Outcome and Clinico-laboratory Profile of pediatric scrub typhus in a tertiary medical college of West Bengal, Eastern India

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Abstract:

Introduction: Scrub typhus is an important cause of acute undifferentiated febrile illness in children which is defined as fever without any evidence of localized infection lasting ≤ 14 days The goal of this study is to provide clinical aspect of the disease in details thus helping to creat a better clinical and laboratory profile of this reemerging disease for clinician to work with.

Methods: it is a prospective observational study done in the department of pediatric medicine NRS medical college between January 2018 to October 2019 with history of acute febrile illness in whom clinical and laboratory features consistant with scrub typhus included in our study Scrub typhus was confirmed by IgM ELISA using INBIOS kit.

Results: A total of 77 cases with median age 8years (range 3months to 12 years) were diagnosed to have scrub typhus with a boys and girls ratio1.4:1. Most case presented during the month of September (34%) followed by August (28%) and October (23%). Fever was the consistant finding (100%) of all cases. Headache(18%), myalgia(!*%), breathlssness(29%), abdominalpain(27%), vomiting(41%), diarrhea(5%), rash(8),

eschar(7%), hepatomegaly(65%), splenomegaly(28%), lymphadenopathy(31%)pedal oedema(10%),altered sensorium,convulsions(21%)etc were other reported symptoms.Among laboratory findings trilineagecellinvolvement, altered liver engymes with hypoalbuminemia ,hyponatremia and increased creatinine level seen in many cases. Scrub encephalitis(21%), MODS(5%),ARDS(2.5%)&AKI(3.8%) were observed complications.Most of cases(87%) discharged after recovery.Case fatality rate was 9%.

Conclusion: Increased awareness of this infection among clinician with background knowledge of endemicity along with reliable rapid diagnostic tool will be the key factor to further reduce the mortality of this deadly disease.

Keywords : Orientiatsutsugamushi; scrub typhus, acute undifferentiated fever, IgM Elisa

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I. Introduction

Scrub typhus is an important cause of acute undifferentiated febrile illness in children which is defined as fever without any evidence of localized infection lasting ≤ 14 days^{1.2}. Orientiatsutsugamushi, the causative agent of this zoonotic disease is an obligatory intracellular bacteria distributed to a part of world known as the "tsutsugamushitriangle" which extends from northern Japan and far-eastern Russia in the north, to northern Australia in the south, and to Pakistan and Afghanistan in the west.³ Of the 29 states in India, 23 have reported the presence of scrub typhus.^{4–7} the infection is transmitted by larval mites or chiggers belonging to the familttrombiculidae. The clinical presentation of scrub typhus ranges from subclinical disease to multiorgan failure and death.⁸ Most patients presents with fever, diffuse lymphadenopathy, myalgia, rash, jaundice, thrombocytopenia, capillary leak syndrome, hepatomegaly, and splenomegaly.11A single painless eschar with a an erythematous rim at the site of chigger bite is seen in 7-68% of cases, and a maculopapular rash is present in 30%, both can be absent.⁹ Severe complications include prominent encephalitis, interstial pneumonia, and ARDS and haemorrhagic features with circulatory collapse. The goal of this study is to provide clinical aspect of the disease in details thus helping to create a better clinical and laboratory profile of this re-emerging disease for clinician to work with.

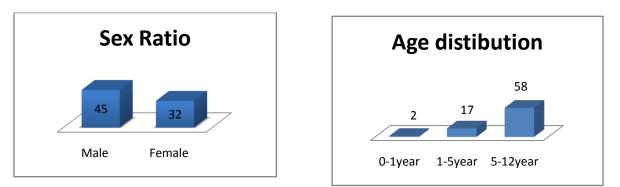
Materials And Methods II.

This was a prospective observational study done in the department of pediatric medicine NRS medical college.All children admitted to pediatric ward between January 2018 to October 2019 with history of acute febrile illness in whom clinical and laboratory features consistant with scrub typhus included in our study. Differential diagnosis such as dengue, malaria, chikungunya, enteric fever, urinary tract infection, respiratory infection, leptospirosis were excludes by history, clinical findings and relevant examinations. Serum electrolytes, Liver function test, renal function test, Widal test, CSF study, chest x-ray and Echocardiogram were done when clinically indicated.

Scrub typhus was confirmed by IgM ELISA using INBIOS kit for scrub typhus which is 90% sensitive and 100% specific. Despite the fact that immunofluorescence test is classically taken as the gold standard for diagnosing rickettsial infections, IgM ELISA is used by many laboratories for its cost, simplicity, and convenience. The collected data were compiled using Microsoft Excel 2010 and then analysed using statistical software Epi Info 7.1.5. The results were expressed in percentages and proportions.

III. Results

A total of 77cases with median age 8years (range 3months to 12 years) were diagnosed to have scrub typhus. Out of 77 children 45 were boys and 32 were girls with a boy and girl ratio 1.4:1. Most case presented during the month of September (34%) followed by August (28%) and October (23%)



Fever was a consistant finding in all scrub cases with 70% having high grade fever. Most of the scrub had fever for more than 7days and minor group (17%) had fever lasting more than typhus (65%) 14days.headache and myalgia (17% each), cough (18%) were other reported clinical features. Breathlessness, abdominal pain, vomiting ,diarroea ,rashes were the found 29%,27%, 41% and 8% cases respectively.21% patients presented with altered sensorium and convulsion.Pedal oedema and ascites found among 10% cases. Multiorgan dysfunction (MODS), ARDS and Acute Kidney Injury(AKI) were observed in 5%, 2.5% and 3.8% cases respectively.

On clinical examination, mild to moderate pallor, severe pallor and lymphadenopathy seen viz. 48%,2% and 31% cases. Eschar which is considered as a pathognomic feature of scrub typhus was seen only 7% cases. Common sites of eschar were groin, buttock followed by axilla and sole. Hepatomegaly (65%) is another common finding followed by splenomegaly (28%).

| Table1: Clinical Manif | estation of Scrub Typhus |
|------------------------|--------------------------|
| | |

| Lusiell' chinese filminestation of Serus Lyphus | | | | | |
|---|-----------|-------------------------------|-----------|--|--|
| Clinical features | Number(%) | Clinical features | Number(%) | | |
| Fever | 77(100%) | Breathlessness | 22(29%) | | |
| Headache | 17(13%) | Lymphadenopathy | 24(31%) | | |
| Myalgia | 17(13%) | Hepatomegaly | 50(65%) | | |
| Rashes | 6(8%) | Spleenomegaly | 21(28% | | |
| Vomiting | 31(41%) | Altered sensorium& convulsion | 16(21%) | | |
| Diarrhoea | 4(5%) | Ascites &pedal oedema | 8(10%) | | |
| Abdominal pain | 21(27%) | Eschar | 5(7%) | | |

| Pallor(Hbgm/dl)(<11gm/dl) | 38(50%) |
|------------------------------|---------|
| Leucocytosis(wbc>11000/cumm) | 27(35%) |
| Leucopenia(wbc<4000/cumm) | 11(14%) |

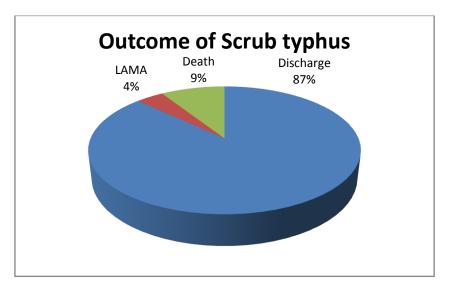
Table2: Laboratory findings

| Leucocytosis(wbc>11000/cumm) | 27(35%) |
|---------------------------------------|----------|
| Leucopenia(wbc<4000/cumm) | 11(14%) |
| Thrombocytopenia(<1.5lacs/cumm) | 61(80%) |
| SGOT/SGPT[>3times of normal (40iu/L)] | 31(41%) |
| Hypoalbuminemia(<2.5gm/dl) | 21(28%) |
| Hyponatremia(<135meq/L) | 12(16%) |
| Serum creatinine (>1mg/dl) | 4(5%) |
| Scrub IgM | 77(100%) |

Among laboratory parameter, anemia and thrombocytopenia found in 50% and 81% children respectively. Leucopenia and leucocytosis seen in 14% and 35% cases. Rest of the cases leucocyte count was normal. Elivated liver engymes(SGOT/SGPT) observed in 41% children. Hypoalbuminemia, hyponatremia and elevated creatinine level found in 28%, 16% &5% cases respectively. All cases were positive for IgM scrub typhus.

Outcome:

Among 77 cases admitted during study period, 67 cases discharged after recovery, only 3 patients left against medical advice. Inspite of our best effort 7 patients could not survive.



IV. Discussion

Present study shows increased incidence of scrub typhus during August to October which coincide with wet season suitable for growth of vegetation and trombiculide mite.

The incidence of scrub typhus is more among males like in previous studies. It may be due to predominance of male children over females as they display greater degree of activity and outdoor exposure.

In our study children of school going age (75%) are mostly affected. It may be due to more exposure to infected chigger. During rainy season chiggers may possibly change their habitat, so disease among infants who are predominantly indoor can be explained.¹⁰⁻¹²

High grade fever was the consistant clinical finding in the present study. This fever symptom is in congruence with most of the previous studies.^{11, 13.} The low incidence of prolonged fever lasting more than two weeks may be due to indiscriminate use of available antibiotics at locality.

Presence of rash is common in spotted fever, extremely rare in scrub typhus. Though rash is considered as hallmark of rickettsial disease ,it was found only 8% cases in our study as compared with22% patients from Vellore,9% patients from Chandigarh and 20% patient from Himalayan region of North India, 30% patients in Thailand, and 55.7% patients fromJeju Islands, South Korea. ¹⁴⁻²³. Rash is initially maculopapular, present in early stage of disease. Low incidence of rash may be due to late seeking of medical advice of these patients.

The pathognomonic eschar was present in 7% patients, a figure corroborative with previous study from the Himalayan region (9.5% patients), but lesser than that reported from south India (43.5%) and Jeju Island in South Korea (75.8%).¹⁷⁻¹⁹ This variation in the presence of eschar may be explained by the geographic distribution of different strains of the organism.

Gastrointestinal manifestations like abdominal pain (27%), vomiting (41%), and diarrhea (5%)occurred in 40% of patients in this study. Vomiting and diarrhea occurred in 54% and 30% cases in a study from south India.^{14,25}. This highlights the fact that febrile patients of scrub typhus can also present with prominent gastrointestinal symptoms. Lymphadenopathy was found in 31% patients in our study which is lesser from South India study(96%)26 and higher than Vellore study(24%)²⁷.

Predominant hepatomegaly was observed in 65% cases as like previous studies.²⁸

Among the laboratory findings thrombocytopenia was observed in 80% cases in our study which is in congruence with other studies like Varghese etal (79%), Sharma et al.(90%) and Sinha et al.(85%).14,24,16Liver engymes (SGOT/SGPT) showed threefold increase in titre among 41% patients. Impaired renal function in the form of elevated creatinine level was noted in 4% cases. This finding is

corroborative with Philomela et al study. ²⁷This research documented hypoalbuminemia as a significant finding (28%) as compared to other study.

The common disease complication in our study were scrub encephalitis (21%),MODS(5%),ARDS(2.5%) and AKI(3.8%).A study on South India on adult population reported MODS 34%14 .ARDS occurred in 25% and Acute kidney injury was noted in 32% in Sharma et al study.²⁴

In our study, the case fatality rate is 9% as compared to previous studies that has ranged 1.2% to as high as 46.3 depending on the complication.²⁹⁻³²

V. Conclusion

With the changing epidemiology Scrub typhus is now a commonest cause of acute febrile illness in children with significant mortality. Mortality is significant in patients with multiorgan dysfunctions.

When a patient presents with fever and elevated liver enzymes with or without the presence of eschar, scrub typhus considered as differential diagnosis and an empirical therapy with doxycycline should be started if there is high index of suspicion. However to further reduce the mortality of this deadly disease, increased awareness of this infection among clinician with background knowledge of endemicity along with reliable rapid diagnostic tool will be the key factor.

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References

- Palanivel S, Nedunchelian K, Poovazhagi V, Raghunadan R, Ramachandran P. Clinical profile of scrub typhus in children. Indian J Pediatr. 2012;79(11):1459-62.
- [2]. Jung H-C, Chon S-B, Oh WS, Lee D-H, Lee H-J. Etiologies of acute undifferentiated fever and clinical prediction of scrub typhus in a non-tropical endemic area. Am J Trop Med Hyg. 2015;92(2):256-61.
- [3]. 3.SharmaParag, KakkarRakesh, Kaore N. Shilpa et al. Geographical distribution, effect of Season and life cycle of scrub Typhus. JK Science 2010; 12:63-65.
- [4]. Kumar K, Saxena VK, Thomas TG, Lal S, 2004. Outbreak investigation of scrub Typhus in Himachal Pradesh (India). J Commun Dis 36: 277–283.
- [5]. Vivekanandan M, Mani A, Priya YS, Singh AP, Jayakumar S, Purty S, 2010. Outbreak of scrub typhus in Pondicherry. J Assoc Physicians India 58: 24–28.
- [6]. Khan SA, Dutta P, Khan AM, Topno R, Borah J, Chowdhury P, Mahanta J, 2012. Re-emergence of scrub typhus in northeast India.Int J Infect Dis 16: e889–e890.
- [7]. Sethi S, Prasad A, Biswal M, Hallur VK, Mewara A, Gupta N, Galhotra S, Singh G, Sharma K, 2014. Outbreak of scrub typhus in north India: a re-emerging epidemic. Trop Doct 44: 156–159.
- [8]. Rathi NB, Rathi AN, Goodman MH, Aghai ZH, 2011. Rickettsial diseases in central India: proposed clinical scoring system for early detection of spotted fever. Indian Pediatr 48: 867–872.
- [9]. kliegman, Stanton, StGeme, Schor 2016. Nelson Textbook of Pediatrics, 20e. 1504
- [10]. Kumar Bhat N, Dhar M, Mittal G, Shirazi N, Rawat A, PrakashKalra B, et al. Scrub typhus in children at a tertiary hospital in north India: clinical profile and complications. Iran J Pediatr. 2014;24(4):387-92.
- [11]. Dass R, Deka NM, Duwarah SG, Barman H, Hoque R, Mili D, et al. Characteristics of Pediatric scrub typhus during an outbreak in the North Eastern region of India: peculiarities in clinical presentation, laboratory findings and complications. Indian J Pediatr. 2011;78(11):1365-70.
- [12]. Sharma PSR, Kakkar, R Kaore SN, Yadav VK. Geographical distribution, effect of season and life cycle of scrub typhus. JK Sci. 2010;12(2):63-4.
- [13]. Basu S, Saha A, Sarkar S, Sinha MK, Das MK, Datta R, et al. Clinical Profile and Therapeutic Response of Scrub Typhus in Children: A Recent Trend from Eastern India. J Trop Pediatr. 2018;23.
- [14]. 23. Varghese GM, Trowbridge P, Janardhanan J, Thomas K, Peter JV, Mathews P, Abraham OC, Kavitha ML, 2014. Clinical profile and improving mortality trend of scrub typhus in south India. Int J Infect Dis 23: 39–43.
- [15]. Kumar V, Kumar V, Yadav AK, Iyengar S, Bhalla A, Sharma N, Aggarwal R, Jain S, Jha V, 2014. Scrub typhus is an underrecognized cause of acute febrile illness with acute kidney injury in India. PLoSNegl Trop Dis 8: e2605.
- [16]. Sinha P, Gupta S, Dawra R, Rijhawan P, 2014. Recent outbreak of scrub typhus in north western part of India. Indian J Med Microbiol 32: 247–250.
- [17]. Griffith M, Peter JV, Karthik G, Ramakrishna K, Prakash JA, Kalki RC, Varghese GM, Chrispal A, Pichamuthu K, Iyyadurai R, Abraham OC, 2014. Profile of organ dysfunction and predictors of mortality in severe scrub typhus infection requiring intensive care admission. Indian J Crit Care Med 18: 497–502.
- [18]. Jang MO, Kim JE, Kim UJ, Ahn JH, Kang SJ, Jang HC, Jung SI, Park KH, 2014. Differences in the clinical presentation and the frequency of complications between elderly and non-elderly scrub typhus patients. Arch GerontolGeriatr 58: 196–200.
- [19]. Mahajan SK, Rolain JM, Kashyap R, Bakshi D, Sharma V, Prasher BS, Pal LS, Raoult D, 2006. Scrub typhus in Himalayas. Emerg Infect Dis 12: 1590–1592.
- [20]. Yoo JR, Heo ST, Koh YS, Kim S, Kim S, 2014. Unusual genotypic distribution of Orientiatsutsugamushi strains causing human infections on Jeju Island. Am J Trop Med Hyg 90: 507–510.
- [21]. Nadjm B, Thuy PT, Trang VD, Dang Ha L, Kinh NV, Wertheim HF, 2014. Scrub typhus in the northern provinces of Vietnam: an observational study of admissions to a national referral hospital. Trans R Soc Trop Med Hyg 108: 739–740.
- [22]. Sirisanthana V, Puthanakit T, Sirisanthana T, 2003. Epidemiologic, clinical and laboratory features of scrub typhus in thirty Thai children. Pediatr Infect Dis J 22: 341–345.
- [23]. Mathai E, Rolain JM, Verghese GM, Abraham OC, Mathai D, Mathai M, Raoult D, 2003. Outbreak of scrub typhus in southern India during the cooler months. Ann N Y AcadSci 990: 359–364.
- [24]. Navneet Sharma, M Biswal, A Kumar, K Zaman, S Jain, and A Bhalla, 2016. Scrub Typhus in a Tertiary Care Hospital in North India. Am. J. Trop. Med. Hyg., 95(2), 2016, pp. 447–451

- [25]. Sethi S, Prasad A, Biswal M, Hallur VK, Mewara A, Gupta N, Galhotra S, Singh G, Sharma K, 2014. Outbreak of scrub typhus in north India: a re-emerging epidemic. Trop Doct 44: 156–159.
- [26]. Lakshmanan S, Sagayaraj BM, Sujatha B, Vasudevan LD. Clinical and laboratory profile of pediatric scrub typhus in a tertiary care teaching hospital in southern India. Int J ContempPediatr 2018;5:2092-7.
- [27]. James Philomena, M Rangaswami, Parthasarathy Prathiba,2016. A study on demographic, clinical profile and outcome of scrub typhus: International Journal of Advances in Medicine . Aug;3(3):586-590
- [28]. Thomas R, Puranik P, Kalal B, Britto C, Kamalesh S, Rego S, et al. Five-year analysis of Rickettsial fevers in children in South India: Clinical manifestations and complications. J Infect DevCtries. 2016;10(6):657-61.
- [29]. Nadjm B, Thuy PT, Trang VD, Dang Ha L, Kinh NV, Wertheim HF, 2014. Scrub typhus in the northern provinces of Vietnam: an observational study of admissions to a national referral hospital. Trans R Soc Trop Med Hyg 108: 739–740.
- [30]. Sirisanthana V, Puthanakit T, Sirisanthana T, 2003. Epidemiologic, clinical and laboratory features of scrub typhus in thirty Thai children. Pediatr Infect Dis J 22: 341–345.
- [31]. Mathai E, Rolain JM, Verghese GM, Abraham OC, Mathai D, Mathai M, Raoult D, 2003. Outbreak of scrub typhus in southern India during the cooler months. Ann N Y AcadSci 990: 359–364.
- [32]. Kumar Bhat N, Dhar M, Mittal G, Shirazi N, Rawat A, PrakashKalra B, Chandar V, Ahmad S, 2014. Scrub typhus in children at a tertiary hospital in north India: clinical profile and complications. Iran J Pediatr 24: 387–392.

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