

ORIGINAL ARTICLE

## Serum IgG, and IgM Levels in Children with Febrile Convulsions

by

GULHIS DEDA\*, UGUR KARAGOL\*\* and YAVUZ GURER\*

(\* Pediatrician and Pediatric Neurologist at the Department  
of Pediatric Neurology, \*\* Associated Professor at the Department  
of Pediatric Neurology, Dr. Sami Ulus Children's Hospital,  
Ankara, Turkiye )

### Abstract

At the Dr. Sami Ulus Children's Hospital Ankara, Turkiye, 20 patients, twelve being between 6 months-2 year old and eight being between 2-4 years, with their first febrile convulsions (FC) were examined for serum IgG, IgA, IgM levels during the period of March 1989-July 1989. Twenty healthy children were used as controls, seven being between 6 months-2 years old and thirteen being between 2-6 years. The serum IgG, IgA and IgM levels of the patients between 6 months-2 years were  $805.000 \pm 307.8984$  mg/dl.,  $49.7167 \pm 27.9807$  mg/dl and  $155.1833 \pm 62.9696$  mg/dl respectively.

The serum IgG, IgA and IgM levels of the patients between 2-4 years were :  $989.1250 \pm 314.5359$  mg/dl.,  $92.6125 \pm 34.5663$  mg/dl. and  $159.8750 \pm 45.6647$  mg/dl respectively. The mean IgA levels of the 12 FC patients between 6 months-2 years were  $49.7167 \pm 27.9807$  mg/dl. and the mean level of IgA in the age matched control group was  $81.0427 \pm 31.3551$  mg/dl. and the difference between them was statistically significant ( $p < 0.005$ ).

We conclude that FC under 2 years of age is associated with low serum IgA levels.

## Introduction

Febrile convulsion (FC) is a type of convulsion that can be seen in children between 6 months and 5 years of age without any acute intracranial infection, meningitis or chronic brain diseases [1,2].

The incidence of FC is 3-4% [2,3,4,5,6,7]. FC is predominantly seen in boys and the ratio of boys to girls is 1.4/1 [3,7].

2-4% of children who experience FC may later develop epilepsy [7]. The type of epilepsy that develops after FC is usu-

ally of the generalized type [8].

In 1975, Seager et al. discussed the relationship between low serum IgA levels and FC [9]. Ariizumi et al. in 1980 detected low serum IgA levels in 33% of patients with FC and in 57% of epileptic patients who had had FC [10].

We studied the serum IgG, IgA and IgM levels in 20 patients with febrile convulsions, 12 being between 0-2 years and 8 being between 2-4 years of age, using age-matched controls.

## Materials and Methods

This study carried out in 20 children with their first FC between March and June 1989 at Dr. Sami Ulus Children's Hospital. The age of the cases ranged between 6 months and 4 years.

The quantitative levels of serum IgG, IgA and IgM on the first or second day were measured in 20 children with FC but we were able to examine IgG, IgA and IgM levels in only 5 patients who came for their follow up examination. Three cc of venous blood was taken and centrifuged for 1 minute at 3000 rpm.

The sera were kept at -20 C. We used Behring Institute Hoechst Nor-Partigen plaques for the quantitative measurement of serum IgG, IgA and IgM levels using radial immuno-diffusion method. The measured values were converted to mg./dl. using Nor-Partigen Reference Value Calculators [11].

The serum IgG, IgA and IgM level measured on the first or second day of FC were compared statistically with values for the age-matched control group using the Chi-Square method [12].

## Results

Of the 20 FC patients 7 were girls (35%) and 13 (65%) were boys. Their ages ranged between 6 months and 48 months with the mean of  $25.16 \pm 14.61$  months.

For the purpose of the research the patients must not have had any previous FC. The duration of the FC varied between 1 and 20 minutes with the mean of  $7.33 \pm 7.35$  minutes.

The patients temperature just before or after FC could only be measured in 14 children. The temperature range was 38-

41 C with a mean of  $39.4 \pm 0.96$  C.

According to their neurologic examinations all the patients had normal status.

The serum IgG, IgA and IgM levels on the first or second day of FC between 6 months-2 years of age were :  $805.0000 \pm 307.8984$  mg./dl.,  $49.7167 \pm 27.9807$  mg./dl. and  $155.1833 \pm 62.9696$  mg./dl, while between 2-4 years of age the serum IgG, IgA and IgM levels were  $989.1250 \pm 314.5339$  mg./dl,  $92.6125 \pm 34.5663$  mg./dl and  $159.8750 \pm 45.6647$  mg./dl. (Table I).

Table I. Mean IgG, IgA and IgM levels of patients with FC between 6 months-2 years of age and the age-matched control

Patients with FC	Control Group	P
IgG (mg./dl.) $805.0000 \pm 397.8984$	$958.4286 \pm 305.5448$	> 0.005
IgA (mg./dl.) $49.7167 \pm 27.9807$	$81.0429 \pm 31.3551$	< 0.005
IgM (mg./dl.) $155.1833 \pm 62.9696$	$158.1571 \pm 36.3420$	> 0.005

The serum IgG, IgA and IgM levels of the age-matched control group between 6 months-2 years were :  $958.4286 \pm 305.5448$  mg./dl.,  $81.0429 \pm 31.3551$  mg./

dl and  $158.1571 \pm 36.3420$  mg./dl where-as between 2-6 years were :  $1117.3846 \pm 246.1194$  mg./dl.,  $94.2692 \pm 30.1359$  mg./dl. and  $148.0769 \pm 40.0260$  md./dl.

## Discussion

Twenty patients with FC were examined at the Department of Pediatric Neurology at the Dr. Sami Ulus Children's Hospital between March and June 1989. Their ages ranged between 6 months and 48 months (mean  $25.16 \pm 14.61$  months). Thirteen (65%) of these 20 children were boys, 7 (35%) of these were girls. In recent reports it was found that boys had FC more than girls and the ratio was 1.4-1.6/1 [3,7]. In our study the ratio of boys to girls was 1.8/1.

The mean fever temperature in our patients that could be measured at the time of FC was  $39.4 \pm 0.96$  C. It is said that one of the predicting factors for FC to develop into afebrile convulsion is the fever temperature. If the fever temperature is under  $38.4$  C during FC the patient will develop afebrile convulsions later in his life [13]. In our series the mean fever temperature was  $39.4 \pm 0.96$  C during FC. We can say that most of our patients will not have this risk, except the one whose fever temperature was  $38^{\circ}$ C.

In 18 patients the duration of the convulsion was less than 15 minutes and in 2 patients it was 20 minutes. The mean du-

ration of the FC was  $7.33 \pm 7.35$  minutes. In some literatures it is said that FC usually lasts less than 15 minutes. If it is longer than 15 minutes the risk of recurrences and epilepsy increase [4,13,14,15].

We found that 5 of our patient's parents were related (26.3%). Three were first degree relatives (15.7%) and two were second degree relatives (10.5%). One patient had a brother with a history of FC and the parents were not related. In 2 of our patients families there was a history of epilepsy. Although there are many reports of genetic transmission of FC, the most common mode is the polygenic transmission [14,16].

The mean serum IgG, IgA and IgM levels in FC patients between 6 months-2 years were  $805.0000 \pm 307.8984$  mg./dl.,  $49.7167 \pm 27.9807$  mg./dl and  $155.1833 \pm 62.9696$  mg./dl. (Table I) while between 2-4 years were :  $989.1250 \pm 314.5339$  mg./dl.,  $92.6125 \pm 34.5663$  mg./dl. and  $159.8750 \pm 45.6647$  mg./dl.

The mean serum IgG, IgA and IgM levels age-matched controls between 6 months-2 years were  $958.4286 \pm 305.5448$  mg./dl.,  $81.0429 \pm 31.3551$  mg./dl. and

158.1571 ± 36.3420 mg./dl. while between 2-6 years were 1117.3846 ± 246.1194 mg./dl, 94.2692 ± 30.1359 mg./dl. and 148.0769 ± 40.0260 mg./dl.

We still do not know much about the causes of FC. The immune mechanism may be involved, but this is still not clearly understood.

Seager et al (1975) and Oloffson et al (1982) reported the probable connection between low serum IgA levels and FC [9,17]. Ariizumi et al (1980) and Bibus & Aarli (1981) reported that 33% of patients with FC had low serum IgA levels and a 57% of epileptic patients who had had FC had low serum IgA levels [10,18].

However Isaacs et al. (1984) reported that 47 FC patients had normal serum IgG, IgA and IgM levels [19].

In our study the mean IgA level of FC patients between 6 months-2 years were 49.7167 ± 27.9807 mg./dl. and the mean level of IgA in the age-matched control groups was 81.0429 ± 31.3551 mg./dl. and the difference between them was statistically significant ( $p < 0.005$ ).

Only 5 of the 20 came for their follow-up examinations after 3 weeks and

serum IgG, IgA and IgM levels were measured in these patients. Four of these 5 patients were on phenobarbital prophylaxis (5 mg./kg./day). The IgA levels of these 4 patients were lower than their first IgA levels but the results were not compared statistically. One patient who did not have phenobarbital prophylaxis had the same serum IgA level as before.

It is well known that some anticonvulsant drugs have immunosuppressive effects. Phenytoin in particular suppresses IgA [19,20]. In one of their studies, Tartara et al. found that patients who were using phenobarbital had low serum IgA levels, while those using phenytoin had normal serum IgA levels [20].

In our study serum IgG and IgM levels were normal in children with FC when compared with the age-matched controls. FC patients between 2-4 years of age also had normal IgA levels when compared with the age-matched controls.

It was concluded that FC under 2 years of age is associated with low serum IgA levels.

## REFERENCES

1. Addy DP . Febrile convulsion. In : Paediatric perspectives on Epilepsy. John Wiley and Sons Ltd, 1985; 73 - 7.
2. Niall V O'Donohoe . Febrile Convulsions . In : Epileptic syndromes in infancy, childhood and adolescence. John Libbey Eurotex, 1985; 34 - 7 .
3. Bell and Mc Cormik . Febrile Seizures. In : Neurologic Infections in Children. WB Saunders 1975, 444 - 7.
4. Erenberg G . Febrile Convulsions; A new look at an old Problem. Cleveland Clinic Quarterly Vol 51, no 2 .
5. Joffe A, Mc Cormik M, De Angelis C . Which children with febrile seizures need lumbar puncture ? Am J Dis Child 1983; 137: 1153 - 6.
6. Lewis HM, Parry JV, Parry RP et al. Role of viruses in febrile convulsions. Arch Dis Child 1979; 54 : 869 - 76 .
7. Nelson K . Febrile Seizures In : The practice of pediatric Neurology. 2nd ed. Mosby 1982; 1070 - 4 .
8. Schmidt D, Isai JJ, Janz D . Febrile seizures in patients with complex partial seizures. Acta Neurol Scand 1985; 72: 68 71 .
9. Seager J, Jamison DL, Wilson J, Hayward AR, Soothill JF . IgA deficiency, epilepsy and phenytoin treatment. Lancet 1975; ii: 632 - 5 .
10. Ariizumi M, Kuromori S, Shiihara H, Eguchi H, Baba K . Febrile c onvulsion and Immunoglobulin Abnormalities. Nihon Univ J Med 1980; 22 : 195 - 201.
11. Fahey JL, Mc Kelvey EM . Quantitative determination of serum immunoglobulins in antibody agar plates. J Immunol 1965; 94 - 84 .
12. Amos JR, Brown FL, Mink OG . Chi-Square Test. In : Statistical Concepts. A Basic program. New York: Harper and Row, 1965; 90 - 2.
13. Takayuki T, Shigeko O . Exogenous causes of seizures in children : A population study. Acta

- Neurol Scand 1985; 71 : 107 - 13 .
14. Verity CM, Butter NR, Colding J . Febrile convulsion in a national cohort follow up from birth. 1-prevalance and recurrence in the first five years of life. Br Med .
15. Viani F, Begni E, Romeo A, Van Lierde A . Infantile febrile status epilepticus : Risk factors and outcome. Develop Med Child Neurol 1987; 29 : 495 - 501.
16. Hauser WA, Annegers JF, Anderson VE, Kurland LT . The risk of seizure disorders among relative of children with febrile convulsions. Neurology 1985; 35 : 1268 - 73 .
17. Olofsson OE, Wigertz A, H Link. Immunoglobulin abnormalities in seizures. Neuropediatrics 1982; 13 : 39 - 41 .
18. Gibus NE, Aarli JA . Immunoglobulin concentration in patients with a history of febrile convulsions prior to the development of epilepsy. Neuropediatrics 1981; 12: 314 - 8 .
19. Isaacs D, Webster ADB, Valman HB . Serum immunoglobulin concentrations in febrile convulsions. Arch Dis Child 1984; 59: 367 - 9 .
20. Tartara A, Verri AP, Nespoli L, Moglia A, Botta MG . Immunological findings in epileptic and febrile convulsion patients before and under treatment. Eur Neurol 1981; 20 : 306 - 11 .