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Review Article

Exercise and Cardiovascular Disease

'Physical exercise should be taken everyday' Susruta; 600 BCE Indian Physician

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ABSTRACT

Cardiovascular diseases are the leading cause of global mortality. Besides imparting a tremendous amount of human suffering, they also inflict huge direct and indirect financial costs on the worldwide society. With the ready availability of affordable therapeutics, and the lack of newer innovations, non-invasive strategies are being developed to halt their worldwide spread. The major emphasis has been on implementing lifestyle changes. Physical exercise is one such prescription. This manuscript briefly discusses the benefits of exercise in the prevention and management of cardiovascular diseases.

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1. Introduction

The American College of Sports Medicine defines cardiovascular exercise as 'any sport or activity that works large groups of muscles, is continually maintained and performed rhythmically'. ¹ It is a subcategory of physical activity, which is defined as any bodily movement produced by skeletal muscles that results in energy expenditure.² Although lack of physical activity or sedentary behavior is deleteriously linked to cardiovascular disease (CVD), 3 this communication is aimed at highlighting the benefits of exercise on CVD. Regular physical exercise provides a multitude of physical and psychological health benefits, 4 including a decrease in premature mortality⁵ and an increase in life expectancy. 6 A curvilinear relationship exists between CVD and exercise, and a dose-dependent reduction in CVD is associated with increasing exercise. 7 In a meta-analysis of 33 studies, Wahid and group noted an 11% CVD risk reduction with low physical activity (0.1-11.5 METs h/week), a 21% risk reduction with medium physical activity (11.5-29.5 METs h/week) and a 25% risk reduction with high physical activity (29.5+ METs h/week). 8 Exercise

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also reduces the risk of CVD mortality, both in healthy individuals ⁹ and in cardiac patients. ¹⁰ It also improves life expectancy. ¹¹ Its importance as a modifiable lifestyle behavior in cardiovascular diseases is being increasingly recognized. ^{12,13}

2. Exercise and Hypertension

Hypertension (HTN) is a major public health problem in the United States. 14 Due to the revision of the threshold for the diagnosis of hypertension to <130/89 mmHg by the ACC/AHA in 2017, there are now 105 million hypertensives in the US (representing 45.4% of the population). 15 HTN is an independent predisposing factor for several other CVDs including, CAD, stroke, HF, and PAD. 16 It is estimated that for every 20 mmHg systolic and 10 mmHg diastolic blood pressure (BP) increase above the threshold the risk of mortality from ischemic heart disease and stroke doubles. 17 Exercise has BP-lowering effects, 18 and is often prescribed as the first step in high BP management. 19 A meta-analyses calculated that aerobic exercise training lowers systolic BP by 5-7 mmHg, which in turn reduces the risk of CVD by 20-30%. ²⁰ Dynamic resistance exercise is also beneficial and lowers the systolic BP by 2-3 mmHg.²¹ These reductions are similar to the reductions seen with many first-line antihypertensive medications. 22

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Exercising as little as 1 day per week also reduces all-cause mortality among those with hypertension. ²³

3. Exercise and Coronary Heart Disease

Coronary heart disease (CHD), due to atherosclerosis, is present in 15.5 million Americans.²⁴ It is characterized by a diseased endothelium, low-grade inflammation, lipid accumulation, and plaque formation within the intima of the vessel wall.²⁵ This can progress into flow-limiting stenosis of large epicardial coronary arteries, resulting in angina. 26 Plaque rupture or erosion can provoke superimposed atherothrombosis and subsequent vessel occlusion, leading to a myocardial infarction, or even death. 27 Exercise helps protect the coronary arteries 28 and the relationship is inverse.²⁹ Sofi and group did a meta-analysis of 26 studies and found that moderate level of leisure-time physical activity was associated with a decreased CHD risk of 0.88 while those engaging in higher levels demonstrated a decreased risk of 0.73.30 In a subsequent meta-analysis of 33 studies, Satelmair and associates reported that involvement in leisure-time physical activity of 150 min/week resulted in a 14% lower CHD risk while those who reached 300 min/week had a 20% lower risk. 31 Secondary cardiovascular protection (re-infarction and cardiac mortality) with cardiac rehabilitation following a myocardial infarction has been repeatedly demonstrated. 32,33 Exercise induces a significant improvement in post-CABG outcomes.³⁴ It also exerts a major salutary effect on CHD mortality. 35

4. Exercise and Stroke

Stroke is also a major cardiovascular disease in the US.³⁶ It is projected that by 2030, there will be a 20.5% increase in stroke prevalence compared to its prevalence in 2012.³⁷ Stroke is the leading cause of serious long-term disability in US adults, ³⁸ and one of the leading causes of death. ³⁹ A large percentage of the stroke burden is attributable to modifiable risk factors, including physical activity. ⁴⁰ In one study, Harvard alumni with a history of athletic involvement in college, demonstrated less than half the risk of fatal stroke when compared with non-athletes. 41 Research data indicates that physical activity is inversely related to the incidence of stroke. 42 Approximately two-thirds of stroke survivors experience long-term impairments in physical, psychosocial, and cognitive function. 43 Regular exercise in stroke patients helps ameliorate these symptoms, with improvements in aerobic fitness, maximal walking speed, balance, and walking endurance. 44 It also helps improve cognitive function and mood, as well as the quality of life, while reducing subsequent cardiovascular events. 45

5. Exercise and Heart Failure

Heart failure (HF) affects more than 37 million individuals in the world. ⁴⁶ It is also pervasive in the USA. ⁴⁷ Projections show that the prevalence of HF will increase by 46% from 2012 to 2030, and this will result in more than 8 million Americans being diagnosed with this disease. ⁴⁸ HF patients are notorious for frequent hospitalizations, with 25% of patients being re-hospitalized within 30 days, and 50% being re-hospitalized within six months. ⁴⁹ Several studies have demonstrated that exercise reduces the risk of future heart failure. ^{50–52} Benefits have been demonstrated in both systolic HF and HF with preserved left ventricular ejection fraction. ⁵³ Exercise reduces HF hospitalizations, ⁵⁴ improves quality of life, ⁵⁵ and reduces mortality ⁵⁶ in these patients. Cardiac rehabilitation is approved by the Center for Medicaid and Medicare Services for HF patients. ⁵⁷

6. Exercise and Cardiac Arrhythmias

Atrial fibrillation (AF) is the most common cardiac arrhythmia in the world and affects about 34 million individuals.⁵⁸ It affects about 1% of the US population and is related to severe prognostic implications and high mortality. 59 Moderate physical exercise reduces the risk of developing AF. 60 However, more strenuous endurance exercise, often practiced by elite athletes and marathon runners, may increase the risk of AF in healthy athletes without organic heart disease. 61 Sudden cardiac death (SCD) is an unexpected death that usually occurs within one hour of symptoms onset. 62 Ventricular arrhythmias are a major cause of SCD and are noted occasionally in healthy elite athletes performing strenuous activity. ⁶³ A recent study by Aune and group suggests that moderate physical activity may reduce the risk of SCD by almost 50% in the general population. 64 Moderate exercise is safe and beneficial in preventing cardiac arrhythmias. 64,65

7. Exercise and Peripheral Artery Disease

It is estimated that more than 8.5 million men and women in the US suffer from lower extremity peripheral artery disease (PAD). 66 Worldwide, it is present in more than 200 million people. 67 It is seen in 7-14% of the general population ⁶⁸ and this increases to approximately 20% in individuals over seventy years of age. 69 Ankle-brachial index (ABI) is obtained by Doppler measurements of the systolic pressures in the lower and upper extremities. ⁷⁰ An ABI < 0.90 is considered highly sensitive and specific for a diagnosis of PAD. 71 PAD is an atherosclerotic disease. 72 In a meta-analysis of sixteen population cohort studies, Fowkes and group noted an approximately 2-fold increased risk of 10-year all-cause mortality, cardiovascular mortality, and coronary event rate, in individuals with an ABI of < 0.90 when compared to those with normal ABI values. 73 People with PAD also have a greater and faster decline in

functional capacity and have a poorer quality of life than people without PAD. ⁷⁴ Morbidity and mortality rates from this disease, unfortunately, continue to rise. ^{75,76} Exercise helps prevent PAD. ⁷⁷ Several exercise programs, including supervised treadmill exercise, significantly improve painfree and maximal walking distance in people with PAD. ^{78–80} Upper body exercises are also beneficial in these patients. ⁸¹ Exercise improves the quality of life. ⁸² Supervised treadmill exercise therapy in patients with PAD is covered in the US by the Center for Medicaid and Medicare Services. ⁸³

8. Exercise and Erectile Dysfunction

Erectile dysfunction (ED) is a common male sexual disorder. Ref. It causes persistent inability to attain and/or maintain an erection sufficient for sexual performance. Epidemiological studies indicate that it affects approximately 37% of men over 70 years old and 11% of men over 30 years. Atherosclerosis of the pelvic and penile vasculature is a major underlying cause. Erectile dysfunction is a strong predictor for other cardiovascular diseases, including coronary artery disease. Aerobic exercise training has been used successfully to treat patients with atherogenic ED. Patients on PDE-5 inhibitors, exercise further improves ED and increases functional capacity.

9. Exercise and DVT

Deep vein thrombosis (DVT) of the lower extremities is a common venous disease and is associated with significant morbidity and a high rate of recurrence. 91,92 Immobilization is an important risk factor for DVT. 93 Ankle exercises help prevent DVT following hospital immobilization or following a long duration air travel. 94 Physical exercises aimed at the leg musculature also help reduce post-thrombotic syndrome and venous ulceration. 95,96

10. Exercise and General Health

Exercise also exerts an important ameliorating effect on major cardiovascular risk factors such as smoking, 97 obesity, 98 diabetes mellitus, 99 hyperlipidemia, 100 metabolic syndrome, 101 alcohol abuse, 102 chronic kidney disease, 103 psychosomatic stress, 104 and depression. 105 Besides the significant benefits in CVD and CVD risk factors, physical exercise also plays a therapeutic role in several non-cardiovascular disorders, including chronic back pain, 106 osteoporosis, 107 several cancers, 108 constipation, 109 anxiety, 110 dementia, 111 inflammatory bowel disease, 112 gall bladder disease, 113 osteoarthritis, 114 rheumatoid arthritis, 115 Parkinson's disease, 116 and multiple sclerosis. 117 Exercise can also provide help in cognition impairment, 118 and drug addiction. 119 The quality of life is improved with exercise, even in healthy

individuals. ¹²⁰ Moderate to vigorous physical activity also helps reduce all-cause mortality and increases life expectancy. ⁹

11. Discussion

CVD includes coronary heart disease (CHD) high blood pressure (HTN), stroke, heart failure (HF), cardiac arrhythmias, peripheral arterial disease (PAD), and deep vein thrombosis (DVT). ¹²¹ Erectile dysfunction is often due to atherosclerosis and may be included under the umbrella of CVD. ¹²²

CVDs impart the greatest non-communicable diseases burden globally. 123 They account for 17.9 million global deaths annually. 124 These deaths represent 31% of the total global mortality 124 and make CVDs the leading factor in worldwide mortality. 125 It is anticipated that in the year 2035, nearly one in four individuals will be over the age of 65. 126 Age is a major non-modifiable risk factor for CVD¹²⁷ and as the world ages, the incidence of CVD is expected to rise globally in the coming years. 128 CVDs cause considerable loss of productivity and health care spending, which results in a huge financial burden worldwide. 129 CVD is also a leading cause of loss of disability-adjusted life years globally. 130 Healthy lifestyles, including recommended physical exercise, are estimated to be able to prevent 80% of premature CVD mortality in the world. ¹³¹ In the USA, CVD is not only common but also its leading cause of death. 132 CVDs are responsible for about 655,000 American deaths each year—that is 1 in every 4 deaths. 133 Despite advances in CVD management and treatment, CVDs still claim more lives than the combination of all cancer forms. 134 Health-related direct and indirect costs of CVDs during the years 2013 to 2014, were estimated at \$329.7 billion and are expected to reach \$1.1 trillion in 2035. 135 CVDs are also associated with a significant reduction in the quality of life. ¹³⁶

In recent years there has been a major push to reduce the CVD burden by encouraging healthy lifestyles. ¹³⁷ Physical exercise, is a major modifiable CVD lifestyle risk factor and generates significant cardiovascular benefits. 8,10-12 Regular exercise can help decrease weight, reduce blood pressure, and improve lipid disorders, including raising HDL, decreasing LDL, and lowering triglycerides. 138-142 It also reduces systemic inflammation, improves glucose tolerance, decreases insulin resistance, and lowers blood coagulation. 138-142 Exercise also helps increase nitric oxide bioavailability, improve endothelial function, reduce heart rate, increase myocardial oxygen supply, improve myocardial contraction and stroke volume, establish electrical stability and increase physiological cardiac hypertrophy. 138–142 The American Heart Association recommends that individuals perform ≥150 min/week moderate or ≥75 min/week vigorous or ≥150 min/week moderate + vigorous-intensity exercise for optimal

cardiovascular health. ¹⁴³ The duration of physical activity appears to be more important than the intensity, ¹⁴⁴ and 40 minutes of moderate to vigorous-intensity aerobic activity, ³ or 4 times a week, also delivers cardioprotection. ¹⁴⁵ Lower degrees of physical activity also generate CVD benefits. ¹³¹

12. Conclusion

Physical activity is now regarded as a major lifestyle intervention in the primary and secondary prevention of cardiovascular diseases. According to the World Health Organization, 1 in 4 adults are non-compliant with the recommended levels of physical activity and experience a 20% -30% higher risk of death when compared to people who are sufficiently active. ¹⁴⁶ In the USA, the numbers are not much different. Approximately 80% of US adolescents and adults are not active enough. ^{147,148} Despite 60.9 million U.S. citizens joining a gym or starting an exercise program every year, 50% drop out or stop exercising after 6 months. ^{149,150} Given the significant benefits of exercise in cardiovascular diseases, health care workers should strongly incorporate exercise counseling in their CVD preventive and therapeutic armamentarium.

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None.

14. Conflict of Interest

The authors declare that there is no conflict of interest.

References

- 2020. Available from: https://healthyliving.azcentral.com/acsm-definition-cardiovascular-exercise-18723.
- Caspersen CJ, Powell KE, Christenson GM. Physical activity, exercise, and physical fitness: definitions and distinctions for healthrelated research. *Public Health Rep.* 1985:100:126–31.
- Pandey A, Salahuddin U, Garg S, Ayers C, Kulinski J, Anand V, et al. Continuous Dose-Response Association Between Sedentary Time and Risk for Cardiovascular Disease. *JAMA Cardiol*. 2016;1(5):575– 83. doi:10.1001/jamacardio.2016.1567.
- Ruegsegger GN, Booth FW. Health Benefits of Exercise.
 Cold Spring Harb Perspect Med. 2018;8(7):a029694.
 doi:10.1101/cshperspect.a029694.
- Jeong SW, Kim SH, Kang SH, Kim HJ, Yoon CH, Youn TJ, et al. Mortality reduction with physical activity in patients with and without cardiovascular disease. *Eur Heart J.* 2019;40(43):3547–55. doi:10.1093/eurheartj/ehz564.
- Mokdad AH, Ballestros K, Echko M. The state of US health, 1990-2016: burden of diseases, injuries, and risk factors among US states. *JAMA*. 2018;319:1444–72.
- Powell KE, Paluch AE, Blair SN. Physical Activity for Health: What Kind? How Much? How Intense? On Top of What? Ann Rev Public Health. 2011;32(1):349–65. doi:10.1146/annurev-publhealth-031210-101151.
- 8. Wahid A. Quantifying the Association Between Physical Activity and Cardiovascular Disease and Diabetes: A Systematic Review and Meta-Analysis. *J Am Heart Assoc.* 2016;5:2495. doi:10.1161/JAHA.115.002495.

- Wen CP, Wai JP, Tsai MK. Minimum amount of physical activity for reduced mortality and extended life expectancy: a prospective cohort study. *Lancet*. 2011;378:1244–53.
- Thijs MH, Eijsvogels S, Molossi, Lee MS, Emery PD, Thompson. Exercise at the Extremes: The Amount of Exercise to Reduce Cardiovascular Events. J Am Coll Cardiol. 2016;67(3):316–29.
- Franco OH, Laet CD, 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.
- Hansen D, Niebauer J, Cornelissen V. Exercise Prescription in Patients with Different Combinations of Cardiovascular Disease Risk Factors: A Consensus Statement from the EXPERT Working Group. Sports Med. 2018;48(8):1781–97.
- Li G, Li J, Gao F. Exercise and Cardiovascular Protection. Adv Exp Med Biol. 2020;1228:205–16.
- 14. Iqbal AM, Jamal SF. Essential Hypertension; 2020.
- Bundy JD, Mills KT, Chen J, Li C, Greenland P, He J. Estimating the Association of the 2017 and 2014 Hypertension Guidelines With Cardiovascular Events and Deaths in US Adults. *JAMA Cardiol*. 2018;3(7):572. doi:10.1001/jamacardio.2018.1240.
- Sawicka K, Szczyrek M, Jastrzębska I, Prasał M, Zwolak A, Daniluk J. Hypertension - The Silent Killer. J Pre Clin Clin Res. 2011;5(2):43–46.
- Lee JH, Kim SH, Kang SH, Cho JH. Blood pressure control and cardiovascular outcomes: real-world implications of the 2017 ACC/AHA Hypertension Guideline. Sci Rep. 2018;8(1):13155. doi:10.1038/s41598-018-31549-5.
- Cornelissen VA, Smart NA. Exercise Training for Blood Pressure: A Systematic Review and Meta-analysis. J Am Heart Assoc. 2013;2(1). doi:10.1161/jaha.112.004473.
- Chobanian AV. The Seventh Report of the Joint National Committee on Prevention, Detection, Evaluation, and Treatment of High Blood Pressure<SUBTITLE>The JNC 7 Report</SUBTITLE>. JAMA. 2003;289(19):2560. doi:10.1001/jama.289.19.2560.
- Pescatello LS, Franklin BA, Fagard R, Farquhar WB, Kelley GA, Ray CA. American college of sports medicine position stand: exercise and hypertension. *Med Sci Sports Exerc*. 2004;36:533–53.
- Cornelissen VA, Fagard RH. Effect of resistance training on resting blood pressure: a meta-analysis of randomized controlled trials. *J Hypertens*. 2005;23(2):251–9. doi:10.1097/00004872-200502000-00003.
- 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.
- Brown RE, Riddell MC, Macpherson AK, Canning KL, Kuk JL. The Joint Association of Physical Activity, Blood-Pressure Control, and Pharmacologic Treatment of Hypertension for All-Cause Mortality Risk. Am J Hypertens. 2013;26(8):1005–10. doi:10.1093/ajh/hpt063.
- Mozaffarian D, Benjamin EJ, Go AS, Arnett DK, Blaha MJ, Cushman M. Heart Disease and Stroke Statistics-2016 Update: A Report from the American Heart Association. Circ. 2016;133(4):38– 360.
- Libby P, Ridker PM, Hansson GK. Progress and challenges in translating the biology of atherosclerosis. N Engl J Med. 2005;473(7347):1685–95.
- Ganz P, Abben RP, Barry WH. Dynamic variations in resistance of coronary arterial narrowings in angina pectoris at rest. *Am J Cardiol*. 1987;59(1):66–70. doi:10.1016/s0002-9149(87)80071-1.
- Bentzon JF, Otsuka F, Virmani R, Falk E. Mechanisms of Plaque Formation and Rupture. Circ Res. 2014;114(12):1852–66. doi:10.1161/circresaha.114.302721.
- 28. Thompson PD, Buchner D, Pina IL. Exercise and physical activity in the prevention and treatment of atherosclerotic cardiovascular disease: a statement from the council on clinical cardiology (subcommittee on exercise, rehabilitation, and prevention) and the council on nutrition, physical activity, and metabolism

- (subcommittee on physical activity). *Arterioscler Thromb Vasc Biol.* 2003;23:42–9.
- Williams PT. Physical fitness and activity as separate heart disease risk factors: a meta-analysis. *Med Sci Sports Exerc*. 2001;33(5):754– 61
- Sofi F, Capalbo A, Cesari F, Abbate R, Gensini GF. Physical activity during leisure time and primary prevention of coronary heart disease: an updated meta-analysis of cohort studies. *Eur J Cardiovasc Prev Rehabil*. 2008;15(3):247–57. doi:10.1097/hjr.0b013e3282f232ac.
- 31. Sattelmair J, Pertman J, Ding EL, Kohl HW, Haskell W, Lee IM. Dose Response Between Physical Activity and Risk of Coronary Heart Disease. *Circ*. 2011;124(7):789–95. doi:10.1161/circulationaha.110.010710.
- 32. 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. *Am Heart J.* 2011;162(4):571–84.e2. doi:10.1016/j.ahj.2011.07.017.
- Rauch B, Davos CH, Doherty P, Saure D, Metzendorf M, Salzwedel A. The prognostic effect of cardiac rehabilitation in the era of acute revascularisation and statin therapy: a systematic review and meta-analysis of randomized and non-randomized studies the Cardiac Rehabilitation Outcome Study (CROS). Eur J Prev Cardiol. 2016;23:1914–39.
- Coyan GN, Reeder KM, Vacek JL. Diet and Exercise Interventions Following Coronary Artery Bypass Graft Surgery: A Review and Call to Action. *Physician Sportsmed*. 2014;42(2):119–29. doi:10.3810/psm.2014.05.2064.
- 35. Keteyian SJ, Brawner CA, Savage PD, Ehrman JK, Schairer J, Divine G, et al. Peak aerobic capacity predicts prognosis in patients with coronary heart disease. *Am Heart J.* 2008;156(2):292–300. doi:10.1016/j.ahj.2008.03.017.
- Go AS, Mozaffarian D, Roger VL. Heart disease and stroke statistics American Heart Association. Circ. 2014;129(3):28–292.
- Ovbiagele B, Goldstein LB, Higashida RT. Forecasting the future of stroke in the United States. *Stroke*. 2013;44:2361–75.
- Roger VL, Go AS, Lloyd-Jones DM, Benjamin EJ, Berry JD, Borden WB. Heart disease and stroke statistics-2012 update: A report from the American Heart Association. *Circ*. 2012;125(1).
- Kochanek KD, Xu JQ, Murphy SL, Minino AM, Kung HC. Deaths: Final data for 2009 National vital statistics reports. Hyattsville, MD; 2011
- Feigin VL, Roth GA, Naghavi M, Parmar P, Krishnamurthi R, Chugh S, et al. Global burden of stroke and risk factors in 188 countries, during 1990-2013: a systematic analysis for the Global Burden of Disease study. *Lancet Neurol*. 2013;15:913–24.
- Paffenbarger RS, Wing AL. Characteristics in youth predisposing to fatal stroke in later years. *Lancet*. 1967;289(7493):753–4. doi:10.1016/s0140-6736(67)91367-0.
- 42. IMLee, Paffenbarger RS. Physical Activity and Stroke Incidence. *Stroke*. 1998;29(10):2049–54. doi:10.1161/01.str.29.10.2049.
- Gresham GE, Fitzpatrick TE, Wolf PA, McNamara PM, Kannel WB, Dawber TR. Residual Disability in Survivors of Stroke The Framingham Study. New Engl J Med. 1975;293(19):954–6. doi:10.1056/nejm197511062931903.
- Pang MYC, Charlesworth SA, Lau RWK, Chung RCK. 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.
- 45. Billinger SA, Arena R, Bernhardt J, Eng JJ, Franklin BA, Johnson CM, et al. American Heart Association Stroke Council; Council on Cardiovascular and Stroke Nursing; Council on Lifestyle and Cardiometabolic Health; Council on Epidemiology and Prevention; Council on Clinical Cardiology. Physical activity and exercise recommendations for stroke survivors: a statement for healthcare professionals from the American Heart Association/American Stroke Association. Stroke. 2014;45(8):2532–53. doi:10.1161/STR.0000000000000022.
- Bui AL, Horwich TB, Fonarow GC, Vos T. Years lived with disability (YLDs) for 1160 sequelae of 289 diseases and injuries 1990-2010: a

- systematic analysis for the Global Burden of Disease Study. *Nat Rev Cardiol*. 2010;8:2163–96.
- 47. National Center for Health Statistics. National Health and Nutrition Examination Survey (NHANES) public use data files. Centers for Disease Control and Prevention website; 2019. Available from: https://www.cdc.gov/nchs/nhanes/.Accessed.
- 48. Heidenreich PA, Albert NM, Allen LA, Bluemke DA, Butler J, Fonarow GC, et al. On behalf of the American Heart Association Advocacy Coordinating Committee; Council on Arteriosclerosis, Thrombosis and Vascular Biology; Council on Cardiovascular Radiology and Intervention; Council on Clinical Cardiology; Council on Epidemiology and Prevention; Stroke Council. Forecasting the impact of heart failure in the United States: a policy statement from the American Heart Association. Circ Heart Fail. 2013;6:606–19.
- Dharmarajan K, Rich MW. Epidemiology, Pathophysiology, and Prognosis of Heart Failure in Older Adults. *Heart Fail Clin*. 2017;13(3):417–26. doi:10.1016/j.hfc.2017.02.001.
- Wang Y, Tuomilehto J, Jousilahti P, Antikainen R, Mähönen M, Katzmarzyk PT, et al. Occupational, Commuting, and Leisure-Time Physical Activity in Relation to Heart Failure Among Finnish Men and Women. *J Am Coll Cardiol*. 2010;56(14):1140–8. doi:10.1016/j.jacc.2010.05.035.
- Djousse L, Driver JA, Gaziano JM. Relation between modifiable lifestyle factors and lifetime risk of heart failure. *JAMA*. 2009;302:394–400.
- Kenchaiah S, Sesso HD, Gaziano JM. Body Mass Index and Vigorous Physical Activity and the Risk of Heart Failure Among Men. Circ. 2009;119(1):44–52. doi:10.1161/circulationaha.108.807289.
- 53. Guazzi M, Myers J, Peberdy MA, Bensimhon D, PChase, Pinkstaff S, et al. Echocardiography with Tissue Doppler Imaging and cardiopulmonary exercise testing in patients with heart failure: A correlative and prognostic analysis. *Int J Cardiol*. 2010;143(3):323–9. doi:10.1016/j.ijcard.2009.03.053.
- 54. 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(2):290–7. doi:10.1016/j.ahj.2014.10.017.
- Sagar VA, Davies EJ, Briscoe S, Coats AJS, Dalal HM, Lough F, et al. Exercise-based rehabilitation for heart failure: systematic review and meta-analysis. *Open Heart*. 2015;2:e000163. doi:10.1136/openhrt-2014-000163.
- 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;2014. doi:10.1002/14651858.CD003331.pub4.
- 57. Forman DE, Sanderson BK, Josephson RA, Raikhelkar J, Bittner V. American College of Cardiology's Prevention of Cardiovascular Disease Section. Heart failure as a newly approved diagnosis for cardiac rehabilitation. Challenges and opportunities. *J Am Coll Cardiol*. 2015;65:2652–9.
- Go AS, Hylek EM, Phillips KA, Chang Y, Henault LE, Selby JV, et al. Prevalence of diagnosed atrial fibrillation in adults: national implications for rhythm management and stroke prevention: the AnTicoagulation and Risk Factors in Atrial Fibrillation (ATRIA) Study. *JAMA*. 2001;285(18):2370–5.
- Malmo V, Nes BM, Amundsen BH, Tjonna AE, Stoylen A, Rossvoll O, et al. Aerobic Interval Training Reduces the Burden of Atrial Fibrillation in the Short Term. Circ. 2016;133(5):466–73. doi:10.1161/circulationaha.115.018220.
- Abdulla J, Nielsen JR. Is the risk of atrial fibrillation higher in athletes than in the general population? A systematic review and meta-analysis. *Europace*. 2009;11(9):1156–9. doi:10.1093/europace/eup197.
- 61. Kuriachan VP, Sumner GL, Mitchell LB. Sudden Cardiac Death. *Curr Problem Cardiol*. 2015;40(4):133–200. doi:10.1016/j.cpcardiol.2015.01.002.
- Priori SG, Blomström-Lundqvist C, Mazzanti A. Scientific Document Group . 2015 ESC Guidelines for the management of

- patients with ventricular arrhythmias and the prevention of sudden cardiac death. *Eur Heart J.* 2015;36:2793–67.
- Aune D, Schlesinger S, Hamer M, Norat T, Riboli E. Physical activity and the risk of sudden cardiac death: a systematic review and meta-analysis of prospective studies. *BMC Cardiovasc Disord*. 2020;20(1):318. doi:10.1186/s12872-020-01531-z.
- 64. Antonio CO, José CC, Karina SE, Laura GB, Judith TM, Marcelo ML. The Association Between Atrial Fibrillation and Endurance Physical Activity: How Much is too Much? *J Atr Fibrillation*. 2019;12(3):2167. doi:10.4022/jafib.2167.
- Benjamin EJ, Blaha MJ, Chiuve SE. Heart Disease and stroke statistics American Heart Association. Circ. 2017;127(1):143–52.
- Fowkes FG, Rudan D, Rudan I. Comparison of global estimates of prevalence and risk factors for peripheral artery disease in 2000 and 2010: a systematic review and analysis. *Lancet*. 2013;382(9901):1329–40.
- 67. Alzamora MT, Forés R, Baena-Díez JM, Pera G, Toran P, Reina MD. The Peripheral Arterial disease study (PERART/ARTPER): prevalence and risk factors in the general population. *BMC Public Health*. 2010;10(1):38. doi:10.1186/1471-2458-10-38.
- Peach G, Griffin M, Jones KG, Thompson MM, Hinchliffe RJ. Diagnosis and management of peripheral arterial disease. *BMJ*. 2012;345:e5208. doi:10.1136/bmj.e5208.
- 69. Aboyans V, Criqui MH, Abraham P. American Heart Association Council on Peripheral Vascular Disease; Council on Epidemiology and Prevention; Council on Clinical Cardiology; Council on Cardiovascular Nursing; Council on Cardiovascular Radiology and Intervention, and Council on Cardiovascular Surgery and Anesthesia. Measurement and interpretation of the ankle-brachial index: A scientific statement from the American Heart Association. Circ. 2012;126:2890–9.
- 71. Aboyans V, Ricco JB, Bartelink M. ESC Guidelines on the Diagnosis and Treatment of Peripheral Arterial Diseases, in collaboration with the European Society for Vascular Surgery (ESVS): document covering atherosclerotic disease of extracranial carotid and vertebral, mesenteric, renal, upper and lower extremity arteriesEndorsed by: the European Stroke Organization (ESO)The Task Force for the Diagnosis and Treatment of Peripheral Arterial Diseases of the European Society of Cardiology (ESC) and of the European Society for Vascular Surgery (ESVS). Eur Heart J. 2017;39:763–816.
- Fowkes FG, Murray GD, Butcher I. Ankle brachial index combined with Framingham Risk Score to predict cardiovascular events and mortality: a meta-analysis. *JAMA*. 2008;300(2):197–208.
- 73. McDermott MM, Ferrucci L, Liu K, Guralnik JM, Tian L, Liao Y, et al. Leg Symptom Categories and Rates of Mobility Decline in Peripheral Arterial Disease. *J Am Geriatr Soc.* 2010;58(7):1256–62. doi:10.1111/j.1532-5415.2010.02941.x.
- Peach G, Griffin M, Jones KG, Thompson MM, Hinchliffe RJ. Diagnosis and management of peripheral arterial disease. *BMJ*. 2012;345(aug14 1):e5208. doi:10.1136/bmj.e5208.
- Layden J, Michaels J, Bermingham S, Higgins B. Diagnosis and management of lower limb peripheral arterial disease: summary of NICE guidance. *BMJ*. 2012;345. doi:10.1136/bmj.e4947.
- Hiatt WR, Regensteiner JG, Hargarten ME, Wolfel EE, Brass EP. Benefit of exercise conditioning for patients with peripheral arterial disease. Circ. 1990;81(2):602–9. doi:10.1161/01.cir.81.2.602.
- Fakhry F, van de Luijtgaarden K, Bax L, den Hoed P, Hunink MGM, Rouwet EV, et al. Supervised walking therapy in patients with intermittent claudication. *J Vasc Surg.* 2012;56(4):1132–42. doi:10.1016/j.jvs.2012.04.046.
- 78. McDermott MM, Liu K, Guralnik JM. Home-Based Walking Exercise Intervention in Peripheral Artery Disease a

- Randomized Clinical Trial. *J Vasc Surg.* 2013;58(5):1423–24. doi:10.1016/j.jvs.2013.09.020.
- McDermott MM. Exercise training for intermittent claudication. J Vasc Surg. 2017;66(5):1612–20. doi:10.1016/j.jvs.2017.05.111.
- Zwierska I, Walker RD, Choksy SA, Male JS, Pockley AG, Saxton JM. Upper- vs lower-limb aerobic exercise rehabilitation in patients with symptomatic peripheral arterial disease: A randomized controlled trial. *J Vasc Surg.* 2005;42(6):1122–30. doi:10.1016/j.jvs.2005.08.021.
- Lane R, Harwood A, Watson L, Leng GC. Exercise for intermittent claudication. *Cochrane Database Syst Rev.* 2017;12(12). doi:10.1002/14651858.cd000990.pub4.
- Jensen TS, Chin J, Ashby L, Schafer J, Dolan D. Proposed national coverage determination for supervised exercise therapy (SET) for symptomatic peripheral artery disease (PAD) Centers for Medicare and Medicaid Services; 2017.
- Muneer A, Kalsi J, Nazareth I, Arya M. Erectile dysfunction. *BMJ*. 2014;348(jan27 7):g129. doi:10.1136/bmj.g129.
- Najari BB, Kashanian JA. Erectile Dysfunction. *JAMA*. 2016;316(17):1838. doi:10.1001/jama.2016.12284.
- 85. Rosen RC, Fisher WA, Eardley I, Niederberger C, Nadel A, Sand M. The multinational Men's Attitudes to Life Events and Sexuality (MALES) study: I. Prevalence oSf erectile dysfunction and related health concerns in the general population. *Curr Med Res Opin*. 2004;20(5):607–17. doi:10.1185/030079904125003467.
- Richardson D, Vinik A. Etiology and treatment of erectile failure in diabetes mellitus. Curr Diabetes Rep. 2002;2(6):501–9. doi:10.1007/s11892-002-0120-4.
- 87. Uddin SMI, Mirbolouk M, Dardari Z, Feldman DI, Cainzos-Achirica M, DeFilippis AP, et al. Erectile Dysfunction as an Independent Predictor of Future Cardiovascular Events: The Multi-Ethnic Study of Atherosclerosis. *Circ.* 2018;138(5):540–2.
- 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
- Maresca L, D'Agostino M, Castaldo L. Exercise training improves erectile dysfunction (ED) in patients with metabolic syndrome on phosphodiesterase-5 (PDE-5) inhibitors. *Monaldi Arch Chest Dis*. 2013;80(4):177–83.
- Heit JA. Epidemiology of venous thromboembolism. Nat Rev Cardiol. 2015;12(8):464–74. doi:10.1038/nrcardio.2015.83.
- Brandjes DPM, Büller HR, Heijboer H, Huisman MV, de Rijk M, Jagt H, et al. Randomised trial of effect of compression stockings in patients with symptomatic proximal-vein thrombosis. *Lancet*. 1997;349(9054):759–62. doi:10.1016/s0140-6736(96)12215-7.
- Sayegh FA, Almahmeed W, Humood S, Marashi M, Bahr A, Mahdi HA. Global Risk Profile Verification in Patients with Venous Thromboembolism (GRIP VTE) in 5 Gulf Countries. *Clin Appl Thromb*. 2009;15(3):289–96. doi:10.1177/1076029608315168.
- 93. Padberg FT, Johnston MV, Sisto SA. Structured exercise improves calf muscle pump function in chronic venous insufficiency: a randomized trial. *J Vasc Surg*. 2004;39(1):79–87. doi:10.1016/j.jvs.2003.09.036.
- 94. Kahn SR, Shrier I, Shapiro S, Houweling AH, Hirsch AM, Reid RD, et al. Six-month exercise training program to treat post-thrombotic syndrome: a randomized controlled two-centre trial. *Can Med Assoc J*. 2011;183(1):37–44. doi:10.1503/cmaj.100248.
- Araki CT, Back TL, Padberg FT, Thompson PN, Jamil Z, Lee BC, et al. The significance of calf muscle pump function in venous ulceration. *Journal of Vascular Surgery*. 1994;20(6):872–879.
 Available from: https://dx.doi.org/10.1016/0741-5214(94)90223-2. doi:10.1016/0741-5214(94)90223-2.
- Allen AM, Abdelwahab NM, Carlson S, Bosch TA, Eberly LE, Okuyemi K. Effect of brief exercise on urges to smoke in men and women smokers. *Addict Behav*. 2018;77:34–7. doi:10.1016/j.addbeh.2017.09.009.
- 97. 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. doi:10.1590/s0102-67202013000600015.
- 98. Wing RR, Bolin P, Brancati FL. for the Look AHEAD Research Group. Cardiovascular effects of intensive lifestyle intervention in type 2 diabetes. *N Engl J Med*. 2013;369:145–54.
- 99. Wang Y, Xu D. Effects of aerobic exercise on lipids and lipoproteins. *Lipids Health Dis.* 2017;16(1):132.
- Roberts CK, Hevener AL, Barnard RJ. Metabolic syndrome and insulin resistance: underlying causes and modification by exercise training. *Compr Physiol*. 2013;3(1):1–58.
- 101. Brown RA, Abrantes AM, Minami H, Read JP, Marcus BH, Jakicic JM, 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.
- 102. St-Jules DE, Marinaro M, Goldfarb DS, Byham-Gray LD, Wilund KR. Managing Hyperkalemia: Another Benefit of Exercise in People With Chronic Kidney Disease? *J Renal Nutr.* 2020;30(5):380–3. doi:10.1053/j.jrn.2019.10.001.
- O'Keefe EL, O'Keefe JH, Lavie CJ. Exercise Counteracts the Cardiotoxicity of Psychosocial Stress. Mayo Clin Proc. 2019;94(9):1852–64.
- Gujral S, Aizenstein H, Reynolds, Butters MA, Erickson KI. Exercise effects on depression: Possible neural mechanisms. *Gen Hosp Psychiatry*. 2017;49:2–10.
- 105. Galán-Martín MA, Montero-Cuadrado F, Lluch-Girbes E, Coca-López MC, Mayo-Iscar A, Cuesta-Vargas A. Pain neuroscience education and physical exercise for patients with chronic spinal pain in primary healthcare: a randomised trial protocol. *BMC Musculoskelet Disord*. 2019;20(1):505. doi:10.1186/s12891-019-2889-1.
- 106. Tong X, Chen X, Zhang S, Huang M, Shen X, Xu J, et al. The Effect of Exercise on the Prevention of Osteoporosis and Bone Angiogenesis. *Biomed Res Int.* 2019;2019:8171897.
- 107. Adamsen L, Quist M, Andersen C, Moller T, Herrstedt J, Kronborg D, et al. Effect of a multimodal high intensity exercise intervention in cancer patients undergoing chemotherapy: randomised controlled trial. *BMJ*. 2009;339(oct13 1):b3410. doi:10.1136/bmj.b3410.
- 108. Harris BP. Exercise and constipation. *Can Fam Physician*. 1981;27:1320–2.
- Stonerock GL, Hoffman BM, Smith PJ, Blumenthal JA. Exercise as Treatment for Anxiety: Systematic Review and Analysis. *Ann Behav Med*. 2015;49(4):542–56. doi:10.1007/s12160-014-9685-9.
- Li X, Guo R, Wei Z, Jia J, Wei C. Effectiveness of Exercise Programs on Patients with Dementia: A Systematic Review and Meta-Analysis of Randomized Controlled Trials. *BioMed Res Int*. 2019;2019:1–16. doi:10.1155/2019/2308475.
- 111. Eckert KG, Abbasi-Neureither I, Köppel M, Huber G. Structured physical activity interventions as a complementary therapy for patients with inflammatory bowel disease a scoping review and practical implications. *BMC Gastroenterol*. 2019;19(1):115. doi:10.1186/s12876-019-1034-9.
- 112. Kwon OS, Kim YK, Her KH, Kim HJ, Lee SD. Physical activity can reduce the prevalence of gallstone disease among males: An observational study. *Med (Baltimore)*. 2020;99(26):e20763. doi:10.1097/MD.0000000000020763.
- Bosomworth NJ. Exercise and knee osteoarthritis: benefit or hazard?
 Can Fam Physician. 2009;55(9):871–8.
- 114. Baillet A, Zeboulon N, Gossec L, Combescure C, Bodin LA, Juvin R, et al. Efficacy of cardiorespiratory aerobic exercise in rheumatoid arthritis: Meta-analysis of randomized controlled trials. *Arthritis Care Res.* 2010;62:984–92. doi:10.1002/acr.20146.
- 115. Rosenthal LS, Dorsey ER. The Benefits of Exercise in Parkinson Disease. *JAMA Neurol*. 2013;70(2):156. doi:10.1001/jamaneurol.2013.772.
- Baird JF, Sandroff BM, Motl RW. Therapies for mobility disability in persons with multiple sclerosis. *Expert Rev Neurotherapeutics*. 2018;18(6):493–502. doi:10.1080/14737175.2018.1478289.
- 117. Bielak AAM, Cherbuin N, Bunce D, Anstey KJ. Preserved Differentiation Between Physical Activity and Cognitive

- Performance Across Young, Middle, and Older Adulthood Over 8 Years. *J Gerontol Series B: Psychol Sci Soc Sci.* 2014;69(4):523–32. doi:10.1093/geronb/gbu016.
- Lynch WJ, Peterson AB, Sanchez V, Abel J, Smith MA. Exercise as a novel treatment for drug addiction: A neurobiological and stagedependent hypothesis. *Neurosci Biobehav Rev.* 2013;37(8):1622–44. doi:10.1016/j.neubiorev.2013.06.011.
- 119. Bruning RS, Sturek M. Benefits of Exercise Training on Coronary Blood Flow in Coronary Artery Disease Patients. *Prog Cardiovasc Dis.* 2015;57(5):443–53. doi:10.1016/j.pcad.2014.10.006.
- 120. 2020. Available from: https://www.who.int/news-room/fact-sheets/detail/physical-activity#:~:text=People%20who%20are%20insufficiently%20active%20have%20a%2020%25,the%20world%27s%20adolescent%20population%20is%20insufficiently%20physically%20active.
- 121. WHO About cardiovascular diseases. World Heal. Organ; 2018.
- 2020. Available from: https://www.who.int/cardiovascular_diseases/ about_cvd/en/.
- 123. 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. *J Am Coll Cardiol*. 1990;70(1):1–25.
- 2020. Available from: www.who.int/news-room/fact-sheets/detail/ cardiovascular-diseases-(cvds).
- 125. Benjamin EJ, Muntner P, Alonso A, Bittencourt MS, Callaway CW, Carson AP, et al. Heart disease and stroke statistics-2019 update: a report from the American Heart Association. *Circ*. 2019;139. doi:10.1161/CIR.0000000000000659.
- 126. Steenman M, Lande G. Cardiac aging and heart disease in humans. *Biophys Rev.* 2017;9(2):131–7. doi:10.1007/s12551-017-0255-9.
- 127. Mozaffarian D, Benjamin EJ, Go AS. Heart Disease and Stroke Statistics-2016 Update: a report from the American Heart Association. *Circ*. 2016;133:38–60.
- Sniderman AD, Furberg CD. Age as a modifiable risk factor for cardiovascular disease. *Lancet*. 2008;371(9623):1547–9. doi:10.1016/s0140-6736(08)60313-x.
- Bloom D, Cafiero E, Jané-Llopis E, Abrahams-Gessel S, Bloom L. The Global Economic Burden of Non-Communicable Diseases; 2011.
- 130. Bourne RR. Global, regional, and national disability-adjusted life-years (DALYs) for 359 diseases and injuries and healthy life expectancy (HALE) for 195 countries and territories, 1990-2017: a systematic analysis for the Global Burden of Disease Study. *Lancet*. 2017;392:1859–1922.
- 131. Piepoli MF, Hoes AW, Agewall S. European Guidelines on cardiovascular disease prevention in clinical practice: The Sixth Joint Task Force of the European Society of Cardiology and Other Societies on Cardiovascular Disease Prevention in Clinical Practice (constituted by representatives of 10 societies and by invited experts) Developed with the special contribution of the European Association for Cardiovascular Prevention & Rehabilitation (EACPR) Eur. Heart J. 2016;37:2315–81.
- 132. Benjamin EJ, Blaha MJ, Chiuve SE. American Heart Association Statistics Committee and Stroke Statistics Subcommittee Heart Disease and Stroke Statistics-2017 Update: A Report from the American Heart Association. Circ. 2017;135(10):146–603.
- 133. Virani SS, Alonso A, Benjamin EJ, Bittencourt MS, Callaway CW, Carson AP. Heart disease and stroke statistics-2020 update: a report from the American Heart Associationexternal icon. *Circ*. 2020;141(9):139–596.
- 134. Mozaffarian D, Benjamin EJ, Go AS, Arnett DK, Blaha MJ, Cushman M, et al. Heart disease and stroke statistics-2015 update: a report from the American Heart Association. 2015;131(4):29–322. doi:10.1161/cir.0000000000000152.
- 135. Benjamin EJ, Virani SS, Callaway CW. American Heart Association Council on Epidemiology and Prevention Statistics Committee and Stroke Statistics Subcommittee. Heart Disease and Stroke Statistics-2018 Update: A Report From the American Heart Association. *Circ*. 2018;137(12):67–492.

- 136. Arija V, Villalobos F, Pedret R, Vinuesa A, Jovani D, Pascual G, et al. Physical activity, cardiovascular health, quality of life and blood pressure control in hypertensive subjects: randomized clinical trial. *Health Qual Life Outcomes*. 2018;16(1):184. doi:10.1186/s12955-018-1008-6.
- 137. Ž Reiner, Laufs U, Cosentino F, Landmesser U. The year in cardiology 2018: prevention. *Eur Heart J.* 2019;40(4):336–44. doi:10.1093/eurheartj/ehy894.
- 138. Mora S, Cook N, Buring JE, Ridker PM, Lee IM. Physical Activity and Reduced Risk of Cardiovascular Events. *Circulation*. 2007;116(19):2110–2118. Available from: https://dx.doi.org/10.1161/circulationaha.107.729939. doi:10.1161/circulationaha.107.729939.
- Myers J. The new AHA/ACC guidelines on cardiovascular risk: When will fitness get the recognition it deserves? *Mayo Clin Proc*. 2014;89:722–726.
- 140. Ellison GM, Waring CD, Vicinanza C, Torella D. Physiological cardiac remodelling in response to endurance exercise training: cellular and molecular mechanisms. *Heart*. 2012;98(1):5–10. doi:10.1136/heartjnl-2011-300639.
- 141. Slentz CA, Bateman LA, Willis LH, Granville EO, Piner LW, Samsa GP, et al. Effects of exercise training alone vs a combined exercise and nutritional lifestyle intervention on glucose homeostasis in prediabetic individuals: a randomised controlled trial. *Diabetol*. 2016;59(10):2088–98. doi:10.1007/s00125-016-4051-z.
- 142. 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.
- 143. Han L, You D, Ma W. National Trends in American Heart Association Revised Life's Simple 7 Metrics Associated With Risk of Mortality Among US Adults. *JAMA Netw Open*. 2019;2(10).

- 144. Cortez MY, Torgan CE, Brozinick JT, Ivy JL. Insulin resistance of obese Zucker rats exercise trained at two different intensities. Am J Physiol Endocrinol Metab. 1991;261(5):E613–9. doi:10.1152/ajpendo.1991.261.5.e613.
- Lobelo F, Young D, Sallis R. Routine assessment and promotion of physical activity in healthcare settings: a scientific statement from the American Heart Association. *Circ.* 2018;137(18):495–522.
- Piercy KL, Troiano RP, Ballard RM, Carlson SA, Fulton JE, Galuska DA, et al. The Physical Activity Guidelines for Americans. *JAMA*. 2018;320(19). doi:10.1001/jama.2018.14854.
- Lobelo F, Young R, Sallis D, Garber MD, Billinger SA, Duperly J, et al. Routine Assessment and Promotion of Physical Activity in Healthcare Settings: A Scientific Statement From the American Heart Association. Circ. 2018;137(18):495–522.
- 148. The 2018 IHRSA Health Club Consumer Report. International Health, Racquet & Sportsclub Association; Web. 25 October 2019.
- Wilson K, Brookfield D. Effect of Goal Setting on Motivation and Adherence in a Six-Week Exercise Program. Int J Sport Exer Psychol. 2009;7(1):89–100. doi:10.1080/1612197x.2009.9671894.
- Berra K, Rippe J, Manson JE. Making Physical Activity Counseling a Priority in Clinical Practice. *JAMA*. 2015;314(24):2617–8. doi:10.1001/jama.2015.16244.

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