

Case Study

Title: Strongyloides hyperinfection in renal transplant - diagnosis by endoscopic intestinal biopsy

Abstract:

Introduction:

Intestinal parasites continue to be a significant health problem in renal transplantation patients. Strongyloides infection is unique that it excretes larvae and can continue life cycle by auto infective cycle also.

Presentation of the case: We present a 26 years old man presented with acute cellular rejection after three months of kidney transplantation. Before transplantation stool of both recipient and donor was negative. He received three doses of intravenous methylprednisolone. After one month he presented with severe epigastric pain and vomiting. On examination he was malnourished, dehydrated and lost two kilograms of weight over one month. His serum albumin was 2.9 mg/dL. pain did not subside with proton pump inhibitors. Stool examination was negative for parasites. As index of suspicion for parasites was high, upper GI endoscopy was done; it showed multiple ulcers in duodenum and jejunum. Biopsy of the ulcer showed strongyloides infection He was treated with ivermectin. Abdomen Pain was subsided soon. He gained weight of 1.5 kilograms over next month.

Conclusion: When transplant patients from developing country with recent increase of immunosuppression presents with severe abdomen pain, the intestinal parasitic infection should be entertained. If stool examination is negative and serology is unavailable, early evaluation by endoscopic biopsy is helpful to diagnose strongyloides. Prolonged treatment with ivermectin and follow-up stool examinations are important for complete cure of strongyloides infection.

Key words:

Strongyloides; intestinal ulcers; renal transplantation; stool examination; endoscopic biopsy

INTRODUCTION:

Strongyloides stercoralis was first identified by Normand in 1876. *Strongyloides* is an intestinal nematode, produces minimal symptoms in immunocompetent individuals, but when occur in persons with impaired T lymphocyte function, leads to life-threatening infection with 60-85% mortality rate^[1]. The *Strongyloides stercoralis* excretes larvae in feces and has autoinfective cycle^[2]. Only female adults are detectable in humans, and subsequent reproduction occurs asexually^[3]. No direct person-to-person transmission occurs. The prevalence of *strongyloides* ranges from 12 to 76%. The disease is estimated to affect 100-200 million people worldwide. Gram-negative sepsis (*Escherichia coli* and *Klebsiella*) is a common consequence of hyperinfection and carries a 50% mortality rate.

Presentation of case:

A 26 year old man hawker with solitary kidney, end stage renal disease underwent live related renal transplantation in March, 2011. His mother was kidney donor. He was given steroids, mycophenolate sodium and tacrolimus. He had good immediate graft function. Pre-transplant routine examination of stool of both recipient and donor was negative for parasites. In May, 2011 he suffered one episode of biopsy proven acute rejection, reversed completely with three daily doses of intravenous methylprednisolone. In June, 2011 he presented with severe epigastric pain and vomiting. Pain was crampy, not radiating to back. It did not subside with injectable pantaprazole (proton pump inhibitors) and injectable antispasmodics (dicyclomine). On examination he was malnourished, dehydrated and lost two kilograms of weight over one month. His serum albumin was 2.9 mg/dL with absence of protein in urine. Stool examination was negative for *strongyloides* larvae. As pain persisted upper gastrointestinal scopy (Olympus- CV150 series) was done, it revealed multiple ulcers with slough seen in first and second part duodenum (Fig 1). Lesions were biopsied and sent for histopathological examination. It showed dysplastic glands with larvae of *strongyloides stercoralis* (Fig 2). He was treated with ivermectin 200 mg/day for 5 days and repeated after 2 weeks. Abdomen Pain was subsided soon. He gained weight of 1.5 kilograms over next month. His stool examination was negative for next 2 months.

DISCUSSION:

When patient presented with severe abdomen pain after treatment of acute rejection, stool examination by wet mount test (with concentration technique) for three consecutive days was done, it was negative for parasites. He did not respond to proton pump inhibitors.

When patient presented with severe epigastric pain with vomiting the following differential diagnosis were considered and excluded by appropriate investigations - acid peptic disease, pancreatitis, biliary colic and left lobe liver abscess. As pain persisted upper GI endoscopy was done, it showed multiple duodeno-jenual ulcers. The Differential diagnosis of ulcers include stress of surgery, steroids, Mycophenolate mofetil, Crohn's disease, ischemia, Zollinger-Ellison syndrome, infections (Mycobacterium, Syphilis, Typhoid, Histoplasmosis, and Cytomegalovirus), medications such as potassium, nonsteroidal anti-inflammatory drugs (NSAIDs) and rarely vasculitis. Biopsy of the ulcer showed strongyloides.

Strongyloides infection may presents as acute, chronic and hyperinfection. Strongyloides hyperinfection represents accelerated autoinfection due to alteration in immune status. The glucocorticoids can transform chronic strongyloidiasis to hyperinfection. Hyperinfection can occur with any dose of steroid and any preparation of steroid. After steroid therapy, strongyloides may present from 20 days to after many years. Cyclosporine does not associate with hyperinfection and may actually have an antihelminthic effect [4]. Exacerbation of gastrointestinal and pulmonary symptoms is seen, and the detection of increased numbers of larvae in stool and/or sputum is the hallmark of hyperinfection.

The gastro-intestinal symptoms in strongyloides infection are common but are non-specific. The gastro-intestinal symptoms includes abdominal pain [5], chronic diarrhea, constipation [5], anorexia, weight loss [5], difficulty swallowing [6], sore throat, nausea [5], vomiting, gastrointestinal bleeding and pruritus ani.

A minimum of three samples of stool is necessary to make a diagnosis of intestinal infection. As excretion of parasite can be intermittent, it is preferable that specimens be sent on different days [7]. Various stool concentration techniques and use of the Baermann method (by which motile larvae are extracted from stool specimens) may increase the diagnostic yield of fecal examination [7]. Several immunodiagnostic assays have been developed, but not readily available.

If the stool examination is negative and serology is not available endoscopy or colonoscopy is appropriate depending on the symptoms. Hussain Issa et al^[8] showed that strongyloides infection was diagnosed by colonoscopic biopsy (later confirmed by stool examination) in a renal transplant patient presented with nausea, vomiting and significant loss of weight. Bobby Kakati et al^[9] reported a case of strongyloides duodenitis in a patient with AIDS, presenting as abdominal pain, intractable nausea and emesis (stool examination without concentration methods were negative). Findings from upper gastrointestinal endoscopy may range from normal-appearing mucosa to severe duodenitis.

Duodenal biopsy histopathological examination identified larvae in 71.4% of immunosuppressed patients^[10]. Thus, in addition to stool analysis, endoscopic observation and biopsies are very important^[11]. Histopathology of the lesions may shows Strongyloides stercoralis larvae embedded in the mucosal lamina propria, where they produce mild to moderate degrees of edema, cellular infiltration, partial villous atrophy, and, ulcerations and fibrosis.

The goal of treatment is total eradication of the parasite, not just symptomatic improvement, as even one viable worm can initiate hyperinfection. Ivermectin is better than thiabendazole^[12] and albendazole^[13] in treatment of strongyloides. Ivermectin dose is 200 µg/kg/day, continue till symptoms resolve and stool tests have been negative for at least 2 weeks (one autoinfection cycle), or longer if patient remains immunosuppressed.

CONCLUSION:

When transplant patients from developing country with recent increase of immunosuppression presents with severe abdomen pain, the intestinal parasitic infection should be entertained. If stool examination is negative and serology is unavailable, early evaluation by endoscopic biopsy is helpful to diagnose strongyloides. Prolonged treatment with ivermectin and follow-up stool examinations are important for complete cure of strongyloides infection.

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COMPETING INTERESTS: NIL

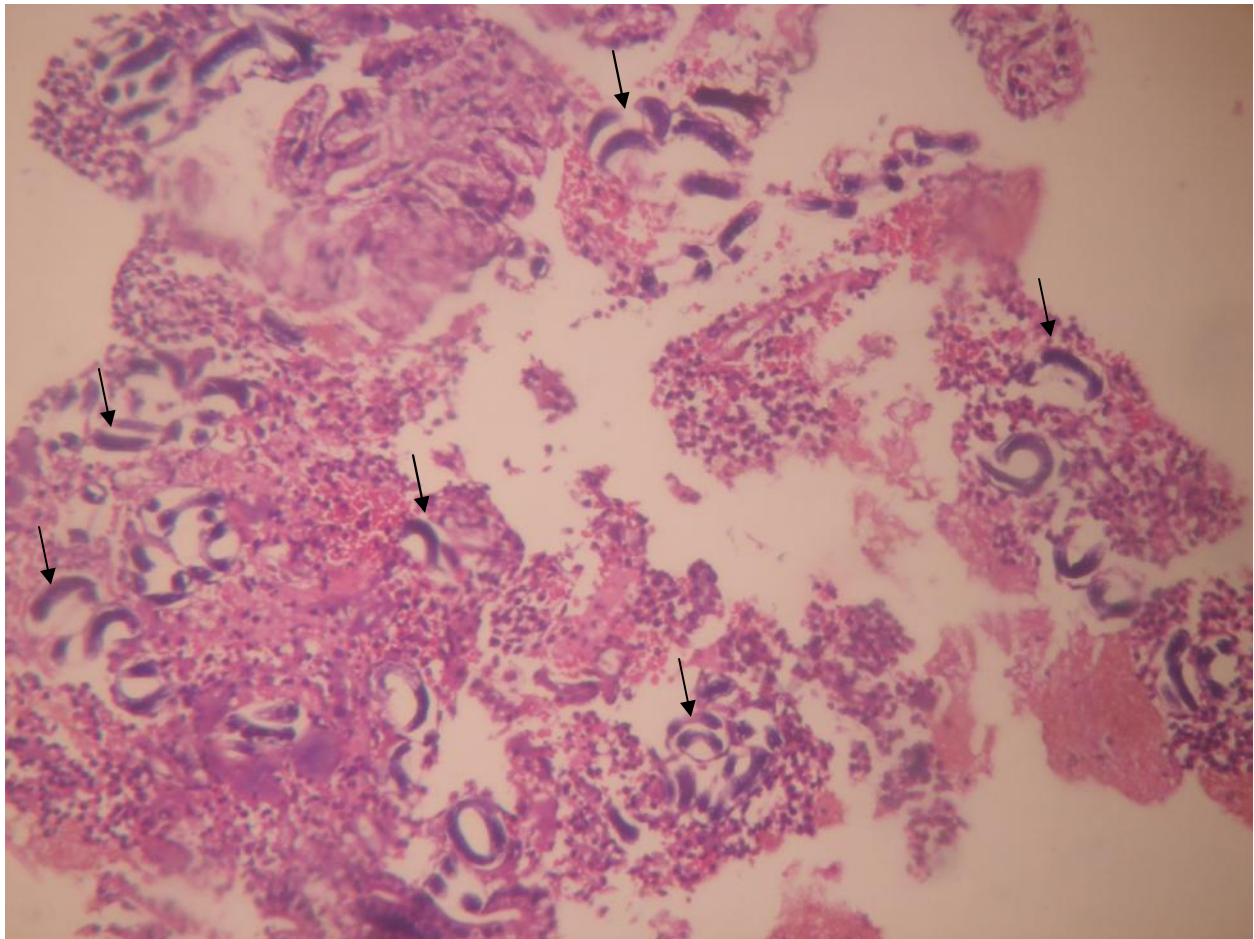
AUTHORS CONTRIBUTION: Author1:He is the main author and corresponding author, he is nephrologist, who transplanted the kidney to the patient and he is the main treating doctor/consultant. Author 2: he is also nephrologist who helped in treating the patient. Author 3: He is pathologist who diagnosed the stronglyloides in biopsy. Author 4: He is medical gastro-enterologist who performed endoscopy to the patient.

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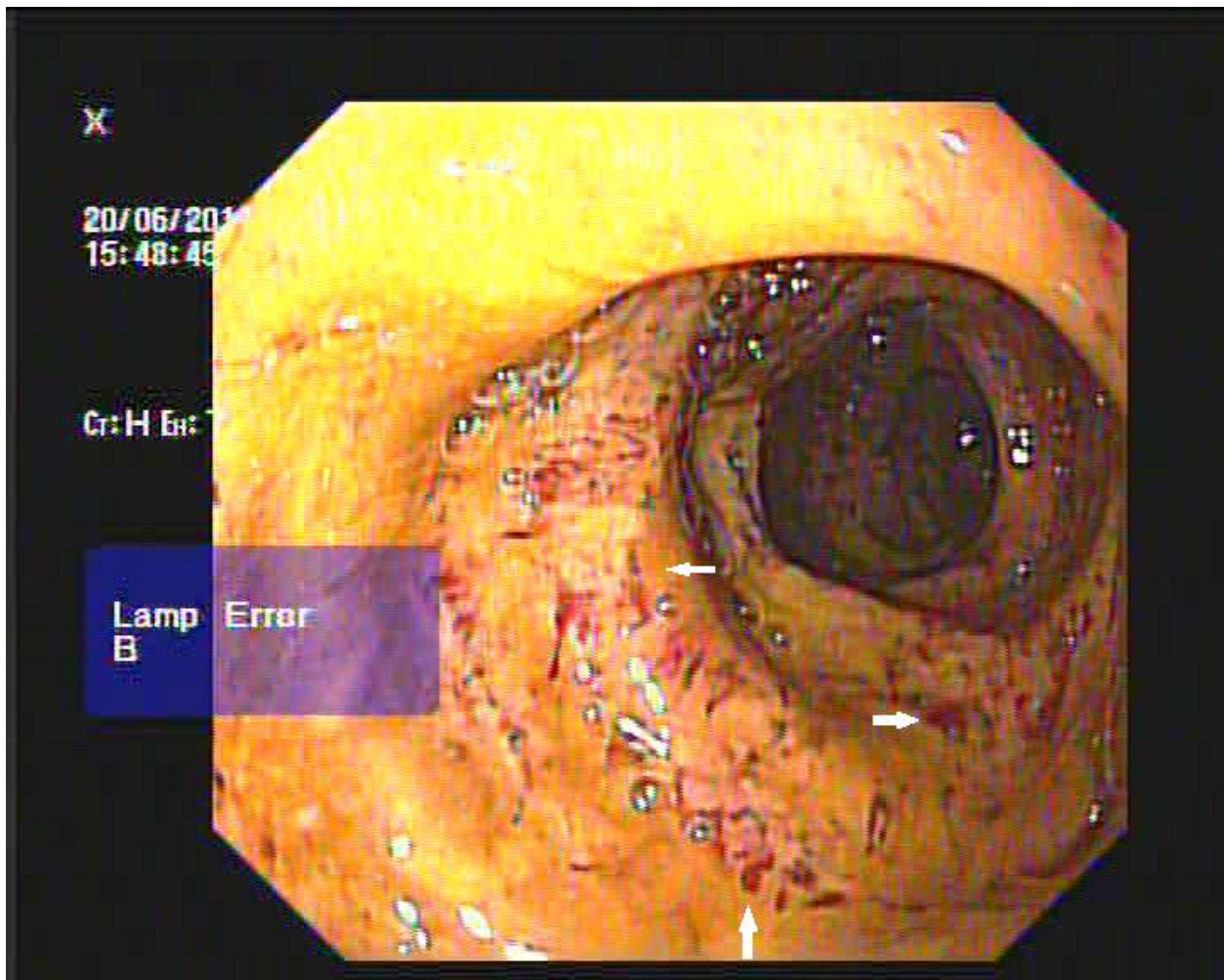
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155 Figure 2: Intestinal Biopsy showing strongyloides larvae (Arrows)



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168 Figure 1: Upper GI endoscopy – multiple ulcers seen in duodenum (arrows)



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