

# Dengue hemorrhagic fever complicated by pancreatitis

## Authors

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## ABSTRACT

Acute pancreatitis is an atypical complication of dengue fever and is rarely described. We are reporting a case of dengue hemorrhagic fever complicated by acute pancreatitis in a patient with history of *diabetes mellitus* type 1 and end stage renal disease on hemodialysis.

**Keywords:** dengue hemorrhagic fever; pancreatitis; *diabetes mellitus* type 1.

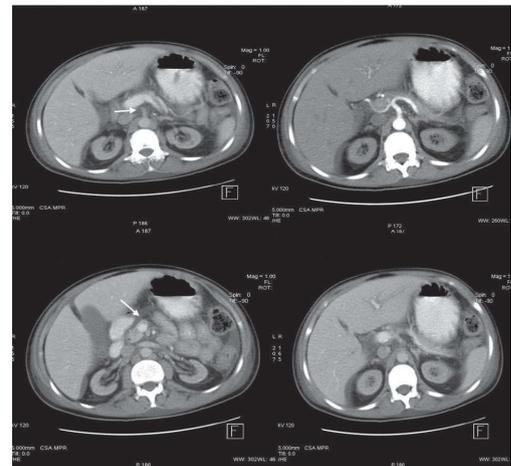
## INTRODUCTION

Dengue has been reported in almost 70 countries, with about five million cases reported between 2000 and 2007.<sup>1</sup> In recent years, the incidence of dengue has increased in the Central and South American region which now accounts for 70% of all cases reported worldwide.<sup>1,2</sup> Pancreatitis is an uncommon manifestation of this infection.

## CASE REPORT

On 14 June 2009, a 27-year-old male was admitted to the Fundación Valle del Lili in Cali, Colombia, with 4 days of fever, chills, loose stools, vomiting, hematemesis and rectal bleeding. The patient has a history of *diabetes mellitus* type 1 diagnosed at the age of 9 years and end stage renal disease (ESRD) secondary to diabetic nephropathy on hemodialysis (HD) over the last 3 years. He was taking amlodipine, verapamil, furosemide, and insulin analogs (Glargine and Lispro) at the time of admission. His vital signs on arrival to the emergency room were: BP: 102/71; HR: 91; BR: 15; T: 36°C; SaO<sub>2</sub>: 97% on FiO<sub>2</sub>: 21%. Physical examination was relevant for epigastric and mesogastric tenderness without rebound or guarding and petechiae on arms and legs. No focal neurologic deficits were detected. A series of laboratory studies revealed lymphopenia, severe thrombocytopenia and hemocentration. Hematologic laboratory values: white cell count  $4.97 \times 10^3/\mu\text{L}$  (normal range  $4.1-10.9 \times 10^3/\mu\text{L}$ ); total lymphocytes  $0.64 \times 10^3/\mu\text{L}$  (normal range  $1.0-4.0 \times 10^3/\mu\text{L}$ ); hemoglobin

22.4 g/dL (normal range 14.0-16.0 g/dL); hematocrit 69.3%(42-48%); platelets  $9.6 \times 10^3/\mu\text{L}$  ( $140-440 \times 10^3/\mu\text{L}$ ). Dengue serological test was performed: on admission dengue virus IgM and IgG were negative; IgM became positive four days later. The chemistry was significant for hyperlipasemia and hypocalcaemia: serum amylase 187.0 U/L (normal range 30-110 U/L); serum lipase 514.0 UI/L (normal range 23.0-300.0 UI/L) and calcium 6.56 mg/dL (normal range 8.4-10.2 mg/dL). Serum lipid profile approximately two years before presentation showed normal triglycerides: 137.6 mg/dL (normal range 0.00-200 mg/dL). Contrast-enhanced abdominal CT reported changes in the peripancreatic fat toward the root of the mesogastrium (Figure 1). Gastrointestinal endoscopy was per-



**Figure 1:** Minimal changes in the peripancreatic fat toward the root of the mesogastrium (see arrows).

Submitted on: 02/09/2011  
Approved on: 03/14/2011

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We declare no conflict of interest.

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formed finding hemorrhagic thrombocytopenic pangastropathy. The patient developed disseminated intravascular coagulation (DIC), was transferred to the intensive care unit where intravenous hydration and transfusion with platelet units, fresh frozen plasma and cryoprecipitate were given. The patient recovered slowly over the next days and was finally discharged at his 13<sup>th</sup> day of hospitalization with significant improvement in his clinical condition.

## DISCUSSION

The dengue virus, a single stranded RNA virus belonging to the Flaviviridae family, has been classified into four serotypes, DENV-1, DENV-2, DENV-3, and DENV-4, which are genetically and antigenically different.<sup>3</sup> The main vector of the dengue virus is the mosquito *Aedes aegypti*. Dengue is classified as dengue fever, dengue hemorrhagic fever (DHF) or dengue shock syndrome (DSS) depending on its severity and presenting features. Our patient met the WHO criteria for a case definition of DHF.<sup>4</sup> Pancreatitis was evident through marked increase in serum lipase and CT inflammatory findings.<sup>5</sup> Although hyperlipasemia and pancreatic enlargement have been reported in acute dengue infection; pancreatitis as a DHF complication is considered an atypical manifestation.<sup>6</sup> The largest described series was in an DHF outbreak in Taiwan in 2002 were pancreatitis (defined by a lipase level 3-fold greater than the upper limit of normal) was diagnosed in three patients with acute DHF,<sup>7</sup> other five isolated cases have been described in Thailand, Indonesia, Noumea (New Caledonia) and Colombia (Table 1).<sup>8-12</sup> Among all the cases it was noted that at least two more had history of diabetes as well. It is possible that diabetes itself was an aggravating condition that might have predisposed those patients to develop pancreatitis. The exact mechanism by which dengue virus induces acute pancreatitis is unclear and might be multifactorial. Based on data from patients with other types of viral pancreati-

tis, several hypotheses for a pathogenic mechanism have been proposed. Among them are viral-associated acute pancreatitis due to direct inflammation and destruction of pancreatic acinar cells by the virus; acute viral infection acting as a trigger for an autoimmune response to pancreatic islet cells, which is induced by the similarity between viral and islet cells antigens, and the development of edema of the ampulla of Vater with obstruction to the outflow of pancreatic fluid.<sup>13-15</sup>

## ACKNOWLEDGEMENTS

To Dr. Mauricio Mejia from the Department of Radiology, Fundación Valle del Lili, for his collaboration with the CT scan interpretation and to Dr. Jose Cadena for his valuable help in the review of this manuscript.

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Table 1. Cases reported of dengue complicated with pancreatitis

Cases	Age/sex	Country	Associated findings	Year	Reference
1	10/F	Thailand	Liver failure	1988	[10]
2	< 13Y	Colombia	None	1992-2002	[12]
3	24/M	Indonesia	Unknown	1998	[11]
4-6	N/A	Taiwan	<i>Diabetes mellitus</i>	2002	[7]
7	66/F	Taiwan	Seizures, <i>diabetes mellitus</i>	2004	[8]
8	29/F	Noumea (New Caledonia)	None described	2005	[9]
9	27/M	Colombia	DIC, <i>diabetes mellitus</i>	2009	Ours

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