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representation of the palms, soles and mucosae, such as lips and genitalia.

# Declaration of patient consent

The authors certify that they have obtained all appropriate patient consent.

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#### **Conflicts of interest**

There are no conflicts of interest.

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# Association of various risk factors with childhood herpes zoster: A case–control study in Indian patients

Sir,

Recent studies from Indian set up are showing a conspicuous rise in cases of childhood herpes zoster in immunocompetent children.<sup>1</sup> The aim was to study the clinical presentation, complications and the association of various risk factors with childhood zoster by conducting age- and sex-matched case– control study in Indian patients.

The study was ethically approved by Institutional Ethics Committee and patients were recruited from the Department of Dermatology, Pandit B. D. Sharma, PGIMS, Rohtak, Haryana. Guardians of patients and controls gave their voluntary informed consent. Convenience sample size was selected which included 30 patients of age  $\leq 14$  years, who were clinically diagnosed as a case of herpes zoster and had multinucleated giant cells on Tzanck smear. Equal number of age- and sex-matched controls were selected from the healthy children who did not have a history of having the disease. The children in the control group were those who were coming along with other patients to our outpatients' department and were healthy. confirmed by Western blot assay. Complete blood count, chest X-ray and ultrasonography of abdomen were done to rule out any underlying malignancy. Anemia was diagnosed on the basis of both peripheral blood smear and mean corpuscular volume.

Anemia was defined as hemoglobin concentration <11 g/ dL for children between 6 and 59 months; <11.5 g/dL for children between 5 and 11 years and <12 g/dL for children aged 12 years according to the WHO.<sup>2</sup> MCV >100 fL was considered as megaloblastic anemia. Statistical Package for the Social Sciences for Windows version 22.0 was used for statistical analysis. Comparisons among groups were performed using the  $\chi^2$  test. Crude odds ratio was calculated.

Thirty patients of childhood herpes zoster were included in the study [Figure 1]. The demographic and clinical data are tabulated in Table 1 [Figures 2 and 3]. None of our patients developed complications except scarring at the site of the lesions in two patients. All patients were treated with oral acyclovir and symptomatic treatment.

Detailed history and clinical examination was done in all the patients. Patient's sera were tested for antibody to HIV which was

Comparison of various risk factors within the study and control groups is given in Table 1. On evaluating patients'

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# Study Letters

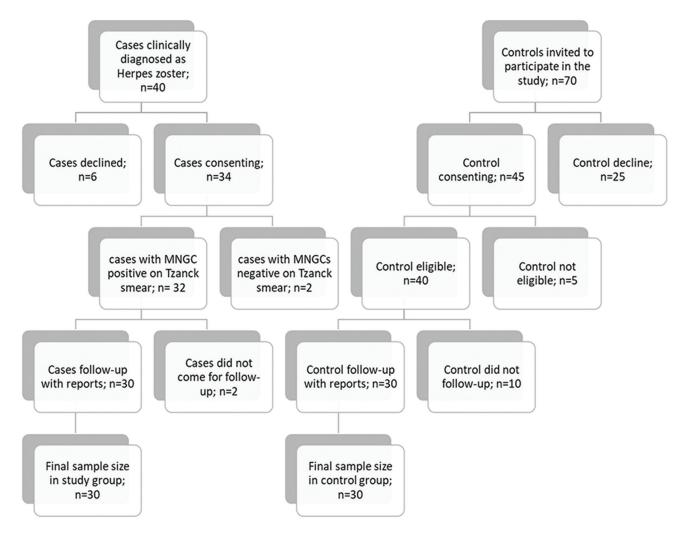


Figure 1: Flowchart of patient recruitment in the study and control groups



Figure 2a: A 12-year-old girl presented with scarring following herpes zoster along T1-T2 dermatome on the right side arm and back

complete blood count, it was found that 21 out of 30 patients had megaloblastic anemia which was significantly associated with occurrence of childhood herpes zoster [Table 2]. Thus,



Figure 2b: A 12-year-old girl presented with scarring following herpes zoster along T1-T2 dermatome on the right side arm and back

megaloblastic anemia had significantly increased odds for occurrence of the condition in childhood (odds ratio: 21; 95% confidence interval: 3.9–112.7).

Table 1: Demographic and clinical data of the study and control patients					
Characteristics	Study patients (n=30)	Control (n=30)	P-value	Odds ratio with confidence interval	
Age (mean±SD)	8.70±4.2 years	8.90±3.7 years	0.9 (insignificant)	-	
Sex (M: F)	M=19, F=11	M=18, F=12	-	-	
Duration of disease (days)	2.5±0.9	-	-	-	
Prodromal symptoms		-	-	-	
Mild burning sensation	3				
Itching	5				
Pain	17				
Itching and pain	2				
Nil	3				
Dermatome involved:		-	-	-	
Thoracic	19				
Lumbar	6				
Head and neck	5				
Contact of chickenpox	Present in six patients	Present in four patients	0.7 (insignificant)	1.6 (0.4-6.5)	
History of chickenpox	Present in eight patients	Present in seven patients	0.766 (insignificant)	1.2 (0.4–3.9)	
Vaccination history	Present in two patients	Present in two patients	1.00 (insignificant)	1 (0.1–7.6)	
History of maternal varicella	Present in two patients $(1^{st}$ patient at the age of 6 years and $2^{nd}$ patient at the age of 5 years)	Present in one patient	1.00 (insignificant)	2.1 (0.18–24.1)	
Evidence of immunosuppression	Present in two patients	Present in none of the patients	0.4 (insignificant)	2.1 (1.6–2.7)	
Presence of megaloblastic anemia	Present in 21 patients	Present in two patients	<0.001 (significant)	21 (3.9–112.7)	



Figure 3: A 1-year-old child presented with herpes zoster of the right T1 dermatome

Vaccination with live-attenuated virus could be one of the reasons for recent rise in childhood herpes zoster according to a few reports.<sup>3</sup> However, our study is in concordance, with those studies which have revealed that varicella vaccination does not increase the incidence of zoster.<sup>4</sup> It is proposed that vaccine strain is attenuated and has less chance of reactivation as compared to wild-type virus. However, the effect of vaccination on occurrence of the disease in childhood cannot be commented on as the trend of vaccination against varicella is not so much prevalent in our country. The previous studies have also shown varicella during early life as an important risk factor.<sup>5</sup> In our study, the difference was statistically insignificant, but a history of chickenpox may be a risk factor as the episode of chickenpox

Table 2: Megaloblastic anemia in the study and control groups

Groups	Megaloblastic anemia present	Megaloblastic anemia absent
Study group	21 Case 1: Hb – 9 g/dl; MCV – 103 fL Case 2: Hb – 9.5 g/dl; MCV – 102 fL Case 3: Hb: 10 g/dl; MCV – 101 fL Case 4: Hb: 8.8 g/dl; MCV – 101 fL Case 5: Hb: 9.8 g/dl; MCV – 102 fL Case 6: Hb: 9.5 g/dl; MCV – 100 fL Case 7: Hb: 8.7 g/dl; MCV – 103 fL Case 8: Hb: 9.4 g/dl; MCV – 102 fL	9
	Case 9: Hb: 9.8 g/dl; MCV - 102 fL Case 10: Hb: 8.8 g/dl; MCV - 103 fL Case 10: Hb: 8.8 g/dl; MCV - 103 fL Case 11: Hb: 8.5 g/dl; MCV - 103 fL Case 12: Hb: 8.2 g/dl; MCV - 103 fL Case 13: Hb: 8.6 g/dl; MCV - 103 fL Case 14: Hb: 9.2 g/dl; MCV - 102 fL Case 15: Hb: 10 g/dl; MCV - 102 fL Case 16: Hb: 9.6 g/dl; MCV - 102 fL Case 17: Hb: 8.9 g/dl; MCV - 102 fL Case 18: Hb: 9 g/dl; MCV - 102 fL Case 19: Hb: 8.5 g/dl; MCV - 104 fL Case 20: Hb: 9 g/dl; MCV - 102 fL Case 21: Hb: 10 g/dl; MCV - 101 fL	
Control group	2	28

is mild in immunocompetent children and may go unnoticed. History of maternal varicella during pregnancy or history of contact with chickenpox patients may also be risk factors.<sup>1</sup> However, in our study, these factors were not significantly associated with the study group when compared with the control group. It may be due to unreliable history given by the patients. Recently, studies have shown that there is no increase in the incidence of malignancy in childhood herpes zoster.<sup>1</sup> A greater number of patients in the study group had immunosuppression with respect to the control group but the difference was statistically insignificant. Therefore, our study is in concordance with the findings of recent studies. However, our study had a very small sample size to conclude these facts.

In our study, we found a strong significant association between megaloblastic anemia and pediatric herpes zoster. The most common cause of megaloblastic anemia is deficiency of vitamin  $B_{12}$  or folic acid. This may be cause of immature immune response leading to low levels of natural killer cells, lymphocytes and cytokines along with virus-specific immunoglobulins which may be the cause of inability to maintain varicella zoster virus latency, leading to appearance of herpes zoster at an early age.<sup>1</sup> However, as our sample size was small, this association cannot be affirmed.

Serology could not be done for detecting antibodies. There may be a possibility of recall and confounding bias. Confounding bias, such as nutritional status of the patients, could have been present. We could not find the underlying cause of megaloblastic anemia.

Despite elaborate literature search, we are unable to find any such study to report the possibility of association of anemia with pediatric herpes zoster. Further prospective populationbased studies with better investigation profile and evaluation of nutritional status of these children are required to confirm our observation.

## Declaration of patient consent

The authors certify that they have obtained all appropriate patient consent.

# Financial support and sponsorship

Nil.

# **Conflicts of interest**

There are no conflicts of interest.

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# Effect of monthly cyclophosphamide pulses on skin sclerosis in systemic sclerosis

### Sir,

Systemic sclerosis is a connective tissue disease affecting the skin, blood vessels and internal organs with significant morbidity and mortality.<sup>1</sup> The reduction in skin sclerosis has a beneficial effect on the quality of life and survival of the patients.<sup>2,3</sup> Only very few controlled clinical trials have been performed for the treatment of skin sclerosis due to the rarity of the disease and its highly variable course.<sup>4</sup>

In the present study, we attempted to study the efficacy of low dose cyclophosphamide for a short duration on skin sclerosis in both limited and diffuse types of systemic sclerosis.

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