

Hiatal Hernia Is Associated With an Increased Prevalence of Atrial Fibrillation in Young Patients

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Abstract

Purpose: Hiatal hernia (HH) causes protrusion of the stomach into the chest cavity, directly impinging on the left atrium and possibly increasing predisposition to atrial arrhythmogenesis. However, such association has not been fully explored. The objective was to determine if an association between HH and atrial fibrillation (AF) exists and whether there are age- and sex-related differences.

Methods: Adult patients diagnosed with HH from 1976 to 2006 at Mayo Clinic Rochester, Minnesota, were evaluated for AF. The number of patients with AF and HH was compared to age- and sex-matched patients with AF reported in the general population. Long-term outcomes were compared to corresponding county and state populations.

Results: During the 30-year period, 111,429 patients were diagnosed with HH (mean age 61.4 ± 13.8 years, 47.9% male) and 7,865 patients (7.1%) also had a diagnosis of AF (mean age 73.1 ± 10.5 years; 55% male). In younger patients (<55 years), the occurrence of AF was 17.5-fold higher in men with HH and 19-fold higher in women with HH compared to the frequency of AF reported in the general population. Incidence of heart failure for patients with AF and HH was worse compared to the overall county population, but better than for those with AF. Similarly, mortality was worse in patients with AF and HH compared to the overall state population, but better than for those with AF in the county.

Conclusions: Hiatal hernia appears to be associated with increased frequency of AF in both men and women of all age groups, but particularly in young patients. Further studies are needed to investigate this possible association and underlying mechanism.

Introduction

Hiatal hernia (HH) causes protrusion of the abdominal contents into the chest cavity, and can directly impinge on the left atrium (Figure 1).¹ The effect of this mechanical compression on the heart is not known. Cardiac compression from atrial masses is associated with various arrhythmias, and HH could potentially result in a similar effect.²⁻⁶ The anatomical proximity of the left atrium to herniated gastric contents raises the possibility of mechanical irritation of the atria, autonomic neural connections or inflammation

that may increase the risk for atrial fibrillation (AF). An association of esophagitis and frequent symptoms related to gastroesophageal reflux disease with the development of AF has been previously reported.⁷⁻¹² The reported reduction in frequency of recurrences of AF with the use of proton-pump inhibitors or following Nissen fundoplication also suggests a link between the gastrointestinal tract and atrial arrhythmias.^{9,12}

In this study, we sought to determine whether the presence of HH is associated with an increased risk for AF, particularly in young patients who lack the significant comorbidities that can otherwise increase predisposition of AF in the elderly. The overall prevalence of AF in patients with HH was compared to previously reported prevalence of AF in the general population.¹³ Clinical outcomes of these patients with AF and HH also were compared to the expected outcomes of the corresponding county's and state's general populations and patients with AF.

Key Words:

Atrial fibrillation; Hiatal hernia; Gastroesophageal reflux disease; Congestive heart failure; Stroke; Mortality

Disclosures:

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Material and Methods

All patients older than 18 years with a Hospital International Classification of Disease Adaptation (HICDA) diagnosis code of HH who were seen at Mayo Clinic in Rochester, MN, in both the ambulatory and inpatient settings from Jan. 1, 1976, to Dec. 31,

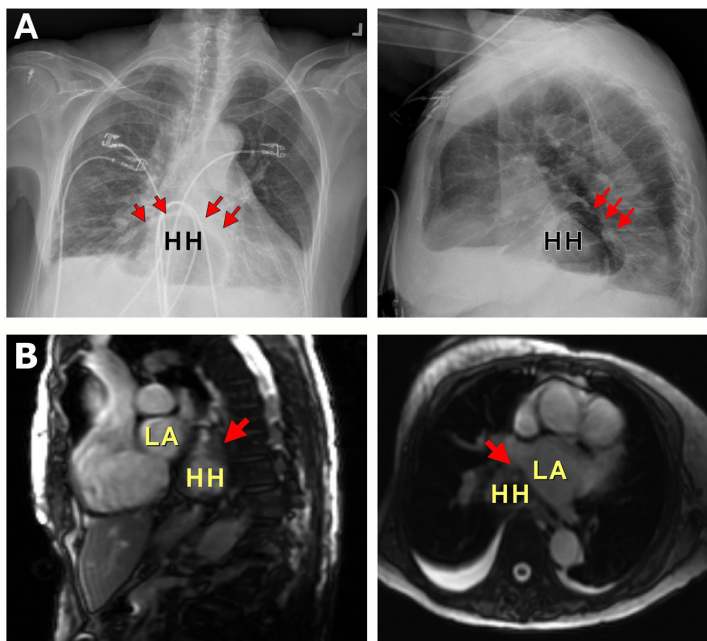


Figure 1: A: Chest X-rays demonstrating large hiatal hernia (HH) (red arrows). Note close proximity to the left atrium (LA). B: Cardiac magnetic resonance imaging illustrating large HH (red arrows) in close proximity to the LA.

2006, were assessed for the presence of AF. An electronic clinical database for patients who were given a HICDA diagnosis code for HH (05515 and 05535) and AF (04163 and 04164) during the specified time period and age group was maintained and reviewed. All patients who were given a coded diagnosis of AF, including lone, paroxysmal, persistent and permanent AF, as well as those who were given a diagnosis of HH by a physician were included in the study. The diagnosis of AF was confirmed by reviewing medical records. Detailed information about the severity of HH was not available. Patients were categorized by age and gender. The dates of initial diagnosis of HH, AF and cardiovascular events including heart failure, stroke and death were obtained. Patients given a diagnosis of AF after the diagnosis of HH were analyzed as well as the overall group of patients with both AF and HH. Risk factors that increase predisposition for AF – hypertension, diabetes mellitus, coronary artery disease, sleep apnea, valvular disease and heart failure at the time of diagnosis of HH – were identified (HICDA and International Classification of Diseases-9 codes 34128430, 02500000, 414.00, 780.58, 03950112, 04279113). The study population was compared to the prevalence of AF reported in the general population by Go et al. based on the similar age and gender breakdown.¹³ Clinical outcomes, including heart failure and transient ischemic attack (TIA) or stroke, in the study population were assessed in a similar fashion using the HICDA codes documented by a physician. These outcomes then were compared to the expected events in the corresponding county's general population. These events for patients with AF and HH then were compared to clinical outcomes reported in patients with AF.¹⁴⁻¹⁶ Mortality was determined from medical records and the Social Security Death Index. All-cause mortality of the study population was compared to the expected death rate in the state population for age- and gender- matched controls. The study was approved by the Mayo Clinic Institutional Review Board.

Statistical Analysis

Data are summarized using frequencies and percentages for categorical variables and mean \pm standard deviation for continuous variables. The Kaplan-Meier method was used to generate survival free-of-endpoint curves for stroke, heart failure and death in patients with AF and HH. Observed curves for each of these endpoints were compared to expected outcomes using one-sample log-rank tests. Expected curves were constructed using population-based studies of each of the outcomes. Expected mortality was constructed from the corresponding state death data.

Results

Hiatal Hernia Is Associated With Increased Frequency of Atrial Fibrillation

A total of 111,429 patients (mean age 61.4 ± 13.8 years, 53,430 males [47.9%]) with a diagnosis of HH were seen during the 30-year study period. Of these, 7,865 patients (7.1%; mean age 73.1 ± 10.5 years) also had a diagnosis of AF (4,337 [55%] men and 3,528 [45%] women). The baseline characteristics of patients with AF and HH compared to those with AF as reported in the general population by Go et al.¹³ are summarized in Table 1. Despite a similar age range (mean age 73.1 ± 10.5 vs. 71.2 ± 12.2 years), the prevalence of common conditions, such as coronary artery disease, heart failure, valvular disease and hypertension that predispose to AF, was lower in the HH and AF group compared to the population with AF.

In both sexes, the number of individuals with AF was higher in patients with HH compared to what has been reported for the general population throughout all age groups (Figure 2).¹³ In young men (<55 years) with HH, AF was present in 3.5%, a 17.5-fold higher prevalence compared to the 0.2% reported in this group by Go et al.¹³ (Figure 2). Other studies have found low prevalence rates for young patients with AF,¹⁷⁻²⁰ similar to Go et al.¹³ Likewise, the occurrence of AF in young women (<55 years) with HH was 19-fold higher than the prevalence of AF reported in the general population in the

Table 1: Comparison of Baseline Characteristics of Patients With AF And HH vs. AF as Reported in the Literature

	AF* (%) n=17,974	AF and HH (%) n=7,865
Mean age (years)	71.2 \pm 12.2	73.1 \pm 10.5
Sex (male)	57	55
Hypertension	49	40
Hyperlipidemia	NR	17
Diabetes	17	16
Coronary artery disease	35	20
Sleep apnea	NR	4
CHF	29	10
Valvular disease	5	2

AF = atrial fibrillation; CHF = congestive heart failure; HH = hiatal hernia; NR = not reported.
*Data taken from ATRIA study, Go et al. JAMA 200113

same age group (1.9% vs. 0.1%) (Figure 2). As illustrated in Figure 2, the higher prevalence of AF in patients with HH was consistently seen across all age groups in both men and women, although the differences diminished in the older age groups. Men with HH aged 55-59 years had a 7.8-fold increase in the prevalence of AF, 60-64 years a 5.2-fold increase, 65-69 years a 3.4-fold increase, 70-74 years a 2.4-fold increase, 80-84 years a 1.2-fold increase and > 85 years a 1.2-fold increase. Similarly, women with HH aged 55-59 years had an 11.7-fold increase in the prevalence of AF, 60-64 years a 5.9-fold increase, 65-69 years a 4-fold increase, 70-74 years a 2.4-fold increase, 75-79 years a 2-fold increase, 80-84 years a 1.6-fold increase and > 85 years a 1.5-fold increase.

When comparing the occurrence of AF in patients with HH over the same one-year period (July 1, 1996, to Dec. 1, 1997) as Go et al.¹³ there was a comparable trend (Figure 3). Men with HH aged < 55 years had a 21-fold increase in the prevalence of AF, 55-59 years a 12-fold increase, 60-64 years an 11-fold increase, 65-69 years a 7-fold increase, 70-74 years a 3-fold increase, 75-79 years a 2-fold increase, 80-84 years a 2-fold increase and > 85 years a 2-fold increase. Women with HH aged < 55 years had a 16-fold increase in the prevalence of AF, 55-59 years a 28-fold increase, 60-64 years a 7-fold increase, 65-69 years a 7-fold increase, 70-74 years a 3-fold increase, 75-79 years a 3-fold increase, 80-84 years a 2-fold increase and > 85 years a 1-fold increase.

Cardiovascular Events in Patients With Atrial Fibrillation and Hiatal Hernia

A total of 1,746 patients had heart failure, 810 patients with AF and HH developed TIA or stroke and 2,810 patients died during the 30-year follow-up period. Although the rate of heart failure development in patients with AF and HH was significantly higher (p<0.0001) when compared to the expected outcomes for the county's general population, it was significantly better (p<0.00001) than for

those with AF in this population (Figure 4).¹⁵ At five years, survival free of stroke was 98.7% in the study population of AF and HH compared with 90.8% in the AF group and 96.6% in the expected county population. Survival free of heart failure at five years was 93.9% in the AF and HH group, 81% in the AF group, and 96.3% in the expected county group. The expected survival at five years was 90% in the AF and HH population, 55.4% in the AF population and 88.1% in the expected state population (Figure 4).

Discussion

The novel finding of our study is demonstration of an association between HH and AF, with a greater likelihood of developing AF in patients with HH. The prevalence of AF in the general population is age-dependent and generally reported as approximately 1%.¹³ The occurrence of AF in patients with HH in our study was 7.1%. This difference was strikingly greater in younger age groups in both men and women. When broken down by age and gender, we found 17.5-fold and 19-fold higher prevalences of AF in men and women younger than 55 years with HH, respectively. A similar trend was found when comparing the occurrence of AF in patients with HH over the same one-year period used by Go et al.¹³ with a 21-fold increase in men < 55 years and a 16-fold increase in women < 55 years, suggesting that secular differences over time with the 30-year follow-up were less of a factor. These differences in the prevalence of AF persisted with advancing age, but the magnitude of the difference between the HH group and the general population diminished, likely due to the presence of other comorbidities and factors in addition to HH that promote the development of AF (Table 1). The higher prevalence of AF in the young HH group, despite reduced prevalence of risk factors for AF such as hypertension, coronary artery disease and congestive heart failure as compared to the AF group, suggests that HH may increase the risk for AF.

Anatomically, the esophagus lies directly behind the left atrium and protrusion of abdominal contents into this limited space can predispose the atria to mechanical compression and stretch that increases the likelihood of developing AF (Figure 1). Hiatal hernias are frequently associated with gastroesophageal reflux and esophagitis,²¹ which could lead to local inflammation of the esophagus and surrounding structures. The proximity of the esophagus, and consequently of the HH to the left atrium, could lead to extension of this inflammation that could contribute to the development of AF by mechanical or chemical/neural influences mediated through vagal or sympathetic nervous systems.^{7,9,10,12,22-26} Huang et al. recently reported an increased risk of AF in patients with gastroesophageal reflux disease (GERD) as an independent predictor.²⁷ They postulated that the increased vagal tone observed in patients with GERD could lead to an increased risk of AF. The presence of a HH may exacerbate GERD and consequently a similar mechanism of vagal stimulation could be attributed to the increased prevalence of AF seen in patients with HH. The presence of cardiac masses has been associated with development of AF and atrial flutter, and similar atrial compression by the presence of HH, associated inflammation of the esophagus or irritation of the autonomic neural plexus present in this region, particularly after heavy meals, can influence the development of AF and other atrial arrhythmias.^{3,5,6,28}

The hypothesis that presence of HH contributes to the development of AF is supported by previous reports. Schilling et al.²⁶ described a case of atrial flutter that resolved in a patient with a large

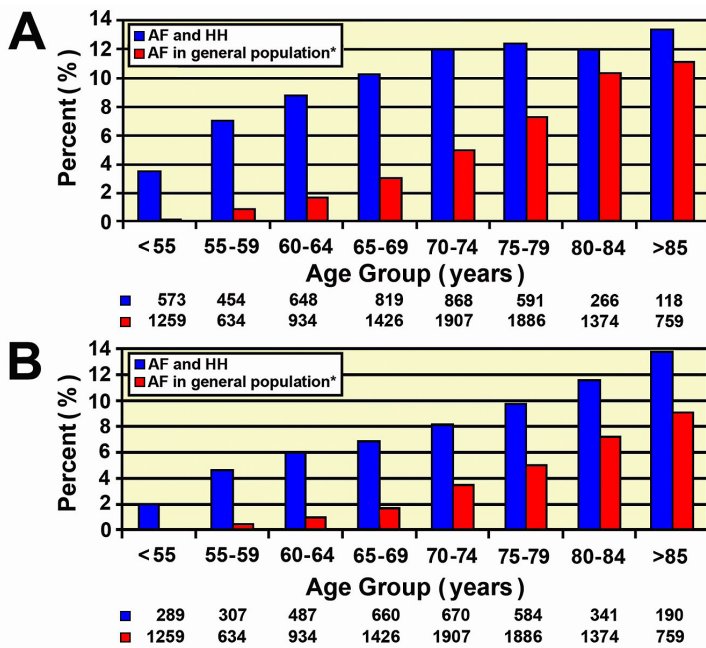


Figure 2: A: Men with atrial fibrillation (AF) and hiatal hernia (HH) compared to AF reported in the general population* B: Women with AF and HH compared to AF reported in the general population*

*Data taken from ATRIA study, Go et al. JAMA 2001 [13]

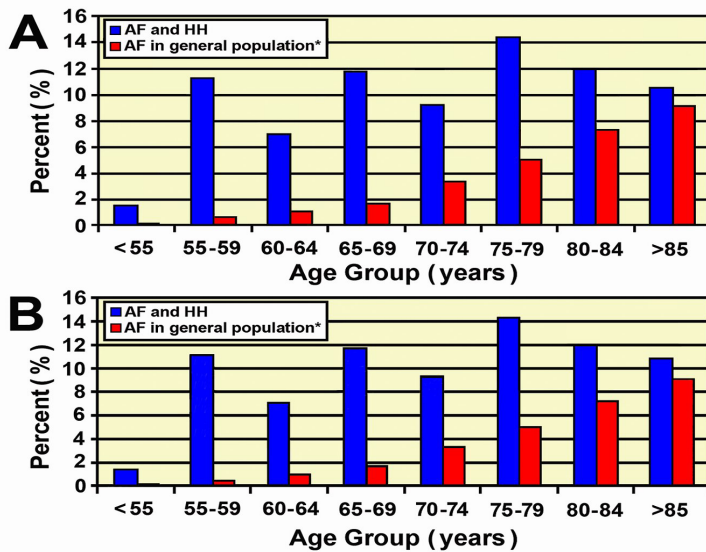


Figure 3: A: Men with atrial fibrillation (AF) and hiatal hernia (HH) compared to AF reported in the general population* from July 1, 1996, to Dec. 1, 1997 B: Women with AF and HH compared to AF reported in the general population* from July 1, 1996, to Dec. 1, 1997

AF = atrial fibrillation; CHF = congestive heart failure; HH = hiatal hernia; NR = not reported.
*Data taken from ATRIA study, Go et al. JAMA 2001;13

paraesophageal hernia after surgical repair of the hernia. Duygu et al.¹² also reported a case of paroxysmal atrial flutter that was refractory to electrical cardioversion and calcium channel blockers in a patient with severe acid reflux and daily heartburn symptoms secondary to a large HH. However, after instituting proton-pump inhibitor therapy, normal sinus rhythm was restored and maintained at one-year follow-up. In another patient with a large paraesophageal HH and postprandial AF, Nissen fundoplication suppressed recurrences of AF.²⁵ Other reports have demonstrated electrocardiographic changes, including ST-segment elevation, T-wave inversion and sinus bradycardia, that can occur in patients with HH.^{4,29} Studies defining a definitive causal relationship with HH and mechanistic insights are warranted.

The occurrence of heart failure, stroke or TIA and death during the five-year follow-up period after the diagnosis of HH was compared to the expected event rates for heart failure and stroke or TIA in the county and the overall mortality in the state. The event rates for patients with AF and HH also were compared to the overall rates in patients with AF in the county population.¹⁴⁻¹⁶ The incidence of these complications was significantly less than that described in the overall AF population, suggesting that patients with AF associated with HH might have a better prognosis than patients with AF without HH (Figure 4). There are a few possibilities for this finding. Patients with AF and HH are more likely to seek medical attention given the presence of a hiatal hernia and related symptoms and, thus, may be treated more aggressively for their AF with medications such as anticoagulation. Another possibility may be that patients with AF and HH represent a unique subgroup of patients with AF that are actually less likely to develop AF-related complications due to a different mechanism for the AF. These patients may be more likely to have lone AF and less structural heart disease but still develop AF due to the mechanical/neural factors from the effect of the HH on the atria and, consequently, have a lower complication rate. Our study suggests that the natural history of AF in patients with HH may be

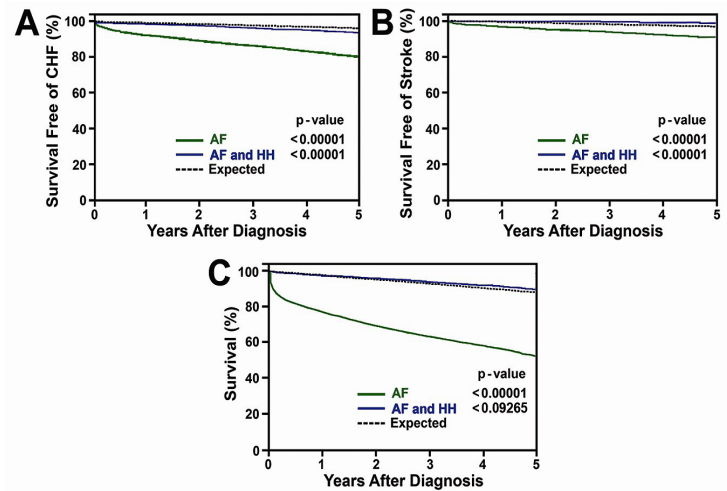


Figure 4: Five-year follow-up comparing patients with atrial fibrillation (AF), AF and HH, and expected population for A: congestive heart failure (CHF), B: stroke and C: mortality

different from AF associated with structural heart disease.

Limitations

The findings of this retrospective study should be considered in the context of the limitations of such a study design that cannot account for other confounding factors predisposing to AF not reported in the medical record. Our study encompassed all types of AF including lone, paroxysmal, persistent and permanent AF, which may affect the degree to which HH played a role in the development of AF. Although the diagnosis of AF was confirmed by reviewing medical records, the frequency and duration of AF episodes could not be assessed. Similarly, the severity of HH could not be assessed from the retrospective review of the records. Echocardiographic assessment was not performed or available in the majority of patients, and, therefore, the impact of HH on atrial function or dimensions or their contribution to the development of AF could not be determined. Further prospective studies investigating patients with HH and minimum risk factors for AF are needed to confirm the novel observations described here and to provide additional evidence to support the association between HH and AF.

Conclusions:

We present a hypothesis-generating study that suggests the presence of hiatal hernia may trigger development of atrial fibrillation, particularly in younger patients. Further prospective investigation is needed to confirm this association and the underlying mechanisms.

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