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Research Article

**RELATIONSHIP BETWEEN HELICOBACTER PYLORI INFECTION
AND UPPER GASTRO-INTESTINAL BLEEDING IN NORTHERN
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of Medical Sciences, Babol, IR Iran⁴ Non-Communicable Pediatric Diseases Research Center, Health Research Institute, Babol University
of Medical Sciences, Babol, IR Iran**Abstract:**

Background: There are controversies regarding relationship between *Helicobacter pylori* (*H. pylori*) and upper gastro-intestinal bleeding (UGIB) in children. The goal of this study was to assess the relationship between *H. pylori* infection and UGIB in Northern Iranian children.

Material and method: One hundred children who had UGIB indicated for upper gastrointestinal endoscopy and 100 children without UGIB who were candidate for upper endoscopy because of chronic abdominal pain considered as their controls. After stabilizing vital sign, nasogastric tube inserted and washing was done then within 24 hours of admission, endoscopy conducted for all children in the case group (under general anesthesia). Upper endoscopy was done for all participants by a children gastroenterologist. A single pathologist reviewed all specimens.

Results: Mean age of cases was 6.2±3.2 years and mean age of controls was 7.1±2.9 years respectively. There was no relationship between *H. pylori* and UGIB. Erosion in fundus was significantly higher in cases (0.001) and erythema in antrum was significantly higher in controls ($p < 0.001$). Inactive gastritis was significantly higher in *H. pylori* negative cases (0.006) while moderate gastritis was present in all *H. pylori* positive ones (0.009).

Conclusion: We found no relationship between *H. pylori* infection and UGIB in children. More studies are needed.

Key words: *H. pylori*, children, Gastro intestinal bleeding

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INTRODUCTION:

Bleeding could occur in any part of gastrointestinal (GI) system which makes discomfort for both children and their caregivers(1). Incidence of upper GI bleeding(UGIB) in children who had Intensive Care Unit (ICU) admission ranged from 6.4 to 10%(2, 3). UGIB mostly presents as melena or hematemesis and rarely as hematochezia. Mucosal lesions, variceal haemorrhage, infections and drugs are common causes of UGIB(4).

A gram negative spiral- bacterium which could be found in the gastric mucous layer or near to the epithelial lining of the stomach is *Helicobacter pylori* (*H. pylori*). Its prevalence in children ranges from 10% to 80%(5). More than 90% of duodenal ulcers and near 80% of gastric ulcers are related to *H. pylori* (6-8).

Literature shows that there is relation between *H. pylori* and UGIB in adults (9-11). In a previous study, El-Mazary reported higher prevalence of *H. pylori* in children with non-variceal bleeding than controls(12).

We designed this study to assess the relationship between *H. pylori* infection and UGIB in Northern Iran children.

MATERIAL AND METHODS:

This cross sectional study conducted in Amikola children hospital (affiliated hospital of Babol university of medical sciences in north of Iran) between 2009 and 2016.

Children with age more than 6 months, who had UGIB indicated for upper gastrointestinal endoscopy considered as case groups while age and sex matched children without UGIB who were candidate for endoscopy because of chronic abdominal pain considered as their controls.

Exclusion criteria were: coagulopathy, bleeding disorders, diabetes mellitus or chronic illness, unstable hemodynamic, foreign body ingestion, oesophageal and gastric varices, caustic ingestion, recent PPI (proton pump inhibitor) and antibiotic consumption.

Informed consent forms were taken from all parents before study entrance.

After stabilizing vital sign, nasogastric tube inserted and washing was done then within 24 hours of admission, then endoscopy conducted for all children in the case group (under general anaesthesia).

Upper endoscopy was done for all participants by a children gastroenterologist. Pentax (EPM 3500) paediatric gastroscope was used for the procedure. Samples from oesophagus, gastric and duodenum collected for all cases. Macroscopic findings of endoscopy were recorded.

A single pathologist reviewed all specimens for microscopic assessment (with Giemsa staining for *H. pylori*). Sydney classification was applied for endoscopic report of gastritis. (13) All data were analyzed using SPSS software version 22 (SPSS Inc., Chicago, IL, USA). Data were presented as Mean \pm SD for continuous or frequencies for categorical variables. Independent sample t test was used for comparison of continuous variables. P value less than 0.05 was considered as significant.

RESULTS:

One hundred cases in bleeding group and 100 in control group enrolled.

Demographic characteristics are summarized in table 1. There was no relationship between *H. pylori* and UGIB (table 1).

Table 1: Demographic characteristics

	Cases	controls	P value
Age (mean \pm SD)(years)	6.2 \pm 3.2	7.1 \pm 2.9	0.07
Sex			
Male	58(58%)	46(46%)	0.1
Female	42(42%)	54(54%)	
H. pylori infection			
Yes	10(10%)	12(12%)	0.8
No	90(90%)	88(88%)	

Mean age in cases who were positive for *H. pylori* was 10.2 \pm 3.3 years and 8.1 \pm 2.7 in controls ($p < 0.001$). Erosion in fundus was significantly higher in cases and erythema in antrum was significantly higher in controls (table 2).

Table 2: Endoscopic findings in two groups

	Cases	controls	P value
Esophagus			
Erythema	33%	68%	0.08
Erosion	7%	0	
Ulcer	0	0	
Nodularity	1%	0	
Mallory Weiss	3%	0	
Fondus			
Erythema	12%	12%	0.001
Erosion	18%	1%	
Ulcer	0	0	
Nodularity	0	0	
Sub epithelial bleeding	18%	0	
Body			
Erythema	8%	17%	0.3
Erosion	12%	2%	
Ulcer	2%	0	
Nodularity	0	0	
Antrum			
Erythema	26%	67%	<0.001
Erosion	14%	2%	
Ulcer	2%	0	
Nodularity	18%	37%	
Bulb duodenum			
Erythema	5%	1%	0.07
Erosion	1%	3%	
Ulcer	3%	0	
Nodularity	6%	11%	
Duodenum			
Erythema	1%	2%	0.08
Erosion	2%	3%	
Ulcer	0	0	
Nodularity	2%	2%	

All of duodenal ulcers were H. pylori positive and antral nodularity was more in H.pylori positive cases (table3).

Table 3: Endoscopic findings in GI bleeding group.

		H.pylori positive	H.pylori negative	P. Value
Esophagus	Normal	13.6 %	86.4 %	0.191
		3 %	97 %	0.159
	Erythema	14.3 %	85.7 %	0.533
		-	-	-
	Erosion	0	100 %	1.000
			100 %	1.000
	Ulcer			
Nodularity				
Mallory Weiss				
Fondus	Normal	12.9 %	87.1 %	0.311
		0	100 %	0.604
	Erythema	5.6 %	94.4 %	0.685
		-	-	-
	Erosion	-	-	-
		5.55 %	94.45 %	0.583
	Ulcer			
Nodularity				
Sub epithelial bleeding				
Body	Normal	11.4 %	88.6 %	0.684
		12.5 %	87.5 %	0.583
	Erythema	0	100 %	0.604
		0	100 %	1.000
	Erosion	-	-	-
	Ulcer			
Nodularity				
Antrum	Normal	2.4 %	97.6 %	0.042
		3.8 %	96.6 %	0.447
	Erythema	0	100 %	0.349
		0	100 %	1.000
	Erosion	56.6%	44.4 %	0.000
		Ulcer		
	Nodularity			
D1(duodenum)	Normal	5.9 %	94.1 %	0.006
		0	100 %	1.000
	Erythema	0	100 %	1.000
		100 %	0	0.001
	Erosion	33.3 %	66.7 %	0.109
		Ulcer		
	Nodularity			
D2(duodenum)	Normal	10.5 %	89.5 %	1.000
		0	100 %	1.000
	Erythema	0	100 %	1.0000
		-	-	-
	Erosion	0	100 %	1.000
		Ulcer		
	Nodularity			

Endoscopic findings were similar between two control groups.(table 4)

Table 4: Endoscopic findings in control group.

		H.pylori positive	H.pylori negative	P. Value
Esophagus	Normal	18.8 %	81.3 %	0.191
		8.8 %	91.2 %	0.191
	Erythema	-	-	-
		-	-	-
	Erosion	-	-	-
	Ulcer	-	-	-
	Nodularity Mallory Weiss			
Fondus	Normal	12.6 %	78.4 %	1.000
		8.3 %	91.7 %	1.000
	Erythema	0	100 %	1.000
		-	-	-
	Erosion	-	-	-
	Ulcer	-	-	-
	Nodularity Sub epithelial bleeding			
Body	Normal	12.3 %	88.7 %	1.000
		11.8 %	88.2 %	1.000
	Erythema	0	100 %	1.000
		-	-	-
	Erosion	-	-	-
	Ulcer Nodularity			
Antrum	Normal	0	100 %	0.208
		11.9 %	88.1 %	1.000
	Erythema	0	100 %	1.000
		-	-	-
	Erosion Ulcer Nodularity	18.9 %	81.1 %	0.120
D1(Duodenum)	Normal	11.8 %	88.2 %	1.000
		0	100 %	1.000
	Erythema	0	100 %	1.000
		-	-	-
	Erosion Ulcer Nodularity	18.2 %	81.8 %	0.618
D2(Duodenum)	Normal	12.5 %	87.5 %	1.000
		0	100 %	1.000
	Erythema	0	100 %	1.000
		-	-	-
	Erosion Ulcer Nodularity	0	100 %	1.000

Esophagitis and inactive gastritis were the most common pathologies in cases and controls (table5).

Table5: Pathology findings in two groups.

Pathology	Cases	Controls
Normal	39 %	23 %
Esophagitis	20 %	35 %
Inactive gastritis	29 %	53 %
Mild gastritis	6 %	3 %
Moderate gastritis	2 %	2 %
Severe gastritis	0	0
Duodenitis	8 %	4 %
Inadequate sample	2 %	2 %

Inactive gastritis was significantly higher in H .pylori negative cases while moderate gastritis was present in all H .pylori positive ones in cases (table 6).

Table6: Pathology in H .pylori positive and negative cases.

Pathology	H.pylori positive	H.pylori negative	P. Value
Normal	0	100 %	0.006
Esophagitis	0	100 %	0.205
Inactive gastritis	24.1 %	75.9 %	0.006
Mild gastritis	16.7 %	83.3 %	0.478
Moderate gastritis	100 %	0	0.009
Severe gastritis	-	-	-
Duodenitis	0	100 %	1.000
Inadequate sample	0	100 %	1.000

Pathologic findings were similar between two control groups.(H. pylori positive and negative) (table7)

Table7: pathology in H .pylori positive and negative control groups.

Pathology	H.pylori positive	H.pylori negative	P. Value
Normal	4.3 %	95.7 %	0.286
Esophagitis	2.9 %	97.1 %	0.052
Inactive gastritis	15.3 %	84.7 %	0.368
Mild gastritis	33.3 %	66.7 %	0.321
Moderate gastritis	50 %	50 %	0.227
Severe gastritis	-	-	-
Duodenitis	0	100 %	1.000
Inadequate sample	0	100 %	1.000

DISCUSSION:

The results of current study showed that there was no relationship between UGIB and H. pylori infection in children between 6 months and 14 years.

We also found that H.pylori was present in 11% of cases with UGIB.

In a study conducted by Usta and Urganci, H.pylori rate in children with GI bleeding reported as 20%(13). El-Mazary et al evaluated 70 children with UGIB and 38 controls and reported higher rate of H. pylori infection in bleeding group(12). They suggested that this could be indicative of strong relationship between H .pylori infection and gastric and duodenal ulcers as the main causes of bleeding which is not in agreement with our findings.

Motamed et al reported prevalence of H. pylori infection as 9% in children suffering from gastrointestinal symptoms(14). They also reported relationship between antral nodularity and H. pylori Infection which was present in current study. Like our findings, gastritis was the most common pathological observation in H. pylori positive group.

Inactive gastritis, esophagitis, duodenitis, and active gastritis were the most common pathological findings in our study groups. In Varanasi et al study, H.pylori infection was found in 30.7% of patients with esophagitis(15). In our study like Motamed et al study, most cases with negative H. pylori result showed gastritis or esophagitis(14). In their study such as ours, there was significant association between H. pylori and antral nodularity.

Three cases in this study who had duodenal ulcer had positive H. pylori infection.

In Javid et al study, 80% of cases with duodenal ulcer had H. pylori infection(16). Antral nodularity was higher in H. pylori positive cases which is consistent with previous studies(14, 17). In Parsad and Luzza's studies nodularity was found in near 40% of H. pylori positive cases (17, 18).

H.pylori and EBV were considered to be associated with abnormal pathology in the stomach(19). Active gastritis and moderate and severe active gastritis were reported in 17.8% and 2.9% of the patients in Cardenas-Mondragon et al's study(19). Nodularity was the most common pathological finding in duodenum in both H.pylori positive and negative cases while in Kori et al study, mild and chronic duodenal inflammation reported in 6.5% of H.pylori positive cases(20).

In 1985, association between H. pylori and peptic ulcer was reported. This infection is acquired during childhood and adolescence both in developing and developed countries. Due to factors such as age, socioeconomic class and geographic distribution, prevalence of H. pylori differs between 10-80%(21, 22).

But there are controversies regarding association between this infection and peptic ulcers or GI bleeding(14, 23).

Larger multi centric studies are needed to evaluate association between H .pylori and GI bleeding or peptic ulcers.

CONCLUSION:

We found no relationship between H. pylori infection and UGIB in children. More studies are needed.

REFERENCES:

1. Dehghani SM, Haghghat M, Imanieh MH, Tabebordbar MR. Upper gastrointestinal bleeding in children in Southern Iran. *Indian journal of pediatrics*. 2009;76(6):635-8.
2. Boyle JT. Gastrointestinal bleeding in infants and children. *Pediatrics in Review*. 2008;29(2):39.
3. Lacroix J, Nadeau D, Laberge S, Gauthier M, Lapierre G, Farrell CA. Frequency of upper gastrointestinal bleeding in a pediatric intensive care unit. *Critical care medicine*. 1992;20(1):35-42.
4. Arora N, Ganguly S, Mathur P, Ahuja A, Patwari A. Upper gastrointestinal bleeding: etiology and management. *Indian journal of pediatrics*. 2002;69(2):155-68.
5. Jafar S, Jalil A, Soheila N, Sirous S. Prevalence of helicobacter pylori infection in children, a population-based cross-sectional study in west iran. *Iran J Pediatr*. 2013;23(1):13-8.
6. Drumm B, Day AS, Gold B, Gottrand F, Kato S, Kawakami E, et al. Helicobacter pylori and peptic ulcer: Working group report of the second world congress of pediatric gastroenterology, hepatology, and nutrition. *Journal of pediatric gastroenterology and nutrition*. 2004;39:S626-S31.
7. Blecker U, Gold B. Gastritis and peptic ulcer disease in childhood. *European journal of pediatrics*. 1999;158(7):541-6.
8. Cox K, Ament ME. Upper gastrointestinal bleeding in children and adolescents. *Pediatrics*. 1979;63(3):408-13.
9. Hsiao FY, Tsai YW, Wen YW, Kuo KN, Tsai CR, Huang WF. Effect of Helicobacter pylori eradication therapy on risk of hospitalization for a major ulcer event. *Pharmacotherapy: The Journal of Human Pharmacology and Drug Therapy*. 2011;31(3):239-47.
10. Tang J-H, Liu N-J, Cheng H-T, Lee C-S, Chu Y-Y, Sung K-F, et al. Endoscopic diagnosis of Helicobacter pylori infection by rapid urease test in bleeding peptic ulcers: a prospective case-control study. *Journal of clinical gastroenterology*. 2009;43(2):133-9.
11. Gisbert JP, Calvet X, Feu F, Bory F, Cosme A, Almela P, et al. Eradication of Helicobacter pylori for the prevention of peptic ulcer rebleeding. *Helicobacter*. 2007;12(4):279-86.
12. El-Mazary A-AM, Elfoly MA, Ahmed MF, Abdel-Hamed WM, Hassan ZM. Helicobacter Pylori Infection in a Group of Egyptian Children With Upper Gastro-Intestinal Bleeding. *Gastroenterology research*. 2013;6(3):95.
13. Usta M, Urganci N. Upper Gastrointestinal Bleeding in Children: The Role of Helicobacter pylori Infection and Non-steroidal Anti-inflammatory Drug Use. *The West Indian Medical Journal*. 2015;64(2):113.
14. Motamed F, Doroudian R, Najafi M, Monajemzade M, Marashi SM, Arastoo L, et al. Helicobacter pylori infection: Clinical, Endoscopic and Pathological findings in Iranian children. *International Journal of Pediatrics*. 2014;2(3.2):9-17.
15. Varanasi RV, Fantry GT, Wilson KT. Decreased prevalence of Helicobacter pylori infection in gastroesophageal reflux disease. *Helicobacter*. 1998;3(3):188-94.
16. Javid G, Zarger SA, Wani MA, Gulzar GM, Singh J, Sodhi MAK, et al. Upper Gastrointestinal Bleeding in Children in Kashmir (India): An Analysis of Causes, Characteristics and Outcome. *Journal of Digestive Endoscopy*. 2010;1(3):145-50.
17. Prasad K, Thapa B, Sharma A, Nain C, Singh K. Reassessment of diagnostic value of antral nodularity for Helicobacter pylori infection in children. *Minerva gastroenterologica e dietologica*. 2008;54(1):1-6.
18. Luzza F, Pensabene L, Imeneo M, Mancuso M, Contaldo A, Giancotti L, et al. Antral nodularity identifies children infected with Helicobacter pylori with higher grades of gastric inflammation. *Gastrointestinal endoscopy*. 2001;53(1):60-4.

19. Cárdenas-Mondragón MG, Carreon-Talavera R, Camorlinga-Ponce M, Gomez-Delgado A, Torres J, Fuentes-Panana EM. Epstein Barr virus and Helicobacter pylori co-infection are positively associated with severe gastritis in pediatric patients. *PLoS One*. 2013;8(4):e62850.
20. Kori M, Gladish V, Ziv-Sokolovskaya N, Huszar M, Beer-Gabel M, Reifen R. The significance of routine duodenal biopsies in pediatric patients undergoing upper intestinal endoscopy. *Journal of clinical gastroenterology*. 2003;37(1):39-41.
21. Brandt LJ, Feldman M, Friedman LS, Sleisenger MH. Sleisenger & Fordtran's Gastrointestinal and Liver Disease: Pathophysiology, Diagnosis, Management: Saunders; 2006.
22. Malcolm C, MacKay W, Shepherd A, Weaver L. Helicobacter pylori in children is strongly associated with poverty. *Scottish medical journal*. 2004;49(4):136-8.
23. Crone J, Gold BD. Helicobacter pylori infection in pediatrics. *Helicobacter*. 2004;9(s1):49-56.