# Environmental Risk Factors Affecting the Prevalence and Transmission of Helicobacter Pylori Infection

# Ву

### Lilian Nabil Naoum Hanna

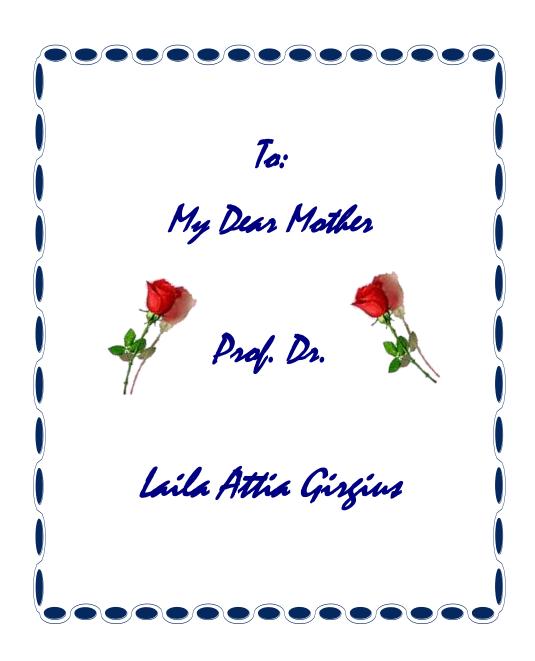
M.B.,B.Ch- Ain shams university,1994 M.Sc of Internal Medicine- Ain shams university, 2001

# **Thesis**

Submitted in Partial Fulfillment of the Requirement for the Doctor of Philosophy in Environmental Science

Department of Medical Science Institute of Environmental Studies & Research Ain Shams University.

2009



# LIST OF CONTENTS

Pag	зe
AcknowledgementI	
List of abbreviationsII	
List of tablesIII	
List of figuresV	
Introduction and aim of work1	
Review of literature:	
Chapter 1: Helicobacter pylori5	
Chapter 2: Diseases associated with Helicobacter pylori36	
Chapter 3: Epidemiological factors affecting <i>Helicobacter</i> pylori infection	
Subjects and methods60	
Results70	
Discussion	
Summary and Conclusion107	
Recommendations111	
References	
Appendix	
Arabic summary	



### ACKNOLEDGMENT



First of all, thanks to **GOD** for all the countless gifts I have been offered, for blessing me this work and for giving me a precious hand so as to be able to fulfill it.

The author would like to express sincere thanks to **Prof. Dr. Moustafa Hassan Said Ragab**, Head of Department of Medical Science, Institute of Environmental Studies &Research, Ain- shams university for suggesting the point, supervising the work and fruitful discussion all through the work. I feel very grateful for his sincere guidance and critical suggestions.

I would also like to direct my deep appreciation to **Dr. Hala Ibrahim Awdalla**, Assistant professor in the Department of Medical Science, Institute of Environmental Studies &Research, Ain-Shams University for supervising the present work, revising the manuscript, invaluable assistance and continuous encouragement.

I would finally like to express my deep appreciation and obligation to my Husband and daughter for their cordial sacrifices and encouragement.

### **ABSTRACT**

**Background**: *Helicobacter pylori (H. pylori)* infection varies remarkably between and within populations suggesting the role of socioeconomic- related environmental factors.

The aim of this study was to determine the prevalence of *Helicobacter pylori* among the studied population, to explore the effect of some environmental risk factors and life style on the transmission of infection, and to study the association between this bacterium and various non contagious diseases.

**Method:** This cross sectional study was conducted over six months. Two hundreds apparently healthy individuals from rural (Unshass) and urban (Nasr-city) areas voluntarily participated in this study. A full detailed clinical assessment in the form of questionnaire was designed for each individual to determine risk factors with specific emphasis to age, sex and residence. Seroprevalence of *H. pylori* IgG antibodies was evaluated using enzyme linked immunosorbent assay (ELISA). ECG, Liver function test, Random blood sugar and Hb level was performed to all participants.

**Results:** 28 % of the urban group was *H. pylori* positive versus 54% of rural group. A significant relation between *H. pylori* and poor sanitary conditions was detected; there was an association between *H. pylori* prevalence and gastric diseases, hepatic diseases and diabetes.

**Conclusion**: *Helicobacter pylori* infection was nearly twice in the rural group than in urban one.

## **List Of Abbreviations**

Cag - A : Cytotoxin Associated Gene **ECG** : Electrocardiograph **ELISA** : Enzyme linked immunosorbent assay H. pylori : Helicobacter pylori Hb level : Hemoglobin level Hsp : Heat shock proteins. **IgG** : Immunoglobulin G : Infantile hypertrophic pyloric stenosis **IHPS LPS** : Lipopolysaccharide **OD** : Optical density. **PAMPS** : Pathogen-associated molecular patterns. **PCR** : Polymerase chain reaction **PPI** : Proton pump inhibitors : Serum glutamic oxaloacetic transaminase **SGOT SGPT** : Serum glutamic pyruvic transaminase. **TLR** : Tol1-lik receptors Vac-A : Vacuolating Cytotoxin **WHO** : World Health Organization **OMP** Outer membrane protein:

# LIST OF TABLES

<u>Page</u>
Table 1: Characteristics of <i>Helicobacter pylori</i> 11
Table 2: Comparative accuracy and availability of tests for Helicobacter pylori infection Error! Bookmark not defined.
Table 3: Dual Therapy with amoxicillin. Error! Bookmark not defined.
Table 4: Dual Therapy with Clarithromycin Error! Bookmark not defined.
Table 5: Low – Dose 1 week triple therapies Error! Bookmark not defined.
Table 6: Triple therapy combination with Amoxicillin and metronidazole Error! Bookmark not defined.
Table 7: Socio-demographic characteristics of the studied population Error! Bookmark not defined.
Table 8: Hygienic environment and habits of the studied population Error! Bookmark not defined.
Table 9: Smoking differences between urban and rural groups Error! Bookmark not defined.
Table 10: Aspirin intake in urban and rural groups Error! Bookmark not defined.
$Table\ 11: Non\ communicable\ diseases\ of\ the\ studied\ population \textbf{Error!}\ \textbf{Bookmark}\ \textbf{not}\ \textbf{defined.}$
Table 12: <i>Helicobacter pylori</i> difference between urban and rural groups Error! Bookmark not defined.
Table 13: Relationship between <i>Helicobacter pylori</i> and age differences in studied population <b>Error! Bookmark not defined.</b>
Table 14: Relationship between <i>Helicobacter pylori</i> and gender in studied population <b>Error! Bookmark not defined.</b>
Table 15: Relationship between <i>Helicobacter pylori</i> and blood group in studied population <b>Error! Bookmark not defined.</b>
Table 16: Relationship between <i>Helicobacter pylori</i> and crowding index in studied population <b>Error! Bookmark not defined.</b>
Table 17: Relationship between <i>Helicobacter pylori</i> and income level in studied population <b>Error! Bookmark not defined.</b>
Table 18: Relationship between <i>H. pylori</i> and education level in studied population Error! Bookmark not defined.
Table 19: Relationship between <i>Helicobacter pylori</i> and occupation in studied population <b>Error! Bookmark not defined.</b>

# LIST OF TABLES (Cont.)

<u>Page</u>
Table 20: Relationship between <i>H. pylori</i> and presence of running water in studied groups <b>Error! Bookmark not defined.</b>
Γable 21: Relationship between Helicobacter pylori and sewage in studied populationError! Bookmark not defined.
Γable 22: Relationship between <i>Helicobacter pylori</i> and hand washing in studied population <b>Error! Bookmark not defined.</b>
Table 23: Relationship between <i>H. pylori</i> and sharing towels and eating utensils in studied population <b>Error! Bookmark not defined.</b>
Table 24: Relationship between <i>Helicobacter pylori</i> and vegetables washing in studied population <b>Error! Bookmark not defined.</b>
Γable 25: Relationship between <i>Helicobacter pylori</i> and house cleaning in studied population <b>Error! Bookmark not defined.</b>
Table 26: Relationship between <i>Helicobacter pylori</i> and pets in studied population
Table 27: Relationship between <i>Helicobacter pylori</i> and flies in studied population
Γable 28: Relationship between <i>Helicobacter pylori</i> and smoking in studied populationError! Bookmark not defined.
Γable 29: Relationship between <i>Helicobacter pylori</i> and aspirin intake in studied population <b>Error! Bookmark not defined.</b>
Table 30: Relationship between <i>Helicobacter pylori</i> and gastric diseases <b>Error! Bookmark not defined.</b>
Γable 31: Relationship between Helicobacter pylori and ischemic heart disease Error! Bookmark not defined.
Γable 32: Relationship between <i>Helicobacter pylori</i> and diabetes mellitus <b>Error! Bookmark not defined.</b>
Table 33: Relationship between <i>Helicobacter pylori</i> and hepatic diseases <b>Error! Bookmark not defined.</b>
Γable 34: Relationship between <i>Helicobacter pylori</i> and

rheumatic diseases ..... Error! Bookmark not defined.

Table 35: Relationship between *Helicobacter pylori* and anemia Error! Bookmark not defined.

# LIST OF FIGURES

<u>Page</u>
Figure 1: A 10.000x computer- aided image of <i>Helicobacter</i> pylori
Figure 2: <i>Helicobacter pylori</i> invade epithelial cells
Figure 3: <i>H. pylori</i> colonized on the surface of regenerative epithelium (image from Warthin-Starry's silver stain)21
Figure 4: A three day culture of <i>HP</i> on blood agar23
Figure 5: <i>Helicobacter pylori</i> positivity between urban and rural groups Error! Bookmark not defined.
Figure 6: Relationship between <i>Helicobacter pylori</i> and age differences in studied population <b>Error! Bookmark not defined.</b>
Figure 7: Relationship between <i>Helicobacter pylori</i> and gender in studied population
Figure 8: Relationship between <i>Helicobacter pylori</i> and blood group in both urban and rural groups <b>Error! Bookmark not defined.</b>
Figure 9: Relationship between <i>Helicobacter pylori</i> and crowding index in studied population <b>Error! Bookmark not defined.</b>
Figure 10: Relationship between <i>Helicobacter pylori</i> and income level in studied population <b>Error! Bookmark not defined.</b>
Figure 11: Relationship between <i>Helicobacter pylori</i> and education level in studied population <b>Error! Bookmark not defined.</b>
Figure 12: Relationship between <i>Helicobacter pylori</i> and occupation in studied population <b>Error! Bookmark not defined.</b>
Figure 13: Relationship between <i>H. pylori</i> and presence of running water studied groups <b>Error! Bookmark not defined.</b>
Figure 14: Relationship between <i>Helicobacter pylori</i> and sewage in studied population
Figure 15: Relationship between <i>Helicobacter pylori</i> and hand washing in studied population <b>Error! Bookmark not defined.</b>

# LIST OF FIGURES (Cont.)

### **Page**

- Figure 16: Relationship between *Helicobacter pylori* and sharing towels and eating utensils in studied population**Error! Bookmark not defined.**
- Figure 17: Relationship between *Helicobacter pylori* and vegetables washing in studied population**Error! Bookmark not defined.**
- Figure 18: Relationship between *Helicobacter pylori* and house cleaning in studied population**Error! Bookmark not defined.**

- Figure 21: Relationship between *Helicobacter pylori* and smoking in studied population**Error! Bookmark not defined.**
- Figure 22: Relationship between *Helicobacter pylori* and aspirin intake in studied population. **Error! Bookmark not defined.**
- Figure 23: Relationship between *Helicobacter pylori* and gastric diseases ...... **Error! Bookmark not defined.**
- Figure 24: Relationship between *Helicobacter pylori* and ischemic heart disease........... Error! Bookmark not defined.
- Figure 25: Relationship between *Helicobacter pylori* and diabetes mellitus ...... **Error! Bookmark not defined.**
- Figure 26: Relationship between *Helicobacter pylori* and hepatic diseases ...... **Error! Bookmark not defined.**
- Figure 27: Relationship between *Helicobacter pylori* and rheumatic diseases ...... **Error! Bookmark not defined.**
- Figure 28: Relationship between *Helicobacter pylori* and anemia**Error! Bookmark not defined.**

### INTRODUCTION

Helicobacter pylori (H. pylori) is a very common bacterium. It is present through out the world (Brown et al., 2002), approximately 50 percent of industrialized countries population and 90 percent of the developing countries populations has been estimated to be infected (Brown, 2000). In rural Egyptian villages the overall prevalence of anti H. pylori immunoglobulin G (IgG) was 91.7% with different degrees of positivity: 40.8% mild, 39.2% moderate and 11.7% severe (Zakaria, 2004).

Helicobacter pylori is a major cause of various gastroduodenal diseases. The infection has an association with chronic histological gastritis, peptic ulcer, gastric cancer and MALT lymphoma in the stomach.

All patients with *H. pylori* infection have histological gastritis and most of them remain asymptomatic for life, but some infected individuals shows ulceration or gastric cancer. Concerning gastric cancer, World Health Organization (WHO) concluded in 1994 that *Helicobacter pylori* is a definite carcinogen in humans on the basis of epidemiological studies (*Sugiyama et al., 2004*).

This bacterium is also involved in the pathogenesis of several extra-gastric diseases, such as coronaritis, gastroesophageal reflux disease, liver adenomas and carcinomas (*Suerbaum et al.*, 2003), iron deficiency anemia, skin disease and Rheumatological conditions (*Luigi and Giuseppe*, 2005), also its association with diabetes mellitus is under study (*Gulcelik et al.*, 2005).

The cure of *Helicobacter pylori* infection prevents recurrence of peptic ulcer without persistent treatments against ulceration (*Sugiyama et al.*, 2004).

The rate of acquisition of *H. pylori* infection varies remarkably between and within populations. This variation suggests that cultured background and socioeconomic- related environmental factors may play a role in its acquisition and transmission (*Wizla-Derambure*, 2001).

There is a great controversy about the mode of transmission of *H. pylori* and the presence of other reservoirs, H. pylori is usually spread by the fecal-oral route but possibly also by the oral-oral route and the spread of contaminated secretions. Thus, in developing countries, individuals catch *H. pylori* at a very young age from other persons (children) in their environment. In developed countries, *H. pylori* is more difficult to acquire and is usually transmitted from one family member to another, possibly by the fecal-oral route, or by the oral-oral route, e.g., kissing, vomitus. On occasion, transmission occurs from person to person via contaminated endoscopes. Other gastric Helicobacter-like organisms have now been observed in a variety of animals, including rodents, primates, swine, and

### Introduction and Aim of the Work

ferrets, but, with the exception of primates and possibly cats, these isolates are clearly different from human isolates. Food borne transmission would not be unusual (*Herrera*, 2004).

The influence of adult socioeconomic status, co-habitation, gender, smoking, coffee and alcohol intake on risk of *H. pylori* infection is uncertain. (*Moayyedi*, 2002).

It also seems that the pathogenesis of *H. pylori* infection is related to the virulence factors of the bacterium, environmental (dietary habits, hygiene, stress) and host factors (age, sex, blood type) (*Srp Arh Celok Lek*, 2004).

# **AIM OF WORK**

The aim of this study is to determine the prevalence of *Helicobacter pylori* in the studied population, to explore the effect of some environmental risk factors and life style on the transmission of infection, and to study the association between this bacterium and various non contagious diseases.