

## Lifestyles and their Role in the Prevention and Management of Cardiovascular Diseases

\*<sup>1</sup>Shashi K Agarwal

\*<sup>1</sup>Director, Center for Contemporary and Complimentary Cardiology, North Brunswick, United States.

### Abstract

Cardiovascular diseases are rampant globally. Coronary artery illness, cardiac attack, stroke, hypertension, and a variety of other disorders are among them. Besides inflicting a huge amount of human pain, they are also responsible for a massive direct and indirect financial cost on the global society. With the widespread attainability of low-cost treatments and the related scarcity of innovative discoveries, increasing research is being focused on the impact of healthy lifestyles on these diseases. Nonsmoking, low consumption of alcohol, a normal BMI, frequently taking exercise, and a wise diet are all examples of healthy lives. Adherence to these helps prevent and attenuate the severity of cardiovascular diseases, decrease their related disabilities, and enhance the health status of life in patients. Cardiovascular diseases are the major cause of mortality worldwide. Healthy lifestyles also help reduce cardiovascular mortality. It is estimated that in 2019, almost 18 million individuals died from cardiovascular diseases all over the world. Over three-quarters of these deaths occurred in low-and average-income countries. Unfortunately, cardiovascular deaths are expected to increase, due to the global growth in population, the continuing Westernization of the developing countries, and the increase in the number of the aged. The World Health Organization estimates that by managing lifestyle risk factors, can stop three-quarters of these deaths. This manuscript will briefly review the influence of these lifestyles on major cardiovascular diseases.

**Keywords:** Cardiovascular diseases, lifestyles, alcohol, exercise, smoking, obesity, diet

### Introduction

Cardiovascular diseases (CVDs) are a heterogeneous group of diseases of the heart and the circulatory system <sup>[1]</sup>. Hypertension (HTN), coronary heart disease (CHD), stroke, heart failure (HF), cardiac arrhythmias, and valvular disease, cardiomyopathies, endocarditis and myocarditis, congenital heart disease, peripheral artery disease (PAD), vasculogenic erectile dysfunction (ED) in men, and venous thromboembolism (VTE) are examples of these conditions <sup>[2]</sup>. Men and women are nearly equally affected <sup>[3]</sup> with the biggest burden being on the elderly <sup>[4]</sup>. CVDs are associated with extremely high morbidity <sup>[5]</sup>. The underlying cause for most CVDs is usually atherosclerosis <sup>[5, 6]</sup>, especially in myocardial infarction, ischemic stroke, and peripheral arterial disease. Cardiovascular diseases are also increasingly being responsible for disability <sup>[7, 8]</sup>. From 17.7 million in 1990 to 34.4 million in 2019, the number of YLDs (years of life lived with disability) for CVDs has two times <sup>[9]</sup>. CVDs continue to be the dominant cause of premature mortality worldwide <sup>[9-11]</sup>. The number of people dying from cardiovascular disease has risen from 12.1 million in 1990 to 18.6 million in 2019 <sup>[9]</sup>. One-third of all deaths worldwide are caused by them <sup>[10]</sup>. Of all the global deaths in 2015, 82% were in low-and average-revenue countries <sup>[11]</sup>. The WHO predicts that by 2030, 23.6 million individuals would pass away from CVDs every year <sup>[12]</sup>.

Hypertension (HTN) is diagnosed if the systolic blood pressure level is 130mmHg or higher, and/or the diastolic level of BP is 80mmHg or higher <sup>[13]</sup>. It's a fairly common illness, with a prevalence between 30-50% in adults

\*Corresponding Author: Shashi K Agarwal

worldwide <sup>[14]</sup>. It is responsible for 22.3% of the global CVD burden <sup>[15]</sup> and accounted for 10.4 million deaths in 2017 <sup>[16]</sup>. Controlling HTN helps reduce several CVDs <sup>[17]</sup>, including coronary artery disease (CAD) <sup>[18, 19]</sup>, stroke <sup>[20]</sup>, heart failure <sup>[21]</sup>, and cardiac arrhythmias <sup>[22]</sup>. Ischemic heart disease (IHD) is known as very usual CVD <sup>[10]</sup>. "It includes acute myocardial infarction, chronic stable angina, chronic IHD, and heart failure due to IHD" <sup>[23]</sup>. The underlying cause is atherosclerotic disease of the arteries <sup>[24]</sup>. In the year 2019, Estimates suggest that 197 million prevalent cases of IHD and 9.14 million IHD deaths worldwide <sup>[9]</sup>. The World Health Organization classifies stroke as ischemic stroke, intracerebral hemorrhage, and subarachnoid hemorrhage <sup>[25]</sup>. 7.63 million ischemic strokes, 3.41 million intracerebral haemorrhage, and 1.18 million subarachnoid haemorrhage were estimated among the 12.2 million incident stroke cases in 2019 <sup>[9]</sup>. The numbers of disability-adjusted life years (DALYs) because of stroke and death have also increased steadily over the past few decades <sup>[26]</sup>. In 2016, there were 116.4 million DALYs and 5.5 million deaths due to stroke <sup>[26]</sup>. HF is a leading cause of morbidity, by which around 26 million people are affected globally <sup>[27, 28]</sup>. Its occurrence is growing. It leads to frequent hospitalizations, poor life quality, and is a notable cause of early death <sup>[27]</sup>. "It is divided into two categories depending on the left ventricular ejection fraction: heart failure with decreased ejection fraction (HFrEF) and heart failure with preserved ejection fraction (HFpEF) (HFpEF)" <sup>[29]</sup>. Most heart failures are a result of hypertension <sup>[30]</sup>. Cardiomyopathy is the result of heart failure caused by primary myocardial disease or myocardial damage caused by toxins such as

alcohol [31]. Alcoholic cardiomyopathy, a non-ischemic dilated cardiomyopathy, resulted in 708,000 cases in 2019 and was responsible for 71,700 deaths [9]. Cardiac arrhythmias may be symptomatic and are usually diagnosed by an electrocardiogram. The most usual arrhythmia to worry about is atrial fibrillation [32], and its incidence is on the rise [33, 34]. It has a high morbidity [35] and mortality rate [36]. Sudden cardiac death occurs when a clinically stable patient, with or without pre-existing heart disease, dies from a cardiac cause within one hour of an abrupt and drastic change in their clinical status [37]. Sudden cardiac deaths are as frequent as deaths from several cancers, such as breast cancer [38]. It is estimated that they are responsible for about 400,000 deaths in the USA, and about 300,000 deaths in Europe, every year [39]. They are predominantly caused by malignant ventricular arrhythmias [40]. PAD is primarily an atherosclerotic disease<sup>41</sup> characterized by claudication [42] and “diagnosed by an ankle-brachial index (ABI) of <0.9” [43]. “The ABI is calculated by measuring the brachial, dorsalis pedis (DP), and posterior tibial (PT) arteries in both legs and utilizing the higher of the DP/PT in each leg as the numerator” [44, 45]. The lower of these ratios represent the person’s ABI. “PAD is also a risk factor for atherosclerotic heart disease” [46, 47] and is accompanied by a poor prognosis [48, 49]. The disability to get or sustain an adequate erection to allow good genital performance is referred to as ED [50]. “It is a prevalent disease that affects 52% of men between the ages of 40 and 70, and >70% of men beyond the age of 70” [51]. Although the causes may be neurogenic, anatomical, hormonal, drug-related, or psychogenic, atherosclerosis also plays an essential role, resulting in arterial insufficiency and causing vasculogenic ED [52]. In men over 50, vasculogenic ED is the most common cause [52]. Although less prevalent, other CVDs have a substantial influence on worldwide morbidity and death.

## Discussion

Lifestyles are day-to-day behaviors, and they have a notable effect on human morbidity and mortality [53, 54]. A mega-analysis of 15 international studies (over 500,000 participants) estimated that unhealthy lifestyles (smoking, inactivity, poor diet, obesity, and excessive alcohol intake) were responsible for over half of the global premature deaths [53]. According to more recent study, Yanping *et al.* estimated that when compared to those with zero low-risk lifestyle variables, it is anticipated that adhering to the five healthy lifestyle-related factors described above might extend life expectancy at 50 years by 14.0yrs for females and 12.2yrs for males in the US [54]. The impact of these five lifestyle behaviors on CVDs is discussed in this manuscript. Exercise is described as “any sport or activity that utilizes large groups of muscles, is sustained continuously, and is performed rhythmically” [55]. “Adults should acquire at least 150 minutes of moderate-intensity aerobic activity or 75 minutes of strenuous aerobic activity (or a balance of both) each week, preferably spaced out throughout the week, according to the AHA” [56]. This should be followed by at least two days of muscle-strengthening exercise [56]. A person's body mass index (BMI) indicates his or her body weight category, “which is calculated by dividing a person's weight by the square of his or her height in meters” [57]. “As per the World Health Organization (WHO), a BMI of 20 to 25 kg/m<sup>2</sup> is regarded normal, a BMI of 25 to 30 kg/m<sup>2</sup> is considered overweight, and a BMI of greater than 30 kg/m<sup>2</sup> is considered obese” [58]. Smoking is the pivotal preventable cause of CVDs and several other serious diseases [59]. Tobacco can be smoked

using cigarettes, cigars, water pipes, bidis, and kreteks [60]. First-hand smoke directly enters into the smoker’s mouth from puffing and is also known as mainstream smoke [61]. Smoke emits from the smoldering ends of a side-stream cigarette. “Second-hand smoke is a combination of side-stream smoke (85%) and the exhaled mainstream smoke” (15%) [62]. “The gas and particle residue from smoked tobacco products that stick to surfaces including hair, skin, clothes, and furniture is known as third-hand smoke” [63]. These pollutants may persist for many days and when airborne, may be inhaled by non-smokers [64]. Diet plays an important role in health [65-67]. Besides caloric restriction to avoid overweight and obesity (maintaining a BMI below 25kg/m<sup>2</sup>) [65]. “A well-balanced diet includes plenty of non-starchy vegetables, fruits, whole grains, and legumes, as well as restricted to moderate amounts of nuts, seafood, lean meats, low-fat dairy products, and vegetable oil, rich in mono and polyunsaturated fats” [66, 67]. “Trans fats, saturated fats, fried meals, excess salt, red meat, refined carbs, and sugar-sweetened drinks are also restricted or eliminated” [66, 67]. Alcohol is a double-edged sword when it comes to CVDs [68]. “A standard drink in the United States comprises 12-15g of pure ethanol. In the US, a standard drink contains 12-15g of pure ethanol and this is present in 100-125ml of wine, 240-300mL of beer, and 30-37.5mL of spirits” [69]. “Two standard drinks each day for males and one standard drink each day for females is considered moderate alcohol consumption” [70, 71]. Heavy level of drinking is explained as a regular consumption of >60g/day for males and >40g/day for females [72]. “Binge drinking is defined as consuming 4 or more drinks in one sitting for women and 5 or more drinks in one sitting for men” [73, 74]. Excessive alcohol use may result in alcohol use disorder [75]. All these health behaviors, if improperly adhered to, have detrimental effects on CVDs [76].

**Exercise:** Physical activity and pro-active physical exercise slow down and may even reverse CVD progression [77-81]. Moderate physical activity is linked to a 26% decrease in CVD risk, whereas high-intensity activity imparts a 42% reduction in risk [80-84]. Patients regularly performing moderate to high-intensity exercise gain 1-3 years of CVD-free life when compared with their sedentary peers [85]. They reduce their mortality and enhance their expected lifespan by 1.3 to 3.7 years [85]. Long-term exercise lowers BP similar in magnitude to those obtained by first-line anti-hypertensive medications [86]. BP is reduced by 5-7 mmHg with aerobic exercise while resistance exercise lowers BP by 2-3 mmHg [87, 88]. Ischemic heart disease is beneficially affected [89] and stroke is reduced [90] with regular exercise. Post-stroke exercise rehabilitation helps improve walking speed and endurance [91]. Its benefits in HF are also significant [92-94]. Exercise, especially through HF rehabilitation, results in reduced hospitalizations [95], improved quality of life [96], and a lower mortality [97]. Benefits have also been noted in atrial fibrillation [98]. Besides its preventive action [99], exercise helps PAD patients walk more 100 and improves their quality of life [101, 102]. Exercising patients also notice a reduction in the risk of ED [103], while established ED patients notice an improvement in performance [104]. Exercise also reduces several other CVD risk components such as smoking [105], obesity [106], diabetes mellitus [107], hyperlipidemia [108], alcohol abuse [109], chronic kidney disease [110], depression [111], loneliness [112], psychosomatic stress [112], sleep disturbances [113], and illicit drug use [114]. The beneficial effects of exercise stem from its ability to reduce body

weight, improve lipid profiles [115], and induce reductions in insulin resistance, blood coagulation, and systemic inflammation [116]. “Lowering heartbeat, increment in the supply of myocardial oxygen supply, and improving myocardial contraction and stroke volume are included as the direct impacts on the heart” [117-119]. Individuals with known CVD should, however, seek medical advice before embarking on an exercise program [120, 121].

### Obesity

Obesity increases CVD risk [122-124] while weight loss reduces it [125, 126]. It is estimated that obesity is thought to be responsible for 70% of HTN in adults [127, 128]. Obese patients have more incident CAD, complex coronary lesions, poor results to interventions, and higher mortality [129-132]. Obesity also deleteriously affects HF [133, 134]. “These patients have poor outcomes with left ventricular assist device implantation or heart transplantation” [135, 136]. On the other hand, they improve clinically with bariatric surgery-induced weight loss [137]. Patients who are obese are also more likely to having a stroke. [138, 139] and “exhibit a >2-fold increase in risk” [140]. “Patients with BMI of  $\geq 35$  kg/m<sup>2</sup> have a three times increase in the risk of having a stroke” [141]. Obesity also increases the risk of AF [142] and SCD [143]. It negatively impacts several other CVDs, including aortic stenosis [144], peripheral artery disease [145], erectile dysfunction [146], and venous thromboembolism [147]. “It also increases several CVD risk factors, such as sleep apnea, diabetes mellitus, and dyslipidemia” [148]. Obesity induces proinflammatory changes and free radical damage, resulting in endothelium dysfunction and atherosclerosis [149].

### Smoking

Tobacco smoking results in an acute rise in systolic and diastolic BP [150]. The long-term effects on HTN are however unclear. Active smoking increases the coronary artery disease risk by about 80%, while passive smoking increases this risk by about 30% [151]. The risk of a stroke and atrial fibrillation is increased by smoking cigarette. A separate risk component for heart failure is smoking also [152]. Smokers also have a higher risk of dying from a heart attack [153]. Smoking is mainly related with PAD and the development of abdominal aortic aneurysms [154, 155]. For the development of ED, it is established as individual risk factor, while also increasing the risk of VTE [156, 157]. When compared to non-smokers, It's been proven that smoking low-tar cigarettes raises the risk of cardiovascular events. “Cigarette smoking enhances atherosclerosis via increased inflammation, pro-thrombotic changes, and oxidation of low-density lipoprotein” [158, 159].

### Diet

Improper diet is a strong modulator of CVDs risk [160-162]. A plethora of scientific studies indicates that red and processed meat [163], salt [164], fried foods [165], sugar-sweetened beverages [166], excess alcohol intake [167], and obesity [168] have detrimental effects on CVDs, while fruits and vegetables [169, 170], whole grains [171], fiber [172], tree nuts [173], chocolate [174], and coffee [175] are cardio-protective. Salt restriction of <1500 mg/day [176] reduces BP by about -5/6mmHg systolic and -2/3mmHg diastolic [177]. A healthy diet as mentioned above, the DASH diet [178], or the Mediterranean diet [179] also helps in BP reduction. Controlling HTN helps reduce several CVDs, including coronary artery disease [181, 182], stroke [183], heart failure [184], and cardiac arrhythmias [185]. 21% risk of CAD is reduced by The DASH diet [186] and is

inversely associated with CAD [187]. Mediterranean diet has been reported to induce a reduction in coronary mortality of 65% over a study period of 46 months [188]. Benefits of a healthy diet have also been recorded with stroke [189], HF [190], cardiac arrhythmias [191, 192], PAD [193], and ED [194]. A cardiovascular healthy diet also helps reduce hypercholesterolemia [195], DM [196], and chronic kidney disease [197], both being CVDs risk factors.

### Alcohol

The relation between cardiovascular disease and alcohol consumption is J-or U-shaped [198]. Alcohol is beneficial for CVD in low to moderate amounts [199] and harmful in heavy amounts [200]. A detrimental association has also been seen with binge drinking [201] and chronic alcohol abuse, as seen in AUD [202]. BP is strongly related to alcohol intake [203, 204], with an increased incidence of HTN of 50% with an intake of 3-4 drinks per day and a 100% increase with an intake of 6-7 drinks per day [205]. Decreasing alcohol consumption to <2 drinks/day helps reduce BP [206]. Low to moderate alcohol consumption is also cardioprotective for CHD morbidity and mortality [207, 208]. A similar pattern has been noted during alcohol consumption and stroke [209]. “Heavy alcohol consumption ( $\geq 5$  drinks/day or  $\geq 35$  drinks/week) considerably increases the risk of HF” [210, 211]. Alcohol also increases the risk of developing AF [212, 213]. “Koskinen *et al* estimated that “5%-10% of all new episodes of AF were related to alcohol consumption” [214]. Alcohol has also been linked to SCD [215, 216]. “The risk of congenital anomalies such as aberrant great vessels, atrial septal defects, and ventricular septal defects is increased in offspring of pregnant women who drink alcohol” [217, 218]. “Heavy alcohol consumption is related with a higher level of risk of PAD” [219]. According to research, light to moderate alcohol use (<21 drinks/week) reduces the incidence of erectile dysfunction [220]. “Many other CVD risk factors, such as sleep disorders [221], chronic kidney disease [222], weight gain [223], depression [224] and smoking” [225] also associated with consumption of alcohol. Low to moderate alcohol intake helps DM [226], increases HDL-C [227-229], provides atherosclerotic plaque stabilization [230], and improves the hemostatic status [231-234].

### Conclusion

CVDs are rampant globally, and they are estimated to increase, due to a high level of population growth and an increase in the aged. Most CVDs can be prevented and progressed by adopting a healthier lifestyle. In addition, healthy behaviors also provide a better life quality and an increment in lifespan, on the other side Structured activities involve rules, can be competitive or cooperative, and typically demand the use of one's cerebral abilities rather than physical strength. To excel in sports and games one should have excellent technique, tactics, training, skill and etcetera. Field Handball performance is heavily influenced by sports science. Field handball is a team sport that necessitates a variety of external and internal criteria such as anthropometrical, physical, physiological, and psychological features in order to compete at the highest level.

### References

1. <https://www.webmd.com/heart-disease/guide/diseases-cardiovascular> - accessed, 2021.
2. [https://www.who.int/news-room/factsheets/detail/cardiovascular-diseases-\(cvds\)](https://www.who.int/news-room/factsheets/detail/cardiovascular-diseases-(cvds))-accessed, 2021.

3. World Health Organization Cardiovascular Disease. [accessed], 2019. Available online: [https://www.who.int/cardiovascular\\_diseases/about\\_cvd/en/](https://www.who.int/cardiovascular_diseases/about_cvd/en/).
4. Yazdanyar A, Newman AB. The burden of cardiovascular disease in the elderly: morbidity, mortality, and costs. *Clin Geriatr Med*. 2009; 25(4):563-77, vii. Doi: 10.1016/j.cger.2009.07.007.
5. Hogas S, Bilha SC, Branisteanu D, Hogas M, Gaipov A, Kanbay M, Covic A. Potential novel biomarkers of cardiovascular dysfunction and disease: Cardiotrophin-1, adipokines and galectin-3. *Arch. Med. Sci*. 2017; 4:897-913. Doi: 10.5114/aoms.2016.58664.
6. Moss JW, Ramji DP. Nutraceuical therapies for atherosclerosis. *Nat Rev Cardiol*. 2016; 13(9):513-32. Doi: 10.1038/nrcardio.2016.103.
7. Roth GA, Mensah GA, Johnson CO *et al*. GBD-NHLBI-JACC Global Burden of Cardiovascular Diseases Writing Group. *Global Burden of Cardiovascular Diseases and Risk Factors*, 1990, 2019.
8. Update From the GBD 2019 Study. *J Am Coll Cardiol*. 2020; 76(25):2982-3021. Doi: 10.1016/j.jacc.2020.11.010.
9. Roth GA, Mensah GA, Johnson CO *et al*. GBD-NHLBI-JACC Global Burden of Cardiovascular Diseases Writing Group. *Global Burden of Cardiovascular Diseases and Risk Factors*, 1990-2019: Update from the GBD 2019 Study. *J Am Coll Cardiol*. 2020 ; 76(25):2982-3021. Doi: 10.1016/j.jacc.2020.11.010.
10. Roth GA, Johnson C, Abajobir A, Abd-Allah F, Abera SF, Abyu G *et al*. Global, regional, and national burden of cardiovascular diseases for 10 causes, 1990 to 2015. *Journal of the American College of Cardiology*. 2017; 70(1):1-25. DOI: 10.1016/j.jacc.2017.04.052.
11. World Health Organization. *World Health Statistics 2018: Monitoring Health for the SDGs: Sustainable Development Goals*. World Health Organization; Geneva, Switzerland: Licence: CC BY-NC-SA 3.0 IGO, 2018.
12. <https://www.who.int/>. - accessed August 14, 2021.
13. Whelton PK, Carey RM, Aronow WS *et al*. ACC/AHA/AAPA/ABC/ACPM/AGS/APhA/ASH/ASPC/NMA/PCNA guideline for the prevention, detection, evaluation, and management of high blood pressure in adults: a report of the American College of Cardiology/American Heart Association task force on clinical practice guidelines. *J Am Coll Cardiol*, 2017, 2018; 71(19):e127-e248. Doi:10.1016/j.jacc.2017.11.006.
14. Zhou B, Danaei G, Stevens G, Bixby H, Taddei C, Carrillo-Larco RM *et al*. Long-term and recent trends in hypertension awareness, treatment, and control in 12 high-income countries: An analysis of 123 nationally representative surveys. *Lancet*. 2019; 394:639-651. Doi: 10.1016/S0140-6736(19)31145-6.
15. Yusuf S, Joseph P, Rangarajan S *et al*. Modifiable risk factors, cardiovascular disease, and mortality in 155 722 individuals from 21 high-income, middle-income, and low-income countries (PURE): A prospective cohort study. *Lancet*. 2020; 395:795-808. Doi: 10.1016/S0140-6736(19)32008-2.
16. Collaborators GBDRF. Global, regional, and national comparative risk assessment of 84 behavioural, environmental and occupational, and metabolic risks or clusters of risks for 195 countries and territories, 1990-2017: A systematic analysis for the Global Burden of Disease Study 2017. *Lancet*. 2018; 392:1923-1994.
17. Lewington S, Clarke R, Qizilbash N, Peto R, Collins R, Prospective Studies C; Prospective Studies Collaboration. Age-specific relevance of usual blood pressure to vascular mortality: a meta-analysis of individual data for one million adults in 61 prospective studies. *Lancet*. 2002; 360(9349):1903-1913. Doi:10.1016/S0140-6736(02)11911-8.
18. Luo D, Cheng Y, Zhang H *et al*. Association between high blood pressure and long term cardiovascular events in young adults: systematic review and meta-analysis. *BMJ*. 2020; 370:m3222. Doi:10.1136/bmj.m3222
19. Kannel WB. Hypertension: reflections on risks and prognostication. *Med Clin North Am*. 2009; 93:541-558
20. Benjamin EJ, Blaha MJ, Chiuve SE, Cushman M, Das SR, Deo R *et al*. Heart Disease and Stroke Statistics-2017 Update: a report from the American Heart Association. *Circulation*. 2017; 135:e146-e603.
21. Slivnick J, Lampert BC. Hypertension and Heart Failure. *Heart Fail Clin*. 2019; 15(4):531-541. Doi: 10.1016/j.hfc.2019.06.007.
22. Varvarousis D, Kallistratos M, Poulimenos L *et al*. Cardiac arrhythmias in arterial hypertension. *J Clin Hypertens* (Greenwich), 2020. Doi: 10.1111/jch.13989.
23. Joseph A, Hill and Eric N, Olson, Editors, Chapter 36- Ischemic Heart Disease, Academic Press, ISBN: 978-0-12-381510-1; <https://doi.org/10.1016/B978-0-12-381510-1.00036-3>.
24. Rognoni A, Cavallino C, Veia A *et al*. Pathophysiology of Atherosclerotic Plaque Development. *Cardiovasc Hematol Agents Med Chem*. 2015; 13(1):10-3. Doi: 10.2174/1871525713666141218163425.
25. [https://www.who.int/healthinfo/statistics/bod\\_cerebrovascular\\_diseases\\_stroke.pdf](https://www.who.int/healthinfo/statistics/bod_cerebrovascular_diseases_stroke.pdf).
26. GBD 2016 Stroke Collaborators. Global, regional, and national burden of stroke, 1990-2016: a systematic analysis for the Global Burden of Disease Study 2016. *Lancet Neurol*. 2019; 18(5):439-458. Doi: 10.1016/S1474-4422(19)30034-1.
27. Savarese G, Lund LH. Global Public Health Burden of Heart Failure. *Card Fail Rev*. 2017; 3(1):7-11. Doi:10.15420/cfr.2016:25:2.
28. Conrad N, Judge A, Tran J *et al*. Temporal trends and patterns in heart failure incidence: a population-based study of 4 million individuals. *Lancet*. 2018; 391:572-580. Doi: 10.1016/S0140-6736(17)32520-5.
29. Ponikowski P, Voors AA, Anker SD *et al*. Authors/Task Force Members. 2016 ESC guidelines for the diagnosis and treatment of acute and chronic heart failure: The Task Force for the diagnosis and treatment of acute and chronic heart failure of the European Society of Cardiology (ESC) Developed with the special contribution of the Heart Failure Association (HFA) of the ESC. *Eur Heart J*. 2016; 37:2129-200. Doi: 10.1093/eurheartj/ehw128.
30. Patel Y, Joseph J. Sodium Intake and Heart Failure. *Int J Mol Sci*. 2020; 21(24):9474. Doi:10.3390/ijms21249474.
31. Piano MR. Alcoholic cardiomyopathy: incidence, clinical characteristics, and pathophysiology. *Chest* 2002; 121:1638-50.
32. Chugh SS, Havmoeller R, Narayanan K, Singh D, Rienstra M, Benjamin EJ *et al*. Worldwide epidemiology of atrial fibrillation: a Global Burden of Disease 2010 Study. *Circulation*. 2014; 129(8):837-847.

33. Colilla S, Crow A, Petkun W, Singer DE, Simon T, Liu X. Estimates of current and future incidence and prevalence of atrial fibrillation in the U.S. adult population. *Am J Cardiol.* 2013; 112(8):1142-1147.
34. Di Carlo A, Bellino L, Consoli D, Mori F, Zaninelli A, Baldereschi M *et al.* Prevalence of atrial fibrillation in the Italian elderly population and projections from 2020 to 2060 for Italy and the European Union: the FAI Project. *Europace.* 2019; 21(10):1468-1475.
35. Guo Y, Lip GY, Apostolakis S. The unmet need of stroke prevention in atrial fibrillation in the far East and South East Asia. *Malays J Med Sci.* 2012; 19:1-7.
36. Stewart S, Hart CL, Hole DJ *et al.* A population-based study of the long-term risks associated with atrial fibrillation: 20-year follow-up of the Renfrew/Paisley study. *Am J Med.* 2002; 113:359-364.
37. [http://www.columbia.edu/itc/hs/medical/pathophys/cardiology/2009/suddendeathColor\\_old.pdf](http://www.columbia.edu/itc/hs/medical/pathophys/cardiology/2009/suddendeathColor_old.pdf).
38. Haissaguerre M, Hocini M, Sacher F, Shah A. La mort subite cardiaque: un défi scientifique majeur [Sudden cardiac death, a major scientific challenge]. *Bull Acad Natl Med.* 2010; 194(6):983-93.
39. Kuriachan VP, Sumner GL, Mitchell LB. Sudden cardiac death. *Curr Probl Cardiol.* 2015; 40(4):133-200. Doi: 10.1016/j.cpcardiol.2015.01.002.
40. John RM, Tedrow UB, Koplán BA, Albert CM, Epstein LM, Sweeney MO *et al.* Ventricular arrhythmias and sudden cardiac death. *Lancet.* 2012; 380(9852):1520-9. Doi: 10.1016/S0140-6736(12)61413-5.
41. Simon F, Oberhuber A, Floros N, Düppers P, Schelzig H, Duran M. Pathophysiology of chronic limb ischemia. *Gefasschirurgie.* 2018; 23(1):13-18.
42. Sontheimer DL. Peripheral vascular disease: diagnosis and treatment. *Am Fam Physician.* 2006; 73(11):1971-6.
43. Hirsch AT, Haskal ZJ, Hertzner NR. ACC/AHA Guidelines for the Management of Patients with Peripheral Arterial Disease (lower extremity, renal, mesenteric, and abdominal aortic): a collaborative report from the American Associations for Vascular Surgery/Society for Vascular Surgery, Society. *J Vasc Interv Radiol.* 2006; 17:1383-1397.
44. Gerhard-Herman MD, Gornik HL, Barrett C *et al.* AHA/ACC Guideline on the Management of Patients With Lower Extremity Peripheral Artery Disease: a report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. *Circulation.* 2016, 2017; 135(12):e726-e779.
45. Aboyans V, Criqui MH, Abraham P *et al.* Measurement and interpretation of the ankle-brachial index: a scientific statement from the American Heart Association. *Circulation.* 2012; 126(24):2890-2909.
46. Trevisan SM, Vittinghoff E, Owens CD, Whooley M, Cohen BE. Peripheral artery disease and risk of cardiovascular events in patients with coronary artery disease: insights from the Heart and Soul Study. *Vascular Medicine.* 2013; 18(4):176-184. Doi: 10.1177/1358863X13493825.
47. US Preventive Services Task Force, Curry SJ, Krist AH, Owens DK, Barry MJ, Caughey AB, Davidson KW *et al.* Screening for Peripheral Artery Disease and Cardiovascular Disease Risk Assessment with the Ankle-Brachial Index: US Preventive Services Task Force Recommendation Statement. *JAMA.* 2018; 320(2):177-183. Doi: 10.1001/jama.2018.8357.
48. McDermott MM. Lower extremity manifestations of peripheral artery disease: the pathophysiologic and functional implications of leg ischemia. *Circ Res.* 2015; 116:1540-50. 10.1161/CIRCRESAHA.114.303517.
49. Zhu J, Tan X, Zhou JZ. Peripheral artery disease and clinical outcomes in patients with atrial fibrillation: A systematic review and meta-analysis. *Clin Cardiol.* 2021; 44(8):1050-1057. Doi:10.1002/clc.23678.
50. Salonia A, Bettocchi C, Carvalho J, Corona G, Jones TH, Kadioglu A. European Association of Urology Guidelines. 2020 ed. European Association of Urology Guidelines Office; Arnhem, The Netherlands: 2020. EAU Guidelines on Sexual and Reproductive Health 2020. Presented at the EAU Annual Congress Amsterdam, 2020.
51. Selvin E, Burnett AL, Platz EA. Prevalence and risk factors for erectile dysfunction in the US. *Am J Med.* 2007; 120:151-157.
52. Wang TD, Lee CK, Chia YC *et al.* HOPE Asia Network. Hypertension and erectile dysfunction: The role of endovascular therapy in Asia. *J Clin Hypertens (Greenwich).* 2021; 23(3):481-488. Doi: 10.1111/jch.14123.
53. Martin Loef, Harald Walach. The combined effects of healthy lifestyle behaviors on all-cause mortality: A systematic review and meta-analysis. *Preventive Medicine.* 2012; 55(3):163-170. <https://doi.org/10.1016/j.ypmed.2012.06.017>.
54. Yanping Li, An Pan, Dong D, Wang *et al.* Impact of Healthy Lifestyle Factors on Life Expectancies in the US Population. *Circulation.* 2018; 138:345-355. <https://doi.org/10.1161/CIRCULATIONAHA.117.032047>.
55. <https://healthyliving.azcentral.com/acsm-definition-cardiovascular-exercise-18723.html> - accessed November 7, 2020.
56. Taylor AE, Johnson DC, Kazemi H. Environmental tobacco smoke and cardiovascular disease: A position paper from the Council on Cardiopulmonary and Critical Care, American Heart Association. *Circulation.* 1992; 86:699-702.
57. Physical status: the use and interpretation of anthropometry. Report of a WHO Expert Committee. World Health Organ Tech Rep Ser. 1995; 854:1-452.
58. WHO Expert Consultation: Appropriate body-mass index for Asian populations and its implications for policy and intervention strategies. *Lancet* 2004; 363:157-163.
59. Samet JM. Tobacco smoking: the leading cause of preventable disease worldwide. *Thorac Surg Clin.* 2013; 23(2):103-12. Doi: 10.1016/j.thorsurg.2013.01.009.
60. Shihadeh A, Schubert J, Klaiany J, El Sabban M, Luch A, Saliba NA. Toxicant content, physical properties and biological activity of water pipe tobacco smoke and its tobacco-free alternatives. *BMJ.* 2015; 24(1):22-30.
61. Pryor WA, Stone K. Oxidants in cigarette smoke: Radicals, hydrogen peroxide, peroxyoxynitrate, and peroxyoxynitrite. *Ann NY Acad Sci.* 1993; 686:12-28.
62. Taylor AE, Johnson DC, Kazemi H. Environmental tobacco smoke and cardiovascular disease: A position paper from the Council on Cardiopulmonary and Critical Care, American Heart Association. *Circulation.* 1992; 86:699-702.
63. Winickoff JP, Friebely J, Tanski SE, Sherrod C, Matt GE, Hovell MF *et al.* Beliefs about the health effects of

- “thirdhand” smoke and home smoking bans. *Pediatrics*. 2009; 123:e74-e79.
64. Acuff L, Fristoe K, Hamblen J, Smith M, Chen J. Third-Hand Smoke: Old Smoke, New Concerns. *J Community Health*. 2016; 41(3):680-7. Doi: 10.1007/s10900-015-0114-1.
  65. Physical status: the use and interpretation of anthropometry. Report of a WHO Expert Committee. World Health Organ Tech Rep Ser. 1995; 854:1-452.
  66. U.S. Department of Health and Human Services and the U.S. Department of Agriculture Dietary Guidelines for Americans, 2015-2020. ed. <https://health.gov/dietaryguidelines/2015/guidelines/>.
  67. Kesteloot H. Nutrition and health. *Eur Heart J*. 1992; 13:120-128.
  68. O’Keefe EL, Di Nicolantonio JJ, O’Keefe JH, Lavie CJ. Alcohol and CV Health: Jekyll and Hyde J-Curves. *Prog Cardiovasc Dis*. 2018; 61(1):68-75. Doi: 10.1016/j.pcad.2018.02.001.
  69. Alcohol Facts and Statistics In: Alcoholism NIAAa, editor, 2017.
  70. Weber MA, Schiffrin EL, White WB *et al*. Clinical practice guidelines for the management of hypertension in the community: A statement by the American Society of Hypertension and the International Society of Hypertension. *Journal of Clinical Hypertension* (Greenwich). 2014; 16(1):14-26.
  71. Veeranki SP, Zhao M, Ma C, Yan Y, Mi J. Relationship of Alcohol Consumption to All-Cause, Cardiovascular, and Cancer-Related Mortality in U.S. Adults. *J Am. Coll. Cardiol*. 2017; 70:913-922. Doi: 10.1016/j.jacc.2017.06.054.
  72. Fernández-Solà J. Cardiovascular risks and benefits of moderate and heavy alcohol consumption. *Nat. Rev. Cardiol*. 2015; 12:576-587. Doi: 10.1038/nrcardio.2015.91.
  73. Kuntsche E, Kuntsche S, Thrul J, Gmel G. Binge drinking: Health impact, prevalence, correlates and interventions. *Psychol. Health*. 2017; 32:976-1017. Doi: 10.1080/08870446.2017.1325889.
  74. Fillmore MT, Jude R. Defining “binge” drinking as five drinks per occasion or drinking to a .08% BAC: Which is more sensitive to risk? *Am. J Addict*. 2011; 20:468-475. Doi: 10.1111/j.1521-0391.2011.00156.x.
  75. Esser MB, Hedden SL, Kanny D, Brewer RD, Gfroerer JC, Naimi TS. Prevalence of Alcohol Dependence Among US Adult Drinkers, 2009-2011. *Prev Chronic Dis*. 2014; 11:140329.
  76. Cunningham SG. Women’s heart health--an integrated approach to prevention. *Can J Cardiovasc Nurs*. 1998; 9(3):28-37.
  77. Paganini-Hill A, Kawas CH, Corrada MM: Activities and mortality in the elderly: the Leisure World Cohort Study. *J Gerontol A Biol Sci Med Sci*. 2011; 66:559-567.
  78. Lavie CJ, Thomas RJ, Squires RW, Allison TG, Milani RV. Exercise training and cardiac rehabilitation in primary and secondary prevention of coronary heart disease. *Mayo Clin Proc*. 2009; 84:373-383.
  79. Lawler PR, Filion KB, Eisenberg MJ. Efficacy of exercise-based cardiac rehabilitation post-myocardial infarction: a systematic review and meta-analysis of randomized controlled trials. *American heart journal*. 2011; 162:571-58400.
  80. Taylor RS, Brown A, Ebrahim S, Jolliffe J, Noorani H, Rees K *et al*. Exercise-based rehabilitation for patients with coronary heart disease: systematic review and meta-analysis of randomized controlled trials. *The American journal of medicine*. 2004; 116:682-692.
  81. Brellenthin AG, Lanningham-Foster LM, Kohut ML *et al*. Comparison of the Cardiovascular Benefits of Resistance, Aerobic, and Combined Exercise (CardioRACE): Rationale, design, and methods. *Am Heart J*. 2019; 217:101-111. Doi:10.1016/j.ahj.2019.08.008.
  82. Whelton SP, Chin A, Xin X, He J. Effect of aerobic exercise on blood pressure: a meta-analysis of randomized, controlled trials. *Ann Intern Med*. 2002; 136:493-503. Doi: 10.7326/0003-4819-136-7-200204020-00006.
  83. Ekelund U, Steene-Johannessen J, Brown WJ *et al*. Does physical activity attenuate, or even eliminate, the detrimental association of sitting time with mortality? A harmonised meta-analysis of data from more than 1 million men and women. *Lancet*. 2016; 388:1302-10.
  84. Wahid A, Manek N, Nichols M *et al*. Quantifying the association between physical activity and cardiovascular disease and diabetes: a systematic review and meta-analysis. *J Am Heart Assoc*. 2018 Physical Activity Guidelines Advisory Committee. 2018 Physical Activity Guidelines Advisory Committee Scientific Report. Washington, DC: U.S. Department of Health and Human Services; 2018, 2016; 5:e002495.
  85. Franco OH, De Laet C, Peeters A, Jonker J, Mackenbach J, Nusselder W. Effects of physical activity on life expectancy with cardiovascular disease. *Arch Intern Med*. 2005; 165:2355-60.
  86. ALLHAT Officers and Coordinators for the ALLHAT Collaborative Research Group Major outcomes in high-risk hypertensive patients randomized to angiotensin-converting enzyme inhibitor or calcium channel blocker vs diuretic: The Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Trial (ALLHAT) JAMA. 2002; 288:2981-97. Doi: 10.1001/jama.288.23.2981.
  87. Cornelissen VA, Smart NA. Exercise training for blood pressure: a systematic review and meta-analysis. *J Am Heart Assoc*, 2013, 2 Doi: 10.1161/JAHA.112.004473.
  88. Cornelissen VA, Fagard RH, Coeckelberghs E, Vanhees L. Impact of resistance training on blood pressure and other cardiovascular risk factors: a meta-analysis of randomized, controlled trials. *Hypertension*. 2011; 58:950-8. Doi: 10.1161/HYPERTENSIONAHA.111.177071.
  89. Bruning RS, Sturek M. Benefits of exercise training on coronary blood flow in coronary artery disease patients. *Prog Cardiovasc Dis*. 2015; 57(5):443-453. Doi:10.1016/j.pcad.2014.10.006.
  90. Lee CD, Blair SN. Cardiorespiratory fitness and stroke mortality in men. *Medicine and Science in Sports and Exercise*. 2002; 34(4):592-5.
  91. Pang MY, Charlesworth SA, Lau RW, Chung RC. Using aerobic exercise to improve health outcomes and quality of life in stroke: evidence-based exercise prescription recommendations. *Cerebrovasc Dis*. 2013; 35(1):7-22. Doi: 10.1159/000346075.
  92. Pandey A, Darden D, Berry JD. Low fitness in midlife: a novel therapeutic target for heart failure with preserved ejection fraction prevention. *Prog Cardiovasc Dis*. 2015; 58:87-93.

93. Khan H, Kunutsor S, Rauramaa R, Savonen K, Kalogeropoulos AP, Georgiopoulou VV *et al.* Cardio-respiratory fitness and risk of heart failure: a population-based follow-up study. *Eur J Heart Fail.* 2014; 16:180-188.
94. Ponikowsky P, Voors AA, Anker SD *et al.* ESC Guidelines for the diagnosis and treatment of acute and chronic heart failure. *Eur Heart J.* 2016; 37:2129-2200.
95. Pandey A, Patel M, Gao A, Willis BL, Das SR, Leonard D *et al.* Changes in mid-life fitness predicts heart failure risk at a later age independent of interval development of cardiac and noncardiac risk factors: the Cooper Center Longitudinal Study. *Am Heart J.* 2015; 169:290-297 e1.
96. Sagar VA, Davies EJ, Briscoe S, Coats AJ, Dalal HM, Lough F *et al.* Exercise-based rehabilitation for heart failure: systematic review and meta-analysis. *Open Heart.* 2015; 2(1):e000163. Doi: 10.1136/openhrt-2014-000163.
97. Taylor RS, Sagar VA, Davies EJ, Briscoe S, Coats AJ, Dalal H *et al.* Exercise-based rehabilitation for heart failure. *Cochrane Database Syst Rev.* 2014, 27(4). CD003331. Doi: 10.1002/14651858.CD003331.
98. Azarbal F, Stefanick ML, Salmoirago-Blotcher E *et al.* Obesity, physical activity, and their interaction in incident atrial fibrillation in postmenopausal women. *J Am Heart Assoc.* 2014; 3:e001127.
99. Creager MA, Loscalzo J. Chapter 275: Arterial Diseases of the Extremities. In: Jameson J, Fauci AS, Kasper DL, Hauser SL, Longo DL, Loscalzo J. eds. *Harrison's Principles of Internal Medicine*, 20e external icon. McGraw-Hill; Accessed, 2020.
100. Fakhry F, Van De Luijtgarden KM, Bax L *et al.* Supervised walking therapy in patients with intermittent claudication. *J Vasc Surg.* 2012; 56(4):1132-1142.
101. Lane R, Ellis B, Watson L, Leng GC. Exercise for intermittent claudication. *Cochrane Database Syst Rev.* 2014; (7):CD000990. Doi: 10.1002/14651858.CD000990.
102. Lane R, Harwood A, Watson L, Leng GC. Exercise for intermittent claudication. *Cochrane Database Syst Rev.* 2017; 12(12):CD000990. Doi: 10.1002/14651858.CD000990.pub4.
103. Cheng JYW, Ng EML, Ko JSN, Chen RYL. Physical activity and erectile dysfunction: meta-analysis of population-based studies. *Int J Impot Res.* 2007; 19(3):245-252. Doi:10.1038/sj.ijir.3901521.
104. Lamina S, Agbanusi E, Nwacha RC. Effects of aerobic exercise in the management of erectile dysfunction: a meta-analysis study on randomized controlled trials. *Ethiop J Health Sci.* 2011; 21(3):195-201.
105. Tosun NL, Allen SS, Eberly LE, Yao M, Stoops WW, Strickland JC *et al.*, Carroll ME. Association of exercise with smoking-related symptomatology, smoking behavior and impulsivity in men and women. *Drug Alcohol Depend.* 2018; 192:29-37. Doi: 10.1016/j.drugalcdep.2018.07.022.
106. Fonseca-Junior SJ, Sá CG, Rodrigues PA, Oliveira AJ, Fernandes-Filho J. Physical exercise and morbid obesity: a systematic review. *Arq Bras Cir Dig.* 2013; 26(1):67-73. English, Portuguese. Doi: 10.1590/s0102-67202013000600015.
107. Wing RR, Bolin P, Brancati FL *et al.* for the Look AHEAD Research Group. Cardiovascular effects of intensive lifestyle intervention in type 2 diabetes. *N Engl J Med.* 2013; 369:145-154.
108. Wang Y, Xu D. Effects of aerobic exercise on lipids and lipoproteins. *Lipids Health Dis.* 2017; 16(1):132. Doi: 10.1186/s12944-017-0515-5.
109. Brown RA, Abrantes AM, Minami H, Read JP, Marcus BH *et al.* A preliminary, randomized trial of aerobic exercise for alcohol dependence. *J Subst Abuse Treat.* 2014; 47(1):1-9. Doi: 10.1016/j.jsat.2014.02.004.
110. Johansen KL, Painter P. Exercise in individuals with CKD. *Am J Kidney Dis.* 2012; 59(1):126-34. Doi: 10.1053/j.ajkd.2011.10.008.
111. Gujral S, Aizenstein H, Reynolds CF 3rd, Butters MA, Erickson KI. Exercise effects on depression: Possible neural mechanisms. *Gen Hosp Psychiatry.* 2017; 49:2-10. Doi: 10.1016/j.genhosppsych.2017.04.012.
112. O'Keefe EL, O'Keefe JH, Lavie CJ. Exercise Counteracts the Cardiotoxicity of Psychosocial Stress. *Mayo Clin Proc.* 2019; 94(9):1852-1864. Doi: 10.1016/j.mayocp.2019.02.022.
113. Wang X, Youngstedt SD. Sleep quality improved following a single session of moderate intensity aerobic exercise in older women: Results from a pilot study. *Journal of Sport and Health Science.* 2014; 3(4):338-342. 10.1016/j.jshs.2013.11.004.
114. Wang D, Wang Y, Wang Y, Li R, Zhou C. Impact of physical exercise on substance use disorders: a meta-analysis. *PLoS One.* 2014; 9(10):e110728. Doi: 10.1371/journal.pone.0110728.
115. Tanasescu M, Leitzmann MF, Rimm EB, Willett WC, Stampfer MJ, Hu FB. Exercise type and intensity in relation to coronary heart disease in men. *JAMA.* 2002; 288:1994-2000.
116. Bethany Barone Gibbs, Marie-France Hivert, Gerald J. Jerome *et al.* Physical Activity as a Critical Component of First-Line Treatment for Elevated Blood Pressure or Cholesterol: Who, What, and How?: A Scientific Statement From the American Heart Association. *Hypertension.* 2021. https://doi.org/10.1161/HYP.000000000000196.
117. Mora S, Cook N, Buring JE, Ridker PM, Lee IM. Physical activity and reduced risk of cardiovascular events: potential mediating mechanisms. *Circulation.* 2007; 116:2110-8.
118. Ellison GM, Waring CD, Vicinanza C, Torella D. Physiological cardiac remodelling in response to endurance exercise training: cellular and molecular mechanisms. *Heart.* 2012; 98:5-10. 10.1136/heartjnl-2011-300639.
119. Wilson MG, Ellison GM, Cable NT. Basic science behind the cardiovascular benefits of exercise. *Heart.* 2015; 101(10):758-65. Doi: 10.1136/heartjnl-2014-306596.
120. Chodzko-Zajko WJ, Proctor DN, Fiatarone Singh MA *et al.* American College of Sports Medicine position stand. Exercise and physical activity for older adults. *Med Sci Sports Exerc.* 2009; 41(7):1510-1530.
121. Canadian Society for Exercise Physiology. PARmed-X: Physical activity readiness medical examination, 2002. <http://uwfitness.uwaterloo.ca/PDF/parmedx.pdf>. Accessed September 10, 2016.
122. Kenchaiah S, Evans JC, Levy D, Wilson PWF, Benjamin EJ, Larson MG, Kannel WB, Vasan RS. Obesity and the risk of heart failure. *N Engl J Med.* 2002; 347(5):305-313.
123. Canoy D, Cairns BJ, Balkwill A, Wright FL, Green J, Reeves G *et al.* Million Women Study Collaborators.

- Body mass index and incident coronary heart disease in women: a population-based prospective study. *BMC Med.* 2013; 11(1):87.
124. Emberson JR, Whincup PH, Morris RW, Wannamethee SG, Shaper AG. Lifestyle and cardiovascular disease in middle-aged British men: the effect of adjusting for within-person variation. *Eur Heart J.* 2005; 26(17):1774-1782.
  125. Gregg EW, Jakicic JM, Blackburn G *et al.* Look AHEAD Research Group. Association of the magnitude of weight loss and changes in physical fitness with long-term cardiovascular disease outcomes in overweight or obese people with type 2 diabetes: a post-hoc analysis of the Look AHEAD randomised clinical trial. *Lancet Diabetes Endocrinol.* 2016; 4(11):913-921.
  126. Lavie CJ, Milani RV. Obesity and cardiovascular disease: the Hippocrates paradox? *J Am Coll Cardiol.* 2003; 42:677-679.
  127. Must A, Spadano J, Coakley EH, Field AE, Colditz G, Dietz WH. The disease burden associated with overweight and obesity. *JAMA.* 1999; 282:1523-9.
  128. Garrison RJ, Kannel WB, Stokes J *et al.* Incidence and precursors of hypertension in young adults: the Framingham Offspring Study. *Prev Med.* 1987; 16:235-51. DOI: 10.1016/0091-7435(87)90087-9.
  129. Ndumele CE, Matsushita K, Lazo M, Bello N, Blumenthal RS, Gerstenblith G *et al.* Obesity and Subtypes of Incident Cardiovascular Disease. *J Am Heart Assoc.* 2016, 5(8).
  130. Garcia-Labbé D, Ruka E, Bertrand OF, Voisine P, Costerousse O, Poirier P. Obesity and coronary artery disease: evaluation and treatment. *Can J Cardiol.* 2015; 31(2):184-94.
  131. Buschmann K, Wrobel J, Chaban R *et al.* Body Mass Index (BMI) and Its Influence on the Cardiovascular and Operative Risk Profile in Coronary Artery Bypass Grafting Patients: Impact of Inflammation and Leptin. *Oxid Med Cell Longev.* 2020, 5724024. DOI: 10.1155/2020/5724024.
  132. Yarnell JW, Patterson CC, Thomas HF, Sweetnam PM. Comparison of weight in middle age, weight at 18 years, and weight change between, in predicting subsequent 14 year mortality and coronary events: Caerphilly Prospective Study. *J Epidemiol Community Health.* 2000; 54:344-8.
  133. Hagg S, Fall T, Ploner A, Magi R, Fischer K *et al.* European Network for Genetic and Genomic Epidemiology Consortium. Adiposity as a cause of cardiovascular disease: a Mendelian randomization study. *Int J Epidemiol.* 2015; 44:578-586.
  134. Kenchaiah S, Evans JC, Levy D *et al.* Obesity and the risk of heart failure. *New England Journal of Medicine.* 2002; 347(5):305-313. DOI: 10.1056/NEJMoa020245.
  135. Lee AY, Tecson KM, Lima B. Durable left ventricular assist device implantation in extremely obese heart failure patients. *Artif Organs.* 2019; 43(3):234-241. DOI: 10.1111/aor.13380.
  136. Lavie CJ, Mehra MR, Ventura HO. Body composition and advanced heart failure therapy: Weighing the options and outcomes. *JACC Hear Fail.* 2016; 4(10):769-771. DOI: 10.1016/j.jchf.2016.07.007.
  137. Yang TWW, Johari Y, Burton PR. Bariatric surgery in patients with severe heart failure. *Obes Surg.* 2020. DOI: 10.1007/s11695-020-04612-2.
  138. Poirier P, Giles TD, Bray GA *et al.* Obesity and cardiovascular disease: pathophysiology, evaluation, and effect of weight loss: an update of the 1997 American Heart Association scientific statement on obesity and heart disease from the obesity committee of the council on nutrition, physical activity, and metabolism. *Circulation.* 2006; 113:898-9181.
  139. Kurth T, Gaziano JM, Berger K, Kase CS, Rexrode KM, Cook NR *et al.* Body mass index and the risk of stroke in men. *Arch Intern Med.* 2002; 162:2557-2562.
  140. Kurth T, Gaziano JM, Rexrode KM, Kase CS, Cook NR, Manson JE *et al.* Prospective study of body mass index and risk of stroke in apparently healthy women. *Circulation.* 2005; 111(15):1992-8. DOI: 10.1161/01.CIR.0000161822.83163.B6. PMID: 15837954.
  141. Tedrow UB, Conen D, Ridker PM *et al.* The long- and short-term impact of elevated body mass index on the risk of new atrial fibrillation: the WHS (women's health study) *Journal of the American College of Cardiology.* 2010; 55(21):2319-2327. DOI: 10.1016/j.jacc.2010.02.029.
  142. Jouven X, Desnos M, Guerot C, Ducimetiere P. Predicting sudden death in the population: the Paris Prospective Study I. *Circulation.* 1999; 99(15):1978-1983. DOI: 10.1161/01.CIR.99.15.1978.
  143. Larsson SC, Wolk A, Håkansson N, Bäck M. Overall and abdominal obesity and incident aortic valve stenosis: two prospective cohort studies. *Eur Heart J.* 2017; 38:2192-2197.
  144. Huang Y, Xu M, Xie L *et al.* Obesity and peripheral arterial disease: a Mendelian randomization analysis. *Atherosclerosis.* 2016; 247:218-224.
  145. Han TS, Tajar A, O'Neill TW, Jiang M, Bartfai G *et al.* Impaired quality of life and sexual function in overweight and obese men: the European Male Ageing Study. *Eur J Endocrinol.* 2011; 164:1003-11.
  146. Lindstrom S, Germain M, Crous-Bou M *et al.* Assessing the causal relationship between obesity and venous thromboembolism through a Mendelian Randomization study. *Hum Genet.* 2017; 136:897-902.
  147. Poirier P, Giles TD, Bray GA *et al.* Obesity and cardiovascular disease: pathophysiology, evaluation, and effect of weight loss: an update of the 1997 American Heart Association Scientific Statement on Obesity and Heart Disease from the Obesity Committee of the Council on Nutrition, Physical Activity, and Metabolism. *Circulation.* 2006; 113(6):898-918. DOI: 10.1161/CIRCULATIONAHA.106.171016.
  148. Lovren F, Teoh H, Verma S. Obesity and atherosclerosis: mechanistic insights. *Can J Cardiol.* 2015; 31:177-183. DOI: 10.1016/j.cjca.2014.11.031.
  149. Nicolozakes AW, Binkley PF, Leier CV. Hemodynamic effects of smoking in congestive heart failure. *Am J Med Sci.* 1988; 296(6):377-80.
  150. Glantz SA, Parmley WW. Passive smoking and heart disease: Epidemiology, physiology, and biochemistry. *Circulation.* 1991; 83:1-12.
  151. Law MR, Morris JK, Wald NJ. Environmental tobacco smoke exposure and ischaemic heart disease: An evaluation of the evidence. *BMJ.* 1997; 315:973-980.
  152. Levitzky YS, Guo CY, Rong J *et al.* Relation of smoking status to a panel of inflammatory markers: the Framingham offspring. *Atherosclerosis.* 2008;



- 201(1):217-224.  
Doi:10.1016/j.atherosclerosis.2007.12.058.
153. Black HR. Smoking and cardiovascular disease. Laragh JH, Brenner BM (Eds.), *Hypertension: Pathophysiology, Diagnosis and Management* (2nd edition), Raven Press Ltd, New York, NY, 1995, 2621-26471.
154. Ding N, Sang Y, Chen J, Ballew SH, Kalbaugh CA *et al.* Cigarette Smoking, Smoking Cessation, and Long-Term Risk of 3 Major Atherosclerotic Diseases. *J Am Coll Cardiol.* 2019; 74(4):498-507. Doi: 10.1016/j.jacc.2019.05.049.
155. Aune D, Schlesinger S, Norat T, Riboli E. Tobacco smoking and the risk of abdominal aortic aneurysm: a systematic review and meta-analysis of prospective studies. *Sci Rep.* 2018; 8(1):14786. Doi: 10.1038/s41598-018-32100-2.
156. Kovac JR, Labbate C, Ramasamy R, Tang D, Lipshultz LI. Effects of cigarette smoking on erectile dysfunction. *Andrologia.* 2015; 47(10):1087-1092. Doi:10.1111/and.12393.
157. Gregson J, Kaptoge S, Bolton T, Pennells L, Willeit P, Burgess S *et al.* Cardiovascular risk factors associated with venous thromboembolism. *JAMA Cardiol.* 2019; 4:163-173.
158. Ambrose JA, Barua RS. The pathophysiology of cigarette smoking and cardiovascular disease: an update. *J Am Coll Cardiol.* 2004; 43(10):1731-7. Doi: 10.1016/j.jacc.2003.12.047.
159. Barua RS, Ambrose JA, Saha DC, Eales-Reynolds LJ. Smoking is associated with altered endothelial-derived fibrinolytic and antithrombotic factors: An *in vitro* demonstration. *Circulation.* 2002; 106:905-908.
160. Mozaffarian D, Fahimi S, Singh GM *et al.* Global sodium consumption and death from cardiovascular causes. *N Engl J Med.* 2014; 371:624-634.
161. Singh GM, Micha R, Khatibzadeh S *et al.* Estimated global, regional, and national disease burdens related to sugar-sweetened beverage consumption in 2010. *Circulation.* 2015; 132:639-666.
162. Wang Q, Afshin A, Yakoob MY, Singh GM, Rehm CD, Khatibzadeh S *et al.* Global Burden of Diseases Nutrition and Chronic Diseases Expert Group (NutriCoDE). Impact of Nonoptimal Intakes of Saturated, Polyunsaturated, and Trans Fat on Global Burdens of Coronary Heart Disease. *J Am Heart Assoc.* 2016; 5(1):e002891. Doi: 10.1161/JAHA.115.002891.
163. Al-Shaar L, Satija A, Wang DD *et al.* Red meat intake and risk of coronary heart disease among US men: prospective cohort study. *BMJ.* 2020; 371:m4141. Doi: 10.1136/bmj.m4141.
164. He FJ, Li J, Macgregor GA. Effect of longer term modest salt reduction on blood pressure: Cochrane systematic review and meta-analysis of randomised trials. *BMJ.* 2013; 346:f1325.
165. Nahab F, Pearson K, Frankel MR, Ard J, Safford MM, Kleindorfer D *et al.* Dietary fried fish intake increases risk of CVD: the REasons for Geographic And Racial Differences in Stroke (REGARDS) study. *Public Health Nutr.* 2016; 19(18):3327-3336. Doi: 10.1017/S136898001600152X.
166. Narain A, Kwok CS, Mamas MA. Soft drinks and sweetened beverages and the risk of cardiovascular disease and mortality: a systematic review and meta-analysis. *Int J Clin Pract.* 2016; 70(10):791-805. Doi: 10.1111/ijcp.12841.
167. Bagnardi V, Zatonski W, Scotti L, La Vecchia C, Corrao G. Does drinking pattern modify the effect of alcohol on the risk of coronary heart disease? Evidence from a meta-analysis. *J Epidemiol Community Health.* 2008; 62:615-9. 10.1136/jech.2007.065607.
168. Kachur S, Lavie CJ, De Schutter A, Milani RV, Ventura HO. Obesity and cardiovascular diseases. *Minerva Med.* 2017; 108(3):212-228. Doi: 10.23736/S0026-4806.17.05022-4.
169. U.S. Department of Health and Human Services and the U.S. Department of Agriculture Dietary Guidelines for Americans 2015-2020. 8th ed. <https://health.gov/dietaryguidelines/2015/guidelines/>.
170. Johnson SA, Figueroa A, Navaei N. Daily blueberry consumption improves blood pressure and arterial stiffness in postmenopausal women with pre- and stage 1-hypertension: a randomized, double-blind, placebo-controlled clinical trial. *J Acad Nutr Diet.* 2015; 115(3):369-377.
171. Aune D, Keum N, Giovannucci E *et al.* Whole grain consumption and risk of cardiovascular disease, cancer, and all cause and cause specific mortality: systematic review and dose-response meta-analysis of prospective studies. *BMJ.* 2016; 353:i2716. Doi: 10.1136/bmj.i2716.
172. McRae MP. Dietary Fiber Is Beneficial for the Prevention of Cardiovascular Disease: An Umbrella Review of Meta-analyses. *J Chiropr Med.* 2017; 16(4):289-299. Doi: 10.1016/j.jcm.2017.05.005.
173. Guasch-Ferré M, Liu X, Malik VS *et al.* Nut consumption and risk of cardiovascular disease. *Journal of the American College of Cardiology.* 2017; 70(20):2519-32.
174. Ludovici V, Barthelmes J, Nägele MP, Enseleit F, Ferri C, Flammer AJ *et al.* Cocoa, Blood Pressure, and Vascular Function. *Front Nutr.* 2017; 4:36. Doi: 10.3389/fnut.2017.00036.
175. Torres-Collado L, Compañ-Gabucio LM, González-Palacios S *et al.* Coffee Consumption and All-Cause, Cardiovascular, and Cancer Mortality in an Adult Mediterranean Population. *Nutrients.* 2021; 13(4):1241. Doi: 10.3390/nu13041241.
176. Arnett DK, Blumenthal RS, Albert MA *et al.* ACC/AHA Guideline on the Primary Prevention of Cardiovascular Disease: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. *Circulation.* 2019; 140:e596-e646.
177. <https://www.ahajournals.org/Doi/10.1161/CIR.0000000000000678> - accessed April 25, 2021.
178. [https://www.nhlbi.nih.gov/files/docs/public/heart/dash\\_brief.pdf](https://www.nhlbi.nih.gov/files/docs/public/heart/dash_brief.pdf).
179. De Pergola G, D'Alessandro A. Influence of Mediterranean Diet on Blood Pressure. *Nutrients.* 2018; 10(11):1700. Doi:10.3390/nu10111700.
180. Luo D, Cheng Y, Zhang H *et al.* Association between high blood pressure and long term cardiovascular events in young adults: systematic review and meta-analysis. *BMJ.* 2020; 370:m3222. Doi:10.1136/bmj.m3222
181. Kannel WB. Hypertension: reflections on risks and prognostication. *Med Clin North Am.* 2009; 93: 541-558.
182. Benjamin EJ, Blaha MJ, Chiuve SE, Cushman M, Das SR, Deo R *et al.* Heart Disease and Stroke Statistics-2017 Update: a report from the American Heart Association. *Circulation.* 2017; 135:e146-e603.

183. Slivnick J, Lampert BC. Hypertension and Heart Failure. *Heart Fail Clin.* 201; 15(4):531-541. Doi: 10.1016/j.hfc.2019.06.007.
184. Varvarousis D, Kallistratos M, Poulimenos L, Triantafyllis A, Tsinirov P, Giannakopoulos A *et al.* Cardiac arrhythmias in arterial hypertension. *J Clin Hypertens* (Greenwich), 2020. Doi: 10.1111/jch.13989.
185. Salehi-Abargouei A *et al.* Effects of Dietary Approaches to Stop Hypertension (DASH)-style diet on fatal or nonfatal cardiovascular diseases-incidence: a systematic review and meta-analysis on observational prospective studies. *Nutrition.* 2013; 29(4):611-618.
186. Djousse L, Ho YL, Nguyen XT *et al.* DASH score and subsequent risk of coronary artery disease: the findings from Million Veteran Program. *J Am Heart Assoc.* 2018, 7. Doi: 10.1161/JAHA.117.008089.
187. De Logeril M, Salen P, Martin JL *et al.* Mediterranean diet, traditional risk factors and the rate of cardiovascular complications after myocardial infarction: final report of the Lyon Diet Heart Study. *Circulation.* 1999; 99:779-785.
188. Tong TYN, Appleby PN, Key TJ *et al.* The associations of major foods and fibre with risks of ischaemic and haemorrhagic stroke: a prospective study of 418 329 participants in the EPIC cohort across nine European countries. *Eur Heart J.* 2020; 41(28):2632-2640. Doi: 10.1093/eurheartj/ehaa007.
189. Alasmre FA, Alotaibi HA. Plant-Based Diet: A Potential Intervention for Heart Failure. *Cureus.* 2020; 12(5):e8282. Published 2020 May 25. Doi:10.7759/cureus.8282.
190. McCully BH, Hasan W, Streiff CT *et al.* Sympathetic cardiac hyperinnervation and atrial autonomic imbalance in diet-induced obesity promote cardiac arrhythmias. *Am J Physiol Heart Circ Physiol.* 2013; 305(10):H1530-7. Doi: 10.1152/ajpheart.00196.2013.
191. Morrow JP, High-fat diet, obesity and sudden cardiac death. *Acta Physiol.* 2014; 211:13-16. <https://doi.org/10.1111/apha.12276>.
192. Nosova EV, Conte MS, Grenon SM. Advancing beyond the "heart-healthy diet" for peripheral arterial disease. *J Vasc Surg.* 2015; 61(1):265-74. Doi: 10.1016/j.jvs.2014.10.022.
193. Bauer SR, Breyer BN, Stampfer MJ, Rimm EB, Giovannucci EL, Kenfield SA. Association of Diet With Erectile Dysfunction Among Men in the Health Professionals Follow-up Study. *JAMA Netw Open.* 2020; 3(11):e2021701. Doi:10.1001/jamanetworkopen.2020.21701.
194. Najjar RS, Moore CE, Montgomery BD. A defined, plant-based diet utilized in an outpatient cardiovascular clinic effectively treats hypercholesterolemia and hypertension and reduces medications. *Clin Cardiol.* 2018; 41(3):307-313. Doi: 10.1002/clc.22863.
195. Utami DB, Findyartini A. Plant-based Diet for HbA1c Reduction in Type 2 Diabetes Mellitus: an Evidence-based Case Report. *Acta Med Indones.* 2018; 50(3):260-267.
196. Gluba-Brzózka A, Franczyk B, Rysz J. Vegetarian Diet in Chronic Kidney Disease-A Friend or Foe. *Nutrients.* 2017; 9(4):374. Doi: 10.3390/nu9040374.
197. O'Keefe EL, DiNicolantonio JJ, O'Keefe JH *et al.* Alcohol and CV health: Jekyll and Hyde J-curves. *Prog Cardiovasc Dis.* 2018; 61:68-75.
198. Goel S, Sharma A, Garg A. Effect of Alcohol Consumption on Cardiovascular Health. *Curr Cardiol Rep.* 2018; 20(4):19. Doi: 10.1007/s11886-018-0962-2.
199. O'Keefe JH, Bhatti SK, Bajwa A *et al.* Alcohol and cardiovascular health: the dose makes the poison...or the remedy. *Mayo Clin Proc.* 2014; 89:382-393.
200. Kuntsche E, Kuntsche S, Thrul J, Gmel G. Binge drinking: Health impact, prevalence, correlates and interventions. *Psychol. Health.* 2017; 32:976-1017. Doi: 10.1080/08870446.2017.1325889.
201. National Institute on Alcohol Abuse Alcoholism NIH. Turning Discovery Into Health © National Institute on Alcohol Abuse and Alcoholism Alcohol Facts and Statistics. Available online at: <https://www.niaaa.nih.gov>. accessed, 2020.
202. Milon H, Froment A, Gaspard P, Guidollet J, Ripoll JP. Alcohol consumption and blood pressure in a French epidemiological study. *Eur Heart J.* 1982; (3)C:59-64.
203. Friedman GD, Klatsky AL, Siegel AB. Alcohol intake and hypertension. *Ann Intern Med.* 1983; 98:846-849.
204. MacMahon S. Alcohol consumption and hypertension. *Hypertension.* 1987; 9:111-121.
205. Roerecke M, Kaczorowski J, Tobe SW, Gmel G, Hasan OSM, Rehm J. The effect of a reduction in alcohol consumption on blood pressure: a systematic review and meta-analysis. *Lancet Public Health.* 2017; 2:e108-e120. Doi: 10.1016/S2468-2667(17)30003-8.
206. Ronksley PE, Brien SE, Turner BJ *et al.* Association of alcohol consumption with selected cardiovascular disease outcomes: A systematic review and meta-analysis. *BMJ.* 2011; 342:d671.
207. Bagnardi V, Zatonski W, Scotti L, La Vecchia C, Corrao G. Does drinking pattern modify the effect of alcohol on the risk of coronary heart disease? Evidence from a meta-analysis. *J Epidemiol Community Health.* 2008; 62:615-9. 10.1136/jech.2007.065607.
208. Larsson SC, Wallin A, Wolk A, Markus HS. Differing association of alcohol consumption with different stroke types: a systematic review and meta-analysis. *BMC Med.* 2016; 14(1):178. Doi:10.1186/s12916-016-0721-4.
209. Wannamethee SG, Whincup PH, Lennon L *et al.* Alcohol consumption and risk of incident heart failure in older men: A prospective cohort study. *Open Heart.* 2015; 2(1):e000266.
210. Fernández-Solà J. The Effects of Ethanol on the Heart: Alcoholic Cardiomyopathy. *Nutrients.* 2020; 12:572. Doi: 10.3390/nu12020572.
211. Lowenstein SR, Gabow PA, Cramer J, Oliva PB, Ratner K. The role of alcohol in new-onset atrial fibrillation. *Arch Intern Med.* 1983; 143(10):1882-5.
212. Csengeri D, Sprünker NA, Di Castelnuovo A *et al.* Alcohol consumption, cardiac biomarkers, and risk of atrial fibrillation and adverse outcomes. *Eur. Heart J.* 2021; 42:1170-1177. Doi: 10.1093/eurheartj/ehaa953.
213. Koskinen P, Kupari M, Leinonen H, Luomanmäki K. Alcohol and new onset atrial fibrillation: a case-control study of a current series. *Br Heart J.* 1987; 57(5):468-73.
214. Hietanen S, Herajärvi J, Junttila J, Pakanen L, Huikuri HV, Liisanantti J. Characteristics of subjects with alcoholic cardiomyopathy and sudden cardiac death. *Heart.* 2020; 106(9):686-690. Doi: 10.1136/heartjnl-2019-315534.
215. Perkiömäki J, Hookana E, Kaikkonen K, Junttila J, Kortelainen ML, Huikuri H. Blood alcohol in victims of

- sudden cardiac death in northern Finland. *Europace*. 2016; 18(7):1006-9. Doi: 10.1093/europace/euv341.
216. Chen Z, Li S, Guo L, Peng X, Liu Y. Prenatal alcohol exposure induced congenital heart diseases: From bench to bedside. *Birth Defects Res*, 2020. Doi: 10.1002/bdr2.1743.
217. Denny L, Coles S, Blitz R. Fetal Alcohol Syndrome and Fetal Alcohol Spectrum Disorders. *Am Fam Physician*. 2017; 96(8):515-522.
218. Xie X, Ma YT, Yang YN *et al*. Alcohol consumption and ankle-to-brachial index: Results from the Cardiovascular Risk Survey. *PLoS One*. 2010; 5(12):e15181.
219. Wang XM, Bai YJ, Yang YB, Li JH, Tang Y, Han P. Alcohol intake and risk of erectile dysfunction: a dose-response meta-analysis of observational studies. *Int J Impot Res*. 2018; 30(6):342-351. Doi: 10.1038/s41443-018-0022-x.
220. Chakravorty S, Chaudhary NS, Brower KJ. Alcohol Dependence and Its Relationship with Insomnia and Other Sleep Disorders. *Alcohol Clin Exp Res*. 2016; 40(11):2271-2282. Doi: 10.1111/acer.13217.
221. Pan CS, Ju TR, Lee CC, Chen YP, Hsu CY, Hung DZ *et al*. Alcohol use disorder tied to development of chronic kidney disease: A nationwide database analysis. *PLoS One*. 2018; 13(9):e0203410. Doi: 10.1371/journal.pone.0203410.
222. Sayon-Orea C, Martinez-Gonzalez MA, Bes-Rastrollo M. Alcohol consumption and body weight: a systematic review. *Nutr Rev*. 2011; 69(8):419-31. Doi: 10.1111/j.1753-4887.2011.00403.x.
223. Boden JM, Fergusson DM. Alcohol and depression. *Addiction*. 2011; 106(5):906-14. Doi: 10.1111/j.1360-0443.2010.03351.x.
224. Stamler J, Caggiula AW, Grandits GA. Relation of body mass and alcohol, nutrient, fiber, and caffeine intakes to blood pressure in the special intervention and usual care groups in the Multiple Risk Factor Intervention Trial. *Am J Clin Nutr*. 1997; 65: 1 suppl: 338s-65s.
225. Knott C, Bell S, Britton A. Alcohol Consumption and the Risk of Type 2 Diabetes: A Systematic Review and Dose-Response Meta-analysis of More Than 1.9 Million Individuals From 38 Observational Studies. *Diabetes Care*. 2015; 38(9):1804-12. Doi: 10.2337/dc15-0710.
226. Mori TA, Burke V, Beilin LJ, Puddey IB. Randomized Controlled Intervention of the Effects of Alcohol on Blood Pressure in Premenopausal Women. *Hypertension*. 2015; 66:517-523. Doi: 10.1161/HYPERTENSIONAHA.115.05773.
227. Gaziano JM, Buring JE, Breslow JL, Goldhaber SZ, Rosner B, Van Denburgh M *et al*. Moderate alcohol intake, increased levels of high-density lipoprotein and its subfractions, and decreased risk of myocardial infarction. *New Engl J Med*. 1993; 329:1829-1834.
228. Rimm EB, Williams P, Fosher K, Criqui M, Stampfer MJ. Moderate alcohol intake and lower risk of coronary heart disease: meta-analysis of effects on lipids and haemostatic factors. *BMJ*. 1999; 319:1523-1528.
229. Gisbertz SS, Derksen WJ, De Kleijn DP, Vink A, Bots ML, De Vries JP *et al*. The effect of alcohol on atherosclerotic plaque composition and cardiovascular events in patients with arterial occlusive disease. *J Vasc Surg*. 2011; 54:123-131.
230. Pace-Asciak CR, Hahn S, Diamandis EP, Soleas G, Goldberg DM. The red wine phenolics trans-resveratrol and quercetin block human platelet aggregation and eicosanoid synthesis: implications for protection against coronary heart disease. *Clin Chim Acta*. 1997; (2):207-219.
231. Dimmitt SB, Rakic V, Puddey IB *et al*. The effects of alcohol on coagulation and fibrinolytic factors: a controlled trial. *Blood Coagul Fibrinolysis, factor VII (FVII), von Willebrand factor (vWF)*. 1998; (1):39-45.
232. Mukamal KJ, Jadhav PP, D'Agostino RB *et al*. Alcohol consumption and hemostatic factors: analysis of the Framingham Offspring cohort. *Circulation*. 2001; (12):1367-1373.
233. Yarnell JW, Sweetnam PM, Rumley A, Lowe GD. Lifestyle and hemostatic risk factors for ischemic heart disease : the Caerphilly Study. *Arterioscler Thromb Vasc Biol*. 2000; (1):271-279. MED: 10634829.