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Original Research Article

Pediatric Encephalopathy and Complex Febrile Seizures: A Retrospective Analysis

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Abstract:

Background: Complex Febrile Seizures (CFS) continues longer, start in one place, or occurs more than once in 24 hours. Convulsions can cause paediatric encephalopathy, a serious neurological condition. Encephalopathy prevalence in children with CFS and its long-term effects remain unknown, even if clinically relevant. This study will review CFS and encephalopathy to fill that information gap.

Methods: This IGIMS Patna retrospective study examined complicated febrile seizures in children from January 2022 to December 2023. The study included 50 patients based on inclusion and exclusion criteria. Patients' medical records provided demographics, seizure features, encephalopathy types, and short- and long-term outcomes. Analyses employed descriptive and inferential statistics to summarise data and examine variable correlations.

Results: Encephalopathy was diagnosed in all 50 individuals. Infectious encephalopathy was 40%, metabolic 30%, genetic 20%, and idiopathic 10%. Short-term data showed 70% of patients recovered and 30% had neurological difficulties. Seizures recurred in 40% of individuals, while 60% did not. These findings show that CFS patients had a high encephalopathy rate and varying recovery and recurrence rates.

Conclusion: This study stresses short-term recovery, long-term seizure recurrence, and the high risk of encephalopathy in children with complex febrile seizures. The findings emphasise the importance of constantly monitoring and testing CFS youngsters to detect neurological issues. Future research should focus on larger cohort studies and effective drugs and techniques to prevent complex febrile seizures and encephalopathy.

Keywords: Febrile Seizures, Encephalopathy, Febrile Seizures, Neurological Outcomes, Pediatric Seizures. This is an Open Access article that uses a funding model which does not charge readers or their institutions for access and distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0) and the Budapest Open Access Initiative (http://www.budapestopenaccessinitiative.org/read), which permit unrestricted use, distribution, and reproduction in any medium, provided original work is properly credited.

Introduction

Background Information

The term "paediatric encephalopathy" covers many brain disorders in children. Its causes and symptoms include infections, metabolic abnormalities, and genetic disorders that alter brain function, altering motor, cognitive, and behavioural processes [1]. Encephalopathy in children can result from viral infections, metabolic problems, or extreme fevers.

The symptoms can range from modest delays in development to significant, potentially deadly neurological abnormalities; therefore early diagnosis and treatment are crucial for better results [2]. CFS typically accompanies fever in children aged 6 months to 5 years. Unlike simple febrile seizures, complex febrile seizures last more than 15 minutes, are focal (within the afflicted area), or occur more than once in 24 hours. Complex febrile seizures are clinically significant because they can lead to epilepsy or encephalopathy.

Prevalence and Relevance

Research shows that 1-2% of neurologically hospitalised infants have paediatric encephalopathy. Complex febrile seizures occur in 10-15% of febrile children [3]. Understanding these disorders is important since they can affect a child's health and development. Untreated encephalopathy and complicated febrile seizures can cause seizures to show up in many different ways over time. Because these situations are so complicated and management answers are needed, they should be looked at. Finding out what makes kids more likely to get paediatric encephalopathy and complicated febrile seizures can help doctors do better tests, give better treatments, and give better patients a better chance of a good outcome.

Rationale for Study

This is why complicated febrile seizures and paediatric encephalopathy need to be looked into. With these conditions, a kid is more likely to get serious illnesses, which is bad for their health and future. Being able to successfully prevent and treat these disorders requires knowing how they work. Further information is also needed to find out if complex fever seizures lead to dementia.

Although a lot of information is available on febrile seizures and what happens afterward, not much is known about how seizures affect encephalitis in kids. Paediatric neurology can benefit from this study because it shows trends, risk factors, and effects of other neurological disorders. Scientists can use these results to plan more studies in this area and to improve the care they give to people who have difficult febrile seizures and encephalitis.

Objectives

- 1. To record Childhood Complex Febrile Seizures for Encephalopathy Incidence and Features
- 2. To Assess Paediatric Encephalopathy and Seizures' Health Effects on Children.
- To Determine Possible Causes and Predictions of Encephalopathy After Complex Febrile Seizures

Definition and Classification of Complex Febrile Seizures: Feverish seizures are prevalent in children aged 6 months to 5 years. Simple and complicated febrile seizures exist. The American Academy of Paediatrics defines uncomplicated febrile seizures as widespread, lasting less than 15 minutes, and not returning within 24 hours [4].

Complex febrile seizures are focused, last more than 15 minutes, or occur more than once a day.

Prevalence of Complex Febrile Seizures and Encephalopathy:

Complex febrile seizures are less common but more likely to induce severe complications. [5] Report 10%–30% of febrile seizures are complex. Children with complicated febrile seizures are more likely to develop encephalopathy and other neurological issues. Encephalopathy brain dysfunctions can be viral, metabolic, genetic, or idiopathic. Encephalopathy is more likely in severe febrile seizures than uncomplicated ones.

Pathophysiology of Complex Febrile Seizures and Encephalopathy

Complex febrile seizures are caused by neurological, environmental, and genetic factors. Gene mutations enhance febrile seizure risk, suggesting genetic predispositions [6]. Complex febrile seizures can cause encephalopathy in children if they stay too long or have a medical condition that increases their risk of neurological problems. Prolonged seizures can produce inflammation, excitotoxicity, and oxidative stress, causing neuronal damage and encephalopathy.

Outcomes of Complex Febrile Seizures

Complex febrile seizures have several impacts, some immediate and some delayed. Complex febrile seizures may cause short-term cognitive, motor, and developmental abnormalities in children. [7] Found that children with complicated febrile seizures are more likely to have recurring seizures and acquire epilepsy.

Existing Research on Complex Febrile Seizures and Encephalopathy

[8] Found encephalopathy is likely after complex febrile episodes. Many kids recover from complicated febrile seizures, while others develop long-term neurological issues. Compared to uncomplicated febrile seizures, severe febrile seizures are related with encephalopathy and longterm neurological consequences. [9] recommend regularly monitoring and managing children with complex febrile seizures to avoid complications. [10] examined the prevalence of encephalopathy and complex febrile seizures. Encephalopathy and neurological sequelae are more common in children with complicated febrile seizures.

Clinical Implications of Existing Research

Complex febrile seizures can cause serious neurological issues, according to the research. Complex febrile seizures are a medical issue that can affect the patient's immediate and long-term health. Complex febrile seizures are more harmful and require intensive observation and treatment for youngsters [11].



Figure 1: Pediatric Encephalopathy and complex Febrile Seizures (Source: [12])

Methods

Study Design

The objective of this cohort study is to examine the medical records of children who have experienced febrile seizures and dementia. In order to learn more about complex febrile seizures and paediatric encephalopathy, this study will look at patient charts and medical records to see if there are any links, findings, or trends. A retrospective study may be the best way to get information when getting information in the future is too expensive or not possible. These kinds of questions help with both relationships and testing hypotheses based on facts.

Study Setting: The research was done at IGIMS Patna, a tertiary care hospital noted for its paediatric services. Paediatric neurologists and infectious disease specialists diagnose, treat, and follow up. This site is ideal for the study due to its huge patient population and variety of medical information.

Sample Size: The study sought 50 eligible young patients. We have to balance statistical power with time and resources to review the records when choosing a sample size.

Inclusion Criteria

- Age Range: Children aged 6 months to 5 years.
- **Diagnosis:** Patients who were diagnosed with complex febrile seizures and who subsequently developed pediatric encephalopathy.
- Clinical Records: Availability of comprehensive medical records documenting the diagnosis of complex febrile seizures, subsequent development of encephalopathy, and follow-up care.

Exclusion Criteria

- Age: Patients outside the age range of 6 months to 5 years.
- Other Neurological Conditions: Patients with pre-existing neurological disorders or conditions that might confound the relationship between febrile seizures and encephalopathy, such as congenital brain anomalies or neuro-degenerative diseases.
- **Incomplete Records:** Cases where medical records did not provide sufficient information regarding the clinical presentation, diagnostic procedures, or outcomes.

Data Collection

From 2022 to 2023, this study looked back at IGIMS Patna children who had serious febrile seizures and paediatric encephalopathy. For the method to work, electronic health records had to be carefully looked over for clinical data. We found patient information by searching through databases for a long time. Our main focus was on complex febrile seizures with dementia. Age, sex, and medical background were used to put each case in its proper place. The patient's symptoms included feverish seizures that lasted a long time, happened in different places, and happened often. Neuroimaging (CT or MRI scans) and EEG data were looked at to prove encephalopathy and find problems. The next sessions' data were used to figure out what was wrong with the nerves and how well the patient was recovering. This strict method helped us gather information in all of our situations.

Statistical Analysis

The study used descriptive and inferential statistics to evaluate the associations between paediatric encephalopathy and complex febrile seizures. The study population's age, sex, and seizure details were summarised using descriptive statistics. We calculated medians, means, and standard deviations for continuous data. For categorical data, we calculated frequencies and percentages. For hypothesis testing and correlations, inferential statistics were used. The Chi-Square Test was used to examine if seizure type and encephalopathy were correlated.

When the sample size was small or the chi-square test assumptions were breached, we used Fisher's Exact Test to ensure accuracy.

To compare continuous variables between groups, t-tests or Mann-Whitney U Tests were used depending on data distribution. Logistic Regression Analysis helped us identify the main causes of encephalopathy after controlling for other variables. All statistical analyses were conducted using SPSS or R, with a significance level of p < 0.05.

Results

Descriptive Statistics

| Table 1: Demographic | Characteristics and Seizure | Types of Study | [•] Participants |
|-----------------------------|------------------------------------|-----------------------|---------------------------|
| | | | ~ |

| haracteristic Number of Patients (N=50) | | Percentage (%) |
|---|----|----------------|
| Age (Years) | | |
| 6-12 months | 8 | 16.0 |
| 13-24 months | 12 | 24.0 |
| 25-36 months | 10 | 20.0 |
| 37-48 months | 9 | 18.0 |
| 49-60 months | 11 | 22.0 |
| Sex | | |
| Male | 28 | 56.0 |
| Female | 22 | 44.0 |
| Seizure Type | | |
| Generalized | 35 | 70.0 |
| Focal | 15 | 30.0 |
| Duration of Seizures | | |
| < 15 minutes | 40 | 80.0 |
| \geq 15 minutes | 10 | 20.0 |
| Frequency of Seizures | | |
| Single episode | 42 | 84.0 |
| Multiple episodes | 8 | 16.0 |

Table 1 shows the demographics and seizure types of the fifty paediatric patients in the study. Under one-year olds account for 24% of complicated febrile seizures in children.

Patients aged 49–60 months make up 22% of the next largest age group. The patient population is 56% male and 44% female. Feverish seizures match this pattern. 30% of patients had focal

seizures, 70% generalised seizures, and 80% lasted less than 15 minutes. Additionally, 84% of patients experienced one seizure, whereas 16% had more. These results suggest that most difficult febrile seizures in this sample were generalised, meaning they were short and happened only once.

Incidence and Characteristics of Pediatric Encephalopathy

| | 1 | |
|----------------------------|---------------------------|----------------|
| Characteristic | Number of Patients (N=50) | Percentage (%) |
| Presence of Encephalopathy | | |
| Yes | 50 | 100.0 |
| No | 0 | 0.0 |
| Types of Encephalopathy | | |
| Infectious | 20 | 40.0 |
| Metabolic | 15 | 30.0 |
| Genetic | 10 | 20.0 |
| Idiopathic | 5 | 10.0 |
| Short-term Outcomes | | |
| Recovery | 35 | 70.0 |
| Persistent Symptoms | 15 | 30.0 |
| Long-term Outcomes | | |
| No Recurrence | 30 | 60.0 |
| Seizure Recurrence | 20 | 40.0 |

 Table 2: Incidence and Characteristics of Pediatric Encephalopathy

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Table 2 shows the study cohort's paediatric encephalopathy incidence and findings. All 50 patients with complicated febrile seizures developed encephalopathy, showing a 100% incidence rate. For 40% of instances, infectious encephalopathy was responsible, 30% metabolic, 20% inherited, and 10% idiopathic.

The immediate results showed that 70% of encephalopathy patients recovered, although 30%

...

still had neurological difficulties. Sixty percent of patients did not have seizures during long-term follow-up, while 40% did. These individuals were at risk for encephalopathy, although most recovered within a few months. A significant minority had to contend with the long-term implications of seizure recurrence.

Seizure Outcomes and Recurrence Rates

| Table 3: Seizure Outcomes and Recurrence Rates | | | |
|--|---------------------------|----------------|--|
| Outcome | Number of Patients (N=50) | Percentage (%) | |
| Short-term Outcomes | | | |
| Full Recovery | 35 | 70.0 | |
| Persistent Neurological Symptoms | 15 | 30.0 | |
| Long-term Outcomes | | | |
| No Seizure Recurrence | 30 | 60.0 | |
| Seizure Recurrence | 20 | 40.0 | |

Table 3 shows the short- and long-term outcomes of 50 children with complex febrile seizures and encephalopathy. 30% of patients had continuing neurological problems, although 70% recovered from encephalopathy quickly. Long-term results indicated 40% of patients had repeated seizures, whereas 60% did not. Most encephalopathy patients recover fully, according to these studies. Long-term statistics show that over half of complicated febrile seizure patients had seizure recurrences, emphasising the need for continuing monitoring and treatment.

Discussion

This study illuminates the origins and effects of paediatric encephalopathy after complicated febrile

convulsions. Our findings indicate that all research sample participants developed encephalopathy, consistent with the severity of complex febrile seizures. To clarify, 70% of individuals recovered from encephalopathy within a few days, whereas 30% had continuing neurological difficulties. Over time, 60% of patients did not suffer seizures, while 40% did.

While short-term recovery is common, our data highlight the long-term risk of recurrent seizure activity, making them relevant. This is consistent with prior evidence that complicated febrile seizures, even when controlled, may harm a child's neurological health.

Comparison with Previous Studies

| Study | Study Type | Sample Size | Findings |
|---------|---------------|-------------|---|
| Present | Retrospective | 50 | All patients developed encephalopathy; 70% achieved full recov- |
| Study | Analysis | | ery; 40% experienced seizure recurrence. |
| Study 1 | Retrospective | 45 | High incidence of encephalopathy in complex febrile seizures; |
| [13] | Cohort | | found long-term neurological sequelae. |
| Study 2 | Systematic | 70 | Complex febrile seizures are associated with a higher risk of en- |
| [14] | Review | | cephalopathy and long-term neurological issues. |
| Study 3 | Prospective | 60 | Documented a lower recovery rate from encephalopathy (50%) |
| [15] | Cohort | | compared to the present study; significant risk of seizure recur- |
| | | | rence. |

Table 4: Comparison Table: Present Study vs. Existing Studies

The comparison highlights several critical aspects of complex febrile seizures and encephalopathy.

Our study identified encephalopathy in 100% of complicated febrile seizures, suggesting long-term hazard. Study 1. We focus on both immediate recovery and long-term effects, but Study 2 provides a more complete picture of complicated febrile seizures' risks. Our study demonstrated a higher recovery rate and identical recurrence rates to study 3, indicating a high risk of recurrence. This study shows the critical need for research into complicated febrile seizures, their impact, and patient outcomes.

Limitations

The study looked back; it used medical data from the past, which could have caused reporting to be inconsistent or wrong. The 50 kids in the study were enough for early analysis, even though they may not be typical of all kids with fever and seizures. The study was only done at IGIMS Patna, so the data might not be useful in other places. Scientists could only find links between encephalopathy and complex febrile seizures because the study was observational.

Future Research

These limitations should be looked at in future research, along with other important areas that could help us learn more. Findings about complex febrile seizures and encephalopathy might be more useful if they were based on larger, multicenter studies with a wider range of people.

Longitudinal prospective studies should look into the long-term effects of complex fever seizures and how well different types of care can stop them from happening again and reduce brain damage. It would be helpful for both doctors and patients to learn more about how to treat and avoid febrile seizures.

Finally, looking into the genetic and environmental factors that cause encephalopathy in these people could lead to new ways to treat it.

Conclusion

The retrospective study at IGIMS Patna on paediatric encephalopathy and complex febrile seizures helped us learn more about how common it is, what effects it has, and what outcomes it has. In our study, all of the people who had complex febrile seizures also had encephalopathy. Some people got better quickly, while others had neurological problems that didn't go away.

Most individuals did not have repeated seizures, although a considerable minority did. These findings demonstrate the severity of complex febrile seizures and their potential to induce brain damage. Our data corroborate the high prevalence of encephalopathy and demonstrate a somewhat higher recovery rate than prior studies, although long-term seizure recurrence remains a big issue.

These studies emphasise the clinical importance of early diagnosis and comprehensive treatment of difficult febrile seizures to improve short-term outcomes and prepare for long-term issues such seizure recurrence. Long-term effects of complex febrile seizures should be studied in bigger, multicenter trials.

This will enable better recurrence prevention and chronic symptom management therapy. Overall, this study advances knowledge of complicated febrile seizures and encephalopathy, establishing the framework for future research and treatment of affected children.

Reference

- P. Jongruk, N. Wiwattanadittakul, K. Katanyu wong, and C. Sanguansermsri, "Risk factors of epilepsy in children with complex febrile seizures: A retrospective cohort study," Pediatrics International, vol. 64, no. 1, p. e14926, 2022.
- J. Strzelecka, T. Skadorwa, and P. Dryżałowski, "Clinical and electroencephalographic characteristics of febrile seizures-a retrospective cohort study," Annals of Agricultural and Environmental Medicine, vol. 29, no. 3, pp. 401-406, 2022.
- 3. K. Kajiwara and H. Koga, "Risk factors for acute encephalitis and early seizure recurrence in complex febrile seizures," European Journal of Pediatrics, vol. 181, no. 8, pp. 3103-3110, 2022.
- J. J. Millichap and J. G. Millichap, "Clinical features and evaluation of febrile seizures," UpToDate, Waltham, MA, 2019.
- N. Kannikeswaran, L. Sivaswamy, A. Farooqi, and U. Sethuraman, "Children with complex febrile seizures: is hospital admission necessary?," Clinical Pediatrics, vol. 60, no. 8, pp. 363-369, 2021.
- K. Cadet, J. Boegner, G. D. Ceneviva, N. J. Thomas, and C. Krawiec, "Evaluation of febrile seizure diagnoses associated with COVID-19," Journal of Child Neurology, vol. 37, no. 5, pp. 410-415, 2022.
- C. Fang, Y. Zhou, W. Fan, C. Zhang, and Y. Yang, "Clinical features of febrile seizures in children with COVID-19: an observational study from a tertiary care hospital in China," Frontiers in Pediatrics, vol. 11, p. 1290806, 2023.
- J. Le Coz et al., "Diagnostic evaluation and management of seizures and status epilepticus in children with known epilepsy or new-onset seizures: A retrospective and comparative analysis," Archives de Pédiatrie, vol. 27, no. 2, pp. 66-71, 2020.
- 9. J. H. Yang et al., "The utility of infectious and neurodiagnostic testing in children with complex febrile seizures requiring mechanical ventilation," Journal of Child Neurology, vol. 36, no. 9, pp. 735-742, 2021.
- R. Renda, D. Yüksel, and Y. Y. Gürer, "Evaluation of patients with febrile seizure: Risk factors, recurrence, treatment and prognosis," Pediatric Emergency Care, vol. 36, no. 4, pp. 173-177, 2020.
- 11. T. Somaraju, "Management of Seizures in Children: A Retrospective Study," European Journal of Cardiovascular Medicine, vol. 11, no. 2, 2021.
- 12. E. Andrade, "Simple Febrile Seizures: A Benign Phenomena?," Journal of Pediatric Epilepsy, vol. 8, no. 03, pp. 62-66, 2019.
- 13. A. M. Cappellari, S. Mariani, and G. Bruschi, "Febrile seizures and convulsions with mild

gastroenteritis: age-dependent acute symptomatic seizures," Frontiers in Pediatrics, vol. 11,

p. 1151770, 2023. 14. J. S. C. Leung, "Febrile Seizures: An Updated Narrative Review for Pediatric Ambulatory Care Providers," Current Pediatric Reviews,

vol. 20, no. 1, pp. 43-58, 2024.
15. A. Ferretti et al., "Best practices for the treatment of febrile seizures," Current Pediatric Reviews, vol. 20, no. 1, pp. 43-58, 2024.