A Case Report on Fish Bile Toxicity- a rare cause of Multiple Organ Dysfunction Syndrome

Farhana Yasmin¹, Shireen Afroz², Tahmina Ferdaus³, Umme Tanjila⁴, Sukriti Baroi⁵

ABSTRACT: A case study on fish bile poisoning is reported. After ingestion of gall bladder of Labeo rohita fish for allergy treatment purpose initially presented with gastrointestinal symptoms such as cramping pain abdomen, nausea and vomiting within 12 hours after ingestion. Subsequently renal, hepatic dysfunctions and cardiac dysfunction were found in that case. The patient recovered fully with conservative treatment and supportive hemodialysis.

KEYWORDS: Labeo rohita, Fish bile, Hemodialysis

INTRODUCTION:

In India, especially in Assam and Chinese people believe that fish gall bladder can improve vision, treat rheumatism, improves eyesight and cure asthma [1]. Due to frequent consumption of fish gall bladder, fish bile poisoning cases are reported more commonly in China, India, Japan, and other Asian countries [2,3,4]. There were many reports about fish gall bladder poisoning leading to acute renal failure (ARF), acute liver injury, and therefore increasing mortality [5]. The incidence of ARF in fish bile poisoning is 55%-100%, while the mortality rate accounts 91.7%. Fish gallbladder contains a heat stable toxin which can severely affect gastrointestinal system, renal, liver, central nervous system, cardiovascular system and leading to multiple organ failure (MODS)[1] This is a case of Rohu fish gall bladder poisoning leading to multi organ involvement in a Bangladeshi child. This report is a whole new perspective on the pathogenesis of acute renal failure and other organ involvement in a case of poisoning by fish gall bladder. This has a positive role in guiding treatment of fish bile poisoning, with obvious effect to improve its prognosis. This case not only focuses over presentations but also guides the management as there was no mortality in our case in spite of severity in presentation. The condition is commonly reversible, and therefore proper history taking is important and prompt biochemical investigations including blood urea and creatinine are needed to enable early diagnosis and fast institution of treatment, which may include HD.

CASE FEATURES

A 10-year-old boy who presented with history of repeated vomiting and diffuse abdominal pain for 3 days and oliguria for 2 days. His father admitted that he convinced his boy to consume raw fish gallbladder for the purpose of treatment of allergy 1 day prior to the presentation. This was followed by diffuse abdominal pain and profuse vomiting and he got admitted into a hospital and treated with IV fluid, antiemetic and antiulcerant medications. He developed oliguria along with puffy face and yellowish discoloration of skin and sclera within 48 hours of presentation. After investigations over that hospital they diagnosed him as a case of Acute Kidney Injury (AKI) and Intermittent peritoneal dialysis (IPD) was started. When there was no improvement, they referred him to our hospital to better management. There was no history of fever, previous history of renal disease or family history of such type of illness.

On general examination, he appeared ill looking, puffy, afebrile but was communicative and oriented. The boy was mildly pale and icteric and on vitals examination pulse rate 82 bpm, blood pressure 130/90 mmHg (above 99th centile), respiratory rate 20/min, and temperature 37.5 ° C. Cardiac exam revealed regular heart sounds and no murmur. His breath sound was vesicular with no wheeze or crackles. Abdomen was soft, mildly tender with IPD catheter in situ. There was no organomegaly but ascites was evident by positive fluid thrill, bowel sound was present. Nervous system with all other systems examination revealed no abnormalities.

His initial Investigations showed in Table 1.

Investigations	Findings	Normal Values
Hb% WBC Neutrophil Lymphocytes Platelet	9.8 gm/dl 12 000/cumm, 78% 15% 245 000/cumm	11 to 14 gm/dl 4 000 to 11 000/cumm 40 to 70% 20 to 40% 150 000 to 40 000/cumm
Serum Creatinine Blood urea Serum Sodium Serum Potassium Serum Chloride Serum Albumin Serum Calcium C-Reactive protein SGOT SGPT Serum Bilirubin PT INR APTT	4.7 mg/dl 18 m mol/L 143 m mol/L 5.2m mol/L 105m mol/L 25 gm/L 2.1 m mol/L 7 mg/L 2196 U/L 7283 U/L 2.5 mg/dl 12 seconds, INR 1 29 seconds	0.3 to 0.8 mg/dl 1.3 to 5.8 m mol/L 135 to 145 m mol/L 3.5 to 5.4 m mol/L 96 to 108 m mol/L 35 to 60 gm/L 2.02 to 2.1 m mol/L < 5 mg/L Up to 37 U/L Up to 40 U/L 0.2 to 1 mg/dl 12 seconds 35 seconds
Serum Ferritin Serum Procalcitonin D-dimer Cardiac Troponin I NT ProBNP RT PCR COVID 19 Antibody (IgG) of Covid 19 Urine RME	288 ng/ml 6.45 ng/ml 1.44 mg/L 0.00 ng/ml 27262 pg/ml Negative Negative Albumin 2+, RBC 10-15/HPF	15 to 300 ng/ml < 0.50 ng/ml < 0.5mg/L 0 to 0.034 ng/ml <125 pg/ml

Chest X ray	Normal	
USG of Whole Abdomen	 Cortical Echogenicity of both kidneys raised with poorly differentiated Cortex and medulla Mild ascites Normal Kidney sizes Right Kidney 8.0 cm Left Kidney 8.1 cm 	
Echocardiography	 Dilated coronaries with loss of distal tapering Mild LV Systolic Dysfunction 	
Blood Culture	No growth	
Urine Culture	No growth	
Complement C3	0.76 g/L 0.75 to 1.65 gm/L	
Complement C4	0.21 g/L 0.15 to 0.45 gm/L	
ANA	Negative	
Anti Ds DNA	Negative	
Serum PTH	78 pg/ml	
ABG	PH 7.35 7.45	
	HCO3 13 18 to 23	
	PO2 138 80 to 100 mmHg	
	BE – 8.2	
Renal Biopsy	Features of Tubular Injury	

After getting three sessions of hemodialysis and conservative management patient was on improving pattern and got discharged. After two weeks of follow up his all-biochemical parameters become normal.

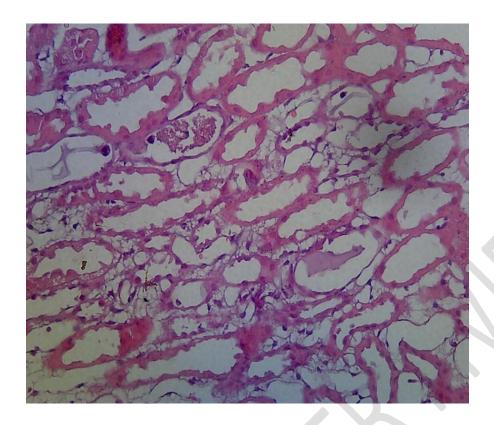


Fig 1: Renal biopsy showing tubular necrosis

DISCUSSION

Fish bile induced renal failure and toxic hepatitis cases have been reported in various parts of Asia & Southeast Asia [5]. Toxicity is attributed to the fishes belonging to the family Cyprinidae. The family includes grass carp, common carp, and silver carp. Amongst these, fish of the grass carp variety has been commonly reported for its toxicity. Rohu (Labeo rohita) the Indian fish carp is commonly consumed in north eastern and eastern region of India. Its bile contains a toxin, sodium cyprinol sulfate, which occurs in three forms: toxins in visceral organs (ichthyosarcotoxic), reproductive organs (ichthyootoxic), or blood (icthyohemotoxic). It is heat stable and alcohol insoluble, so cases are reported even after consumption of cooked bile[6,7]. Toxicity is directly proportional to the size and quantity of gall bladder or bile consumed [8]. After ingestion, initial manifestations include abdominal pain, nausea, vomiting and watery diarrhea, followed by oliguria and renal failure. The hepatic impairment and cardiac dysfunction usually precedes renal dysfunction, but may be concomitant with kidney injury. Kidney biopsy reveals proximal tubular cell damage on light microscopy. Electron microscopy shows decreased mitochondria crista in the proximal tubular epithelial cells, swollen glomerular cells and partially fused podocytes. The toxin is believed to damage lysosomes and inhibit cytochrome oxidase enzyme, thus blocking cellular metabolism and causing necrosis of the proximal tubular epithelial cells. In addition, loss of fluid due to vomiting, diarrhea can lead to decreased effective circulating blood volume and eventually leads to oliquric or the non-oliquric form of acute renal failure, usually within 48-72 hours after toxin ingestion [9, 10]. BichHuyen Nguyen Xuanet al. Al from Vietnam have conducted a large study and showed the effects of certain freshwater fish

bile associated acute tubular necrosis. In India, Dwijen Das *et al.* have published a case series on fish bile toxicity causing lethal renal failure and hepatic dysfunction. Fish bile can also damage other organs causing multiple organ dysfunction syndrome (MODS) [11,12]. However, acute renal failure after fish gallbladder ingestion has an excellent prognosis, though death from fulminant hepatic failure can occur. Proper management comprises of hemodialysis and supportive management, is essential to save lives of these patients [13]. Eliciting a proper clinical history in such cases is of paramount importance as many different substances can produce simultaneous renal, hepatic and cardiac damage. This includes variety of toxins like carbon tetrachloride, trichloroethylene, chloroform, copper sulfate and chromium, mushroom poisoning and drugs including paracetamol overdose and fluorinated anesthetic agents such as methoxyfluorane and fluoxene. Hence these more common etiological agents should be first ruled out by detailed history taking before considering uncommon fish bile as the injurious agent [14].

CONCLUSION

This case report should help the general physicians in developing countries to be aware of the fact that various types of food poisoning can cause AKI and fish bile can be a possible but rare cause of reversible acute renal & hepatic failure. Proper history taking is important in these circumstances and prompt biochemical investigations are needed for an early diagnosis and institution of proper treatment for the purpose of fruitful outcome.

Footnotes: Authors' Contribution: Study concept and design: FY; Acquisition of data: FY; Analysis and interpretation of data: FY; Drafting of the manuscript: FY; Critical revision of the manuscript for important intellectual content: SA,TF; Administrative, technical, and material support: FY,SA,UT,SB. Conflict of Interests: There are no conflicts of interest to declare. Funding/Support: No funding was needed for this study. Informed Consent: We obtained informed written consent from the father.

Author's information:

- 1.Farhana Yasmin, Resident Medical Officer, Critical Care Nephrology and Dialysis Unit, Department of Pediatric Nephrology, Bangladesh Shishu (Children) Hospital & Institute. First author contact person.
- 2. Professor (Dr.) Shireen Afroz, Professor & Head of department of Pediatric Nephrology, Bangladesh Shishu Hospital & Institute.
- 3.Tahmina Ferdaus, Register, Critical Care Nephrology and Dialysis Unit, Department of Pediatric Nephrology, Bangladesh Shishu (Children) Hospital & Institute.
- 4.Umme Tanjila, Resident Medical Officer, Critical Care Nephrology and Dialysis Unit, Department of Pediatric Nephrology, Bangladesh Shishu (Children) Hospital & Institute.
- 5.Sukriti Baroi, Resident Medical Officer, Critical Care Nephrology and Dialysis Unit, Department of Pediatric Nephrology, Bangladesh Shishu (Children) Hospital & Institute.

References:

- 1.X. L. Cheng, Z. I. Wang, and L. M. Reng, "Fish gall bladder poisoning damage liver, renal and heart," *Chinese Journal of Integrative Medicine*, vol. 3, pp. 238–239, 1991.
- 2. D. S.W. Cahn, C. K. Yeung, and M. K. Chan, "Acute renal failure after eating raw fish gall bladder," *British Medical Journal*, vol.290, no. 6472, article 897, 1985.
- 3. Y. Yamamoto, O. Wakisaka, S. Fujimoto et al., "Acute renal failure caused by ingestion of the carp gall bladder-a report of 3 cases, with special reference to the reported cases in Japan," *Nihon Naika Gakkai Zasshi*, vol. 77, no. 8, pp. 1268–1273, 1988.
- 4.R. N. Sahoo, M. K. Mohapatra, B. Sahoo, and G. C.Das, "Acute renal failure associated with freshwater fish toxin," *Tropical and Geographical Medicine*, vol. 47, no. 2, pp. 94–95, 1995.
- 5. Singh NS, Singh LK, Khaidem I, Singh G, Reddy VS, Bawi NS, Singh YI. Acute renal failure following consumption of raw fish gall-bladder from Manipur. JOURNAL-ASSOCIATION OF PHYSICIANS OF INDIA. 2004 Sep;52:743-5.
- 6. Xuan BH, Nguyen Thi TX, Nguyen ST, Goldfarb DS, Barry Stokes MS, Rabenou R.IchthyotoxicARFAfter Fish Gallbladder Ingestion: A Large Case Series From Vietnam. American Journal of Kidney Diseases. 2003(41):220-224.
- 7. Yip LL, Chow CL, Yung KH, Chiu KW. Toxic material from the gallbladder of the grass carp(Ctenopharyngodonidellus). Toxicon. 1981 Jan1;19(4):567-9.
- 8. Xuan BH, Thi TX, Nguyen ST, Goldfarb DS, Stokes MB, Rabenou RA. Ichthyotoxic ARF after fish gallbladder ingestion: a large case series from Vietnam. American journal of kidney diseases. 2003 Jan 1;41(1):220-4.
- 9. Lim PS, Lin JL, Hu SA, Huang CC. Acute renal failure due to ingestion of the gallbladder of grass carp: report of 3 cases with review of literature. Renal failure. 1993 Jan 1;15(5):639-44.
- 10. Deng Y, Xiao G, Jin Y, Luo X, Meng X, Li J, Ao Z, Xiao J, Zhou L. Multiple organ dysfunction syndrome due to ingestion of fish gall bladder. Chinese medical journal. 2002 Jul;115(7):1020-2.
- 11. Bich Huyen Nguyen Xuan, Tan Xuan Nguyen Thi, Su Tan Nguyen, David S.Goldfarb, M.Barry Stokes, Rahmin A. Rabenou. American Journal of Kidney Diseases. January 2003; 41(1): 220–224.
- 12. Dwijen Das, Kallol Bhattacharjee, Amit Kr. Kalwar, Bhaskar Debnath. Journal of Evidence based Medicine and Healthcare. 2015; 2(33), 5073-5076.
- 13. Tao FW, Liao FT, Xu YZ. A case of fish gall bladder poisoning leading to MODS. Chin J Intern Med. 1990;29:119-20.
- 14. Bhaumik P, Lakshmanan KP. Fish gallbladder consumption almost costing life. Global journal of medicine and public health. 2016(5):32-35.