

Can We Consider *H. Pylori* as A Risk Factor for Acute Coronary Syndrome? A Proposal Study for A Sample of Egyptian Population

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Abstract: *Helicobacter pylori* (*H. pylori*) is a Gram negative spiral bacterium which colonizes gastric mucosa of nearly half of human population was described by Marshall and Warren in 1984. A characteristic feature of *H. Pylori* infection is an excessive inflammatory response. The majority of *H. pylori* infections remain asymptomatic. However, still it leads to the development of histological gastritis with the recruitment of immune cells. About 10% of infected subjects develop symptomatic gastritis, erosions or peptic ulcer. Gastric cancer is the most severe consequence of *H. pylori* infection. Recently, a possible association between chronic infections with *H. pylori* and extragastric disorders - including coronary heart disease, has been intensively investigated. Here we will shed light on a possible association between chronic *H.Pylori* infection and one of the commonest Heart disease; Acute Coronary Syndrome (ACS).

Keywords: *H.Pylori*, ACS, Heart diseases, stomach.

1 Introduction

Coronary artery diseases (CADs) are leading cause of mortality and morbidity in the modern world, a major public health problem. Acute coronary syndrome is a crucial stage of the clinical manifestation of CAD and results in substantial morbidity and mortality [1-10]. However, research in ACS has propelled the field from one driven by anecdote to one guided by scientific evidence. For instance, pioneer findings identified the relationship of chronic infections influence on ACS and CADs [10-19]. The role of virus and bacterial pathogens including *helicobacter pylori* (*H.pylori*) are now considered as factors implicated in development of ACS [20-25]. Recently, possible association between *H.pylori* infection and extragastric disorders has been suggested. Knowledge on the etiology of atherosclerosis together with current findings in the area of *H. pylori* infections constitute the background for the newly proposed hypothesis that those two processes may be related. Many research studies confirm the indirect association between the prevalence of *H. pylori* and the occurrence of CHD. According to majority of findings the involvement of *H. pylori* in this process is based on the chronic inflammation which might facilitate the CHD related-pathologies. It needs to be elucidated, if the infection initiates or just

accelerates the formation of atheromatous plaque. *H.pylori* is the most common chronic bacterial infection of the human upper gastrointestinal tract. Recent evidence from Taiwan's national retrospective cohort study identified a greater relationship between *H.pylori* and ACS. However, a higher percentage of 91.7% seropositive *H.pylori* antibodies identified among Egyptian population [26-45]. Similarly, a national representative survey in Egypt found an adjusted overall prevalence of CAD of 8.3% [9]. Further investigating this among Egyptian sample may provide an interesting insight to clarify the relationship between *H.pylori* and ACS. Therefore, we aimed to investigate the correlation between ACS and chronic infection of *H.pylori* in a sample of Egyptian population.

2 Patients & Methods

100 patients presented with ACS with Positive *H.Pylori* test both by Eliza; IgM and IgG and Stool antigenicity will be enrolled to the current study. Smearing from infected coronaries or even thrombi aspirates will be examined for detection of Gram negative *H.Pylori* bacterium. Another 20 patients with ACS without *H.Pylori* infection, their coronaries 'thrombi will be examined for presence of other bacteria (Control group).

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3 Goals

- 1-If there is a direct correlation between Chronic H.Pylori infection and ACS (Dysfunction of coronary vascular endothelium by H.Pylori)
- 2-If there is cross antigenicity that make ACS related-chronic H.Pylori infection
- 3-Considering remodeling of risk factors related-ACS in the presence of H.Pylori.

4 Conclusions

Since classic risk factors do not explain all cases of ACS it has been suggested that chronic infections and even commensal microorganisms may affect the development or maintenance of ACS. Among various pathogens possibly involved in atherogenesis *H. pylori* is particularly interesting, since it induces chronic long-term infection within gastric epithelium which leads to not only local but systemic inflammation. Recent knowledge on the pathogenesis of atherosclerosis together with current findings in the field of *H. pylori* related diseases constitute the background for the newly proposed hypothesis that those two processes may be related, cross antigenicity or remodeling or risk factors would be new risk factors for ACS.

References

- [1] Libby P, Theroux P. Pathophysiology of coronary artery disease. *Circulation* 2005; 111: 3481-3488.
- [2] Kutuk O, Basaga H. Inflammation meets oxidation: NF-kappaB as a mediator of initial lesion development in atherosclerosis. *Trends Mol Med* 2003; 9: 549-557.
- [3] Rodella LF, Rezzani R. Endothelial and vascular smooth cell dysfunction: a comprehensive appraisal. In: Parthasarathy S. *Atherogenesis*. InTech under CC BY 3.0 licence, 2012: 105-134
- [4] Lai CY, Yang TY, Lin CL, Kao CH. Helicobacter pylori infection and the risk of acute coronary syndrome: a nationwide retrospective cohort study. *Eur J Clin Microbiol Infect Dis*. 2015;34(1):69-74.
- [5] Sharma V, Aggarwal A. Helicobacter pylori: Does it add to risk of coronary artery disease. *World J Cardiol*. 2015;7(1):19.
- [6] Budzyński J, Koziński M, Kłopotcka M, Kubica JM, Kubica J. Clinical significance of Helicobacter pylori infection in patients with acute coronary syndromes: an overview of current evidence. *Clin Res Cardiol*. 2014;103(11):855-86.
- [7] Eskandarian R, Ghorbani R, Shiyasi M, Momeni B, Hajifathalian K, Madani M. Prognostic role of Helicobacter pylori infection in acute coronary syndrome: a prospective cohort study: cardiovascular topics. *Cardiovasc J Afr*. 2012;23(3):131-5.
- [8] Khedmat H, Karbasi-Afshar R, Agah S, Taheri S. Helicobacter pylori Infection in the general population: A Middle Eastern perspective. *Caspian J Intern Med*. 2013;4(4):745.
- [9] Hassanin N, Gharib S, El Ramly MZ, Meged MA, Makram A. Metabolic syndrome and coronary artery disease in young Egyptians presenting with acute coronary syndrome. *Kasr Al Ainy Med J* 2015;21:27-33
- [10] Brevetti G, Giugliano G, Brevetti L, Hiatt WR. Inflammation in peripheral artery disease. *Circulation* 2010; 122: 1862-1875.
- [11] Sun J, Rangan P, Bhat SS, Liu L. A meta-analysis of the association between Helicobacter pylori infection and risk of coronary heart disease from published prospective studies. *Helicobacter*. 2016;21:11-23.
- [12] Said Mohammad Zade M, Eishi A, Behrozian R, Rahimi E. Relationship between Helicobacter pylori infection and cardiac syndrome X. *J Shahrekord University Med Sci* 2009;11(1):58-63.
- [13] Eskandarian R, Malek M, Mousavi SH, Babaei M. Association of Helicobacter pylori infection with cardiac syndrome X. *Singapore Med J* 2006;47(8):704.
- [14] Sung KC, Rhee EJ, Ryu SH, Beck SH. Prevalence of Helicobacter pylori infection and its association with cardiovascular risk factors in Korean adults. *Int J Cardiol* 2005;102:411-417.
- [15] Shin A, Shin HR, Kang D, Park SK, Kim CS, Yoo KY. A nested case-control study of the association of Helicobacter pylori infection with gastric adenocarcinoma in Korea. *Br J Cancer*. 2005;92:1273-1275.
- [16] Leja M, Cine E, Rudzite D, Vilkoite I, et al. Prevalence of Helicobacter pylori infection and atrophic gastritis in Latvia. *Eur J Gastroenterol Hepatol*. 2012;24:1410-1417.
- [17] Sasazuki S, Inoue M, Iwasaki M, Otani T, et al. Effect of Helicobacter pylori infection combined with caga and pepsinogen status on gastric cancer development among Japanese men and women: A nested case-control study. *Cancer Epidemiol Biomarkers Prev*. 2006;15:1341-1347.
- [18] Ferreccio C, Rollan A, Harris PR, Serrano C, et al. Gastric cancer is related to early Helicobacter pylori infection in a high-prevalence country. *Cancer Epidemiol Biomarkers Prev*. 2007;16:662-667.
- [19] Wang X, Terry P, Yan H. Stomach cancer in 67 Chinese counties: evidence of interaction between salt consumption and Helicobacter pylori infection. *Asia Pac J Clin Nutr*. 2008;17:644-650.
- [20] Torres J, Leal-Herrera Y, Perez-Perez G, Gomez A, et al. A community-based seroepidemiologic study of Helicobacter pylori infection in Mexico. *J Infect Dis*. 1998;178:1089-1094.
- [21] Chey WD, Leontiadis GI, Howden CW, Moss SF. ACG clinical guideline: treatment of Helicobacter pylori infection. *Am J Gastroenterol* 2017;112:212-238.
- [22] Proton Pump Inhibitors Information. 2015; <http://www.fda.gov/Drugs/DrugSafety/InformationbyDrugClass/ucm213259.html>. Accessed February 26, 2017.

- [23] Melloni C, Washam JB, Jones WS, Halim SA, Hasselblad V, Mayer SB, Heidenfelder BL, Dolor RJ. Conflicting results between randomized trials and observational studies on the impact of proton pump inhibitors on cardiovascular events when coadministered with dual antiplatelet therapy: systematic review. *Circ Cardiovasc Qual Outcomes*. 2015;8(1):47-55.
- [24] Kowalski M, Konturek PC, Pieniazek P, Karczewska E, Kluczka A, Grove R, Kranig W, Nasser R, Thale J, Hahn EG, Konturek SJ. Prevalence of *Helicobacter pylori* infection in coronary artery disease and effect of its eradication on coronary lumen reduction after percutaneous coronary angioplasty. *Dig Liver Dis* 2001;33(3):222-229
- [25] Budzynski J. The favourable effect of *Helicobacter pylori* eradication therapy in patients with recurrent angina-like chest pain and non-responsive to proton pump inhibitors: a preliminary study. *Arch Med Sci* 2011;7(1):73-80.
- [26] Jafarzadeh A, Nemati M, Tahmasbi M, Ahmadi P, Rezayati MT, Sayadi AR. The association between infection burden in Iranian patients with acute myocardial infarction and unstable angina. *Acta Med Indones* 2011;43(2):105-111
- [27] Chiu B. Multiple infections in carotid atherosclerotic plaques. *Am Heart J* 1999; 138: S534-S536
- [28] Espinola-Klein C, Rupprecht HJ, Blankenberg S, Bickel C, Kopp H, Victor A, Hafner G, Prellwitz W, Schlumberger W, Meyer J. Impact of infectious burden on progression of carotid atherosclerosis. *Stroke* 2002; 33: 2581-2586
- [29] Calabrese F, van der Wal AC, Levi M. Infection and inflammation in the cardiovascular system. *Cardiovasc Res* 2003; 60: 1-4
- [30] Jafarzadeh A, Nemati M, Tahmasbi M, Ahmadi P, Rezayati MT, Sayadi AR. The association between infection burden in Iranian patients with acute myocardial infarction and unstable angina. *Acta Med Indones* 2011; 43: 105-111.
- [31] Turnbaugh PJ, Ley RE, Hamady M, Fraser-Liggett CM, Knight R, Gordon JI. The human microbiome project. *Nature* 2007; 449:804-810
- [32] Hattori M, Taylor TD. The human intestinal microbiome: a new frontier of human biology. *DNA Res* 2009; 16: 1-12 .
- [33] Manco M, Putignani L, Bottazzo GF. Gut microbiota, lipopolysaccharides, and innate immunity in the pathogenesis of obesity and cardiovascular risk. *Endocr Rev* 2010; 31: 817-844
- [34] Rabizadeh S, Sears C. New horizons for the infectious disease specialist: how gut microflora promote health and disease. *Curr Infect Dis Rep* 2008; 10: 92-98.
- [35] Sanches, P. L. et al. Association of nonalcoholic fatty liver disease with cardiovascular risk factors in obese adolescents: the role of interdisciplinary therapy. *J. Clin. Lipidol.* 8, 265-272 (2014).
- [36] Torun, E. et al. Carotid intima-media thickness and flow-mediated dilation in obese children with non-alcoholic fatty liver disease. *Turk. J. Gastroenterol.* 25 (Suppl 1), 92-98 (2014).
- [37] Musso, G. et al. Dietary habits and their relations to insulin resistance and postprandial lipemia in nonalcoholic steatohepatitis. *Hepatology* 37, 909-916 (2003).
- [38] Ginsberg, H. N. New perspectives on atherogenesis: role of abnormal triglyceride-rich lipoprotein metabolism. *Circulation* 106,2137-2142 (2002).
- [39] Choe, Y. G. et al. Apolipoprotein B/AI ratio is independently associated with non-alcoholic fatty liver disease in nondiabetic subjects. *Gastroenterol. Hepatol.* 28, 678-683 (2013).
- [40] Huang, B. et al. CagA-positive *Helicobacter pylori* strains enhanced coronary atherosclerosis by increasing serum OxLDL and HsCRP in patients with coronary heart disease. *Dig. Dis. Sci.* 56, 109-114 (2011).
- [41] Sung, K. C., Rhee, E. J., Ryu, S. H. & Beck, S. H. Prevalence of *Helicobacter pylori* infection and its association with cardiovascular risk factors in Korean adults. *Int. J. Cardiol.* 102, 411-417 (2005).
- [42] Kucukazman, M. et al. The relationship between updated Sydney system score and LDL cholesterol levels in patients infected with *Helicobacter pylori*. *Dig. Dis. Sci.* 54, 604-607 (2009).
- [43] Hatziagelaki, E. et al. Predictors of impaired glucose regulation in patients with non-alcoholic fatty liver disease. *Exp. Diabetes Res.* 2012, 351974, doi: 10.1155/2012/351974 (2012).
- [44] Cakir, E., Ozbek, M., Colak, N., Cakal, E. & Delibaşı, T. Is NAFLD an independent risk factor for increased IMT in T2DM? *Minerva Endocrinol.* 37, 187-193 (2012).
- [45] Hsieh, M. C. et al. *Helicobacter pylori* infection associated with high HbA1c and type 2 diabetes. *Eur. J. Clin. Invest.* 43, 949-956 (2013).