



BEYOND THE BURDEN: THE IMPACT OF ATRIAL FIBRILLATION IN ASIA PACIFIC

2019 Report

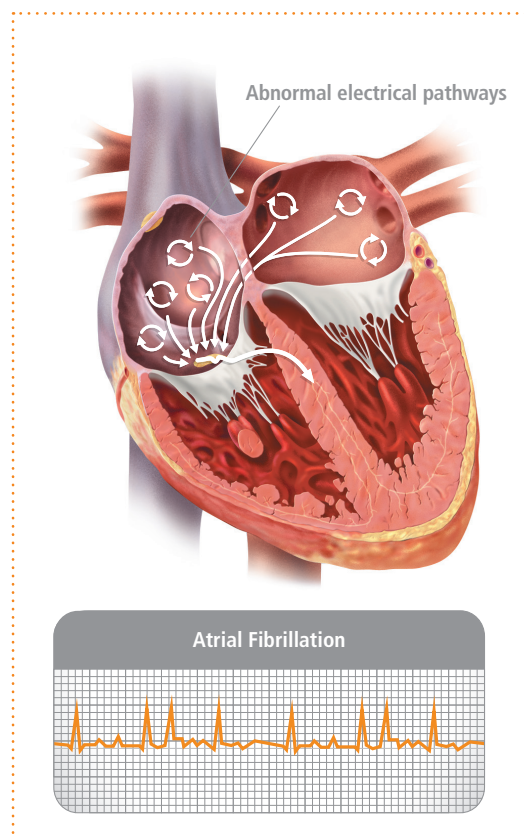


Atrial Fibrillation is fast becoming one of the world's most significant health issues that places a critical burden on healthcare systems

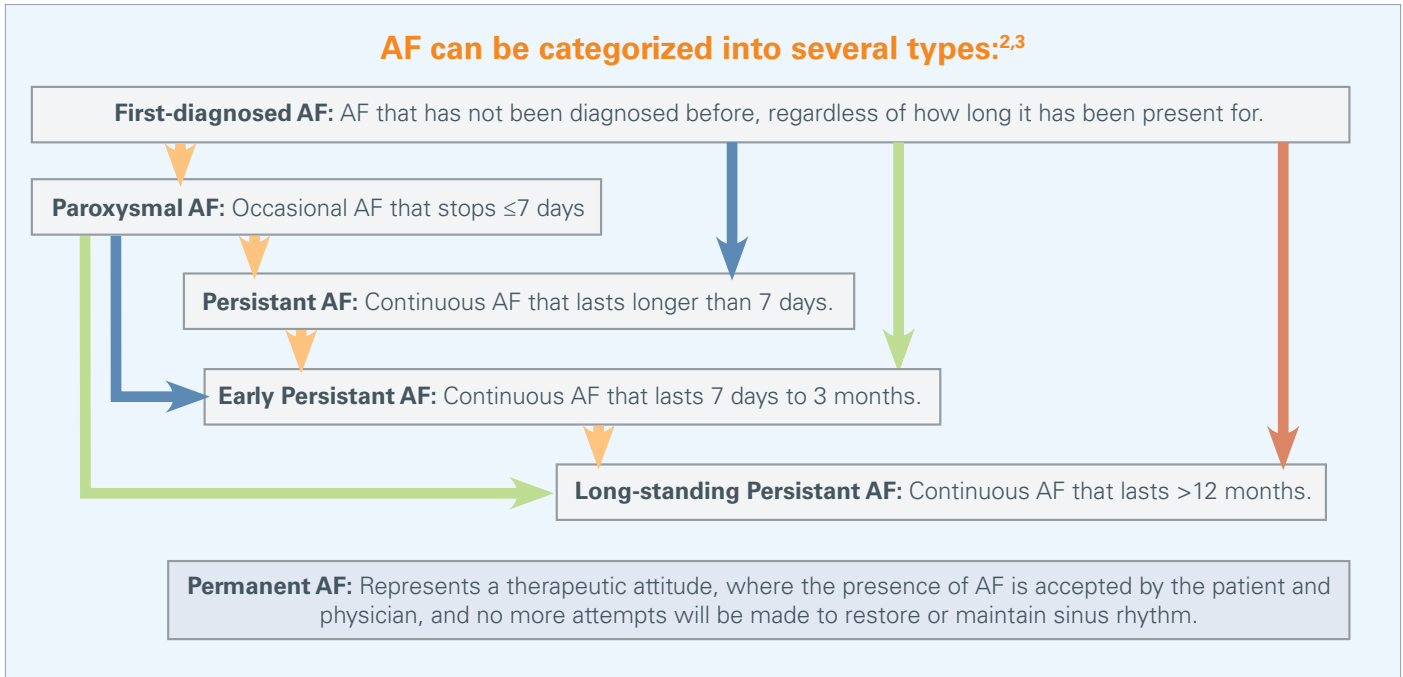
ATRIAL FIBRILLATION

What is Atrial Fibrillation and why is it important?

- Atrial fibrillation (AF) is **characterized by an irregular and often fast heartbeat** that results in uncoordinated contraction of the top 2 chambers of the heart (i.e., atria).¹

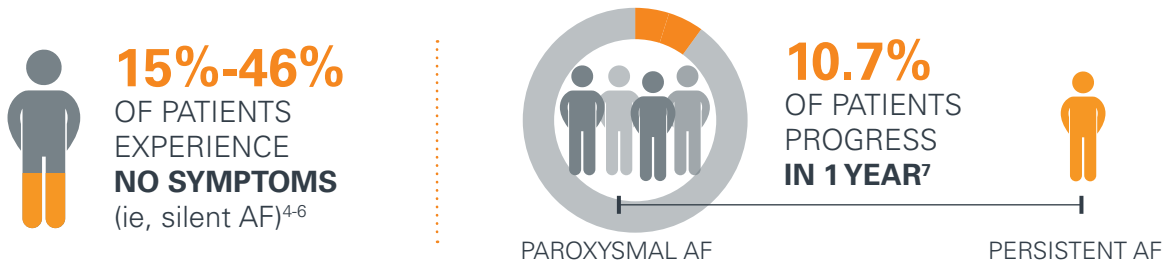


AF can be categorized into several types:^{2,3}



■ Patients may have episodes of AF that fall into one or more of the above categories; patients are categorized based on their most frequent pattern of AF.^{2,3}

■ Early detection and diagnosis of AF may help improve patient outcomes, since a long history and an extended duration of AF have been associated with recurrence and may worsen cardiac function.²



Patients with AF have an increased risk for life-threatening complications and other diseases:⁸⁻¹³



■ AF worsens quality of life for patients, which can be burdensome to caregivers.^{4,6,14,15}

■ AF increasingly places a **critical financial burden** on the healthcare system with a **1.8 to 5.6x increase in healthcare costs** every 10 years across APAC.¹⁶⁻²⁰

AF is a new millennium epidemic that affects **over 16M people** in APAC, primarily the middle-aged and the elderly.²¹



How common is AF?

AF is the most common type of cardiac arrhythmia. Within APAC, the percentage of affected population varies from **0.28% in India** to **1.48% in Australia**.²¹ However, AF is progressing rapidly across all APAC countries, affecting over

1.4M NEW PEOPLE EACH YEAR.²¹

HOW WILL AF AFFECT THE APAC REGION IN THE FUTURE?

The number of patients is rapidly rising due to an **aging population** and **increasing lifestyle risk factors**.²²⁻²⁸

With more patients suffering with atrial fibrillation, **rates for strokes, hospitalizations, and doctor visits** are expected to rise.



It is estimated there will be **72 MILLION** AF patients in Asia by 2050^{28,30}



2.9 MILLION ASIAN PATIENTS anticipated to suffer from AF-ASSOCIATED STROKE IN 2050^{30,31}



HOSPITALIZATIONS FOR AF have been increasing by **5-42%** annually across APAC^{16,19,28}

DEMOGRAPHICS OF AF

Who is at risk for AF?

AF is a common age-related arrhythmia; it generally affects people over 30 years old and becomes more prevalent with advancing age.^{28,32-36}



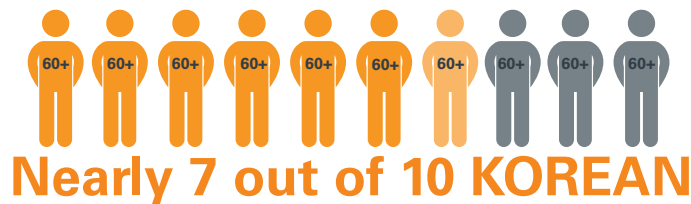
1 in 5 CHINESE adults

Estimated lifetime risk for AF³²

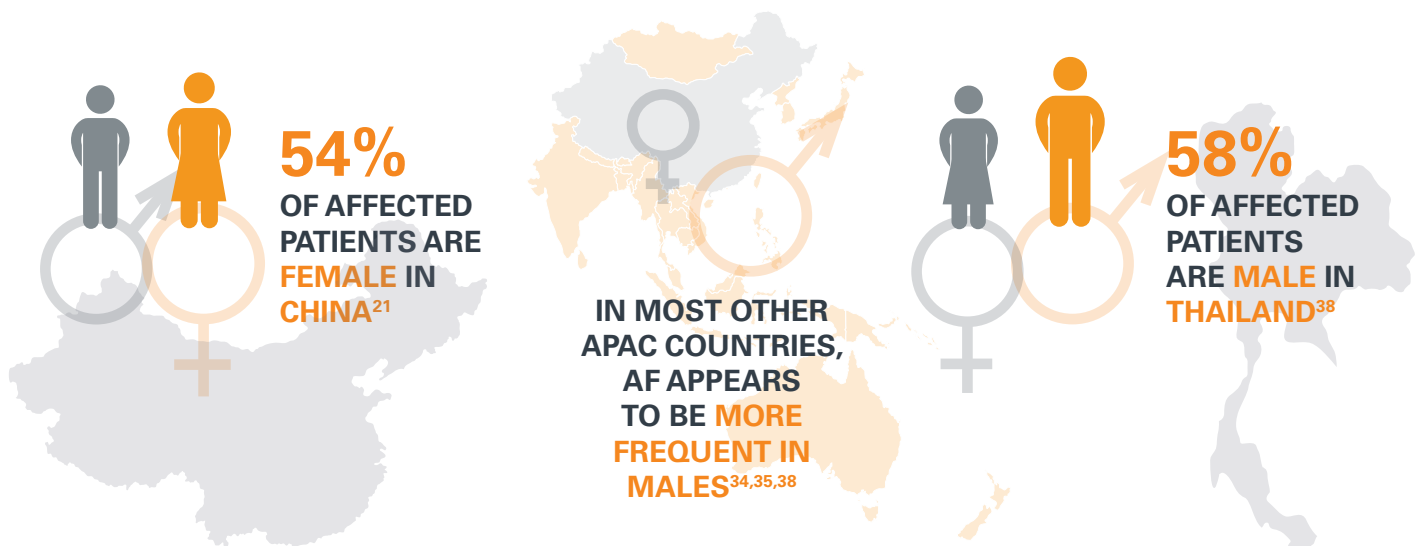


1 in 4 AUSTRALIAN adults

Estimated lifetime risk for AF^{23,37}



Nearly 7 out of 10 KOREAN adults suffering from AF ARE AT LEAST 60 YEARS OLD³⁴



CAUSES AND RISK FACTORS OF AF

AF develops from structural changes to the heart due to lifestyle, other chronic conditions, and non-modifiable factors.

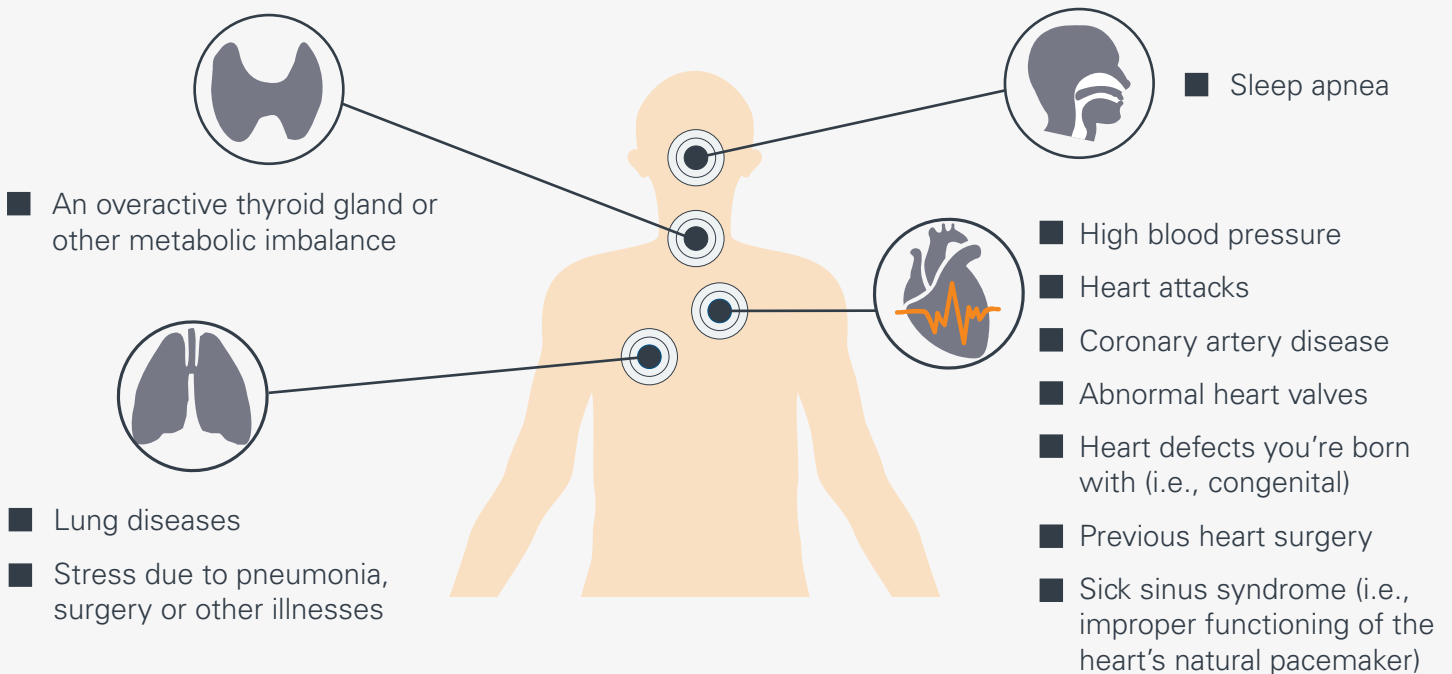
WHAT CAUSES AF?

AF is most often caused by structural changes to the heart due to other conditions and lifestyle factors.^{1,3}

- AF is an **irregular and often rapid heartbeat** that occurs when there are extra, uncoordinated electrical signals in the atria.¹

Common causes of AF

Abnormalities or damage to the heart's structure are the most common cause of AF, and this can be caused by:^{3,35,40,41}



Other factors that cause AF:



- Exposure to stimulants, such as medications, caffeine, tobacco or alcohol

WHAT FACTORS LEAD TO AF?

Lifestyle factors, other health conditions, and non-modifiable factors increase the risk of developing AF.



LIFESTYLE FACTORS

- Obesity
- Alcohol consumption
- Risks for cardiovascular disease: smoking, stress, caffeine and other stimulants
- Activity level

OTHER CONDITIONS

- Rheumatic heart disease (cardiac murmur)
- Dilated cardiomyopathy
- Heart failure
- Hyperthyroidism
- Coronary artery disease
- Chronic kidney disease, Diabetes
- Hypertension

NON-MODIFIABLE FACTORS

- Older age
- Congenital heart defects
- Family history or other genetic factors
- Male sex

- Rheumatic heart disease is one of the most important risk factors in lower-income APAC countries.^{25,28,41}
- Dilated cardiomyopathy, heart failure, hyperthyroidism, coronary artery disease, and ageing are the other strongest risk factors in Asian populations.^{25,35,40,41}
- Obesity, chronic kidney disease, diabetes, hypertension, heart failure, coronary artery disease, male sex, and aging are the strongest risk factors in Australia.^{42,43}

The symptoms and clinical consequences of AF disrupt patient quality of life and increase the risk of mortality.

WHAT ARE THE SYMPTOMS OF AF?

Symptoms of AF disrupt daily life and range from mild to debilitating.^{14,49}
The most common symptoms are:^{5,6,44}



Up to **65%**
PALPITATIONS

50%
FATIGUE

Up to **50%**
SHORTNESS
OF BREATH

Up to **35%**
MALAISE



Up to **35%**
DIZZINESS

12%
ANXIETY

Up to **41%**
CHEST PAIN

55%
MORE
THAN 1
SYMPTOM



OVER
50% of AF PATIENTS
have a reduced ability to exercise

■ The frequency and severity of symptoms varies greatly from patient to patient and, within a patient, symptoms can fluctuate widely over time.⁴⁴

- **Patients who do not** experience symptoms of AF may be at **greater risk** of complications and disease severity due to lack of treatment:



15-46%
SILENT AF^{4-6,44,45}



21-59%
**PATIENTS
DIAGNOSED
WITH AF AFTER
SUFFERING
A STROKE**⁴⁶⁻⁴⁸



**PATIENTS WITH
SILENT AF
EXPERIENCE
POORER GENERAL
HEALTH & QUALITY
OF LIFE**⁴⁹

AF increases the risk of:

- **Mortality:**

AF is independently associated with a significantly greater risk of mortality.^{50,51}



**45-
50%**

**ALL-CAUSE
MORTALITY**⁽¹⁾

- **Stroke:**

a serious complication of AF that is associated with long-term disability and mortality.



**170-
490%**

**ANY
STROKE**^{(1),9,50,54-55}

- **Heart attack:**

a serious complication of AF that also significantly increases the risk of stroke and mortality.³²

- **Heart failure and left ventricular dysfunction:**

a common complication of AF and the most common cause of death among Asian patients with AF.^{52,53}



**500-
800%**

**HEART
FAILURE**^{50,54}

- **Cognitive dysfunction or vascular dementia:**

a complication of AF that causes a decline in memory and thinking skills, which can interrupt daily life and independent function.



**40-
60%**

**DEMENTIA OR
COGNITIVE
IMPAIRMENT**^{(1),32,56}

- **Obstructive sleep apnea:**

is common in AF patients and may increase the rate of arrhythmia recurrence.⁵⁷

(1) Relative increased risk based on the relative risk of morbidity and mortality when compared to patients without AF, adjusted for other confounding factors.

AF worsens the quality of life for patients, which can be burdensome to caregivers.

WHY DO PEOPLE WITH AF SEEK MEDICAL TREATMENT?

AF symptoms and repeated recurrence increase unplanned medical visits and hospitalizations.



SYMPTOMS OCCUR IN 54-69%

PATIENTS WITH AF, EVEN IF THEY ARE BEING TREATED^{4-6,44}

- Clinical decision-making can be challenging because symptoms related to AF can differ a lot between patients and within patients at different time points.⁴⁴
- AF and its related symptoms are a major therapeutic challenge and burden to healthcare systems.⁴⁴

AF is currently THE MOST COMMON CAUSE of cardiovascular hospitalization in Australia.¹⁶ From 1993-2013, AF admissions increased by:

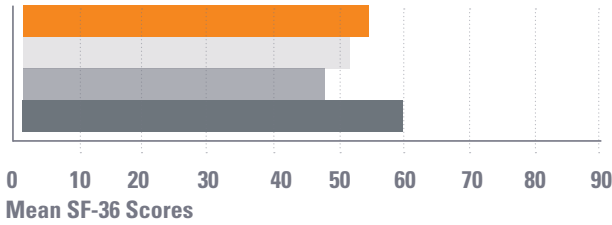
- 295% for Atrial Fibrillation
- 73% for Myocardial Infarction
- 39% for Heart Failure



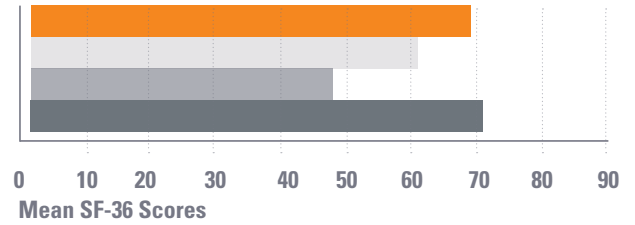
Comparison of Quality of Life between Patients with AF and other Cardiovascular Conditions⁵⁸

SF-36 Quality of Life Subscales

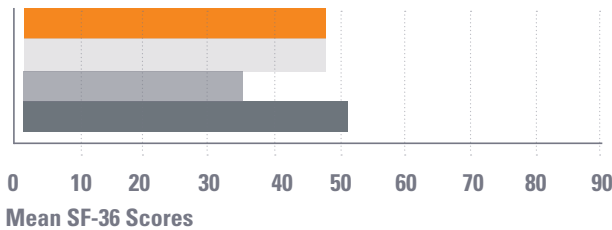
General Health



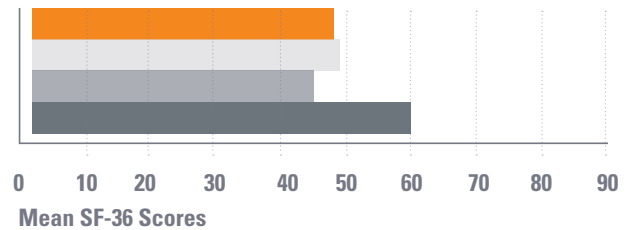
Physical Functioning



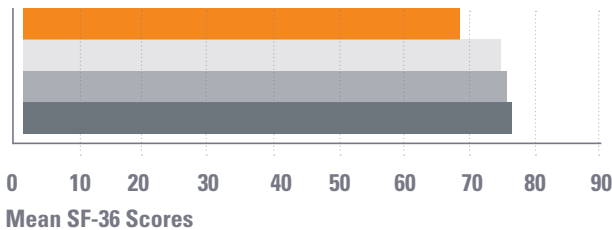
Role Physical



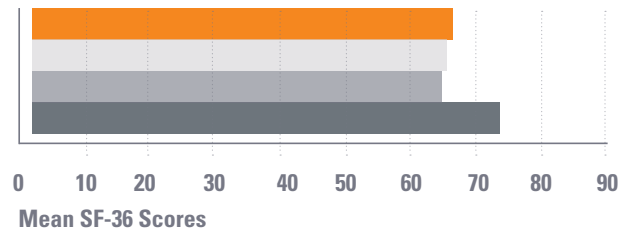
Vitality



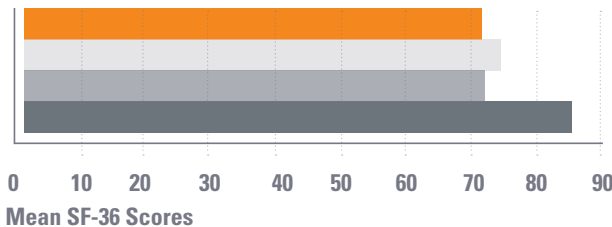
Mental Health



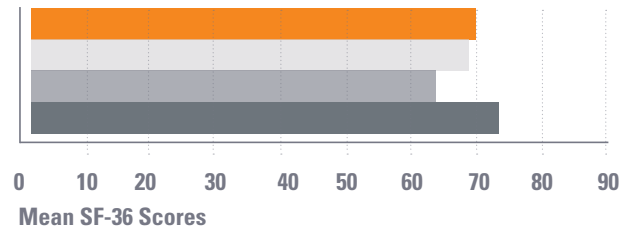
Role Emotional



Social Functioning



Bodily Pain



■ AF Patients (n=152)
 ■ PTCA Patients (n=69)
 ■ CHF Patients (n=216)
 ■ Post-Heart Attack Patients (n=69)

Abbreviations: AF = atrial fibrillation; CHF = congestive heart failure; PTCA = percutaneous transluminal coronary angioplasty in patients with coronary artery disease.

■ AF type has been associated with perceived symptom severity and reductions in quality of life.^{52,59}

■ Patients with intermittent AF (paroxysmal and early persistent AF) **had worse impairment of quality of life** than those with chronic AF (persistent and permanent AF).⁵⁹

AF increasingly places a critical financial burden on healthcare systems.

WHAT ARE THE DIRECT AND INDIRECT COSTS OF AF?

The direct cost for the management of AF is highly variable across the Asia Pacific region.

Costs for AF management can be divided into 2 groups:

DIRECT COSTS

- Hospitalization
- Outpatient and Physicians Visits
- Prescriptions
- Laboratory Testing
- Long-term Care

INDIRECT COSTS

- Work Productivity Losses
- Support Provided By Caregivers

The direct costs of AF have been rising exponentially over the past two decades across APAC and are projected to continue to rise.



479% INCREASE IN AF HOSPITALIZATION COSTS BETWEEN 1993-2003 IN AUSTRALIA. This is twice that of myocardial infarction and heart failure.¹⁶



KRW 8.79B → KRW 49.8B INCREASE IN AF-RELATED COST OF CARE FROM 2006-2015. This was equivalent to 0.78% of the Korean national health insurance system (NHIS) expenditure.¹⁹



In 2018, it was estimated that Chinese AF patients over 35 had lost more than 620,000 years of healthy life due to AF-related disability,⁶⁰ which suggests high indirect costs due to caregiver support and productivity loss.

HOW DO STROKE AND HEART FAILURE AFFECT THE COST OF AF?

The cost for the treatment of stroke and heart failure in AF is high, contributing substantially to the total cost of AF management.

STROKE

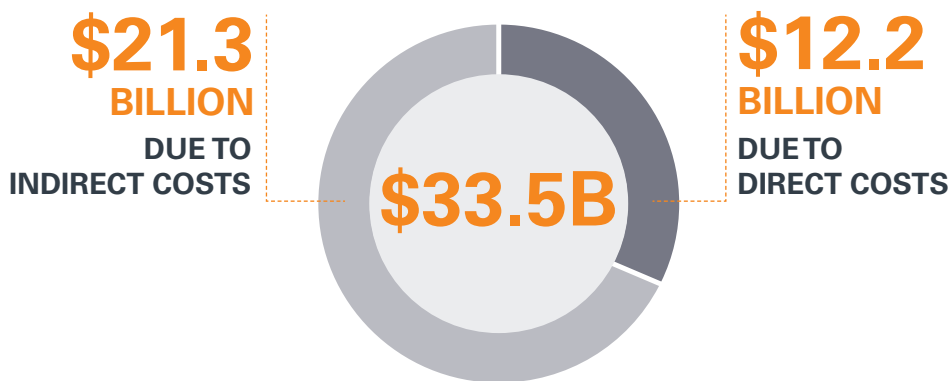
A Chinese study found that the COST of STROKE TREATMENT IN PATIENTS WITH AF is

19.5%

HIGHER than in PATIENTS WITHOUT AF^{(1),61}



In 2016, the overall cost of STROKE across the region composing of China, Australia, Hong Kong, Japan, South Korea, Taiwan, and Thailand was estimated to be USD 33.5 BILLION.⁶²



HEART FAILURE

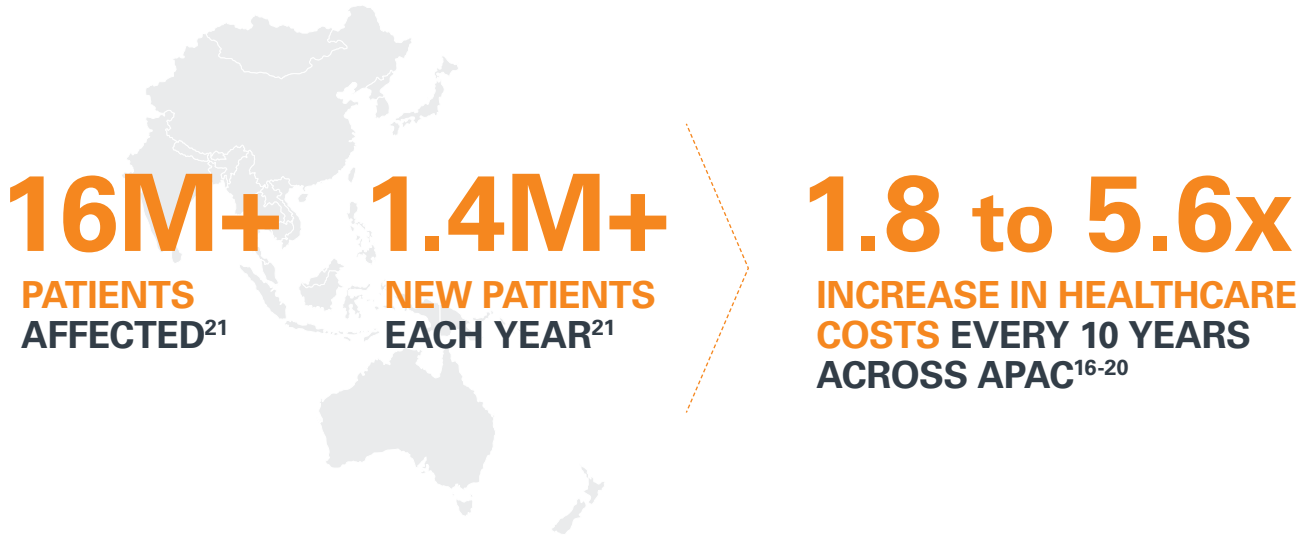
HEART FAILURE is the most common cause of mortality among patients with AF.^{52,53} About 1 out of 5 heart failure patients in Asia has concomitant atrial fibrillation.⁶³

HEART FAILURE PATIENTS in Asia spend an average of 5 to 12.5 days per YEAR in hospitals, incurring COSTS of USD 2400 IN TAIWAN, USD 3600 IN THAILAND, and nearly USD 9000 IN KOREA.⁶³



(1) Relative increased cost based on the comparison to patients without AF, and adjusted for other confounding factors.

AF places a high and increasing financial burden on healthcare systems across the Asia Pacific region.



Given the increasing age, population, and incidence of risk factors for AF in APAC, the **burden** of AF is expected to be far **greater than in any other region in the world.**

BY 2050 ASIA-PACIFIC IS PROJECTED TO HAVE MORE THAN TWICE AS MANY AF-PATIENTS AS EUROPE AND NORTH AMERICA COMBINED.^{28,30,64} This is expected to increase the number of stroke events, hospitalizations, and physician visits, ultimately raising the cost to national healthcare systems.^{16,18-20}

■ LIMITATIONS AND FUTURE DIRECTIONS

To date, summaries about the epidemiology of AF have been disproportionately derived from North American and European populations. There is comparatively limited data in the Asia Pacific region.

Interpretation of the existing data is difficult due to the variation in studied regions, ethnic differences, diagnostic methods, and clinical settings.^{28,29} Additionally, a large part of the evidence is based on data collected over 10 years ago. Findings on prevalence and healthcare-related costs are likely to be underestimated.

Epidemiologists note that additional high-quality data is required. Key areas for exploration and analysis include the following:

- Data harmonization across countries and improvements in data collection⁶⁵
- Validation of country-specific observations in order to identify generalizability of previously established research findings, including risk stratification and outcomes, to a multinational AF population⁶⁵

Additional epidemiological studies in urban and rural areas should also be undertaken in order to:

- Capture a more accurate and updated picture of the incidence and prevalence of AF,^{28,66} especially in remote areas, to guide prevention and management strategies
- Fill existing gaps in the understanding of morbidity, treatment, and its consequent direct and indirect costs, especially in low-income APAC countries where information is scarce
- Clarify risk factors, including the impact of genetic disparities in APAC



1. Laizzo PA. *Handbook of Cardiac Anatomy, Physiology, and Devices*. 2015. Springer Science+Business Media. LLC: Switzerland
2. Ogawa H, An Y, Ikeda S, et al. Progression From Paroxysmal to Sustained Atrial Fibrillation Is Associated With Increased Adverse Events. *Stroke*. 2018;49(10):2301-2308.
3. Staerk L, Sherer JA, Ko D, Benjamin EJ, Helm RH. Atrial Fibrillation: Epidemiology, Pathophysiology, and Clinical Outcomes. *Circ Res*. 2017;120(9):1501-1517.
4. Ikemura N, Kohsaka S, Kimura T, et al. Assessment of Sex Differences in the Initial Symptom Burden, Applied Treatment Strategy, and Quality of Life in Japanese Patients With Atrial Fibrillation. *JAMA Netw Open*. 2019;2(3):e191145.
5. Streur M, Ratcliffe SJ, Ball J, Stewart S, Riegel B. Symptom Clusters in Adults With Chronic Atrial Fibrillation. *J Cardiovasc Nurs*. 2017;32(3):296-303.
6. Wang KL, Wu CH, Huang CC, et al. Complexity of atrial fibrillation patients and management in Chinese ethnicity in routine daily practice: insights from the RealiseAF Taiwanese cohort. *J Cardiol*. 2014;64(3):211-217.
7. Im SI, Chun KJ, Park SJ, Park KM, Kim JS, Oh YK. Long-term Prognosis of Paroxysmal Atrial Fibrillation and Predictors for Progression to Persistent or Chronic Atrial Fibrillation in the Korean Population. *J Korean Med Sci*. 2015;30(7):895-902.
8. Chao TF, Wang KL, Liu CJ, et al. Age Threshold for Increased Stroke Risk Among Patients With Atrial Fibrillation: A Nationwide Cohort Study From Taiwan. *J Am Coll Cardiol*. 2015;66(12):1339-1347.
9. Goto S, Tokai University K, Japan, Oh S, et al. Regional differences in use of antithrombotic therapy for stroke prevention in atrial fibrillation and associated outcomes: European and Asian insights. *European Heart Journal*. 2019;34(suppl_1).
10. Jung YH, Kim YD, Kim J, Han SW, Lee KY. Atrial fibrillation in patients with first-ever stroke: Incidence trends and antithrombotic therapy before the event. *PLoS One*. 2018;13(12):e0209198.
11. Kongbunkiat K, Kasemsap N, Travanichakul S, Thepsuthammarat K, Tiamkao S, Sawanyawisuth K. Hospital mortality from atrial fibrillation associated with ischemic stroke: a national data report. *Int J Neurosci*. 2015;125(12):924-928.
12. Li YG, Lee SR, Choi EK, Lip GY. Stroke Prevention in Atrial Fibrillation: Focus on Asian Patients. *Korean Circ J*. 2018;48(8):665-684.
13. Son MK, Lim NK, Kim HW, Park HY. Risk of ischemic stroke after atrial fibrillation diagnosis: A national sample cohort. *PLoS One*. 2017;12(6):e0179687.
14. Son YJ, Seo EJ. Impact of Sociodemographic and Clinical Factors on Health-Related Quality of Life in Older Adults With Atrial Fibrillation. *Res Gerontol Nurs*. 2018;11(4):207-215.
15. Thrall G, Lane D, Carroll D, Lip GY. Quality of life in patients with atrial fibrillation: a systematic review. *Am J Med*. 2006;119(5):448.e441-419.
16. Gallagher C, Hendriks JM, Giles L, et al. Increasing trends in hospitalisations due to atrial fibrillation in Australia from 1993 to 2013. *Heart*. 2019.
17. Hu S, Zhan L, Liu B, et al. Economic Burden of Individual Suffering from Atrial Fibrillation-Related Stroke in China. *Value Health Reg Issues*. 2013;2(1):135-140.
18. Joung B, Lee JM, Lee KH, et al. 2018 Korean Guideline of Atrial Fibrillation Management. *Korean Circ J*. 2018;48(12):1033-1080.
19. Kim D, Yang PS, Jang E, et al. Increasing trends in hospital care burden of atrial fibrillation in Korea, 2006 through 2015. *Heart*. 2018;104(24):2010-2017.
20. Lee H, Kim TH, Baek YS, et al. The Trends of Atrial Fibrillation-Related Hospital Visit and Cost, Treatment Pattern and Mortality in Korea: 10-Year Nationwide Sample Cohort Data. *Korean Circ J*. 2017;47(1):56-64.
21. Global Burden of Disease Collaborative Network (2017) Global Burden of Disease Study 2017 (GBD 2017) Results. Seattle, United States: Institute for Health Metrics and Evaluation (IHME), 2017. Accessed 2019-07-16. Available from <http://ghdx.healthdata.org/gbd-results-tool>.
22. Ball J, Thompson DR, Ski CF, Carrington MJ, Gerber T, Stewart S. Estimating the current and future prevalence of atrial fibrillation in the Australian adult population. *Med J Aust*. 2015;202(1):32-35.
23. Ball J, et al. *Int J Cardiol*. 2013;167(5):1807-24.
24. Deng H, Guo P, Zheng M, et al. Epidemiological Characteristics of Atrial Fibrillation in Southern China: Results from the Guangzhou Heart Study. *Sci Rep*. 2018;8(1):17829.
25. Guo Y, Tian Y, Wang H, Si Q, Wang Y, Lip GYH. Prevalence, incidence, and lifetime risk of atrial fibrillation in China: new insights into the global burden of atrial fibrillation. *Chest*. 2015;147(1):109-119.
26. Wang X, Fu Q, Song F, et al. Data on prevalence of atrial fibrillation and its association with stroke in low-, middle-, and high-income regions of China. *Data Brief*. 2018;19:1822-1827.
27. Wang X, Fu Q, Song F, et al. Prevalence of atrial fibrillation in different socioeconomic regions of China and its association with stroke: Results from a national stroke screening survey. *Int J Cardiol*. 2018;271:92-97.
28. Wong CX, Brown A, Tse HF, et al. Epidemiology of Atrial Fibrillation: The Australian and Asia-Pacific Perspective. *Heart Lung Circ*. 2017;26(9):870-879.
29. Bai Y, Wang YL, Shantsila A, Lip GYH. The Global Burden of Atrial Fibrillation and Stroke: A Systematic Review of the Clinical Epidemiology of Atrial Fibrillation in Asia. *Chest*. 2017;152(4):810-820.
30. Chiang CE, Wang KL, Lip GY. Stroke prevention in atrial fibrillation: an Asian perspective. *Thromb Haemost*. 2014;111(5):789-797.
31. Bai Y, Guo SD, Shantsila A, Lip GYH. Modelling projections for the risks related with atrial fibrillation in East Asia: a focus on ischaemic stroke and death. *Europace*. 2018;20(10):1584-1590.
32. Chao TF, Liu CJ, Tuan TC, et al. Lifetime Risks, Projected Numbers, and Adverse Outcomes in Asian Patients With Atrial Fibrillation: A Report From the Taiwan Nationwide AF Cohort Study. *Chest*. 2018;153(2):453-466.
33. Chei CL, Raman P, Ching CK, et al. Prevalence and Risk Factors of Atrial Fibrillation in Chinese Elderly: Results from the Chinese Longitudinal Healthy Longevity Survey. *Chin Med J (Engl)*. 2015;128(18):2426-2432.
34. Kim D, Yang PS, Jang E, et al. 10-year nationwide trends of the incidence, prevalence, and adverse outcomes of non-valvular atrial fibrillation nationwide health insurance data covering the entire Korean population. *Am Heart J*. 2018;202:20-26.
35. Koretsune Y, Etoh T, Katsuda Y, et al. Risk Profile and 1-Year Outcome of Newly Diagnosed Atrial Fibrillation in Japan – Insights From GARFIELD-AF –. *Circulation Journal*. 2019;83(1):67-74.
36. Li CY, Lin CP, Lin YS, Wu LS, Chang CJ, Chu PH. Newly diagnosed atrial fibrillation is an independent factor for future major adverse cardiovascular events. *PLoS One*. 2015;10(4):e0123211.

37. The L. Atrial fibrillation and stroke: unrecognised and undertreated. *Lancet*. 2016;388(10046):731.
38. Krittayaphong R, Winijkul A, Methavigul K, et al. Risk profiles and pattern of antithrombotic use in patients with non-valvular atrial fibrillation in Thailand: a multicenter study. *BMC Cardiovasc Disord*. 2018;18(1):174.
39. Guo Y, Wang H, Tian Y, Wang Y, Lip GY. Multiple risk factors and ischaemic stroke in the elderly Asian population with and without atrial fibrillation. An analysis of 425,600 Chinese individuals without prior stroke. *Thromb Haemost*. 2016;115(1):184-192.
40. Kokubo Y, Watanabe M, Higashiyama A, Nakao YM, Kusano K, Miyamoto Y. Development of a Basic Risk Score for Incident Atrial Fibrillation in a Japanese General Population- The Suita Study. *Circ J*. 2017;81(11):1580-1588.
41. Li Y, Pastori D, Guo Y, Wang Y, Lip GYH. Risk factors for new-onset atrial fibrillation: A focus on Asian populations. *Int J Cardiol*. 2018;261:92-98.
42. Briffa T, Hung J, Knuiman M, et al. Trends in incidence and prevalence of hospitalization for atrial fibrillation and associated mortality in Western Australia, 1995-2010. *Int J Cardiol*. 2016;208:19-25.
43. Diouf I, Magliano DJ, Carrington MJ, Stewart S, Shaw JE. Prevalence, incidence, risk factors and treatment of atrial fibrillation in Australia: The Australian Diabetes, Obesity and Lifestyle (AusDiab) longitudinal, population cohort study. *Int J Cardiol*. 2016;205:127-132.
44. Rienstra M, Lubitz SA, Mahida S, et al. Symptoms and functional status of patients with atrial fibrillation: state of the art and future research opportunities. *Circulation*. 2012;125(23):2933-2943.
45. Seow SC, How AK, Chan SP, et al. High Incidence of Occult Atrial Fibrillation in Asian Patients with Cryptogenic Stroke. *J Stroke Cerebrovasc Dis*. 2018;27(8):2182-2186.
46. Hsieh CY, Lee CH, Wu DP, Sung SF. Characteristics and outcomes of ischemic stroke in patients with known atrial fibrillation or atrial fibrillation diagnosed after stroke. *Int J Cardiol*. 2018;261:68-72.
47. Li J, Luo W. Hospitalization expenses of acute ischemic stroke patients with atrial fibrillation relative to those with normal sinus rhythm. *J Med Econ*. 2017;20(2):114-120.
48. Nakamura A, Kuroda J, Ago T, et al. Causes of Ischemic Stroke in Patients with Non-Valvular Atrial Fibrillation. *Cerebrovasc Dis*. 2016;42(3-4):196-204.
49. Savelieva I, Paquette M, Dorian P, Luderitz B, Camm A. Quality of life in patients with silent atrial fibrillation. *Heart*. 2001;85(2):216-217.
50. Odotayo A, Wong CX, Hsiao AJ, Hopewell S, Altman DG, Emdin CA. Atrial fibrillation and risks of cardiovascular disease, renal disease, and death: systematic review and meta-analysis. *Bmj*. 2016;354:i4482.
51. Senoo K, An Y, Ogawa H, et al. Stroke and death in elderly patients with atrial fibrillation in Japan compared with the United Kingdom. *Heart*. 2016;102(23):1878-1882.
52. Lim TW. Baseline, treatment and outcomes with ACS and HFin patients with newly diagnosis AF from Asia. Paper presented at: GARFIELD-AF Asia Pacific Regional Symposium; June 2-3 2018, 2018; Hong Kong.
53. An Y, Ogawa H, Yamashita Y, et al. Causes of death in Japanese patients with atrial fibrillation: The Fushimi Atrial Fibrillation Registry. *Eur Heart J Qual Care Clin Outcomes*. 2019;5(1):35-42.
54. Ohsawa M, Okamura T, Tanno K, et al. Risk of stroke and heart failure attributable to atrial fibrillation in middle-aged and elderly people: Results from a five-year prospective cohort study of Japanese community dwellers. *J Epidemiol*. 2017;27(8):360-367.
55. Healey JS, Oldgren J, Ezekowitz M, et al. Occurrence of death and stroke in patients in 47 countries 1 year after presenting with atrial fibrillation: a cohort study. *Lancet*. 2016;388(10050):1161-1169.
56. Liao JN, Chao TF, Liu CJ, et al. Risk and prediction of dementia in patients with atrial fibrillation--a nationwide population-based cohort study. *Int J Cardiol*. 2015;199:25-30.
57. Zhang L, Hou Y, Po SS. Obstructive Sleep Apnoea and Atrial Fibrillation. *Arrhythm Electrophysiol Rev*. 2015;4(1):14-18.
58. Dorian P, Jung W, Newman D, et al. The impairment of health-related quality of life in patients with intermittent atrial fibrillation: implications for the assessment of investigational therapy. *J Am Coll Cardiol*. 2000;36(4):1303-1309.
59. Nazlı C, Eren NK, Tülüce SY, Yağız İ GK, Kılıçaslan B, Kocabaş U. Impaired quality of life in patients with intermittent atrial fibrillation. In: *Anatol J Cardiol*. Vol 16.2016:250-255.
60. Wang Z, Chen Z, Wang X, et al. The Disease Burden of Atrial Fibrillation in China from a National Cross-sectional Survey. *Am J Cardiol*. 2018;122(5):793-798.
61. Wen L, Wu J, Feng L, Yang L, Qian F. Comparing the economic burden of ischemic stroke patients with and without atrial fibrillation: a retrospective study in Beijing, China. *Curr Med Res Opin*. 2017;33(10):1789-1794.
62. Dalai R. The Cost of Silence: Cardiovascular disease in Asia. *The Economist*. 2019.
63. Reyes EB, Ha JW, Firdaus I, et al. Heart failure across Asia: Same healthcare burden but differences in organization of care. *Int J Cardiol*. 2016;223:163-167.
64. Chiang CE, Okumura K, Zhang S, et al. 2017 consensus of the Asia Pacific Heart Rhythm Society on stroke prevention in atrial fibrillation. *J Arrhythm*. 2017;33(4):345-367.
65. Hsu JC, Akao M, Abe M, et al. International Collaborative Partnership for the Study of Atrial Fibrillation (INTERAF): Rationale, Design, and Initial Descriptives. *J Am Heart Assoc*. 2016;5(11).
66. Zulkify H, Lip GYH, Lane DA. Epidemiology of atrial fibrillation. *Int J Clin Pract*. 2018;72(3):e13070.

Manufacturer | Biosense Webster

33 Technology Drive, Irvine | California 92618, USA

Tel: +1-909-839-8500 | Tel: +1-800-729-9010 | Fax: +1-909-468-2905

www.biosensewebster.com

