

The Relationship between *Helicobacter pylori* Infection and Dyslipidemia in Hypertension: Case-Control Model

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Abstract

Background: Hypertension and/or *Helicobacter pylori* infections have an increasing prevalence in populations. *H. pylori* normally colonizes the stomach due to its ability to resist high acidity. Under abnormal conditions, it can cause infection through various virulence factors. **Objective:** The current study aims to determine the relationship between *H. pylori* IgG prevalence and lipid profile in hypertension using a case-control model. **Materials and Methods:** A total of 68 cases (34 hypertension and 34 healthy) were attended to at Al-Hakeem General Hospital in AL-Najaf Province, Iraq, during the period from July 2022 to September 2022, with an age range of 35–65 years. The blood specimens were aspirated and tested for *H. pylori* IgG using a rapid test (Lungene, China) and for lipid profiles using RX Daytona Analyzer (Randox, United Kingdom), following the instructions of manufacturing companies. **Results:** The mean age was significantly higher in hypertension than in control ($P = 0.033$), with the predominant age group in hypertension patients being 56–65 years. This study revealed that *H. pylori* was insignificantly higher in hypertension than in control group ($P = 0.089$). The lipid profile (cholesterol, triglycerides, and low-density lipoprotein) in hypertension patients was significantly $P = 0.000$, $P = 0.000$, and $P = 0.000$ higher in both the presence and absence of anti-*H. pylori* IgG ($P = 0.000$) than in control group, respectively. However, high-density lipoprotein level was insignificantly different between patients and controls in both presence ($P = 0.394$) and absence ($P = 0.133$) of anti-*H. pylori* IgG. **Conclusion:** Lipid profile abnormalities are related to hypertension, without difference in the presence and absence of anti-*H. pylori* IgG by case-control model.

Keywords: Dyslipidemia, *Helicobacter pylori*, hypertension, Iraq, lipid profile

INTRODUCTION

Helicobacter pylori is a gram-negative, motile, microaerophilic bacterium that colonizes the stomach as a part of normal flora. However, it can lead to a wide range of disorders, including autoimmune diseases (inflammatory bowel disease, autoimmune metabolic diseases, autoimmune liver diseases, rheumatic diseases, and autoimmune dermatosis),^[1] gastrointestinal diseases (dyspepsia, gastritis, peptic ulcer disease, gastric cancer, and mucosa-associated lymphoid tissue lymphoma), and extra-gastric manifestations (iron deficiency anemia and idiopathic thrombocytopenic purpura).^[2] Moreover, it plays a role in other *H. pylori* extra-gastric disorders, including typhoid fever,^[3] dyslipidemia disease,^[4] and hypertension.^[5]

This wide range of different diseases may be due to the large number of virulence factor; *H. pylori* has 22 reported virulence factors such as urease, flagellum, catalase, heat shock proteins, and lipopolysaccharide.^[6] “Dyslipidemia is defined as the imbalance of lipids such as cholesterol, low-density lipoprotein cholesterol (LDL-C), triglycerides (TG), and high-density lipoprotein (HDL), resulting from primary (genetic) or secondary (environmental) factors”.^[7] It is classified according to clinical practice, phenotype, and pathogenesis.^[8]

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Submission: 14-Aug-2023 **Accepted:** 21-Jan-2024 **Published:** 23-Jan-2026

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How to cite this article: Hussein AR, Almayali EJB. The relationship between *Helicobacter pylori* infection and dyslipidemia in hypertension: Case-control model. Med J Babylon 2025;22:1334-8.

Access this article online

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DOI:

10.4103/MJBL.MJBL_1187_23

A meta-analysis has shown that *H. pylori* infection is positively related to LDL-C, total cholesterol, and TG and negatively to HDL-C.^[9] Moreover, Yue *et al.*'s study^[5] suggested that *H. pylori* infection increases the risk of hypertension. This evidence implies that *H. pylori* infection may be correlated with dyslipidemia and hypertension.^[10]

However, most published studies are performed in the form of a cross-sectional model. Moreover, in Iraq, there is no available case-control report on the role of *H. pylori* seropositivity and lipid profile in hypertension. Therefore, this study aims to evaluate the relationship between *H. pylori* IgG seropositivity (for increased sensitivity) and lipid profile in hypertensive Iraqi individuals.

MATERIALS AND METHODS

Collection of specimens

A total of 68 cases (34 hypertension and 34 healthy) were attended to at Al-Hakeem General Hospital in AL-Najaf Province, Iraq, during the period from July 1, 2022 to September 30, 2022, with an age range of 35–65 years. Patients with signs and symptoms of hypertension were selected only if they had high blood pressure readings; those with normal blood pressure were excluded. They were then directed for lipid profile and *H. pylori* IgG assessment. Control individuals with normal blood pressure and without any clinical signs and symptoms of hypertension were selected, and those found to have high blood pressure were excluded. Furthermore, any subjects (patients and controls) who were supposed to have any other disorder were excluded from this study. The blood specimens from both the patient and control groups were collected, centrifuged, and transferred into two tubes as follows:

- One tube was used for the detection of *H. pylori* by a rapid test.
- Another tube was frozen at -20°C in a deep freeze for the determination of lipid profile levels at a later time.

H. pylori diagnosis

H. pylori was serologically diagnosed by a rapid test, which includes immobilized anti-human IgG and was performed according to the instructions of the manufacturing company (Lungene, China). If the specimen contains

H. pylori antibodies, a colored band will appear in the test line region, indicating a positive result..

Lipid profile

The lipid profile was studied using the RX Daytona Analyzer following the instructions provided by the manufacturing company (Randox, United Kingdom).

Statistical analysis

Statistical analysis was carried out using IBM SPSS Statistics 26 software. The statistical differences in qualitative parameters among different groups were detected using the Chi-square test. For quantitative data, mean $\pm$ SE were calculated, and differences were tested using *t* test for two means. Statistical significance was measured based on probability of $P < 0.05$.

Ethical approval

The study was performed in agreement with the ethical basics that have their origin in the Declaration of Helsinki. It was carried out with the verbal and analytical authorization of the patients before samples were taken. The study protocol, subject information, and consent form were evaluated and approved by a local ethics committee under document number 50,843 (issued in December 2021) to get this approval.

RESULTS

Hypertension and sex

The current study exhibited that out of 34 hypertension patients, 19 (55.9%) were females and 15 (44.1%) were males. For the 34 apparently healthy controls, 18 (52.9%) were females and 16 (47.1%) were males. The proportion of females in hypertension was insignificantly higher than that of males ($P = 0.808$), as shown in Table 1.

Hypertension and age

The age range of hypertension patients in the present study is 35–65 years, with a mean of 51.65 ± 1.4 years. For the control group, the age range is 35–64 years, with a mean of 47.02 ± 1.6 years. The mean age was significantly higher in hypertension than in control group ($P = 0.033$).

The subjects in this study were divided into three age sets (35–35, 46–55, and 56–65). The predominant age group in hypertension patients is 56–65, followed by 46–55, and then 35–45, whereas in control group, it is 35–45, followed

Table 1: Distribution of hypertension patient rendering to sex

Sex	Patient		Control		Total		P value
	No.	%	No.	%	No.	%	
Female	19	55.9	18	52.9	37	54.4	0.808
Male	15	44.1	16	47.1	31	45.6	
Total	34	100	34	100	68	100	

by 46–55, and then 56–65 [Table 2]. However, there are no significant differences ($P = 0.80$).

Hypertension and *H. pylori*

This study depended on a rapid test in qualitative (positive/negative) detection of *H. pylori* IgG in serum. This study revealed that 58.8% (20/34) of hypertension patients were *H. pylori* positive, and 41.2% (14/34) were *H. pylori* negative in serum. In the control group, 38.2% (13/34) were *H. pylori* positive, and 61.8% (21/34) were *H. pylori* negative. *H. pylori* is insignificantly higher in hypertension ($P = 0.089$) [Table 3].

Lipid profile in hypertension and control

Cholesterol level is significantly ($P = 0.000$) higher in hypertension (269.06 ± 7.7) than in control group (145.06 ± 4.4). Additionally, the TG level is significantly ($P = 0.000$) higher in hypertension (246.06 ± 14.2) than in control group (100.6 ± 2.9). However, HDL level is insignificantly ($P = 0.186$) lower in hypertension (38.8 ± 1.4) than in control group (41.2 ± 1.03). In respect of HDL, LDL level is significantly ($P = 0.000$) higher in hypertension (119.8 ± 4.3) than in control group (80.71 ± 1.4) [Table 4].

Lipid profile according to *H. pylori*

H. pylori-infected patients' serum showed a cholesterol level significantly ($P = 0.000$) higher in hypertension

(262.7 ± 7.95) than in control group (145.8 ± 7.3). The TG level was also significantly ($P = 0.000$) higher in hypertension (247 ± 17.99) than in control group (2.706 ± 0.440). However, the HDL level was insignificantly ($P = 0.394$) lower in hypertension (39.1 ± 2) than in control group (40.2 ± 1.96). The LDL level was significantly ($P = 0.000$) higher in hypertension (114.5 ± 5.5) than in control group (77.9 ± 195).

H. pylori non-infected patients' serum showed a cholesterol level significantly ($P = 0.000$) higher in hypertension (278.2 ± 14.8) than in control group (144.6 ± 5.7). The TG level was significantly ($P = 0.000$) higher in hypertension (246.5 ± 23.7) than in control group (104 ± 4.07). However, HDL level was insignificantly ($P = 0.133$) lower in hypertension (38.36 ± 1.8) than in control group (41.7 ± 1.2). The LDL level was significantly ($P = 0.000$) higher in hypertension (127.4 ± 6.8) than in control group (82.4 ± 1.8) [Table 5].

DISCUSSION

This study reports an insignificant relationship between the development of hypertension and gender. Alternatively, females appear to be more susceptible to hypertension than males. There is an insignificantly higher level of *H. pylori* IgG in patients, but a significantly higher lipid profile is observed in hypertension. The lipid profile is significantly higher in both subjects, irrespective of the presence or absence of anti-*H. pylori* IgG. The more valuable insight from the current study is the

Table 2: Distribution of hypertension patients according to age groups

Age	Patient		Control		Total		P value
	No.	%	No.	%	No.	%	
35-45	8	23.5	16	47.1	24	35.3	0.80
46-55	12	35.3	11	32.3	23	33.8	
56-65	14	41.2	7	20.6	21	30.9	
Total	34	100	34	100	68	100	

Table 3: Distribution of *H. pylori* seropositivity in hypertension and control

<i>H. pylori</i> infection	Patient		Control		Total		P value
	No.	%	No.	%	No.	%	
<i>H. pylori</i> +ve	20	58.8	13	38.2	33	48.5	0.089
<i>H. pylori</i> -ve	14	41.2	21	61.8	35	51.5	
Total	34	100	34	100	68	100	

Table 4: Lipid profile in hypertension and control

Lipid profile	Cholesterol	TG	HDL	LDL
Patient	269.06 ± 7.7	246.06 ± 14.2	38.8 ± 1.4	119.8 ± 4.3
Control	145.06 ± 4.4	100.6 ± 2.9	41.2 ± 1.03	80.71 ± 1.4
P value	0.000	0.000	0.186	0.000

TG: triglycerides, HDL: high-density lipoprotein, LDL: low-density lipoprotein

Table 5: Lipid profile according to *H. pylori*

Lipid profile		Cholesterol	TG	HDL	LDL
<i>H. pylori</i> +ve	Patient	262.7 ± 7.95	247 ± 17.99	39.1 ± 2	114.5 ± 5.5
	Control	145.8 ± 7.3	93.9 ± 3.2	40.2 ± 1.96	77.9 ± 195
	P value	0.000	0.000	0.69	0.000
<i>H. pylori</i> -0ve	Patient	278.2 ± 14.8	246.5 ± 23.7	38.36 ± 1.8	127.4 ± 6.8
	Control	144.6 ± 5.7	104.8 ± 4.07	41.7 ± 1.2	82.4 ± 1.8
	P value	0.000	0.000	0.133	0.000

TG: triglycerides, HDL: high-density lipoprotein, LDL: low-density lipoprotein

absence of any role of *H. pylori* in elevated lipid profiles in hypertensive cases.

The study by Xiong *et al.*^[11] revealed a significant susceptibility of females to hypertension. Moreover, Blood *et al.*,^[12] in the USA, found that 56% of 10,803 hypertension patients were female. The current study reports the mean age of patients as 51.65 years old, while a meta-analysis^[13] revealed that the mean age across different populations is between 42.58 and 68.05 years old. Another study^[11] also reports significantly older hypertensive subjects than control subjects.

The current case-control study revealed insignificantly higher *H. pylori* IgG in hypertension subjects than in control group, as reported by Fang *et al.*^[13] Their study indicated that *H. pylori*-infected individuals had a 13.4% higher risk for hypertension than uninfected individuals. The previous non-case-control study reports a significant relationship between *H. pylori* and hypertension. This is evident in a meta-analysis by Huang *et al.*,^[14] as well as in cross-sectional studies conducted by Xiong *et al.*^[11] and Soomro *et al.*^[15]

Kountouras *et al.*^[16] proposed several pathogenic mechanisms of hypertension induced by *H. pylori*, including metabolic and liver disorders such as augmented oxidative stress, risen insulin resistance, gastrointestinal dysbiosis, raised vasoconstriction, and decreased vasodilation, alongside genetic and epigenetic alterations. *H. pylori* can affect a variety of biomarkers such as the lipid profile, Th1/Th2 balance, activation of platelet, and autoimmunity, which are essential for cardiovascular disease.^[17] In other words, there is a relationship between the intake of salt, which is a critical agent of hypertension, and *H. pylori* infection.^[18]

The variations in results compared to other studies may be attributed to differences in study design, sample size, differences in diagnosis tests, or the possibility of false positives. It is worth considering that hypertension patients, who are typically older, may have undergone chemotherapy with potential side effects on the stomach, rendering them more susceptible to *H. pylori* infection.^[19]

The current case-control study shows higher lipid profile results in both cases with the presence and absence of anti-*H. pylori* IgG between hypertension

and control, as follows: cholesterol, TG, and HDL were significantly higher in hypertension, and HDL level was insignificantly higher in control group. There is a study^[20] that shows *H. pylori* infection is linked with an high risk of dyslipidemia in the Cameroon community. This study also indicates that *H. pylori* infection is an independent risk factor for coronary heart disease and reveals that the pathogen might directly invade the cardiovascular wall. There is evidence indicating that *H. pylori* can lead to altered lipid profiles.^[21] Hashim *et al.*^[22] registered that more than two-thirds of the *H. pylori* seropositive sets revealed dyslipidemia in at least one lipid profile.

Most research on the relationship between *H. pylori* and dyslipidemia has been conducted on random samples using a cross-sectional method. In contrast, the current case-control study focused on the hypertension population. Thus, there is a consensus that the evidence supporting the role of *H. pylori* in the development of coronary artery disease is misleading, and no definite signs of benefit have been shown in eliminating the organism or preventing cardiovascular consequences.^[23]

The current study recommended performing more studies to confirm if there is a role of *H. pylori* in hypertension and dyslipidemia.

CONCLUSIONS

This study concludes that *H. pylori* is insignificantly higher in hypertension than in control. Additionally, the lipid profile (cholesterol, TG, and LDL) is significantly higher in both cases, regardless of the presence and absence of anti-*H. pylori* IgG, in hypertension patients.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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