



Prevalence of *Helicobacter pylori* in Southern Part of Iran

Ramin Niknam,¹ Mohammad Reza Fattahi,¹ Masood Sepehrimanesh,^{2,*} and Alireza Safarpour¹

¹Gastroenterohepatology Research Center, Shiraz University of Medical Sciences, Shiraz, IR Iran

²Gastrointestinal and Liver Diseases Research Center, Guilan University of Medical Sciences, Rasht, IR Iran

*Corresponding author: Masood Sepehrimanesh, Gastrointestinal and Liver Diseases Research Center, Guilan University of Medical Sciences, Rasht, IR Iran. Tel: +98-1333535116, Fax: +98-1333534951, E-mail: sepehrimaneshmasood@gmail.com

Received 2017 October 02; Revised 2018 March 30; Accepted 2018 April 07.

Abstract

Background: There are clear variations between and within countries regarding the prevalence of *Helicobacter pylori*. In addition, there are no estimations of its prevalence in the general population in the south of Iran.

Objectives: We aimed to evaluate the prevalence of *H. pylori* in the rural population of Kavar, a southern city of Iran.

Methods: A random sample of 500 individuals were selected from the Kavar cohort study. Serum total IgG against *H. pylori* was checked. The serum positive patients recalled for stool sampling and the *H. pylori* stool antigen was checked using sandwich ELISA. Moreover, age and gender were recorded for all patients.

Results: Among the remained 441 who participated, 254 patients (57.6%) were positive for serum IgG. A total of 14 patients (5.5%) gave up due to stool sampling unsatisfaction. Overall, the rate of *H. pylori* infection based on both serum IgG and stool Ag of *H. pylori* was 41.5%. There were no significant association between gender and *H. pylori* stool antigen positivity ($P = 0.776$). Furthermore, no differences were detected between different ages categorized in positivity for stool antigen of *H. pylori* ($P = 0.327$). The mean and SD of age for negative and positive groups were 40.98 ± 15.42 and 43.65 ± 15.65 years, respectively ($P = 0.267$).

Conclusions: The prevalence of the *H. pylori* infection in our study based on serum IgG and stool Ag positivity was 41.5%, which are lower than many previous reported rates. Further studies in larger sample size and in different rural populations are needed.

Keywords: *Helicobacter pylori*, Rural Population, Stool Antigen, Cohort

1. Background

Helicobacter pylori is a common infectious organism, which affected humans, and has been known as the major cause of gastroduodenal disease include peptic ulcer, nodular gastritis, gastric premalignant lesions, gastric adenocarcinoma, and mucosa-associated lymphoid tissue (MALT) lymphoma (1, 2). In addition, the role of *H. pylori* infection in gastroesophageal reflux disease (GERD) is controversy (3). Approximately half of the world population was infected with *H. pylori* at the start of the 21st century (4, 5).

Today, the worldwide prevalence of *H. pylori* infection is ranged from 30% to 80% and the incidence of infection increases with age (6). In addition, the prevalence of *H. pylori* can be differing between geographical regions and populations. It can be 35% or less in developed countries of Europe and North America and reaches to 70% or more in developing countries of Asia and Latin America due to a difference in hygiene, living conditions, the spread of *H. pylori*, and the utilization of antimicrobial therapy (7-10). Despite of between countries variations, some studies showed the

difference in *H. pylori* prevalence within countries based on age and race differences (11, 12).

In Iran, as a developing country, the rate of the *H. pylori* infection is approximately high. It ranged from 19.2% in the city of Sari (capital of Mazandaran province in North of Iran) to 74.3 in Tehran (13). In addition, there is a significant difference in prevalence of *H. pylori* between different cities and provinces of Iran. This can be due to sampling from villages and cities, difference in age and gender, and some other demographic characteristics. In addition, difference in tests, which used for screening, may have an impact on the final reported prevalence. Serum antibody to *H. pylori* is tested in mass screening; however, stool antigen tests have been developed as a new non-invasive test (14). In addition, it has been reported that stool antigen test is a sensitive and specific mean to diagnose and screen the *H. pylori* infection (15). Different therapeutic regimen for eradication of *H. pylori* has been suggested due to antibiotic resistance of this infection (6, 16).

2. Objectives

The number of studies regarding the prevalence of *H. pylori* infection in the rural population is scarce. Therefore, the aims of the present study were to evaluate the prevalence of *H. pylori* infection in the rural population of Kavar region in the south of Iran based on both serum immunoglobulin and stool antigen tests.

3. Methods

3.1. Ethics Statement

The Ethical Committee of Shiraz University of Medical Sciences, Shiraz, Iran, approved the protocol of this study (95-01-13-12234).

3.2. Patients

Our patients were selected from the Kavar cohort study (KCS), which started from 2006 in Kavar with a population of about 71856 and a rural nature of lifestyle. For more information regarding this cohort study, you can read the following two studies, which were published by Fattahi and his team (17, 18). Based on the complete available data and entrance date, we randomly selected 500 subjects from the database of KCS. All selected subjects received oral information concerning the study and gave their written consent. Our exclusion criteria were (a) history of *H. pylori* eradication, (b) history of use of proton pump inhibitors, H₂-receptor antagonists, or antibiotics within the last 4 weeks prior to study, (c) history of gastric or esophageal surgery, and (d) patients with poor cooperation. Finally, 441 participants were enrolled in this study.

3.3. Serum Anti-*H. pylori* IgG Assay

Blood samples were taken from the cubical vein of each participant into sterile vacutainers without any anticoagulants. Then, the samples were centrifuge at 3500 × g for 10 min and obtained serum stored at -20 °C until used. Serum samples were analyzed by sandwich enzyme linked immunosorbent assay (ELISA) test for anti-*H. pylori* IgG antibody using a commercial test kit (AccuBind TM ELISA, Monobind, USA) according to instructions of the manufacturer. This test has a sensitivity of 0.1424 U/ml.

3.4. Stool Antigen of *H. pylori* Assessment

For confirmation of *H. pylori* infection, the feces samples were taken from seropositive patients and checked for stool Ag of *H. pylori*. We defined *H. pylori* as positive if both of these test results (serum anti-*H. pylori* IgG and stool antigen) were positive.

3.5. Statistical Analysis

The raw data were inserted into and analyzed by IBM SPSS Statistics version 23 (IBM Corp., Armonk, NY, USA). Kolmogorov-Smirnov test was used to check normal distribution of patients. Age difference between *H. pylori* positive and negative patients was checked using two independent sample t-test. The relationship between gender and *H. pylori* infection and between different age categories and *H. pylori* infection were analyzed using the Chi-square test. P < 0.05 was considered as a significant difference or relationship. To draw graphs, the data were inserted as mean, SD, and N or frequency into the GraphPad Prism 6.01 for Windows (GraphPad Software, Inc., CA, USA).

4. Results

The schematic chart of what happened and what we found in this study are presented in Figure 1.

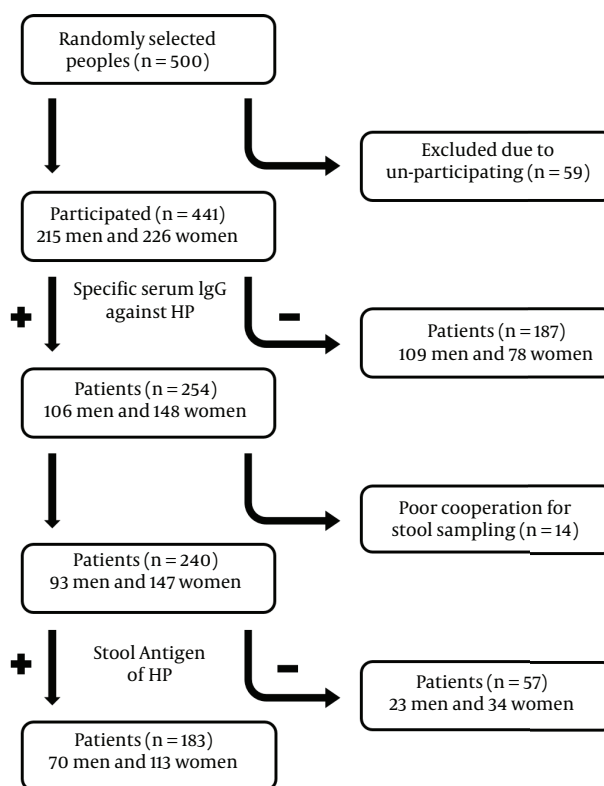


Figure 1. The fellow chart of the setting of this study and overall findings

In total, 57.6% of included patients were IgG positive against *H. pylori* and among them, 76.2% were also positive for stool Ag of *H. pylori*. Cooperation rate of patients was 94.5% and just 5.5% was not cooperated due to certain

problems for stool sampling. Overall, the rate of *H. pylori* infection based on both serum IgG and stool Ag of *H. pylori* was 41.5% (183 of 441 participated ones).

Comparison of the age of two groups of stool Ag negative and positive patients is presented in Figure 2. As shown there were no significant differences between these two groups regarding the age ($P = 0.267$).

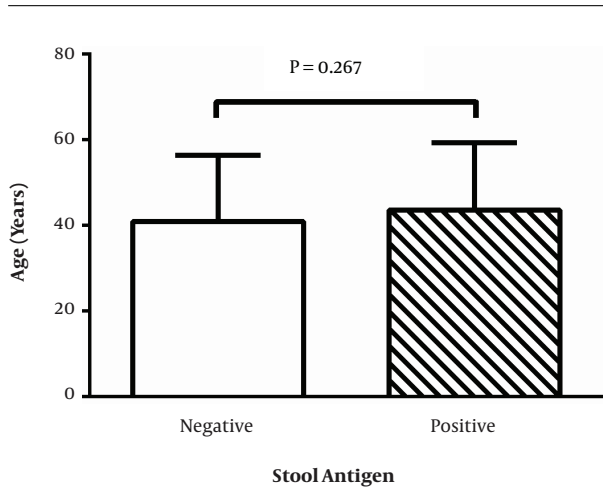


Figure 2. The mean and SD of the age in stool Ag negative and positive of *Helicobacter pylori*

The association of gender and infection of *H. pylori* based on stool Ag status is presented in Figure 3. As demonstrated, there is no significant relationship between gender and *H. pylori* infection and both gender showed a similar positive percentage of stool Ag of *H. pylori* ($P = 0.776$).

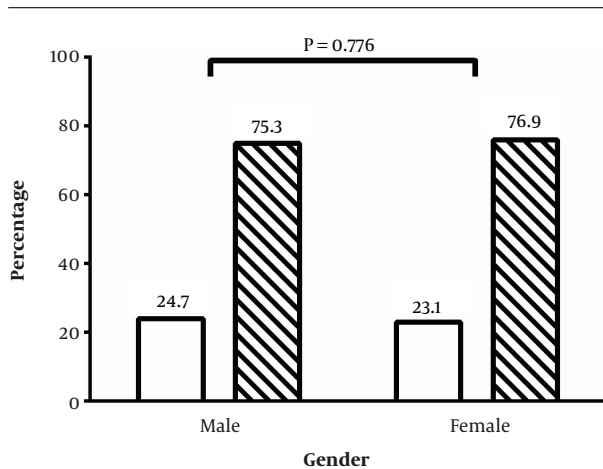


Figure 3. Association of gender and status of stool antigen of *H. pylori*. White and sketch bars are *H. pylori* stool Ag negative and positive groups, respectively

Finally, we categorized the patients into 7 age categories, ≤ 20 , 21 - 30, 31 - 40, 41 - 50, 51 - 60, 61 - 70, and ≥ 71 years.

Then, the percentage of two status of stool Ag of *H. pylori*, negative or positive, were compared between these age categories (Figure 4). Although, the most positive *H. pylori* patients belonged to the 41 - 50 years age category, no significant association was detected between age categories and of *H. pylori* infection ($P = 0.327$). Most patients were in the age category of 41 - 50 years (52 patients, 21.8%) and this category showed the most stool Ag positive patients (41 patients, 78.8%).

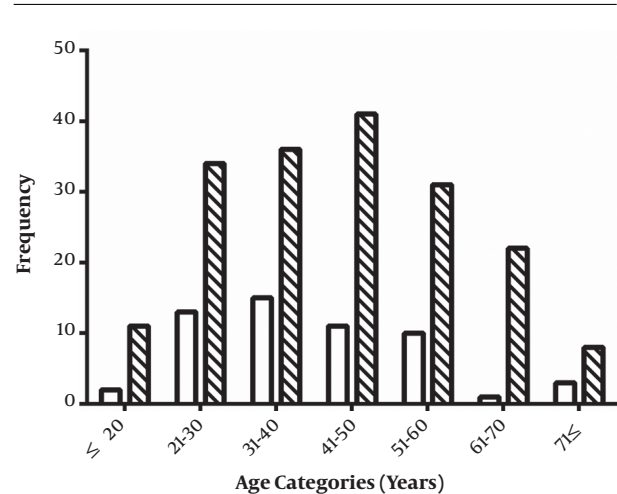


Figure 4. Frequency of stool antigen negative and positive patients in different age categories. White and sketch bars are *H. pylori* stool Ag negative and positive groups, respectively

5. Discussion

In the present study, the prevalence of the *H. pylori* infection based on positivity of anti-*H. pylori* IgG and stool Ag of *H. pylori* was evaluated in the rural population of the Kavar region, south Iran. The prevalence of the *H. pylori* infection in our study based on serum IgG and stool Ag positivity was 41.5%, which are lower than many previous reported rates. The reported values from developed and developing countries and from rural and urban populations regarding the prevalence of *H. pylori* infection show severe variations. A seropositivity of about 50% and near 90% in adults of developed and developing countries were reported previously (19, 20). In addition, the rate of rural the 13C-urea breath test based *H. pylori* infection of 57% was reported by Cheng et al., for rural Pinggu of Beijing, China (21). However, Dorji and colleagues reported that the overall prevalence of *H. pylori* infection based on the assessment of *H. pylori*-IgG using ELISA in Bhutan, remote Himalayan country between India and Tibet (China) with 70% of rural and agriculture based, was about 87% (22).

Despite the socioeconomic state, especially for countries with homogenous social class, the density of living is the most significant risk factor of *H. pylori* infection (23, 24), due to the fact that oral-oral and fecal-oral routes are the most common modes of *H. pylori* transmission. This factor, along with other above-mentioned factors, lead to higher prevalence of *H. pylori* infection in the rural population as other previous studies (21-24). In an old study in Shiraz, Iran, the infection rate of more than 85% of adults based on the measuring of IgG against *H. pylori* using ELISA was reported (25). However, it has been reported that *H. pylori* prevalence has now started to decline in Asia and the Middle East (26-28).

In a recent study using culturing, Gram staining, and rapid urease test, we reported the *H. pylori* positivity of 31% in 548 dyspeptic patients of Fars province with both rural and urban nature of lifestyle, Iran, which was surprisingly lower than other local reports (29). Regarding these differences, it can be said that high prevalence of *H. pylori* infection is related to socioeconomic status, poor living conditions, and poor management of drinking water (30, 31). Moreover, higher prevalence of *H. pylori* infection in rural against urban population may be due to the nature of the lifestyle, the source of the consumed water and lower socioeconomic state. As shown, it seems that the *H. pylori* prevalence is decreasing in both populations and this may be due to treatment strategies and increasing the knowledge of individuals regarding this bacterium.

Nonetheless, the stool Ag of *H. pylori* is qualitative test and the results are expressed as negative or positive, however, its results can be used as semi-quantitative measurement along with other parameters. As another result found in this study, the association of the positive stool Ag of *H. pylori* and age category was detected. The stool Ag positivity of *H. pylori* was related with age, increasing with age up to 41 - 50 years category and then decreasing. *Helicobacter pylori* is typically acquired in early childhood in developing countries and usually persist during lifetime if left untreated (29), however, the *H. pylori* infection rate increases with age in adults (32). Although, serum assaying of anti-*H. pylori* IgG or IgA antibodies could be used to determine prevalence of infection (33), stool antigen test, as a non-invasive test, could be used both safely and cost effectively to screen patients (34, 35). Regarding the decline in the prevalence of *H. pylori* in elderly population, a marked reduction in the *H. pylori* prevalence in elderly people with age more than 85 years are also previously reported (36, 37). There are several reasons for this observation, which were mentioned previously as existence of chronic atrophic gastritis in higher age, plus the severe use of antibiotics and other related drugs in this population age group (37).

6. Conclusions

Among the limitations of the current study, the following can be mentioned: 1) lack of any molecular confirmation tests of *H. pylori* existence such as PCR of specific genes; 2) no endoscopic evaluation of these patients and performing no antral biopsies for *H. pylori* checking by rapid urease test or culture. Despite these limitations, data from the present study show that the prevalence of *H. pylori* infection in rural population is still high and this may be related to the socioeconomic state of this population. However, no significant difference was detected between two genders in prevalence of *H. pylori* infection. Performing further studies on the gastrointestinal complications related to *H. pylori* infection in this rural population, in larger sample size, and in different rural populations is highly recommended as we direct in our center.

Acknowledgments

The authors would like to thank the Center for Development of Clinical Research of Nemazee Hospital, Shiraz University of Medical Sciences, Shiraz, Iran for editorial assistance. The authors also thank the staff of the Gastroenterohepatology Research Center for cooperation in the obtaining data and performing the laboratory tests. In addition, we would gratefully like to thank all the patients who contributed to this study.

References

1. Niknam R, Manafi A, Maghbool M, Kouhpayeh A, Mahmoudi L. Is endoscopic nodular gastritis associated with premalignant lesions? *Neth J Med*. 2015;73(5):236-41. [PubMed: 26087803].
2. Niknam R, Manafi A, Fattahi MR, Mahmoudi L. The association between gastric endoscopic findings and histologic premalignant lesions in the Iranian rural population. *Medicine (Baltimore)*. 2015;94(17). e715. doi: 10.1097/MD.0000000000000715. [PubMed: 25929902]. [PubMed Central: PMC4603049].
3. Niknam R, Manafi A, Khazforoosh S, Mahmoudi L. Management of gastroesophageal reflux disease in adults. *Trend Pharmaceutical Sci*. 2015;1(2):65-74.
4. Bode G, Brenner H, Adler G, Rothenbacher D. Dyspeptic symptoms in middle-aged to old adults: the role of *Helicobacter pylori* infection, and various demographic and lifestyle factors. *J Intern Med*. 2002;252(1):41-7. [PubMed: 12074737].
5. Ortiz D, Cavazza ME, Rodriguez O, Hagel I, Correnti M, Convit J. Prevalence of *Helicobacter pylori* infection in Warao lineage communities of Delta Amacuro State, Venezuela. *Mem Inst Oswaldo Cruz*. 2003;98(6):721-5. [PubMed: 14595445].
6. Mousavi S, Pourabbas B, Niknam R. Susceptibility testing of *Helicobacter pylori*: Comparison of E-test and disk diffusion for metronidazole and mutations in *rdxA* gene sequences of *Helicobacter pylori* strains. *Trend Pharmaceutical Sci*. 2015;1(4):235-42.
7. Cave DR. How is *Helicobacter pylori* transmitted? *Gastroenterol*. 1997;113:S9-S14. [PubMed: 9394753].

8. Brown LM. Helicobacter pylori: epidemiology and routes of transmission. *Epidemiol Rev*. 2000;**22**:283-97. [PubMed: [11218379](#)].
9. Peleteiro B, Bastos A, Ferro A, Lunet N. Prevalence of Helicobacter pylori infection worldwide: a systematic review of studies with national coverage. *Dig Dis Sci*. 2014;**59**(8):1698-709. doi: [10.1007/s10620-014-3063-0](#). [PubMed: [24563236](#)].
10. Malaty HM. Epidemiology of Helicobacter pylori infection. *Best Pract Res Clin Gastroenterol*. 2007;**21**(2):205-14. doi: [10.1016/j.bpg.2006.10.005](#). [PubMed: [17382273](#)].
11. Nguyen T, Ramsey D, Graham D, Shaib Y, Shiota S, Velez M, et al. The Prevalence of Helicobacter pylori Remains High in African American and Hispanic Veterans. *Helicobacter*. 2015;**20**(4):305-15. doi: [10.1111/hel.12199](#). [PubMed: [25689684](#)].
12. Grad YH, Lipsitch M, Aiello AE. Secular trends in Helicobacter pylori seroprevalence in adults in the United States: evidence for sustained race/ethnic disparities. *Am J Epidemiol*. 2012;**175**(1):54-9. doi: [10.1093/aje/kwr288](#). [PubMed: [22085628](#)]. [PubMed Central: [PMC3244610](#)].
13. Sayehmiri F, Darvishi Z, Sayehmiri K, Soroush S, Emaneini M, Zarrilli R, et al. A Systematic Review and Meta-Analysis Study to Investigate the Prevalence of Helicobacter pylori and the Sensitivity of its Diagnostic Methods in Iran. *Iran Red Crescent Med J*. 2014;**16**(6). e12581. doi: [10.5812/ircmj.12581](#). [PubMed: [25068041](#)]. [PubMed Central: [PMC4102974](#)].
14. Shimoyama T, Oyama T, Matsuzaka M, Danjo K, Nakaji S, Fukuda S. Comparison of a stool antigen test and serology for the diagnosis of Helicobacter pylori infection in mass survey. *Helicobacter*. 2009;**14**(2):87-90. doi: [10.1111/j.1523-5378.2009.00672.x](#). [PubMed: [19298335](#)].
15. Megraud F, European Paediatric Task Force on Helicobacter P. Comparison of non-invasive tests to detect Helicobacter pylori infection in children and adolescents: results of a multicenter European study. *J Pediatr*. 2005;**146**(2):198-203. doi: [10.1016/j.jpeds.2004.10.044](#). [PubMed: [15689908](#)].
16. Mahmoudi L, Farshad S, Seddigh M, Mahmoudi P, Ejtehad F, Niknam R. High efficacy of gemifloxacin-containing therapy in Helicobacter Pylori eradication: A pilot empirical second-line rescue therapy. *Medicine (Baltimore)*. 2016;**95**(42). e4410. doi: [10.1097/MD.0000000000004410](#). [PubMed: [27759625](#)]. [PubMed Central: [PMC5079309](#)].
17. Fattahi MR, Niknam R, Safarpour A, Sepehrimanesh M, Lotfi M. The Prevalence of Metabolic Syndrome In Non-alcoholic Fatty Liver Disease; A Population-Based Study. *Middle East J Dig Dis*. 2016;**8**(2):131-7. doi: [10.15171/mejdd.2016.18](#). [PubMed: [27252820](#)]. [PubMed Central: [PMC4885612](#)].
18. Fattahi MR, Safarpour A, Sepehrimanesh M, Hosseini Asl SM, Mohamaddoust F. The prevalence of hepatitis C virus infection and its related risk factors among the rural population of fars province, southern iran. *Hepat Mon*. 2015;**15**(2). e24734. doi: [10.5812/hepatmon.24734](#). [PubMed: [25788957](#)]. [PubMed Central: [PMC4350250](#)].
19. Siavoshi F, Malekzadeh R, Daneshmand M, Ashktorab H. Helicobacter pylori endemic and gastric disease. *Dig Dis Sci*. 2005;**50**(11):2075-80. doi: [10.1007/s10620-005-3010-1](#). [PubMed: [16240218](#)].
20. Murakami K, Kodama M, Fujioka T. Latest insights into the effects of Helicobacter pylori infection on gastric carcinogenesis. *World J Gastroenterol*. 2006;**12**(17):2713-20. [PubMed: [16718758](#)]. [PubMed Central: [PMC4130980](#)].
21. Cheng H, Hu F, Zhang L, Yang G, Ma J, Hu J, et al. Prevalence of Helicobacter pylori infection and identification of risk factors in rural and urban Beijing, China. *Helicobacter*. 2009;**14**(2):128-33. doi: [10.1111/j.1523-5378.2009.00668.x](#). [PubMed: [19298340](#)].
22. Dorji D, Dendup T, Malaty HM, Wangchuk K, Yangzom D, Richter JM. Epidemiology of Helicobacter pylori in Bhutan: the role of environment and Geographic location. *Helicobacter*. 2014;**19**(1):69-73. doi: [10.1111/hel.12088](#). [PubMed: [24102940](#)].
23. Mitchell HM, Li YY, Hu PJ, Liu Q, Chen M, Du GG, et al. Epidemiology of Helicobacter pylori in southern China: identification of early childhood as the critical period for acquisition. *J Infect Dis*. 1992;**166**(1):149-53. [PubMed: [1607687](#)].
24. Malaty HM, Paykov V, Bykova O, Ross A, Graham DP, Anneger JF, et al. Helicobacter pylori and socioeconomic factors in Russia. *Helicobacter*. 1996;**1**(2):82-7. [PubMed: [9398883](#)].
25. Massarrat S, Saberi-Firooz M, Soleimani A, Himmelmann GW, Hitzges M, Keshavarz H. Peptic ulcer disease, irritable bowel syndrome and constipation in two populations in Iran. *Eur J Gastroenterol Hepatol*. 1995;**7**(5):427-33. [PubMed: [7614105](#)].
26. Farshad S, Japoni A, Alborzi A, Zarenezhad M, Ranjbar R. Changing prevalence of Helicobacter pylori in south of Iran. *Arch Clin Infect Dis*. 2010;**5**(2):65-9.
27. Tan HJ, Goh KL. Changing epidemiology of Helicobacter pylori in Asia. *J Dig Dis*. 2008;**9**(4):186-9. doi: [10.1111/j.1751-2980.2008.00344.x](#). [PubMed: [18959588](#)].
28. Yim JY, Kim N, Choi SH, Kim YS, Cho KR, Kim SS, et al. Seroprevalence of Helicobacter pylori in South Korea. *Helicobacter*. 2007;**12**(4):333-40. doi: [10.1111/j.1523-5378.2007.00504.x](#). [PubMed: [17669107](#)].
29. Niknam R, Seddigh M, Fattahi MR, Dehghanian A, Mahmoudi L. Prevalence of Helicobacter pylori in Patients With Dyspepsia. *Jundishapur J Microbiol*. 2014;**7**(10). e12676. doi: [10.5812/jjm.12676](#). [PubMed: [25632327](#)]. [PubMed Central: [PMC4295317](#)].
30. Webb PM, Knight T, Greaves S, Wilson A, Newell DG, Elder J, et al. Relation between infection with Helicobacter pylori and living conditions in childhood: evidence for person to person transmission in early life. *BMJ*. 1994;**308**(6931):750-3. [PubMed: [8142828](#)]. [PubMed Central: [PMC2539652](#)].
31. Perez-Perez GI, Rothenbacher D, Brenner H. Epidemiology of Helicobacter pylori infection. *Helicobacter*. 2004;**9** Suppl 1:1-6. doi: [10.1111/j.1083-4389.2004.00248.x](#). [PubMed: [15347299](#)].
32. Mishra S, Singh V, Rao GR, Dixit VK, Gulati AK, Nath G. Prevalence of Helicobacter pylori in asymptomatic subjects—a nested PCR based study. *Infect Genet Evol*. 2008;**8**(6):815-9. doi: [10.1016/j.meegid.2008.08.001](#). [PubMed: [18771754](#)].
33. Jafarzadeh A, Rezayati MT, Nemati M. Specific serum immunoglobulin G to H pylori and CagA in healthy children and adults (south-east of Iran). *World J Gastroenterol*. 2007;**13**(22):3117-21. [PubMed: [17589930](#)]. [PubMed Central: [PMC4172621](#)].
34. Soltani J, Amirzadeh J, Nahedi S, Shahsavari S. Prevalence of Helicobacter pylori infection in children, a population-based cross-sectional study in west of Iran. *Iran J Pediatr*. 2012;**23**(1):13-8. [PubMed: [23550042](#)]. [PubMed Central: [PMC3574986](#)].
35. Malfertheiner P, Megraud F, O'Morain C, Hungin APS, Jones R, Axon A. Current concepts in the management of Helicobacter pylori infection—The Maastricht 2-2000 Consensus Report. *Aliment Pharmacol Therapeut*. 2002;**16**(2):167-80.
36. Ford AC, Axon AT. Epidemiology of Helicobacter pylori infection and public health implications. *Helicobacter*. 2010;**15** Suppl 1:1-6. doi: [10.1111/j.1523-5378.2010.00779.x](#). [PubMed: [21054646](#)].
37. Salles-Montaudon N, Dertheil S, Broutet N, Gras N, Monteiro L, De Mascarel A, et al. Detecting Helicobacter pylori infection in hospitalized frail older patients: the challenge. *J Am Geriatr Soc*. 2002;**50**(10):1674-80. [PubMed: [12366621](#)].