuellar Lobo M, Cárdenas Castro V, Meza Cely N. Neuropsychiatric manifestations of Posterior Reversible Encephalopathy Syndrome (PRES) after a severe obstetric complication. *Acta Neurol Colomb.* 2024;40(3):e1806. <https://doi.org/10.22379/anc.v40i3.1806>



Introduction

Posterior reversible encephalopathy syndrome (PRES), or reversible posterior leukoencephalopathy syndrome (RPLS), represents a clinical and radiological condition with diverse underlying causes, all characterized by shared neuroimaging features. The syndrome was first introduced in 1996 by Hinchey et al. (1). Although, the precise global incidence of PRES is unknown, it notably tends to manifest during pregnancy, especially among patients who develop pre-eclampsia/eclampsia, those experiencing acute elevation in blood pressure, or undergo immunosuppressive therapy (2–3). Psychiatric manifestations related to PRES are poorly described in the medical literature; they are atypical, with few case reports during and after the illness. Documented symptoms are diverse, ranging from confusion, agitation, and even hallucinations (4).

Here, we present the case of a 29-year-old woman who experienced cardiac arrest, requiring 25 minutes of advanced cardiac life support. After 15 days, she experienced depressive symptoms, hallucinations, seizures, and hypertension. A brain-computer tomography (CT) scan findings of parieto-occipital vasogenic edema suggested the diagnosis of PRES.

Case report

A 29-year-old woman, in her first pregnancy at 9.6 weeks gestation, was admitted with a chief complaint of widespread abdominal pain that had persisted for the past 20 hours. The patient had no prior medical or personal psychiatric history, with a family history of her mother attempting suicide.

Upon admission, her vital signs were as follows: hypotension (69/43 mmHg), tachycardia (120 beats per minute), delayed capillary refill (4 seconds), and signs of peritoneal irritation. Notably, her mental state remained unchanged at this stage. Shortly after admission, the patient experienced cardiac arrest, requiring immediate cardiopulmonary resuscitation (CPR) for 25 minutes. Imaging studies, including abdominal ultrasound and transvaginal obstetric ultrasonography, indicated a likely ruptured ectopic pregnancy. Paraclinical assessments revealed severe anemia (hemoglobin: 4.8 g/dl), consumption coagulopathy, elevated lactic acid levels (12.34 mmol/l), metabolic acidosis, and elevated creatinine (2.82

mg/dl). Subsequently, the patient underwent an exploratory laparotomy, during which 3500 ml of hemoperitoneum was drained, and a left salpingectomy was performed.

She was admitted to the intensive care unit (ICU), where she required mechanical ventilation care and hemodialysis after acute kidney injury. After a two-week ICU stay, she was transferred to the general wards. In the last week, she has been experiencing loss of appetite, sadness, irritability, anhedonia, hopelessness, expressing thoughts like 'I'm not going to recover,' insomnia, feelings of worthlessness, death thoughts without any suicidal intentions or plans, and poor introspection. Additionally, she presented complex visual hallucinations described as 'a black male who visits me every night and puts himself on top of the bed,' causing her to feel fear. No other abnormalities were noted on the mental examination. Psychiatry initiated treatment with olanzapine. Later, she experienced a 50-second episode of generalized tonic-clonic seizure followed by a 1-minute postictal state, and two hours later, another epileptic crisis occurred; the CT scan (figure 1) showed parieto-occipital vasogenic edema. The next day, she developed a severe holocranial headache with blurred vision; during the physical examination, a hypertension crisis (186/121 mmHg) was found, fundus examination was regular as well as cranial nerves, but there was brachio-cranial hemiparesis (0/5), hypoesthesia, augmented reflexes (+++/++++) and a left foot extensor plantar response. A new CT scan (figure 2) showed subarachnoid hemorrhage, intracranial hemorrhage, and vasogenic edema that may be associated with posterior reversible encephalopathy syndrome. The hospital did not count on access to magnetic resonance, so it was not performed. Infections and other autoimmune diseases were ruled out as differential diagnoses.

After admission to the ICU, a pan-angiography revealed normal blood vessels, neurocritical goals were met, hypertension was successfully treated, and there were no new epileptic crises.

After one week, hemodialysis was discontinued, and without needing any additional intervention, she was again transferred to general hospitalization wards. Following her neurological condition's improvement, forty days after her admission, she was discharged with complete resolution of her neurological state.

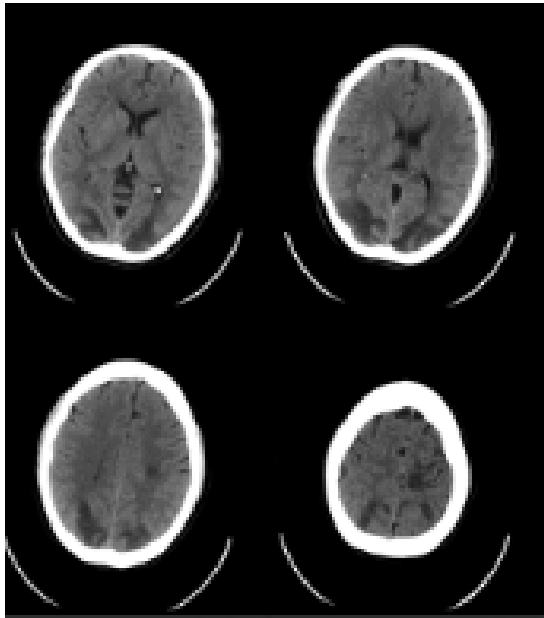


Figure 1. Initial axial brain CT scan

Parietooccipital vasogenic edema, potentially related to reversible posterior leukoencephalopathy syndrome

Source: Image provided by authors.

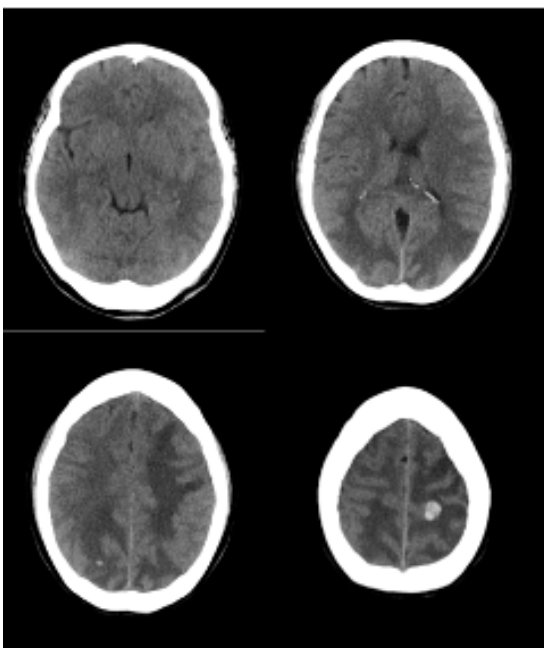


Figure 2. Subsequent axial brain CT scan

Subarachnoid hemorrhage observed in the left frontal convexity, accompanied by intraparenchymal hemorrhagic foci. Vasogenic edema, potentially related to posterior reversible leukoencephalopathy syndrome

Source: Image provided by authors.

Discussion

A case of PRES in a patient with obstetric complications who developed neuropsychiatric manifestations is presented. We will discuss the neuropsychiatric manifestations of PRES and the relevance of this case in shedding light on the significance of these symptoms. PRES is conventionally defined as a clinical and imaging syndrome marked by symptoms such as headaches, encephalopathy, visual disturbances, and seizures (1). Nevertheless, one of the most critical features of this disorder is the potential for symptom reversibility. Moreover, psychiatric comorbidity is less frequently reported. Among the numerous potential causes of PRES, preeclampsia, and eclampsia are commonly identified as the primary triggers (5). Hence, seizures, vision loss, depressive symptoms, and suicide ideas in a full-term primigravida can yield a complex scenario with multiple potential explanations.

A retrospective case series of 47 patients over nine years in England revealed that psychiatric symptoms were reported in 26% of PRES cases. Interestingly, these symptoms could manifest before, during, or after the onset of PRES (4). The most frequently observed clinical features included speech disturbance, confusion, agitation, hallucinations, disinhibition, low mood, delusions, vivid dreams, religious preoccupations, self-harm tendencies, and anxiety (4). Precisely, this patient experienced depressive symptoms and visual hallucinations in the days leading up to the first seizure episode. A literature review of case reports summarizing psychiatric symptoms is presented in table 1. Of these, it must be remarked that the most common symptoms were catatonia, agitation, memory impairment, and psychosis. Moreover, 75% of these cases were females, and half of the patients required antipsychotics or benzodiazepines for symptom relief (6–13). It is worth noting that there is a scarcity of studies that specifically address the psychiatric aspects of PRES morbidity, which is noteworthy as it leaves a gap in our understanding of how psychiatric symptoms may manifest in patients with PRES and what implications this may have for their clinical management and long-term outcomes.

While no specific diagnostic criteria exist, PRES is increasingly recognized as a disorder. Even if it is not common, in the presentation of psychiatric symptoms, the clinician may promptly become suspicious and proceed with diagnostic assessments. In the ap-

Table 1. Comparative overview of psychiatric symptoms in PRES case reports

Author, year and country	Sex and age (years)	History of psychiatric disease	History of organic disease	Neuropsychiatric manifestation of PRES	Psychopharmacology drug use	Vital state (Alive or death)
Gómez S et al., 2024, Colombia (this case)	Female, 29 years old	Family history	No	Irritability, anhedonia, insomnia, worthlessness ideas, death thoughts complex visual hallucinations	Olanzapine	Alive
Anđelo J et al., 2016, Croatia (6)	Female, 26 years old	Non reported	Non reported	Short- and long-term amnesia	Non reported	Alive
Spiegel D et al., 2011, United States (7)	Female, 48 years old	Non reported	Arterial hypertension and cerebrovascular disease	Catatonia, grimacing, stereotypy, visual hallucinations, blunted affect	Olanzapine	Alive
Sarala Kumari D et al., 2020, India (8)	Female, 20 years old	Non reported	Porphyria	Irritability, agitation, psychosis	Clonazepam	Alive
Fernández-Sotos P et al., 2019, Spain (9)	Male, 75 years old	Non reported	Parkinson disease, arterial hypertension, kidney transplantation	Delirium, expansive mood, confusion, visual hallucinations	Non reported	Alive
Shen FC et al., 2008, Taiwan (10)	Male, 60 years old	Non reported	Porphyria	Psychosis, disorientation	Non reported	Alive
Takaoka Y. et al., 2020, Japan (11)	Female, 26 years old	Schizophrenia	Non reported	Attention disorder, decreased visual memory, delusions, catatonia	Non reported	Alive
Tsai S et al., 2016, Australia (12)	Female, 57 years old	Non reported	Non reported	Delusional infestation	Aripiprazole	Alive
Klingensmith K et al., 2017, United States (13)	Female, 61 years old	Schizoaffective disorder	Non reported	Catatonia, auditory manifestations, acute-onset confusion	Lorazepam	Alive

Source: Data generated by authors.

appropriate clinical setting, neuroimaging is crucial for diagnosis. In bilateral cerebral regions supplied by the posterior circulation, edema is apparent in non-contrast computed tomography (CT) brain scans (1). Magnetic resonance imaging (MRI), when available, is preferred for its sensitivity to vasogenic edema through fluid-attenuated inversion recovery and T2-weighted sequences; it typically reveals signal

changes in the bilateral white-matter regions, often within the occipital lobes (14). This patient's CT neuroimaging revealed bilateral symmetrical hypodensities in the white matter of the parieto-occipital areas. The characteristic involvement of the occipital and posterior parietal lobes, which is a defining feature of posterior reversible encephalopathy syndrome (PRES), may be attributed to lower sympathetic

tic innervation in these posterior cerebral regions, compared to the anterior circulation (15). This lower sympathetic innervation renders these areas more susceptible to injury during blood pressure fluctuations (16–17).

An association can be established between neuropsychiatric symptoms and the temporal–limbic network lesions commonly found in PRES (18–19). Neuropsychiatric symptoms often stem from disruptions in specific limbic networks, widely distributed throughout the brain (17). Despite their diffuse nature, these symptoms remain recognizable as manifestations of temporolimbic lesions, albeit lacking neuroanatomical specificity (19). There is an association between hippocampal sclerosis and psychosis in epilepsy that has been well–documented (20).

Additionally, individuals with schizophrenia often display changes in the hippocampi, including gray matter loss, reduced glutamate and disarray of hippocampal cells (21). Moreover, delusions, emotional behaviors, and other neuropsychiatric symptoms may occasionally be secondary to PRES, due to the middle and posterior cerebral arteries supplying regions of the temporal lobe (17). Despite its label as “reversible,” PRES can lead to residual infarcts and subsequent leukomalacia affecting areas of the mesolimbic system, potentially resulting in possible sequelae of neuropsychiatric symptoms (18).

Although PRES is generally considered a reversible condition, it is essential to acknowledge that residual infarcts and leukomalacia can occur as potential consequences, which may align with the possibility of long–term psychiatric symptoms in some patients (4). PRES typically leads to favorable outcomes when managed appropriately, with patients often recovering from neurological deficits within a consistently observed two–week timeframe in subsequent studies (15). The recovery of the presented case aligns with these findings, as the gradual reduction of blood pressure often results in a marked improvement in patients' conditions (22). She also had improvement in psychomotor agitation, and the hallucinations ceased. Nevertheless, it is crucial to recognize that it may lead to residual brain damage, which may be associated with the development of permanent severe neurological impairments and long–term psychiatric symptoms.

Future research and case series on PRES should aim to comprehensively document psychiatric comorbi-

dity, including the period preceding the condition's onset. This approach will help clarify the true significance of these findings and their connection with the neuropsychiatric trajectory of such illnesses.

Conclusions

This case sheds light on the importance of recognizing psychiatric symptoms as a manifestation of PRES, which should not be underestimated. While they are not clearly understood, there are apparent anatomical relationships between PRES pathogenesis and limbic circuits. Consider this possibility as a diagnosis in the appropriate clinical scenario when evaluating psychiatric symptoms. Further investigations could expand our understanding of the neurobiology of this correlation and the short– and long–term clinical outcomes that a neuropsychiatric presentation in PRES could generate in patients.

Data confidentiality. The project adheres to Resolution 8430 of October 4, 1993, of Colombia and had the verbal and written consent of the patient for publication as well as approval from the Hospital Research Committee.

Ethical statement. The authors declare that the procedures were followed according to the regulations established by the Clinical Research and Ethics Committee and to the Helsinki Declaration of the World Medical Association updated in 2013. Also, the project adheres to Resolution 8430 of October 4, 1993, of Colombia and had the verbal and written consent of the patient for publication as well as approval from the Hospital research committee.

Author contributions. Santiago Gomez Jordan: design, conception, acquisition of information, draft and, reviewing for important intellectual content; Maira Polo Trujillo: design, conception, acquisition of information, draft and, reviewing for important intellectual content; Marcela Cuellar Lobo: analysis, acquisition of information, and draft; Valentina Cárdenas Castro: analysis, acquisition of information, and draft; Nohemi Meza Cely: interpretation of data, analysis, reviewing for important intellectual content.

Conflict of interests. The authors declare that they have no conflict of interest in carrying out this document.

Funding source. No institution contributed

financially to the study; the expenses were assumed by the researchers.

Acknowledgements. To the Universidad del Norte.

References

1. Hinchey J, Chaves C, Appignani B, Breen J, Pao L, Wang A, et al. A reversible posterior leukoencephalopathy syndrome. *N Engl J Med.* 1996;334(1):494–500. <https://doi.org/10.1056/NEJM19960223340803>
2. Shaikh N, Nawaz S, Ummunisa F, Shahzad A, Hussain J, Ahmad K, et al. Eclampsia and posterior reversible encephalopathy syndrome (PRES): A retrospective review of risk factors and outcomes. *Qatar Med J.* 2021;1:4. <https://doi.org/10.5339/qmj.2021.4>
3. Kwon S, Koo J, Lee S. Clinical spectrum of reversible posterior leukoencephalopathy syndrome. *Pediatr Neurol.* 2001;24(5):361–4. [https://doi.org/10.1016/S0887-8994\(01\)00265-X](https://doi.org/10.1016/S0887-8994(01)00265-X)
4. Keynejad RC, David AS. Psychiatric morbidity and its prognosis in posterior reversible encephalopathy syndrome. *J Neuropsychiatry Clin Neurosci.* 2020;32(4):385–8. <https://doi.org/10.1176/appi.neuropsych.19080184>
5. Long TR, Hein BD, Brown MJ, Rydberg CH, Wass CT. Posterior reversible encephalopathy syndrome during pregnancy: Seizures in a previously healthy parturient. *J Clin Anesth.* 2007;19(2):145–8. <https://doi.org/10.1016/j.jclinane.2006.07.004>
6. Borovac JA, Božić J, Žaja N, Kolić K, Hrboka V. A global amnesia associated with the specific variant of posterior reversible encephalopathy syndrome (PRES) that developed due to severe preeclampsia and malignant hypertension. *Oxf Med Case Rep.* 2016;4:76–80. <https://doi.org/10.1093%2Fomcr%2Fomw016>
7. Spiegel DR, Varnell C. A case of catatonia due to posterior reversible encephalopathy syndrome treated successfully with antihypertensives and adjunctive olanzapine. *Gen Hosp Psychiatry.* 2011;33(3):302.e3–5. <https://doi.org/10.1016/j.genhosppsy.2011.01.007>
8. Daram SK, Arumilli MN, Reddy LS, Reddy DN, Motor R. Acute intermittent porphyria presenting with posterior reversible encephalopathy syndrome: A rare cause of abdominal pain and seizures. *Indian J Crit Care Med.* 2020;24(8):724–6. <https://doi.org/10.5005/jp-journals-10071-23532>
9. Fernandez-Sotos P, Lopez-Arcas Calleja P, Lozano-Vicario L, Garcia-Tercero E, Del Yerro-Alvarez MJ, Lopez-Alvarez J. Síndrome de leucoencefalopatía posterior reversible en un paciente con enfermedad de Parkinson y sintomatología inicial psiquiátrica: una presentación clínica compleja. *Rev Neurol.* 2019;68(10):426–30. <https://doi.org/10.33588/rn.6810.2018435>
10. Shen F, Hsieh C, Gung C, Hospital M, Lui C, Chuang Y. Acute intermittent porphyria presenting as acute pancreatitis and posterior reversible encephalopathy syndrome. *Acta Neurol Taiwan.* 2008;17(3):177–83.
11. Takaoka Y, Akaho R, Inada K, Muraoka H, Hokama C, Inoue A, et al. Posterior reversible encephalopathy syndrome due to acute water intoxication in a patient with schizophrenia. *Int Med Case Rep J.* 2020;13:117–21. <https://doi.org/10.2147%2FIMCRJ.S237430>
12. Tsai SJ, Yeh CB, Wang CW, Mao WC, Yeh TC, Tai YM, et al. Delusional infestation in a patient with posterior reversible encephalopathy syndrome. *Aust N Z J Psychiatry.* 2016;50(12):1212–1213. doi: <https://doi.org/10.1177/0004867416656259>
13. Klingensmith K, Sanacora G, Ostroff R. Co-occurring catatonia and posterior reversible encephalopathy syndrome responsive to electroconvulsive therapy. *J ECT.* 2017;33(3):e22. <https://doi.org/10.1097/yct.0000000000000407>
14. Bartynski WS, Boardman JF. Distinct imaging patterns and lesion distribution in posterior reversible encephalopathy syndrome. *Am J Neuroradiol.* 2007;28(7):1320–1327. <https://doi.org/10.3174/ajnr.A0549>.
15. Fugate JE, Rabinstein AA. Posterior reversible encephalopathy syndrome: clinical and radiological manifestations, pathophysiology, and outstanding questions. *Lancet Neurol.* 2015;14(9):914–925. [https://doi.org/10.1016/s1474-4422\(15\)00111-8](https://doi.org/10.1016/s1474-4422(15)00111-8)
16. Lassen NA. Regulation of cerebral circulation. *Acta Anaesthesiol Scand Suppl.* 1971;15(S45):78–80. <https://doi.org/10.1111/j.1399-6576.1971.tb00661.x>
17. Bartynski WS. Posterior reversible encephalopathy syndrome, part 2: Controversies surrounding pathophysiology of vasogenic edema. *Am J Neuroradiol.* 2008;29(6):1043–9. <https://doi.org/10.3174/ajnr.A0929>.

18. Lee VH, Wijdicks EF, Manno EM, Rabinstein AA. Clinical spectrum of reversible posterior leukoencephalopathy syndrome. *Arch Neurol*. 2008;65(2):205–10. <https://doi.org/10.1001/archneurol.2007.46>
19. Trimble MR, Mendez MF, Cummings JL. Neuropsychiatric symptoms from the temporolimbic lobes. *J Neuropsychiatry Clin Neurosci*. 1997;9(3):429–38. <https://doi.org/10.1176/jnp.9.3.429>.
20. Erwin LG, Fortune DG. Risk factors for psychosis secondary to temporal lobe epilepsy: a systematic review. *J Neuropsychiatry Clin Neurosci*. 2014;26(1):5–23. <https://doi.org/10.1176/appi.neuropsych.12120403>
21. Singh S, Khushu S, Kumar P, Goyal S, Bhatia T, Deshpande SN. Evidence for regional hippocampal damage in patients with schizophrenia. *Neuroradiology*. 2018;60(2):199–205. <https://doi.org/10.1007%2Fs00234-017-1954-4>
22. Roth C, Ferbert A. Posterior reversible encephalopathy syndrome: Long-term follow-up. *J Neurol Neurosurg Psychiatry*. 2010;81(7):773–7. <https://doi.org/10.1136/jnnp.2009.189647>