

# Endoscopic Findings and Associated Risk Factors in Primary Health Care Settings in Havana, Cuba

Enrique Galbán MD PhD, Enrique Arús MD PhD, Ulises Periles MD

## ABSTRACT

**INTRODUCTION** Upper gastrointestinal endoscopy, traditionally performed in Cuba in specialized hospitals, was decentralized to the primary health care level in 2004 to make it more patient-accessible.

**OBJECTIVES** Describe frequency and distribution of the principal symptomatic diseases of the upper gastrointestinal tract and their relation to the main risk factors associated with each in a sample of urban adults who underwent upper gastrointestinal endoscopy in primary care facilities in Havana in selected months of 2007.

**METHODS** A multicenter cross-sectional study was conducted, including 3556 patients seen in the primary health care network of Havana from May through November 2007. The endoscopies were performed at the 22 polyclinics (community health centers) providing this service. Diagnostic quality and accuracy were assessed by experienced gastroenterologists using a validated tool. Patients responded to a questionnaire with clinical, epidemiologic, and sociodemographic variables. Univariate and multivariate analyses (unconditional logistical regression) were used to identify associated risk factors. The significance level was set at  $p < 0.05$  (or confidence interval excluding 1.0).

**RESULTS** The diagnoses were: gastritis (91.6%), duodenitis (57.8%), hiatal hernia (46.5%), esophagitis (25.2%), duodenal

ulcer (15.8%), gastric ulcer (6.2%) and malignant-appearing lesions (0.4%). Overall prevalence of *Helicobacter pylori* infection was 58.4%. The main risk factors for duodenal ulcer were *H. pylori* infection (OR 2.70, CI 2.17–3.36) and smoking (OR 2.08, CI 1.68–2.58); and for gastric ulcer, *H. pylori* (OR 1.58, CI 1.17–2.15) and age  $\geq 60$  years (OR 1.78, CI 1.28–2.47). *H. pylori* infection was the main risk factor for gastritis (OR 2.29, CI 1.79–2.95) and duodenitis (OR 1.58, CI 1.38–1.82); and age  $\geq 40$  years for hiatal hernia (OR 1.57, CI 1.33–1.84). External evaluation was “very good” or “good” for 99.3% of endoscopic procedures and 97.9% of reports issued.

**CONCLUSIONS** Gastrointestinal endoscopy performed in primary care yielded high quality results and important information about prevalence of the most common diseases of the upper GI tract and associated risk factors. This study provides a reference for new research and can inform objective recommendations for community-based interventions to prevent and control these diseases. The existence of a network of universally accessible diagnostic endoscopy services at the primary care level, will contribute to conducting further research.

**KEYWORDS** Endoscopy, gastrointestinal diseases, upper GI tract, prevalence, risk factors, primary care, Cuba

## INTRODUCTION

In Cuba and most countries, prevalence studies of upper gastrointestinal (GI) tract diseases in the general population are virtually nonexistent; reports generally refer to frequency of these diseases in case series on endoscopies and hospital admissions, which are not representative of what actually occurs in the general population. In such studies, the diseases most commonly found are gastritis, peptic ulcer (both duodenal and gastric), hiatal hernia and gastric cancer.[1–5]

Availability of endoscopy at a primary care facility provides greater population access and thus enables earlier diagnosis of digestive tract diseases.[6] The international literature contains some evidence regarding endoscopic diagnosis performed at the primary care level—for example, in the United States and the United Kingdom, where the majority of procedures were done at the initiative of physicians in the public sector, but not as part of an integrated population-wide program.[7–10] In this respect, the Cuban experience appears to be unique.

In the early years of the 21st century, critical changes in the health sector have been introduced at all levels in Cuba. Of particular interest are changes in primary health care (PHC), where new technologies have been introduced to make diagnostic procedures more accessible, among them upper GI endoscopy.[6]

In 2004 the decision was made to offer upper GI endoscopy for diagnostic purposes in community-based polyclinics throughout Havana, to be extended later to those in the rest of the country. However, there were not enough endoscopists to materialize this

idea: at the time, there were some 200 gastroenterologist–endoscopists, only enough for performing the procedure in hospital. As a solution, family physicians were trained as endoscopists, creating a network of endoscopy services in PHC (Nodarse PO. Personal communication).

This study’s objective is to describe the prevalence of the main upper GI tract diseases and their principal risk factors in patients seen in primary care facilities in Havana.

## METHODS

**Study design** A multicenter cross-sectional study was conducted that included 3556 patients undergoing upper GI endoscopy in 22 endoscopy services at polyclinics in the 15 municipalities of Havana City province from May through November 2007. The National Gastroenterology Institute (IGE, its Spanish acronym) served as the study’s coordinating center.

In the preparatory stage, training workshops were held on research methods and procedures to standardize diagnostic criteria and procedures. Participants included endoscopists, interviewers and senior management from each of the polyclinics involved.

The minimum sample size necessary for a simple proportion was calculated using the EpiTable calculator from EpiInfo 6.1, yielding a requirement for 3082 persons undergoing upper GI endoscopy for the first time that year. Calculations assumed an estimated 10,000 endoscopies per year in Havana’s PHC facilities, approximately 8000 of which would be performed on patients for the first

time that year; a 5% prevalence of the least frequent disease; a 2% design effect; desired precision of 1.5%; and a 95% confidence level (CI).

Participant selection began in all settings on the same randomly-chosen date in May 2007, and patients were consecutively added to the study over the next six months. The target number for each participating unit was 140.

**Inclusion criteria** Patients aged  $\geq 18$  years who had not previously undergone upper GI endoscopy that year.

**Exclusion criteria** Prior endoscopy that year; being monitored endoscopically or with conditions that could result in emergency endoscopies (e.g., gastroenterologic hemorrhage and ingestion of caustic substances or foreign bodies).

**Study variables** Dependent variables were clinical diagnoses of esophagitis, hiatal hernia, gastric ulcer, duodenal ulcer, gastritis, duodenitis, and malignant-appearing lesion. Independent variables are listed in Table 1. Height and weight were measured for calculation of body mass index (BMI).

**Endoscopy** The procedure was performed with Olympus (Japan) fiberoptic endoscopes, model GIF-E. Patient preparation consisted of a 6-hour fast prior to endoscopy. A local pharyngeal anesthetic was administered (2% xylocaine spray) prior to the procedure. The patient was placed in the left lateral decubitus position and a mouth guard inserted. The endoscopist passed the extreme distal end of the endoscope through the mouth guard, observing through the eyepiece or monitor its passage over the tongue, then the epiglottis and vocal cords, the interior of the esophagus, continuing in this manner until reaching the second segment of the duodenum. On reaching this point, the endoscope was slowly retracted, the endoscopist examining the entire trajectory through which the instrument had advanced.[11]

**Helicobacter pylori infection** To assess prevalence of *H. pylori* infection (overall and specific by age and disease), a mucus specimen was taken from the gastric antrum at the *incisura angularis* of every participant, regardless of endoscopic indication or diagnosis. *H. pylori* was identified with a urease test using Urepyl (a reagent produced and standardized at the research laboratory of Cuba's IGE and distributed to all endoscopy services in the country).[12,13]

**Quality** Endoscopists' competence was evaluated with a validated questionnaire covering technical quality of the endoscopy and diagnostic accuracy.[14] Assessment was by an external committee of evaluators assembled for this purpose, comprising 10 professors of gastroenterology with recognized expertise, who evaluated *in situ* at all of the polyclinics a simple random sample of 280 patients, or 8% of study participants. A five-point scale—very good, good, fair, poor, very poor—was used to classify endoscopists' work on six dimensions considered fundamental in upper GI endoscopy: (1) appropriateness of the indication; (2) time between patient arrival and procedure; (3) procedure explanation given the patient by the attending physician; (4) technical quality of the endoscopy; (5) quality of diagnostic report; and (6) equipment cleaning and disinfection.

**Information gathering** General information about the participants, their clinical and epidemiologic data, information on expo-

**Table 1: Study variables**

Variable	Description
Age group (years)	<20
	20–29
	30–39
	40–49
	50–59
	60–69
	70–79
	$\geq 80$
Sex	Male, female
Skin color	Black, mestizo, white
Marital status	Single
	With a partner
	Married
	Divorced
	Widowed
Education completed	Illiterate
	Primary school
	Middle school
	Secondary (high) school University
Occupation	Professional
	Technical
	Laborer
	Student
	Homemaker
	Retiree
	Military
Other	
Obesity (BMI in kg/m <sup>2</sup> )	Overweight or obese $\geq 25$
	Not obese $< 25$
Family history of disease of segment studied	<b>Diagnosis in a first-degree relative of:</b>
	Gastrointestinal cancer
	Ulcer
	Gastritis Hiatal hernia
Risk factors	<b>Self-reported:</b>
	Current smoking
	Current alcohol use
	Coffee >2 cups a day
	ASA, current use
	Other NSAIDs, current use
	Steroids, current use <i>H. pylori</i>
Psychosocial factors	<b>Self-report in the previous six months of:</b>
	Death of a close relative
	Problems with spouse or partner
	Legal charges
	Job change Loneliness

ASA: Acetylsalicylic acid

BMI: Body Mass Index

NSAIDs: Nonsteroidal anti-inflammatory drugs

sure to risk factors and endoscopy results were recorded on standardized forms, which were sent to IGE for processing and analysis once completed in the polyclinics.

**Data analysis** Information from the forms was entered in a database using EpiInfo 6.04 and statistical analysis performed using the same software and its EPI 2000 version for Windows. Prevalences of dependent and independent variables in the study population were calculated as percentages. Analysis of potential sociodemographic, hereditary and psychosocial predictors, as

## Original Research

well as others such as *H. pylori* infection and use of medications, and their association with positive endoscopic diagnoses, was performed using univariate statistical techniques (chi-square test and Fisher's exact tests); and resulting independent variables associated ( $p < 0.05$ ) were included in a model for multivariate analysis (unconditional logistic regression). The measure of association strength was the odds ratio (OR), with significance level set at  $p < 0.05$  (or 95% confidence interval excluding 1.0).

**Ethical considerations** All patients provided written informed consent for participation. The study was approved by the IGE Research Ethics Committee.

### RESULTS

The external evaluation committee assessed as "very good" or "good" the quality of 99.3% of endoscopies and 97.9% of reports.

The study sample used was 115% of the minimum necessary for estimates of the desired precision and confidence, representing approximately 50% of all people undergoing upper GI endoscopy for the first time in 2007 in Havana polyclinics. Seventeen polyclinics surpassed their sample goal and the remaining five achieved a sample greater than 80% of target.

Of the 3556 participants, 63% (2239) were women and 37% (1317) men; 72% (2560) were of working age (18–59 years) (Table 2). Of the total, 55.6% (1976) had completed secondary school; 59% (2098) were married or cohabiting; 55.5% (1974) were white; and 24.4% (867) were laborers.

In the study population, 32.8% were smokers; 28.5% drank alcohol; 58.7% drank more than two cups of coffee a day; 2.6% took aspirin, 10% other nonsteroid anti-inflammatories (NSAIDs), and 1.2% steroids. Some 30.1% reported having experienced gastritis at some point in their life, 18.3% peptic ulcers, and 3.1% esophagitis; while 24.3% had a first-degree family history of gastritis, 22.4% peptic ulcers, 10.5% hiatal hernias and 7.9% GI cancers.

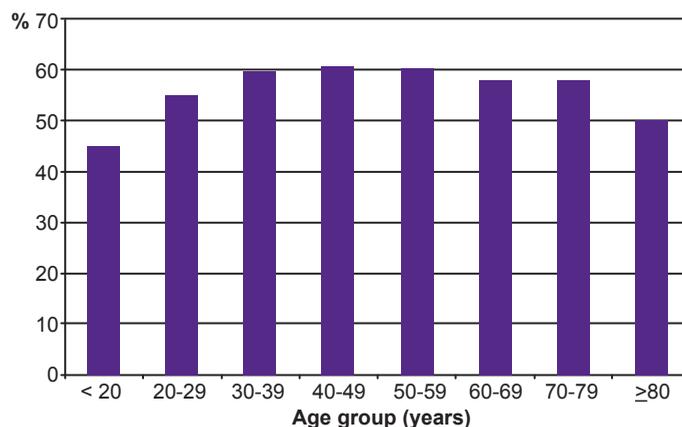
Slightly over half (57.1%) complained of being "stressed," and 21.8% said that they had experienced important life changes in the previous six months. Specifically, 19.9% said they felt lonely; 13.9% had recently lost a close relative; 12.2% were having prob-

lems with spouses or partners; 11.1% had recently switched jobs; and 1% had recently faced legal charges.

Most commonly reported symptoms were epigastric pain, 71.4% (2540); acid indigestion, 56.8% (2019); dyspepsia—assorted symptoms, problems, or discomfort related to the upper GI tract—43.7% (1555); heartburn, 24.2% (859); abdominal pain, 17.1% (608); and reflux, 14.1% (502). A triad of symptoms consisting of epigastralgia, acid indigestion and dyspepsia (nonspecific symptoms or a sensation of discomfort) was observed quite frequently.

The urease test for detecting *H. pylori* infection was performed on specimens from all patients; analysis was based on 3444 satisfactory specimens for which there were complete data, 96.9% of total; of these, 2011 (58.4%) were positive. *H. pylori* positivity was significantly more frequent ( $p < 0.001$ ) in men (62%) than in women (56.2%), and prevalence increased in the fourth to sixth decades of life, declining slightly in those aged  $\geq 70$  years (Figure 1). Prevalences of *H. pylori* infection by diagnosis were: duodenal ulcer, 77.7%; gastric ulcer, 69.4%; duodenitis, 63%; gastritis, 60.1%; and esophagitis, 55.2%.

**Figure 1: Endoscopic prevalence of *H. pylori* by age group in Havana polyclinics, May–November, 2007 (n=3444)**



**Frequency and distribution of upper GI diseases and associated risk factors** The most frequent diagnoses were: gastritis (91.6%), duodenitis (57.8%) and hiatal hernia (46.5%) (Table 3). In general, duodenitis, duodenal ulcer and gastric ulcer were significantly more frequent in men ( $p < 0.001$ ), while esophagitis ( $p = 0.03$ ) and hiatal hernia ( $p = 0.02$ ) were more frequent in women. Prevalence of gastritis was similar in both sexes. Table 3 displays the total number of diagnoses for each condition in men and women, average age, the population subgroups with higher prevalence and the frequency of associations among the different diagnoses.

**Esophagitis.** Some 73.3% of the cases were considered reflux esophagitis; 1.9% were classified as monilial and in 24.8%, no probable etiology was specified. None of the variables included in the model behaved as a predictor for esophagitis, and only coffee drinking turned out to be a protective factor (Table 4).

**Hiatal hernia.** Family history was reported by 13.9%. Risk of hiatal hernia almost doubled in the presence of family history (OR 1.96, CI 1.56–2.45). This effect disappeared in multivariate analysis.

**Table 2: Age and sex distribution of patients receiving endoscopy in primary care in Havana polyclinics, May–November, 2007 (n=3556)**

Age group (years)	Women		Men		Total	
	No.	%	No.	%	No.	%
<20	59	2.6	28	2.1	87	2.4
20–29	217	9.7	146	11.1	363	10.2
30–39	362	16.2	268	20.3	630	17.7
40–49	555	24.8	297	22.6	852	24.0
50–59	455	20.3	239	18.1	694	19.5
60–69	366	16.3	195	14.8	561	15.8
70–79	168	7.5	110	8.4	278	7.8
$\geq 80$	33	1.5	22	1.7	55	1.6
Age unknown	24	1.1	12	0.9	36	1.0
<b>TOTAL</b>	<b>2239</b>	<b>100.0</b>	<b>1317</b>	<b>100.0</b>	<b>3556</b>	<b>100.0</b>

Women 63% of total; Men 37%; Ratio F/M = 1.7  
Average age: W 48.3 years; M 47.4 years

**Table 3: Common upper GI tract conditions in patients receiving endoscopy in Havana polyclinics, May–November, 2007 (n=3556)**

Diagnosis	No.	Prevalence (%)			Age (years)			Subgroups with highest prevalence (%)	GI comorbidities (%)		
		Total	M	F	Mean	Extreme values					
						Min	Max				
Esophagitis	895	25.2	23.1	26.4	43.7	18	89	<20 years	50.0		
								Students <sup>b1</sup>	42.1	Gastritis	92.5
								Single <sup>b2</sup>	28.6	Duodenitis	60.4
								University educated <sup>b3</sup>	27.4	Duodenal ulcer	12.5
								Mestizo	26.5	Gastric ulcer	4.6
								Women	23.1		
Hiatal hernia	1653	46.5	44.0	47.3	50.0	18	89	Coffee <sup>b4</sup>	63.2	Gastritis	93.0
								Women <sup>b5</sup>	47.3	Duodenitis	61.0
								Family history <sup>b6</sup>	13.9	Esophagitis	27.5
										Duodenal ulcer	15.8
										Gastric ulcer	5.9
Gastritis	3256	91.6	92.5	91.0	44.6	18	96	Widow(er)s	96.1		
								Retirees	93.8	Duodenitis	59.0
								Men	92.5	Esophagitis	25.4
								White skin	92.2	Duodenal ulcer	15.5
								Primary school	92.0	Gastric ulcer	5.7
								Coffee drinkers	59.3		
Gastric ulcer	220	6.2	7.7	5.3	54.4	19	87	Coffee drinkers	67.7		
								Stress	53.2		
								Smokers <sup>b7</sup>	41.8	Gastritis	84.1
								Alcohol	30.9	Duodenitis	55.0
								Family history	22.7	Hiatal hernia	44.5
								≥70 years	12.5	Esophagitis	18.6
								NSAIDs	10.9	Duodenal ulcer	10.0
								Retirees	10.9		
								Men <sup>b8</sup>	7.7		
Duodenal ulcer	562	15.8	23.4	11.3	50.0	18	85	Coffee drinkers	59.3		
								Smokers <sup>b7</sup>	47.9	Gastritis	90.0
								Men <sup>b8</sup>	23.4	Duodenitis	77.4
								Alcohol <sup>b9</sup>	35.1	Hiatal hernia	46.4
								Stress	56.5	Esophagitis	20.0
								Family history <sup>b6</sup>	29.5	Gastric ulcer	3.9
Duodenitis	2056	57.8	63.5	54.4	49.0	18	96	Military	69.4		
								Primary school <sup>b2</sup>	66.5	Gastritis	93.8
								Men <sup>b8</sup>	63.5	Hiatal hernia	49.1
								Coffee drinkers	59.1	Esophagitis	26.3
								Married	58.3	Duodenal ulcer	21.2
								Smokers <sup>b7</sup>	35.6	Gastric ulcer	5.9
70–79 years	20.0										

<sup>a</sup> Excluding malignant-appearing lesions; N Total= 3556; Women 2239; Men 1317

<sup>b</sup> Prevalence in specific subgroup differs significantly from the rest

<sup>b1</sup> Occupation: Students vs. other occupations for esophagitis ( $p < 0.001$ ); Military vs. other for duodenal ulcer ( $p < 0.001$ )

<sup>b2</sup> Marital status: Single and divorced people vs. married people and people living with a partner for esophagitis ( $p = 0.017$ )

<sup>b3</sup> Education completed: People with a university education vs. people with lower levels of education for esophagitis ( $p < 0.001$ ) and people with primary level for duodenitis ( $p < 0.001$ )

<sup>b4</sup> Coffee: as a protective factor for hiatal hernia ( $p < 0.001$ ); gastritis ( $p = 0.023$ )

<sup>b5</sup> Sex: Women vs. men for esophagitis ( $p = 0.03$ ) and hiatal hernia ( $p = 0.017$ )

<sup>b6</sup> Family history: Duodenal ulcer and hiatal hernia ( $p < 0.001$ )

<sup>b7</sup> Smoking: Gastric ulcer, duodenal ulcer and duodenitis ( $p < 0.001$ )

<sup>b8</sup> Sex: Men vs. women for duodenal ulcer, gastric ulcer and duodenitis ( $p < 0.001$ )

<sup>b9</sup> Alcohol use: Duodenal ulcer ( $p = 0.007$ )

Overweight–obesity (BMI > 24.99) did not behave as a predictor of hiatal hernia ( $p > 0.05$ ); univariate analysis found discrete positive associations with *H. pylori* infection (OR 1.24, CI 1.08–1.42) and coffee drinking (OR 1.42, CI 1.24–1.63). Multivariate analysis reaffirmed *H. pylori* infection as an independent predictor, as well as age  $\geq 40$  years, female and white skin color (Table 4).

**Gastritis.** Some 63.8% of the gastritis cases were classified as erythematous, 18.2% as erosive and 14.7% as reflux; type was not

reported in the remaining 3.3%. Regarding location, 55.5% of diagnoses were limited exclusively to the gastric antrum, 5.4% to the corpus and 33.4% affected the entire stomach; for 5.7% there was no indication.

In those with gastritis, 59.3% were coffee drinkers and 33.1% smokers; 28.9% drank alcohol; 10.3% took NSAIDs with certain regularity, 2.7%, aspirin and 1.3%, steroids; and 60.1% of all tests were positive for *H. pylori*.

Univariate analysis showed that *H. pylori* infection was a significant risk factor for gastritis (OR 2.28, CI 1.77–2.94) and there was a slight association with coffee drinking (OR 1.33, CI 1.04–1.69). Using multivariate analysis to study all these independent variables yielded similar results for *H. pylori*, and over age 40 appeared as another associated risk factor; however, coffee drinking disappeared as a potential predictor of gastritis (Table 4).

**Duodenitis.** Exposure to potential predictive or risk factors was as follows: 59.1% drank coffee; 35.6% smoked, and 28.7% drank alcohol; 10% regularly took NSAIDs, 2.9% aspirin and 1.1% steroids; and 63.1% tested positive for *H. pylori*.

Univariate analysis showed that *H. pylori* infection (OR 1.58, CI 1.38–1.82) and smoking (OR 1.46, CI 1.17–1.56) behaved as risk factors, and this was confirmed in multivariate analysis. Moreover, being male and aged  $\geq 60$  years also had an effect. Alcohol consumption behaved as a discrete protective factor (Table 4).

**Duodenal ulcer.** The triad of symptoms most commonly associated with diagnosis of duodenal ulcer was epigastric pain (77.9%), acid indigestion (64.4%) and dyspepsia (40.4%); however, other symptoms were also present to a lesser extent, among them heartburn (26.7%), nonspecific abdominal pain (21.5%) and reflux (13.2%).

This diagnosis was accompanied by gastritis in 90% of cases, duodenitis in 77.4%, hiatal hernia in 46.4% and esophagitis in 20% (Table 3).

**Table 4: Factors associated with upper GI tract conditions in patients receiving endoscopy in Havana polyclinics, May–November, 2007 (multivariate analysis, unconditional logistic regression)**

Condition	Factor (comparison)	OR	CI 95%	p Value
Esophagitis	Coffee consumption	0.59	0.35–0.90	0.020
	<i>H. pylori</i> (+/-)	1.24	1.08–1.43	0.002
Hiatal hernia	Skin color (M+B/W)	0.86	0.75–0.99	0.040
	Age ≥40 years (Y/N)	1.57	1.33–1.84	<0.001
	Sex (M/F)	0.83	0.71–0.97	0.020
	Coffee (Y/N)	1.29	1.11–1.51	0.001
Gastritis	Age ≥40 years (Y/N)	1.51	1.14–2.00	0.004
	<i>H. pylori</i> (+/-)	2.29	1.79–2.95	<0.001
Gastric ulcer	Age ≥60 years	1.78	1.28–2.47	<0.001
	Male sex	1.42	1.04–1.95	0.030
	Smoking	1.39	1.01–1.90	0.042
	<i>H. pylori</i> (+/-)	1.58	1.17–2.15	0.003
Duodenitis	Alcohol (Y/N)	0.80	0.67–0.96	0.014
	Age ≥60 years (Y/N)	1.28	1.06–1.54	0.009
	Sex (M/F)	1.46	1.24–1.70	<0.001
	Smoking (Y/N)	1.38	1.17–1.62	<0.001
	<i>H. pylori</i> (+/-)	1.58	1.38–1.82	<0.001
Duodenal ulcer	Coffee (Y/N)	0.78	0.63–0.97	0.026
	Skin color (M+B/W)	1.22	1.01–1.48	0.043
	Age ≥40 years (Y/N)	1.46	1.16–1.85	0.002
	Sex (M/F)	2.30	1.87–2.83	<0.001
	Smoking (Y/N)	2.08	1.68–2.58	<0.001
	<i>H. pylori</i> (+/-)	2.70	2.17–3.36	<0.001

M: mestizo B: black W: white

One hundred percent of ulcers were located in the first part of the duodenum (bulb). There was a single lesion in 84.5% of patients, two lesions in 10% and multiple lesions in 5.5%. Some 3.9% (22 persons) also had gastric ulcer. Round shapes predominated (59.2%), followed by linear shapes (21.3%), “salami ulcers” (13.2%), star shapes (4.8%) and less often, “kissing ulcers” (1.5%). Some 70% were superficial and 16.2% deep (this information was not provided in 13.8% of patients). Based on their progression, the vast majority (72.2%) were considered active ulcers.

Potential risk factors considered predictive of duodenal ulcer were as follows: 59.3% of patients were coffee drinkers and 47.9% smokers; 35.1% drank alcohol; 8.7% regularly took NSAIDs; 3% took aspirin and 1.1% steroids. Among the psychosocial factors involved in the etiology of peptic ulcers (especially ulcers in the duodenum), most acting as co-factors or triggers of their clinical manifestation, it was found that of the 562 patients with duodenal ulcer, 56.5% indicated that they were under “considerable” stress, 22.4% indicated that they had experienced major life changes in the six months prior to the endoscopy, 15.1% had recently lost a close relative, 13.4% were having problems with their spouse or partner and 1.3% said they had recently faced legal charges. Despite the relatively high frequency of some of these variables, statistical analysis did not show a significant association between diagnosis of duodenal ulcer and any of them.

Family history of peptic ulcer was present in 29.5% of patients with duodenal ulcer and behaved as a potential predictor (OR 1.57, CI 1.28–1.93), but when persons with duodenal ulcers positive for *H. pylori* were disaggregated from those negative for this agent, fam-

ily history of peptic ulcer remained predictive for *H. pylori*-positive (OR 1.52, CI 1.19–1.95), but not *H. pylori*-negative cases (OR 1.51, CI 0.97–2.33).

In univariate analysis, the strongest associations with duodenal ulcer were for *H. pylori* (OR 2.76, CI 2.23–3.44) and smoking (OR 2.14, CI 1.77–2.58) and, to a lesser extent, alcohol consumption (OR 1.44, CI 1.18–1.75). Multivariate analysis using a model including these variables reaffirmed *H. pylori* infection as the most important predictor of duodenal ulcer (OR 2.70, CI 2.17–3.36). Alcohol no longer behaved as a risk factor, and coffee appeared to have a slight protective effect (Table 4).

**Gastric ulcer.** Symptoms most often mentioned were epigastric pain (81.8%), acid indigestion (57.3%) and dyspepsia (42.3%). Fifty percent of patients reported epigastric pain and acid indigestion combined, and the epigastric pain–acid indigestion–dyspepsia symptom triad was present in 25% of patients.

Antral location (81.4%), round shape (56.6%) and a single lesion were the descriptions most often reported. Some 69.4% were superficial, 56.1% were <1 cm in any dimension, 76.4% were classified as “active ulcers,” and only four cases (1.8%) were described as “appearing malignant.”

Exposure to potential factors predictive of gastric ulcer behaved as follows: 67.7% were coffee drinkers; 41.8% were smokers; 30.9% drank alcohol; 10.9% regularly took NSAIDs, 3.6% aspirin, and 0.5% steroids; and 69.4% were positive for *H. pylori*.

Of the 220 with gastric ulcer, 53.2% stated that they were under stress; 17.9% said they had experienced major life changes in the six months prior to endoscopy; 14.2% had recently lost a close relative; 11.5% were having problems with their spouse or partner and 1.4% indicated they had recently faced legal charges. Moreover, one in five patients with gastric ulcers (19.3%) said they felt lonely all the time and 8.7% indicated they no longer had anyone to provide emotional support in highly stressful situations. None of these variables showed a significant association with the diagnosis of gastric ulcer.

A history of peptic ulcer in a first-degree relative was present in 22.7% of persons with gastric ulcer, but estimating an increased risk for the subgroup of persons with a family history was not possible (OR 1.02, CI 0.73–1.43).

Through univariate analysis the following were identified as independent variables for gastric ulcer: *H. pylori* infection (OR 1.66, CI 1.22–2.28), smoking (OR 1.51, CI 1.13–2.01) and coffee drinking (OR 1.51, CI 1.12–2.05), while multivariate analysis showed *H. pylori* infection and smoking retained their effect, and age and male sex became important (Table 4).

**Cancer and malignant-appearing lesions in the upper GI tract.** Since positive diagnosis of malignant lesions is histological, this section includes only endoscopic suspicions of cancer and malignant-appearing lesions.

A total of three patients (0.1%) received a presumptive diagnosis of esophageal cancer, eight (0.2%) of stomach cancer and four (0.1%) of malignant-appearing gastric ulcers—in all, 15 persons with an endoscopic diagnosis of “malignant-appearing lesion,” for

a prevalence of 0.4%; that is, one presumptively malignant lesion was found per 237 endoscopies.

The three cases identified as presumptive esophageal cancer were men aged 41, 58, and 80; among the presumptive cases of stomach cancer, four were men and four women with ages ranging from 42 to 75. Of the latter, two were diagnosed in the early stage and six in advanced stages. The risk for GI cancer was estimated as three times higher for men than for women (OR 3.42, CI 1.06–12.8).

## DISCUSSION

The decision of the Ministry of Public Health to provide GI endoscopy services in the primary health care network in Cuba made this study possible, and with it, acquisition of information on the frequency of and risk factors associated with the most common diseases of the digestive tract in the health services level most accessible to patients. This resulted in indicators that were closer to their real distribution in the population, in comparison with those obtained until now in hospitals, where a more narrowly-defined subset of patients is seen, often those more seriously ill.

In this regard, our literature review yielded only a few studies from the United States and the United Kingdom on GI endoscopies performed by physicians at the primary care level and on nurses working at that level; however, none of them described results on the frequency and distribution of the principal diseases of the upper GI tract.[7–9] The unique characteristics of Cuba's primary health care subsystem made it possible to obtain this information more readily and precisely[15] and to provide rates for a specific geographically-defined population.

Studies in the United States[7,16] conclude that it is probable that endoscopies performed at the primary care level do not always comply with the national technical recommendations; thus, the number of inappropriate indications tends to be higher in comparison with that of specialized health services. These studies conclude that allowing primary care physicians to perform endoscopies should be reevaluated. In contrast, some studies in the United Kingdom conclude that there are no significant differences in results of upper GI endoscopies done at the primary care level, even when performed by nurses instead of physicians.[8,17]

In Cuba, PHC endoscopy services are coordinated by and closely connected with hospital gastroenterology services where upper GI endoscopy is not only diagnostic, but prognostic and therapeutic. Thus, specialist competition with PHC services is not a factor in patient care, since the two are complementary.[18]

The relative frequencies of diagnosis of the principal diseases of the upper GI tract obtained in primary care were the ones expected when compared with a series of consecutive endoscopies performed at IGE in 2007. The frequency of diagnosis in a series of 1575 patients seen consecutively in IGE between September 2007 and May 2008 showed that gastritis was also the most frequent (82%), followed by hiatal hernia (31.6%), duodenitis (31.1%), duodenal ulcer (9.7%), esophagitis (9.4%), gastric ulcer (3.4%) and malignant lesions (0.95%).[18] In general, there is a higher prevalence of inflammation and benign ulcers in patients seen in primary care, while malignant-appearing lesions are seen almost twice as often in IGE, consistent with

the types of patients that should be seen in primary care and tertiary care levels.

The prevalence of *H. pylori* infection varies with population socioeconomic status, with lower-income countries exhibiting higher rates than higher-income countries.[1] Infection is very high in Africa, Asia, and many parts of Central and South America ( $\geq 90\%$ ), while it is relatively low in northern and eastern Europe, North America, and Australia ( $< 50\%$ ).[1] This infection is contracted early in life; over 50% of children worldwide are infected by the age of 10,[19–22] and, once again, the most important risk factor for contracting *H. pylori* infection appears to be the low socioeconomic status of the child's family.[23]

Studies by Gutiérrez et al. on *H. pylori* prevalence in Cuba in the normal population and dyspeptic patients in Havana, using a set of serologic diagnostic reagents for the detection of *H. pylori* IgG antibodies, show prevalences of infection ranging from 75% to 100%, although data were obtained from relatively small population samples. The prevalence values for *H. pylori* infection found in each of the diseases in this study are within usual range observed with the urease test; values may vary with the location of the mucosa, the number of specimens taken and the waiting time for observing the change in color.[1,2]

Our study presents the first epidemiologic data on diseases of the upper GI tract for a geographically-defined population in Cuba. Gastritis, the diagnosis most frequently encountered in PHC in Havana, is also the most common diagnosis in the majority of upper GI endoscopy series published in the literature. In a study that included patients from four countries on three continents (Japan, China, Tanzania, and the Dominican Republic), Aoki *et al.* reported gastritis prevalence rates ranging from 23.5% to 96.5%.[22]

The prevalence of esophagitis found in our study is quite similar to that reported by Du in China,[23] who found a rate of 20.8% among 2231 patients. This in turn was higher than the 15.8% also reported in China by Tseng[24] and the 12% reported by Kim in a major study conducted in South Korea with 25,536 subjects[25] and by Chen in Taiwan (12%).[26]

A Swedish endoscopy study of peptic ulcer[27] in a random sample of 1001 persons from two communities found a relatively low prevalence (4.1%). However, other non-population-level studies involving series of patients spontaneously presenting in hospitals found higher prevalences: 7% in Finland,[28] 8.3% in Peru,[4] 11.5% in Kuwait,[29] 15% in the United States (Alaska),[30] 17.2% in China[31] and 31.8% in Poland.[32] The prevalence found in Havana is within the range of reported values.

Risk factors associated with diseases of the upper GI tract in this study generally coincide with those reported by other authors. A selection of published articles on studies in Europe, Asia, and the Americas shows that *H. pylori* infection is the main predictor of peptic ulcer disease,[4,25,31–34] followed by smoking,[31,32,35–37] although some authors do not consider the latter an independent predictor but rather a synergistic co-factor with *H. pylori* infection.[33,35] Other risk factors associated with peptic ulcer found in the literature are: male sex,[23,36] advanced age,[23,28] prior history of ulcer,[21] use of aspirin or other NSAIDs[28,32,36,37] and being divorced.[23] Alcohol consumption is reported as a risk factor in some studies[23] and as a protective factor in others.[36] Moderate

physical exercise, a variable not included in our study, has been cited as an independent protective factor against peptic ulcer.[31]

Drugs such as aspirin and other NSAIDs are considered predictors of significant risk for gastric ulcer,[31,32] but this study was not able to show that association.

Without a doubt, *H. pylori* infection is the main predictor of gastritis,[21,34] just as hiatal hernia, male sex, smoking, drinking alcohol, and overweight–obesity are for esophagitis.[26]

Limitations of this study include use of a diagnostic test for *H. pylori* that, while highly sensitive and specific with positive predictive values of >90% when compared with histological diagnosis, has not been validated against another similar, commercially available test; thus, the existence of a certain degree of underreporting of *H. pylori* prevalence cannot be ruled out. Another limitation is that pathology confirmation was not obtained for some diagnoses, especially those related to malignant-appearing lesions. The failure to show the use of aspirin or other NSAIDs as an important factor for development of gastric ulcers may be associated with the low sensitivity of the questionnaire employed in terms of how questions about their use were formulated, and perhaps greater use of buffered varieties of these types of drugs in our environment in recent years.

### CONCLUSIONS

This study's greatest merit is that it contributes information on the frequency of a group of diseases whose prevalence is virtually unknown in Cuba and most countries. Its implementation in a sample population residing in a well-defined territory with good and similar guarantees of access to diagnostic services and prompt data acquisition offer greater guarantees of accuracy and precision of epidemiologic results, compared to traditional studies

published in the international literature that focus on the population seeking care at hospitals.

For the first time in Cuba, important population-based data were obtained on frequency of the most common diseases of the upper GI tract and their main risk factors. This information, together with a network of patient-accessible diagnostic endoscopy services at the primary care level, lays the foundation for future studies and facilitates objective recommendations on community-based interventions for prevention and control of these diseases.

### ACKNOWLEDGMENTS

The authors are grateful to the following colleagues who collaborated in study design, organization or data management:

Drs Héctor Hernández Garcés, Pedro Gil González, William Haedo Quiñones, Carlos Fábregas Rodríguez, Rolando Martínez López, Mirta Infante Velásquez, Miguel González-Carvajal Pascual, Orlando Nodarse Pérez, Juan Antonio Mas Páez, Hermidio Hernández Mulet and Amada Palomino Besada; and technician Fabiola Medina Colina.

We also thank the primary care personnel who interviewed patients and performed endoscopies:

Drs Nuvia Figueredo Zayas, Armando Orellana Molina, Eduardo Morandera Abreu, Libia Isela González Hernández, Juan Castellanos Safonts, Elsa Tamara Chávez Rivero, Gustavo Veitía, Yanet Valdés Viera Corrales, Ileana Pacho Saavedra, Jaqueline Vega Bello, Humberto Soto Smat, Elena Roseaux Mola, Osmary Lemagne Lugo, Ana Rosa Piñero Santana, Moraima Hernández Sáez, Mirian Pérez Frías, Jaime Lázaro Candelario Fuentes, María Yudith Monzón Rodríguez, Raúl Mederos Salgado, Lina Martínez Acosta, Sirsi Guilarte Díaz and Martha Bernal Pereira. 

### REFERENCES

- Gutiérrez B, Cavazza ME, Ortiz D, Correnti M, Vidal T, Mégraud F, et al. Seroprevalencia de la infección por *Helicobacter pylori* en pacientes con Gastritis Crónica, Úlcera Duodenal y Gástrica: Primer estudio de corte retrospectivo [Internet]. Rev Cubana Invest Bioméd. 2008 Apr–Jun [cited 2011 Jan 16];27(2). Available from: [http://scielo.sld.cu/scielo.php?script=sci\\_arttext&pid=S0864-03002008000200011&lng=es](http://scielo.sld.cu/scielo.php?script=sci_arttext&pid=S0864-03002008000200011&lng=es). Spanish.
- Gutiérrez B, Vidal T, Valmaña CE, Camou-Juncas C, Santos A, Mégraud F, et al. Infección por *Helicobacter pylori* en Ciudad de La Habana, Cuba. Prevalencia de las cepas cagA positivas [Internet]. Vaccinmonitor. 2005 Jul–Dec [cited 2011 Jan 16];14(2):15–9. Available from: [http://scielo.sld.cu/scielo.php?script=sci\\_arttext&pid=S1025-028X2005000200003&lng=en](http://scielo.sld.cu/scielo.php?script=sci_arttext&pid=S1025-028X2005000200003&lng=en). Spanish.
- Wang YR, Richter JE, Dempsey DT. Trends and Outcomes of Hospitalizations for Peptic Ulcer Disease in the United States, 1993 to 2006 [Internet]. Ann Surg. 2010 [cited 2011 Jan 16];1:51–8. Available from: [http://journals.lww.com/annalsurgery/Abstract/2010/01000/Trends\\_and\\_Outcomes\\_of\\_Hospitalizations\\_for\\_Peptic.8.aspx](http://journals.lww.com/annalsurgery/Abstract/2010/01000/Trends_and_Outcomes_of_Hospitalizations_for_Peptic.8.aspx)
- Montes Teves P, Salazar Ventura S, Monge Salgado E. Cambios en la Epidemiología de la Úlcera Péptica y su Relación con la Infección con *Helicobacter pylori*. Hospital Daniel Carrion 2000–2005. Rev Gastroenterol Perú. 2007 Oct–Dec [cited 2011 Jan 16];27(4):382–8. Available from: <http://www.scielo.org.pe/pdf/rgp/v27n4/a07v27n4.pdf>. Spanish.
- Piñol Jiménez F, Paniagua Estévez M. Cáncer gástrico: factores de riesgo [Internet]. Rev Cuba Oncol. 1998 Sep–Dec [cited 2011 Jan 16];14(3):171–9. Available from: <http://bases.bireme.br/cgi-bin/wxislind.exe/iah/online/?IscScript=iah.iah.xis&src=google&base=LILACS&lang=p&nextAction=ink&exprSearch=299713&indexSearch=ID>. Spanish.
- Luna Morales EC, Sierra Pérez DC, Gandul Salabarría L. La transformación del policlínico en Cuba de cara al siglo XXI [Internet]. Rev Cubana Med Gen Integr. 2009 Jul–Sep [cited 2011 Jan 16];25(2):1–10. Available from: [http://scielo.sld.cu/scielo.php?script=sci\\_arttext&pid=S0864-21252009000200016&lng=es](http://scielo.sld.cu/scielo.php?script=sci_arttext&pid=S0864-21252009000200016&lng=es). Spanish.
- Charles RJ, Cooper GS, Wong RC, Sivak MV Jr, Chak A. Effectiveness of open-access endoscopy in routine primary-care practice. Gastrointest Endosc. 2003 Feb;57(2):183–6
- Smale S, Bjarnason I, Forgacs I, Prasad P, Mukhood M, Wong M, et al. Upper gastrointestinal endoscopy performed by nurses: scope for the future? Gut. 2003 Aug;52(8):1090–4.
- Williams J, Russell I, Durai D, Cheung WY, Farrin A, Bloor K, et al. What are the clinical outcome and cost-effectiveness of endoscopy undertaken by nurses when compared with doctors? A Multi-Institution Nurse Endoscopy Trial (MINuET). Health Technol Assess. 2006 Oct;10(40):iii–iv, ix–x, 1–195.
- Hernández Gómez LC. Presencia de la mujer en la Salud Pública Cubana [Internet]. Rev Cub Salud Pública. 2009 Mar [cited 2011 Jan 16];35(1). Available from: [http://www.scielosp.org/scielo.php?script=sci\\_arttext&pid=S0864-34662009000100010&lng=en](http://www.scielosp.org/scielo.php?script=sci_arttext&pid=S0864-34662009000100010&lng=en). doi: 10.1590/S0864-34662009000100010. Spanish.
- Hernández Garcés HR. Manual de endoscopia digestiva superior diagnóstica. Havana: Editorial Ciencias Médicas; 2008. 232 p. Spanish.
- Buesa F, Sotto A, Quintero M. Primer reporte del aislamiento de *Campylobacter pyloridis* en Cuba, 21 de marzo de 1986. Bol Epidemiol Hosp Doc Gen Calixto García. 1986;2:3. Spanish.
- Mandado S, Gra B, González-Carvajal M, Paniagua Piñol F, Domínguez C. Diagnóstico morfológico de *Helicobacter pylori* mediante citología gástrica por cepillado. Rev Cubana Med. 2003 Jan–Mar;42(1):27–33. Spanish.
- Periles U. Tesis de Terminación de Residencia en Gastroenterología. Impacto de la Endoscopia Digestiva Superior en la Atención Primaria de Salud [thesis]. [Havana]: Facultad de Ciencias Médicas Calixto García Iñiguez; 2009.
- Arús Soler E, Fábregas Rodríguez C, Haedo Quiñones W, Martínez López R, Nodarse Pérez O, Ruiz Torres J, et al. Proyecciones de la Especialidad de Gastroenterología en Cuba para el 2015 [Internet]. Havana: Ministry of Public Health (CU); 2006 [cited 2011 Jan 16]. 28 p. Available from: [http://www.sld.cu/galerias/pdf/sitios/gastroenterologia/informe\\_finala.pdf](http://www.sld.cu/galerias/pdf/sitios/gastroenterologia/informe_finala.pdf). Spanish.
- Zuccaro G Jr, Provencher K. Does an open access system properly utilize endoscop-

- ic resources? *Gastrointest Endosc.* 1997 Jul;46(1):15–20.
17. Meaden C, Joshi M, Hollis S, Higham A, Lynch D. A Randomized Controlled Trial Comparing the Accuracy of General Diagnostic Upper Gastrointestinal Endoscopy Performed by Nurse or Medical Endoscopists. *Endoscopy.* 2006 Jun;38(6):553–60.
  18. Sistema ProGastro [Internet]. [place unknown]: Rincón Grupo Sapiem; 2008. Available from: <http://gruposapiem.wordpress.com/windowsapp/sistema-progastro/>. Spanish.
  19. Gutiérrez B, Vidal T, Valmaña CE, Santiesteban N, González N, Leonard I, et al. Primer Informe sobre el Aislamiento de *Helicobacter pylori* asociado a enfermedades digestivas en Ciudad de La Habana. *Vaccimonitor.* 2001 Jan–Mar;10(1). Spanish.
  20. Camargo MC, Yepez MC, Ceron C, Guerrero N, Bravo LE, Correa P, et al. Age at Acquisition of *Helicobacter pylori* Infection: Comparison of Two Areas with Contrasting Risk of Gastric Cancer. *Helicobacter.* 2004 Jun;9(3):262–70.
  21. González-Carbajal M, Rojas Zurita F, Grá Oramas B, Ávalos García R. Prevalencia de la infección por *Helicobacter pylori* en pacientes dispépticos. *Rev Panam Infect.* 2004;6(4):8–14. Spanish.
  22. Aoki K, Kihai PE, Wenyuan Z, Xianghang Z, Castro M, Disla M, et al. Comparison of Prevalence of Chronic Atrophic Gastritis in Japan, China, Tanzania, and the Dominican Republic. *Ann Epidemiol.* 2005 Sep;15(8):598–606.
  23. Du J, Liu J, Zhang H, Yu CH, Li YM. Risk factors for gastroesophageal reflux disease, reflux esophagitis and non-erosive reflux disease among Chinese patients undergoing upper gastrointestinal endoscopic examination. *World J Gastroenterol.* 2007 Dec 7;13(45):6009–15.
  24. Tseng PH, Lee YC, Chiu HM, Huang SP, Liao WC, Chen CC, et al. Prevalence and Clinical Characteristics of Barrett's Esophagus in a Chinese General Population. *J Clin Gastroenterol.* 2008 Nov–Dec;42(10):1074–9.
  25. Kim N, Lee SW, Cho SI, Park CG, Yang HC, Kim HS, et al. The prevalence of and risk factors for erosive oesophagitis and non-erosive reflux disease: a nationwide multicentre prospective study in Korea. *Aliment Pharmacol Ther.* 2008 Jan;15;27(2):173–85.
  26. Chen TS, Chang FY. The prevalence and risk factors of reflux esophagitis among adult Chinese population in Taiwan. *J Clin Gastroenterol.* 2007 Oct;41(9):819–22.
  27. Aro P, Storskrubb T, Ronkainen J, Bolling-Sternvald E, Engstrand L, Vieth M, et al. Peptic Ulcer Disease in a General Adult Population: the Kalixanda Study: A Random Population-based Study. *Am J Epidemiol.* 2006 Jun 1;163(11):1025–34.
  28. Voutilainen M, Mäntynen T, Färkkilä M, Juhola M, Sipponen P. Impact of Non-Steroidal Anti-Inflammatory Drug and Aspirin Use on the Prevalence of Dyspepsia and Uncomplicated Peptic Ulcer Disease. *Scand J Gastroenterol.* 2001 Aug;36(8):817–21.
  29. Abahussain EA, Hasan FA, Nicholls PJ. Dyspepsia and *Helicobacter pylori* infection: Analysis of 200 Kuwaiti patients referred for endoscopy. *Ann Saudi Med.* 1998 Nov–Dec;18(6):502–5.
  30. Sacco F, Bruce MG, McMahon BJ, Bruden D. A prospective evaluation of 200 upper endoscopies performed in Alaska Native persons. *Int J Circumpolar Health.* 2007 Apr;66(2):144–52.
  31. Li Z, Zou D, Ma X, Chen J, Shi X, Gong Y, et al. Epidemiology of Peptic Ulcer Disease: Endoscopic Results of the Systematic Investigation of Gastrointestinal Disease in China. *Am J Gastroenterol.* 2010 Dec;105(12):2570–7.
  32. Konturek SJ, Bielański W, Plonka M, Pawlik T, Pepera J, Konturek PC, et al. *Helicobacter pylori*, non-steroidal anti-inflammatory drugs and smoking in risk pattern of gastroduodenal ulcers. *Scand J Gastroenterol.* 2003 Sep;38(9):923–30.
  33. Halter F, Brignoli R. *Helicobacter pylori* and smoking: two additive risk factors for organic dyspepsia. *Yale J Biol Med.* 1998 Mar–Apr;71(2):91–9.
  34. Weck MN, Brenner H. Association of *Helicobacter pylori* infection with chronic atrophic gastritis: Meta-analyses according to type of disease definition. *Int J Cancer.* 2008 Aug 15;123(4):874–81.
  35. Moshkowitz M, Brill S, Konikoff FM, Averbuch M, Arber N, Halpern Z. Additive deleterious effect of smoking on gastroduodenal pathology and clinical course in *Helicobacter pylori*-positive dyspeptic patients. *Isr Med Assoc J.* 2000 Dec;2(12):892–5.
  36. Schubert TT, Bologna SD, Nensley Y, Schubert AB, Mascha EJ, Ma CK. Ulcer risk factors: interactions between *Helicobacter pylori* infection, nonsteroidal use, and age. *Am J Med.* 1993 Apr;94(4):413–8.
  37. Yamada M, Wong FL, Fujiwara S, Tatsukawa Y, Suzuki G. Smoking and alcohol habits as risk factors for benign digestive diseases in a Japanese population: the radiation effects research foundation adult health study. *Digestion.* 2005;71(4):231–7.

## THE AUTHORS

**Enrique Galbán García** (Corresponding author: [galban@infomed.sld.cu](mailto:galban@infomed.sld.cu)), physician specializing in epidemiology with a doctorate in public health. Chief of epidemiology, National Gastroenterology Institute (IGE), Havana, Cuba.

**Enrique Arús Soler**, gastroenterologist with a doctorate in medical sciences. Director, IGE, Havana, Cuba.

**Ulises Periles Gordillo**, gastroenterologist, IGE, Havana, Cuba.

Submitted: October 14, 2010

Approved for publication: January 10, 2012

Disclosures: None

  
vacci  
pharma  
2012

Second International Congress on the  
Pharmacology of Vaccines  
June 16–20, 2012  
Barceló Cayo Santa María Hotel  
Cayo Santa María, Villa Clara, Cuba



Sponsored by  
 **Pan American  
Health  
Organization**  
Regional Office of the  
World Health Organization



The Cuban Society of Pharmacology invites you to VacciPharma 2012 to learn about the latest developments in human and veterinary vaccines, exchange experiences with other experts and help accelerate progress in vaccine development.

### Three workshops will be offered:

- Prophylactic vaccines
- New technologies in prophylactic vaccine development
- Therapeutic vaccines

### Related events:

- Symposium on veterinary vaccines
- Regional meeting on DTP/HB/Hib/IPV combined vaccine

For more information:  
Mario Landys Chovel Cuervo MD  
[mlandys@finlay.edu.cu](mailto:mlandys@finlay.edu.cu)

Rolando Ochoa Azze MD  
[vaccipharma@finlay.edu.cu](mailto:vaccipharma@finlay.edu.cu)

or visit [www.vaccipharmacuba.com](http://www.vaccipharmacuba.com)