

EPIDEMIOLOGICAL AND CLINICAL PROFILE OF STRONGYLOIDIASIS - EXPERIENCE FROM A TERTIARY CARE CENTRE

RATNA HARIKA DUSI^{1*}, SUBBARAYUDU BODA², NITIN MOHAN¹, RAJYALAKSHMI CHEPURU³,
JYOTHI PADMAJA INDUGULA¹

¹Department of Microbiology, GITAM Institute of Medical Sciences and Research, GITAM Deemed to be University, Rushikonda, Visakhapatnam, Andhra Pradesh, India. ²Department of General Medicine, NRI Institute of Medical Sciences, Visakhapatnam. ³Department of Community Medicine, GITAM Institute of Medical Sciences and Research, GITAM Deemed to be University, Rushikonda, Visakhapatnam, Andhra Pradesh, India. Email: harika.dusi@gmail.com

Received: 04 July 2022, Revised and Accepted: 20 August 2022

ABSTRACT

Objectives: The objective of the study was to study the clinical presentations, predisposing factors, and underlying conditions associated with Strongyloidiasis.

Methods: A prospective observational study was conducted from 2018 to 2021 on patients who presented with medical complaints in a tertiary care hospital, and 19 were diagnosed with strongyloidiasis by stool wet mount examination. Other relevant details were collected to analyze the risk factors.

Results: A total of 19 cases were found positive for strongyloidiasis. Males 13 (68.4%) were more and females 6 (31.6%), and most of them were above 50 years age group (73.7%). Among the cases, respiratory symptoms (42.1%) were predominantly observed, followed by gastrointestinal (31.6%). Multiple predisposing factors such as chronic obstructive pulmonary disease, corticosteroid usage, TB, diabetes, alcohol, and asthma have been identified in strongyloidiasis cases. Peripheral eosinophilia is a frequent finding in the complete blood picture.

Conclusions: Strongyloidiasis should be strongly suspected in every immune compromised patient presenting with gastrointestinal, respiratory manifestations, or peripheral eosinophilia, and asymptomatic immune competent patients with comorbid conditions.

Keywords: *Strongyloides stercoralis*, Epidemiology, Neglected disease, Parasite, Nematode, Helminth.

© 2022 The Authors. Published by Innovare Academic Sciences Pvt Ltd. This is an open access article under the CC BY license (<http://creativecommons.org/licenses/by/4.0/>) DOI: <http://dx.doi.org/10.22159/ajpcr.2022v15i11.45733>. Journal homepage: <https://innovareacademics.in/journals/index.php/ajpcr>

INTRODUCTION

Strongyloidiasis is a worldwide disease caused by parasitic nematodes of the genus *Strongyloides*, with high frequency in the Caribbean, Latin America, Europe, Asia, and Sub-Saharan Africa [1]. Although it is underdiagnosed due to low parasite load and uncertain clinical symptoms, around 370 million people worldwide are found to be infected [2]. *Strongyloides stercoralis* is unique among the nematodes infectious for humans in that rhabditiform larva passed out in the feces can either become infective filariform larvae or a free-living generation of worms which, in turn, give rise to infective larvae or can persist in the host by autoinfection. This so-called heterogenic development process as an amplification mechanism allows for increased numbers of infective larvae in the external environment. Infection results when these free-living *S. stercoralis* in the soil enter humans through intact skin penetration (e.g., occupational exposure to soil and walking barefoot) [3]. Various factors can influence the prevalence of *S. stercoralis* such as the immune status of the host, socioeconomic status, sanitary and hygiene practices of individuals, associated comorbid conditions, levels of endogenous corticoid, alcohol intake, and the level of heat and humidity, which affects parasite development in soil. The prevalence of strongyloidiasis in a population can be divided into three categories: sporadic (<1%), endemic (1–5%), and hyperendemic (>5%) [4]. The clinical manifestations in people infected with *Strongyloides stercoralis* range from asymptomatic to a potentially fatal disseminated infection [5]. Chronic strongyloidiasis can progress to disseminated illness with a 70% fatality rate. When the immune system is compromised, *S. stercoralis* hyperinfection occurs, speeding up the pace of autoinfection. Disseminated strongyloidiasis occurs when larvae spread to various organs, including the liver, lungs, and central nervous

system, due to hyperinfection. Gram-negative sepsis is prevalent, caused by enteric bacteria translocation when significant numbers of larvae migrate through the intestinal wall [6]. Strongyloidiasis is a major medical concern since it can lead to death in people who are immunosuppressed or immunocompromised due to steroid use, coinfection with human T-lymphotropic virus-1, HIV, or alcoholism [4,7]. Strongyloidiasis generally presents as diffuse, nonspecific gastrointestinal (mild abdominal pain, intermittent, or persistent diarrhea), respiratory (cough, wheezing and asthma, and chronic bronchitis), dermatologic symptoms (pruritus, rash), or systemic symptoms (weight loss and cachexia) leading to delayed treatment, medical complications, septicemia, and finally, death [3,8]. Misdiagnoses of strongyloidiasis in patients often lead to expensive, nonspecific, invasive diagnostic techniques, including endoscopy, barium swallow, cancer biopsies, chest X-rays, and computed tomography (CT) scans. Therefore, a high index of suspicion and thorough clinical and laboratory workup is required to diagnose and treat every case to prevent chronic and life-threatening forms. The present work was performed to study the various clinical presentations, predisposing factors, and underlying conditions associated with strongyloidiasis.

MATERIALS AND METHODS

A prospective observational study was done in a tertiary care hospital in Coastal Andhra Pradesh. The study was done from 2018 to 2021 on 1801 patients who underwent stool examination (wet mount preparation) for parasites, and 19 were diagnosed with strongyloidiasis. An informed written consent form was taken from all the study participants. Larvae were identified as strongyloides rhabditiform (~300 µm in length, short buccal cavity and double bulb esophagus

with a prominent genital primordium) or filariform (~500 µm in size, long esophagus, and a notched tail) stages of differentiating them from hookworm larvae [9,10]. Data were collected for details such as age, gender, presenting complaints, personal history, occupation, TLC, and absolute eosinophil count.

RESULTS

A series of 19 cases of strongyloidiasis who presented with medical complaints were analyzed in the study. Thirteen males and six females aged 30–70 years with a mean age of 57.61±11.45 years. Most of the patients belong to the >50 years age group (Table 1). Most of them presented with (Table 2) respiratory symptoms such as cough, and shortness of breath, followed by gastrointestinal symptoms such as pain abdomen, diarrhea, vomiting, and anorexia. Few others were diagnosed in patients who presented with diverse symptoms.

In the present study, multiple predisposing and risk factors (Table 3) among the patients, such as alcohol use, diabetes, asthma, retroviral infection, and steroid use (for arthritis, myasthenia gravis, asthma, and chronic obstructive pulmonary disease [COPD]), led to immunosuppression have been identified, the frequent ones observed were the usage of steroids for acute exacerbation of COPD. Peripheral eosinophilia is frequent in complete blood pictures (Table 4) and Table 5 shows the conditions associated with strongyloidiasis in our cases and the % age of eosinophil count. The rhabditiform larvae of *S. stercoralis* were identified in an unstained wet mount of stool at 10× and 40× magnification depicted in Fig. 1a and b.

DISCUSSION

Most of the patients in this study were over the age of 50 years, and the majority took corticosteroids. Eosinophilia is more common in persons infected with *S. stercoralis* than in other parasite infections [11]. This condition is induced by parthenogenetic females residing in the intestinal submucosa rather than the lumen, resulting in a stronger

eosinophilic response. The eosinophil-dependent mechanisms are also engaged in the death of *Strongyloides* by filarial larvae [12,13]. As a result, the absence of eosinophils is a poor predictor of *S. stercoralis* infection, especially in immunocompromised patients. In healthy adults, eosinophils make up only 2–5% of peripheral leukocytes. The eosinophil fraction in the blood increases with active helminthic infections like *Strongyloides*, ranging from 25–35% in acute cases to 6–8% in chronic cases. However, eosinophil counts are reduced in immunosuppressive situations like corticosteroid therapy, and their absence in patients signals a poor prognosis [14]. Most patients were immune compromised, on corticosteroid medication, or had *Strongyloidiasis* for various causes. Under rare circumstances, strongyloidiasis infection can develop into a lethal fulminant disease, compromising the host's immunological system. [10,15]. As steroids predispose to immunosuppression in the host, female worms of *Strongyloides* produce more eggs in the host body. This further facilitates worm growth and development in the host [16]. It is an important helminthic disease due to its peculiar auto-infective cycles and risk of hyperinfection syndrome in immunocompromised patients. However, strongyloidiasis is not always associated with immunocompromised states and occurs in an immunocompetent host, mostly remaining asymptomatic [17]. It stimulates the hypothalamic-pituitary-adrenal (HPA) axis to produce excessive endogenous cortisol levels that further aggravate the immunosuppression by inhibiting the Th2 response. Cortisol metabolites resemble the parasite hormone ecdysone, which regulates the fertility of parthenogenetic *Strongyloides*, causing the transformation of rhabditiform larva to filariform larva and, together with the presence of a steroid receptor on *S. stercoralis* further exacerbate the condition [4]. Therefore, alcoholics were at higher

Table 1: Age-wise distribution of cases

Age	Number and %age of cases
<30	1 (5.2%)
31–40	1 (5.2%)
41–50	3 (15.8%)
>50	14 (73.7%)

Table 2: Distribution of cases according to clinical manifestation at the time of presentation to the hospital

Presenting complaints	Number and %age of cases
Respiratory symptoms	8 (42.1%)
Gastrointestinal symptoms	6 (31.6%)
Co-incident finding	5 (26.3%)

Table 3: Distribution of predisposing factors among the cases

Predisposing factor	Number of cases
Corticosteroid usage for:	
COPD	7
Asthma	1
Inflammatory arthritis	2
Myasthenia gravis	1
Diabetes	6
Alcohol	5
Old tuberculosis	2
HIV	1
Hepatitis B	1
Malignancy	1

COPD: Chronic obstructive pulmonary disease

Table 4: Distribution of absolute Eosinophil count among Strongyloidiasis patients

Absolute Eosinophil count	Number and %age of cases
<500/mm ³	6 (31.6%)
>500–1500/mm ³	5 (26.3%)
1500–5000/mm ³	5 (26.3%)
>5000/mm ³	3 (15.8%)

Table 5: Conditions associated with Strongyloidiasis in our cases and % age of Eosinophil count

Diagnosis	Percentage of Eosinophil count
Anaemia with COPD	52
Rheumatoid arthritis with ILD with old	49
Pulmonary Koch's	
Gastroduodenal ulcer, chronic hepatitis-B, UTI	22
COPD, CAD, seronegative arthritis	20
Anemia with hypothyroidism with T2DM	17
Acute exacerbation of COPD	16
COPD with pneumonia	10
Carcinoma of stomach	5
Cerebrovascular accident, hypertension, cellulitis, type II diabetes	5
CLD with varicella with T2DM	5
Ulcerative colitis with T2DM	4
Hypertension with anemia with pneumonia	3
Acute exacerbation of COPD with Cor	3
Pulmonale	
Myasthenia Gravis, Thymoma, Acute Gastroenteritis	2
Pneumonia with T2DM with HIV	2
COPD with LRTI, Mild Anemia	1
Old Pulmonary Koch's with COPD	1
Bronchial Asthma with Anemia	1
Anemia with T2DM	1

COPD: Chronic obstructive pulmonary disease, T2DM: Type II diabetes mellitus, HIV: Human immunodeficiency virus, ILD: Interstitial lung disease

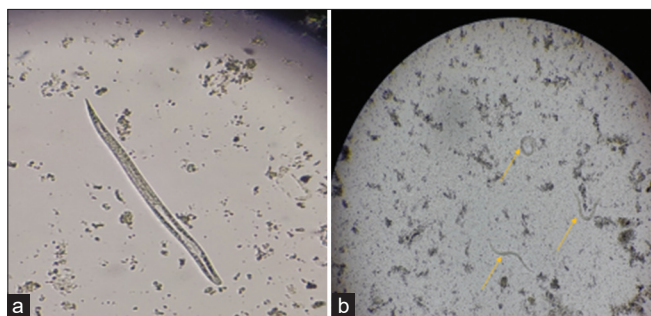


Fig. 1: *Strongyloides stercoralis*. (a) A single rhabditiform larva of *S. stercoralis* in unstained wet mount of stool (40× magnification). (b) A single rhabditiform larva of *S. stercoralis* in unstained wet mount of stool (10× magnification)

risk of autoinfection due to invasion of the intestinal mucosa by the helminth larvae and, if untreated, would progress to hyperinfection and dissemination syndrome [18]. In most case scenarios, gastrointestinal and pulmonary symptoms exacerbate because of immunocompromised states, leading to increased detection of the number of larvae in stool and sputum. Patients may present with increasing cough, dyspnea, or wheezing mimicking COPD exacerbation [19,20]. In the present study, all the cases were treated with Ivermectin, which is currently the gold standard for treating Strongyloidiasis, affecting both adults and larvae [21]. Around 42% of the cases presented with respiratory complaints. In patients presenting with COPD exacerbation with peripheral eosinophilia, strongyloidiasis should be ruled out before starting treatment with steroids, and stool specimens should be sent to examine eggs and larvae in current clinical settings. According to a systematic review conducted [22] on published Strongyloides infection rates and considering the sensitivity of the used diagnostic methods, the prevalence of Strongyloides in Africa varied from 0.1% in the Central African Republic to up to 91.8% in Gabon. In South America and Central America, Haiti reported a prevalence of 1.0%, while in Peru, the infection rate was as high as 75.3%. Several countries reported infection rates within a comparably small range in South-East Asia. The infection rate in Cambodia, Thailand, and Lao PDR is 17.5%, 23.7%, and 26.2%. According to community-based surveys, the infection rate in India was 6.6%, and hospital-based surveys were 11.2% [22]. The authors observed the infection rate in the present study as 1.05%. Most of the information available over the internet regarding the parasite *S. stercoralis* was either discussed with other soil-transmitted helminths [23-25] or as case reports [26,27]. Very few studies across the globe have focused exclusively on *S. stercoralis* epidemiology and risk factors for acquisition [3,22], infection rates [22,28], and clinical presentation [10,18,29]. The present study includes the rate, clinical presentations, predisposing factors, and underlying conditions associated with Strongyloidiasis in our clinical settings.

CONCLUSION

Our findings show a high infection prevalence rate in risk groups. Strongyloidiasis should be strongly suspected in every immune compromised patient presenting with gastrointestinal, respiratory manifestations, peripheral eosinophilia, and asymptomatic immune competent patients with comorbid conditions.

ACKNOWLEDGMENT

AUTHORS' CONTRIBUTION

Ratna Harika Dusi: Contributed conceptual design, literature collection, data collection, data analysis, and drafted the manuscript. Subbarayudu Boda: Contributed conceptual design, guided the work, literature collection and drafted the manuscript. Nitin Mohan collected the literature, data collection, data analysis and corrected the manuscript.

Rajyalakshmi C: Contributed to the data collection, data analysis and corrected the manuscript. I. Jyothi Padmaja: Guided the work, literature collection and corrected the manuscript.

CONFLICTS OF INTERESTS

The authors declared, "No conflict of interest."

AUTHORS' FUNDING

The work was not supported by any kind of funds.

REFERENCES

1. Puthiyakunnon S, Boddu S, Li Y, Zhou X, Wang C, Li J, et al. Strongyloidiasis--an insight into its global prevalence and management. PLOS Negl Trop Dis 2014;8:e3018.
2. Kaminsky RL, Reyes-García SZ, Zambrano LI. Unsuspected *Strongyloides stercoralis* infection in hospital patients with comorbidity in need of proper management. BMC Infect Dis 2016;16:98. doi: 10.1186/s12879-016-1424-3, PMID 26923091
3. Keiser PB, Nutman TB. *Strongyloides stercoralis* in the immunocompromised population. Clin Microbiol Rev 2004;17:208-17. doi: 10.1128/CMR.17.1.208-217.2004, PMID 14726461
4. Teixeira MC, Pacheco FT, Souza JN, Silva ML, Inês EJ, Soares NM. *Strongyloides stercoralis* infection in alcoholic patients. BioMed Res Int 2016;2016:4872473. doi: 10.1155/2016/4872473, PMID 28105424
5. Rajamanickam A, Munisankar S, Bhootra Y, Dolla CK, Nutman TB, Babu S. Elevated systemic levels of eosinophil, neutrophil, and mast cell granular proteins in *Strongyloides stercoralis* infection that diminish following treatment. Front Immunol 2018;9:207. doi: 10.3389/fimmu.2018.00207, PMID 29479356
6. Yates J. Parasitic infections: Do not neglect Strongyloidiasis. Am Fam Physician 2021;104:224-5. PMID 34523890
7. Olsen A, van Lieshout L, Marti H, Polderman T, Polman K, Steinmann P, et al. Strongyloidiasis--the most neglected of the neglected tropical diseases? Trans R Soc Trop Med Hyg 2009;103:967-72. doi: 10.1016/j.trstmh.2009.02.013, PMID 19328508
8. Nutman TB. Human infection with *Strongyloides stercoralis* and other related *Strongyloides species*. Parasitology 2017;144:263-73. doi: 10.1017/S0031182016000834, PMID 27181117
9. Chordia P, Christopher S, Abraham OC, Muliyl J, Kang G, Ajjampur SS. Risk factors for acquiring *Strongyloides stercoralis* infection among patients attending a tertiary hospital in south India. Indian J Med Microbiol 2011;29:147-51. doi: 10.4103/0255-0857.81797, PMID 21654109
10. Nagpal S, Oberoi A. *Strongyloides stercoralis*-an underdiagnosed parasitic infection?-a study from a tertiary care hospital in North India. J Evol Med Dent Sci 2018;7:1468.
11. Khanna V, Tilak K, Mukhopadhyay C, Khanna R. Significance of diagnosing parasitic infestation in evaluation of unexplained eosinophilia. J Clin Diagn Res 2015;9:DC22-4. doi: 10.7860/JCDR/2015/12222.6259, PMID 26393130
12. Negrão-Corrêa D. Importance of immunoglobulin E (IgE) in the protective mechanism against gastrointestinal nematode infection: Looking at the intestinal mucosae. Rev Inst Med Trop Sao Paulo 2001;43:291-9. doi: 10.1590/s0036-46652001000500011, PMID 11696854
13. Maruyama H, Yabu Y, Yoshida A, Nawa Y, Ohta N. A role of mast cell glycosaminoglycans for the immunological expulsion of intestinal nematode, *Strongyloides venezuelensis*. J Immunol 2000;164:3749-54. doi: 10.4049/jimmunol.164.7.3749, PMID 10725734
14. Dawson-Hahn EE, Greenberg SL, Domachowski JB, Olson BG. Eosinophilia and the seroprevalence of schistosomiasis and strongyloidiasis in newly arrived pediatric refugees: An examination of centers for disease control and prevention screening guidelines. J Pediatr 2010;156:1016-8.e1. doi: 10.1016/j.jpeds.2010.02.043, PMID 20400098
15. Spencer JV, Lockridge KM, Barry PA, Lin G, Tsang M, Penfold ME, et al. Potent immunosuppressive activities of *Cytomegalovirus*-encoded interleukin-10. J Virol 2002;76:1285-92. doi: 10.1128/jvi.76.3.1285-1292.2002, PMID 11773404
16. Hunter CJ, Petrosyan M, Asch M. Dissemination of *Strongyloides stercoralis* in a patient with systemic lupus erythematosus after initiation of albendazole: A case report. J Med Case Rep 2008;2:156. doi: 10.1186/1752-1947-2-156, PMID 18479527
17. Tiwari S, Rautaraya B, Tripathy KP. Hyperinfection of *Strongyloides*

- stercoralis* in an immunocompetent patient. Trop Parasitol 2012;2:135-7. doi: 10.4103/2229-5070.105182, PMID 23767024
18. Marcos LA, Terashima A, Canales M, Gotuzzo E. Update on Strongyloidiasis in the immunocompromised host. Curr Infect Dis Rep 2011;13:35-46. doi: 10.1007/s11908-010-0150-z, PMID 21308453
 19. Romero MD, Martínez MD, Pérez MA, Cuesta AA, Sánchez AC, Martínez JH. *Strongyloides stercoralis* as an unusual cause of COPD exacerbation. Arch Bronconeumol 2008;44:451-3. doi: 10.1016/s1579-2129(08)60079-8, PMID 18775258
 20. Liu HC, Hsu JY, Chang KM. *Strongyloides stercoralis* hyperinfection presenting with symptoms mimicking acute exacerbation of chronic obstructive pulmonary disease. J Chin Med Assoc 2009;72:442-5. doi: 10.1016/S1726-4901(09)70403-4, PMID 19687002
 21. Henriquez-Camacho C, Gotuzzo E, Echevarria J, White AC Jr., Terashima A, Samalvides F, et al. Ivermectin versus albendazole or thiabendazole for *Strongyloides stercoralis* infection. Cochrane Database Syst Rev 2016;2016:CD007745. doi: 10.1002/14651858.CD007745.pub3, PMID 26778150
 22. Schär F, Trostorf U, Giardina F, Khieu V, Muth S, Marti H, et al. *Strongyloides stercoralis*: Global distribution and risk factors. PLOS Negl Trop Dis. 2013;7:e2288. doi: 10.1371/journal.pntd.0002288, PMID 23875033
 23. Silver ZA, Kaliappan SP, Samuel P, Venugopal S, Kang G, Sarkar R, et al. Geographical distribution of soil transmitted helminths and the effects of community type in South Asia and South East Asia-a systematic review. PLOS Negl Trop Dis 2018;12:e0006153. doi: 10.1371/journal.pntd.0006153, PMID 29346440
 24. Salam N, Azam S. Prevalence and distribution of soil-transmitted helminth infections in India. BMC Public Health 2017;17:201. doi: 10.1186/s12889-017-4113-2, PMID 28209148
 25. Devi U, Borkakoty B, Mahanta J. Strongyloidiasis in Assam, India: A community-based study. Trop Parasitol 2011;1:30-2. doi: 10.4103/2229-5070.72110, PMID 23508997
 26. Pradhan G, Behera P, Panigrahi MK, Bhuniya S, Mohapatra PR, Turuk J, et al. Pulmonary strongyloidiasis masquerading as exacerbation of chronic obstructive pulmonary disease. Tuberc Respir Dis (Seoul) 2016;79:307-11. doi: 10.4046/trd.2016.79.4.307, PMID 27790284
 27. Reddy PR, Thomas SM, Rajalakshmi A, Vijayan D, Raman M. A rare case of *Strongyloides* hyperinfection from hypogammaglobulinemia. Indian J Crit Care Med 2017;21:466-8. doi: 10.4103/ijccm.IJCCM_139_17, PMID 28808370
 28. Beknazarova M, Whiley H, Ross K. Strongyloidiasis: A disease of socioeconomic disadvantage. Int J Environ Res Public Health 2016;13:517. doi: 10.3390/ijerph13050517, PMID 27213420
 29. Paul M, Meena S, Gupta P, Jha S, Rekha US, Kumar VP. Clinico-epidemiological spectrum of strongyloidiasis in India: Review of 166 cases. J Family Med Prim Care 2020;9:485-91. doi: 10.4103/jfmpc.jfmpc_1182_19, PMID 32318369