# Review Article

## Loneliness, Lifestyles, and Lockdowns: Effects on Health

#### **Abstract**

Mother Teresa said: "the greatest disease in the West today is not TB or leprosy; it is being unwanted, unloved, and uncared for. We can cure physical diseases with medicine, but the only cure for loneliness, despair, and hopelessness is love..." Loneliness is a global phenomenon. Although more prevalent in the elderly, it is seen in all age groups. Contrary to logic, social media appears to have increased loneliness. Loneliness has several detrimental effects on human health. It is associated with a wide array of chronic diseases, contributing to their development, and often worsening their prognosis. Although there are several mechanisms involved in this harmful association, lower adherence to five healthy lifestyles is a major factor. These lifestyles include not smoking, not drinking alcohol in excess, maintaining an adequate body weight, prudent diet, and regular physical activity. Unfortunately, unhealthy lifestyles have been aggravated by the recent COVID-19 pandemic. These relationships are reviewed in this manuscript.

**Keywords:** loneliness, social isolation, lockdown, smoking, obesity, alcohol intake, physical activity, diet

#### Introduction

Loneliness and social isolation are generating increasing attention, primarily because of the significant harm they inflict on human health<sup>1,2</sup>. Loneliness is a feeling that one's own social network is smaller than desired while social isolation is more representative of a feeling that one does not belong to or is being avoided by society<sup>3,4</sup>. In other words, one may feel lonely despite having good social relationships, while one may not feel lonely despite being socially isolated. In this manuscript, loneliness and social isolation are lumped together and referred to as 'loneliness'. Loneliness, a usually silent but distressful feeling, is ubiquitous in our world<sup>5</sup>. Its prevalence is on the rise<sup>6</sup>. It has been noted in all population groups and all age groups<sup>7-9</sup>. Periods of loneliness may affect as many as 80% of those under 18 years of age. Further, about one-third of those aged 60 years and over experience loneliness in later life <sup>10,11</sup>. A recent study in the United States of America (US) by Cigna, an insurance company. of adults aged 18 and older found that 46 percent reported "sometimes or always feeling alone" 12. Loneliness is associated with poor health <sup>13,14</sup>. Loneliness is closely linked to cardiovascular diseases (CVDs) <sup>15</sup>. Cardiovascular disease remains the number one cause of mortality in the world 16. Data from 16 longitudinal studies indicate that the relative risk for incident coronary heart disease and stroke in individuals with high versus low levels of loneliness was 1.29<sup>17</sup>. It has also been linked with obesity<sup>18</sup>, several cancers<sup>19,20</sup>, chronic obstructive pulmonary disease (COPD)<sup>21</sup>, arthritis<sup>22</sup>,

depression<sup>23</sup>, dementia<sup>24</sup>, and a host of other chronic diseases. Lonely individuals are also more susceptible to infectious diseases<sup>26</sup>. They usually report poor self-rated health<sup>27</sup> and a poor quality of life<sup>28</sup>. Their overall mortality is often premature<sup>29,30</sup>. In 2010, a meta-analysis revealed that the odds ratio for increased mortality for loneliness was 1.45, which is approximately double the odds ratio for increased mortality for obesity and quadruple the odds ratio for mortality related to air pollution<sup>31</sup>. Other studies have shown that loneliness confers a mortality risk that is greater or equivalent to that seen with smoking or alcohol abuse disorder<sup>32</sup>. Studies have also found loneliness-related rates of mortality like that from lack of physical activity<sup>33</sup>. Loneliness is also associated with a host of psychiatric disorders which include depression<sup>34</sup>, anxiety<sup>35</sup>, alcoholism<sup>36</sup>, suicidal ideation<sup>37</sup>, and aggressive behavior and impulsivity<sup>38</sup>.

Diagnosis of loneliness is easy. The University of California, Los Angeles, Loneliness Scale-Version 3 (UCLA 3-item) is the most frequently used questionnaire to diagnose loneliness<sup>39.</sup> Results are also considered reliable with a single question, 'Do you feel lonely?" <sup>40</sup>. The negative effects of this global scourge have prompted several international associations to recommend remedial actions <sup>41-44</sup>. These include the Monalisa initiative in France <sup>41</sup>, the RISE campaign in Canada <sup>42</sup>, and the Australian coalition to end loneliness <sup>43</sup>. In the US, the American Association of Retired Persons (AARP) has launched a campaign called Connect2Affect <sup>44</sup> while the Institute on Aging offers a 24-hour toll-free Friendship Line (1-800-971-0016) for adults 60 and older to call when they're feeling lonely. Individuals that are socially integrated are healthier, happier, and live longer than those who are isolated <sup>45</sup>.

## **Discussion**

Five healthy lifestyle factors are: never smoking, a body mass index 18.5-24.9, participating in moderate to vigorous physical activity (≥30 minutes/day), a moderate intake of alcohol (women: 5-15 g/day; men 5-30 g/day), and a higher quality of diet<sup>46</sup>. Healthy lifestyles are important for good health<sup>47-49.</sup> They help increase disease-free lifespan<sup>50-52</sup>. Stenholm et al. noted that a disease-free life (without chronic diseases cardiovascular disease, cancer, respiratory disease, and diabetes) was increased in individuals with zero unhealthy lifestyles when compared to those with at least two unhealthy behaviors<sup>50</sup>. The CHANCES study also demonstrated an increase in disease-free life<sup>50</sup> in people following a healthy lifestyle<sup>51</sup>. This longer life expectancy free of major chronic diseases at midlife in these individuals was recently confirmed by Li and colleagues<sup>52</sup>. Lifestyles also impact mortality. Studies have shown that smoking, inactivity, poor diet quality, and heavy alcohol consumption contribute up to 60% of all premature deaths<sup>53-58</sup>. A recent study that looked at the data from the Nurses' Health Study (1980-2014; n=73,196) and the Health Professionals Follow-Up Study (1986-2014; n=38,366), confirmed that poor lifestyles accounted for a significant number of deaths<sup>59</sup>. In this study, 70% of cardiovascular mortality, and 50% of cancer mortality resulted from an unhealthy lifestyle. Since many of these deaths are premature, healthy lifestyles also increase longevity<sup>46</sup>. Li et al. estimated that the life expectancy at age 50 years was 79.0 years for women and 75.5 years for men with unhealthy lives. Adopting all five healthy lifestyles changed this to 93.1 years for women and 87.6 years for men $^{46}$  – a marked increase in longevity.

To date, several psychological, biological, and behavioral pathways leading to poor health and increased mortality from loneliness have been identified<sup>60,61</sup>. Lonely individuals are under chronic stress<sup>62</sup>. They are often depressed<sup>23</sup> and demonstrate high anxiety levels<sup>63</sup>. This triggers several neuroendocrine responses that culminate in harm. These include HPA activation<sup>64</sup>, sympathetic system hyperactivity<sup>65</sup>, parasympathetic system dysfunction<sup>66</sup>, increased blood levels of catecholamines<sup>67</sup>, decreased glucocorticoid receptor sensitivity with higher cortisol levels<sup>68</sup>, an alteration in the vascular resistance<sup>69</sup>, and a proinflammatory immune response<sup>70</sup>. In some cases, there may be a genetic contribution<sup>71</sup>. These changes contribute towards the development or worsening of several chronic diseases<sup>18-24</sup>.

Lonely individuals often have diminished self-regulation - an incapacity to adequately control their feelings, emotions, and behavior<sup>72</sup>. This enhances their noncompliance with healthy lifestyles<sup>73-76</sup>. Lacking support from social networks, family members, and neighbors, they also tend to underutilize/overutilize health care services<sup>77</sup> and often fail to follow health recommendations adequately<sup>78</sup>. On the flip side, lonely individuals that maintain good lifestyle behaviors demonstrate lower morbidity and mortality<sup>79-81</sup>. The worldwide lockdowns associated with COVID-19 have also impacted healthy behaviors. It has not only enhanced loneliness but also decreased compliance with a healthy lifestyle<sup>82</sup>.

## **Smoking**

Tobacco smoking, either with cigars, cigarettes, water pipes, electronic cigarettes, bidis, or by other means <sup>83-85</sup> is associated with considerable harm <sup>86</sup>. Smoking increases the risk of a wide range of chronic diseases, including coronary artery disease <sup>87</sup>, stroke <sup>88</sup>, type 2 diabetes mellitus (T2DM)<sup>89</sup>, cancer <sup>90</sup>, chronic kidney disease (CKD)<sup>91</sup>, and COPD<sup>92</sup>. Besides this increase in morbidity, smokers also face higher disability <sup>93</sup> and premature death <sup>94</sup>. It has been estimated that the smokers' life expectancy is on average is 10 years shorter than that of non-smokers <sup>95</sup>. Smoking exposes the smoker and others (second-hand and third-hand smoke inhalers) to several thousand chemicals such as carbon monoxide, gaseous aldehydes, nicotine, polycyclic aromatic compounds, nitrosamines, and heavy metals <sup>96</sup>. Many of these are toxic and carcinogenic <sup>97,98</sup>. The estimated economic cost associated with smoking is almost 1.8% of global gross domestic product <sup>99</sup>.

Lonely individuals were more likely to be smokers <sup>100,101</sup>. In a study of older English individuals, Kobayashi and Steptoe found that lonely individuals were more likely to smoke <sup>102</sup>. In this study of 3,392 men and women aged ≥52 years (English Longitudinal Study of Ageing from 2004/2005 to 2014/2015), they found that when compared to non-lonely individuals, the risk ratio for smoking in lonely individuals at any time was high (RR=1.46). And the higher the level of loneliness, the more the likelihood of them being smokers <sup>103</sup>. Once addicted, lonely individuals also have a harder time giving up smoking <sup>102</sup>. The relationship between loneliness and smoking appears to be bidirectional <sup>104-106</sup>. Smokers tend to suffer more loneliness than non-smokers <sup>104</sup>. This was also reported recently by Zhang et al, in a study of 1,452 freshmen university students in Wuhan, China <sup>105</sup>. The COVID-19 related lockdown has increased both loneliness and smoking. A study involving 6,003 Italian adults aged 18-74 years, revealed that the lockdown increased cigarette consumption by 9.1% <sup>106</sup>.

## **Obesity**

Bodyweight is generally classified according to the body mass index (BMI)<sup>107</sup>. This is calculated in individuals by dividing their weight in Kg by the square of their height in meters. A body mass index should ideally be 18.5 kg/m<sup>2</sup> to 24.9 kg/m<sup>2</sup>. A BMI of 25 kg/m<sup>2</sup> to 29,9 kg/m<sup>2</sup> is considered overweight, while at or >30 kg/m<sup>2</sup> is considered obese<sup>108</sup>. The Centers for Disease Control and Prevention of USA further subdivides obesity into several categories: Class 1: BMI of 30 to < 35, Class 2: BMI of 35 to < 40, and Class 3: BMI of 40 or higher <sup>109</sup>. The World Health Organization (WHO) estimates that there were 1.9 billion overweight adults and 650 million obese adults globally in 2016<sup>110</sup>. Central or visceral obesity may exist even if the BMI is normal and is typically more harmful<sup>111</sup>. Being overweight or obese results in more deaths than being underweight<sup>112</sup>. Chronic diseases resulting from excess body weight include CVDs<sup>113</sup>, metabolic syndrome<sup>114</sup>, T2DM<sup>115</sup>, CKD<sup>116</sup>, nonalcoholic fatty liver disease<sup>117</sup>, several cancers<sup>118</sup>, obstructive sleep apnea<sup>119</sup>, osteoarthritis<sup>120</sup>, dementia<sup>121</sup>, and depression<sup>122</sup>. Obesity also worsens the course of many of these diseases and results in premature mortality 123. For example, in heart failure patients it increases emergency room visits by 57%, hospitalization by 68%, and the risk of death by almost 4 times <sup>124</sup>. Negative attitudes towards obese people also result in significant socio-cultural harm<sup>125</sup>. Obese individuals appear to have a 30% higher medical cost than those with a normal BMI, inflicting a major financial burden on society<sup>126</sup>.

The relationship between loneliness and obesity is complex <sup>127</sup>. Loneliness enhances obesity <sup>128</sup>. Lonely people eat more energy-dense foods, have decreased physical activity 129, and have poor daytime functioning<sup>130</sup>. Lonely people tend to sleep poorly, which further increases their tendency to get obese<sup>131</sup>. In 2006, Morse et al. studied 714 patients and found that 63.8% of those who reported being lonely (40.2%), had obesity inducing nighttime-eating behaviors<sup>132</sup>. A 2010 study found that each 1-unit increase in loneliness was associated with a 10% increase in the odds of a person meeting the criteria for metabolic syndrome<sup>133</sup>. Obesity in turn is associated with higher levels of loneliness in the general population <sup>134</sup>. Obese individuals face isolation due to several factors including not being non-compliant with societal norms<sup>135</sup>. Obesity-related stigma enhancing loneliness includes lower rates of hire and promotion and higher rates of wrongful dismissal<sup>136</sup>. Stigmatization is especially evident in children with obesity or in young adults<sup>137</sup>. Furthermore, a higher BMI may also decrease participation in religious activities/church services 138. Internalizing the stigma decreases self-esteem and further worsens loneliness <sup>139</sup>. And finally, the increase in loneliness and added stress during COVID-19 lockdown has negatively affected weight-related behaviors among adults with higher BMI and eating disorders <sup>140</sup>.

## **Alcohol Intake**

Alcohol is a popular psychoactive substance that may result in dependence <sup>141</sup>. According to WHO, in 2016, the total alcohol per capita consumption in the world (age 15 years and older) was 6.4 liters annually <sup>142</sup>. This approximates to 13.9 grams of pure alcohol per person per day. To put this amount in perspective, in the UK, a 'unit' of alcohol is classified as 8 g of pure alcohol, while a standard drink in other countries may contain as much as 20 g of alcohol <sup>143</sup>. In the United States, a 'standard drink' contains 14 g of pure alcohol <sup>144</sup>. Several studies have shown

a consistent relationship between low/moderate alcohol consumption and several health benefits, including a reduced risk of T2DM<sup>145</sup>, coronary heart disease<sup>146</sup>, ischemic stroke<sup>147</sup>, chronic renal disease<sup>148</sup>, and all-cause mortality<sup>149</sup>. However, excessive alcohol consumption increases all-cause and cause-specific morbidity and mortality<sup>150</sup>. Excess alcohol intake has been associated with harm to literally every bodily system<sup>151-165</sup>. It has repeatedly been shown to cause/aggravate cardiovascular diseases<sup>151</sup>, obesity<sup>152</sup>, cancer<sup>153</sup>, T2DM<sup>154</sup>, chronic respiratory diseases<sup>155</sup>, CKD, depression<sup>157</sup>, dementia<sup>158</sup>, GI tract disorders<sup>159</sup>, liver diseases<sup>160</sup>, sleep disorders<sup>161</sup>, congenital heart disease<sup>162</sup>, sexual disorders<sup>163</sup>, trauma/violence<sup>164</sup>, and even infections<sup>165</sup>. Its association with cancer is especially alarming. The International Agency for Research on Cancer classifies both alcohol and its acetaldehyde metabolite as type 1 carcinogens<sup>166</sup>. The World Heart Association recently opined that despite the recommendations of several organizations that low to moderate intake may be beneficial for certain conditions, even a single drink of alcohol is not safe for overall health<sup>167</sup>.

Lonely people are more likely to drink<sup>168</sup>. This has been noted in all age groups. In a study of school children, lonely females were 2.9 times as likely to engage in alcohol consumption when compared to their male peers<sup>169</sup>. A positive relationship between loneliness and alcoholism has also been noted in adolecents<sup>170,171</sup> and adults<sup>172</sup>. Loneliness not only contributes to alcohol abuse but also helps in maintaining it<sup>173</sup>. It confers a poor prognosis in these individuals<sup>173</sup>. Many people reported that they were experiencing more social isolation and loneliness due to COVID restrictions<sup>174</sup> and this was accompanied by a 54% surge in alcohol purchase<sup>175</sup> and consumption<sup>176</sup>. Lonely people often lack social support, and this may aggravate the tendency for addiction<sup>177</sup>. The relationship has also been noted the other way around. Alcoholics tend to experience more loneliness<sup>173</sup>. They often have a more negative attitude towards themselves and their social interactions<sup>173</sup>. They have a feeling of being dissatisfied with life and are overall more lonely<sup>178</sup>.

## **Physical Activity**

Physical activity refers to any activity performed that expends energy more than I metabolic equivalent (MET)<sup>179</sup>. Sedentary behavior is typically referred to as an energy expenditure of fewer than 1.5 METs<sup>180</sup>. Exercise, on the other hand, is a planned, repetitive, and purposive workout, resulting in an energy expenditure of more than 1.5 METs<sup>181</sup>. Light-intensity activities expend less than 3 METs, moderate-intensity activities expend 3 to 6 METs, while vigorous activities expend 6 or more METs<sup>182</sup>. Sedentary behaviors and lack of regular physical exercise, on the other hand, is associated with an increased incidence and worse progression of a wide range of diseases<sup>183,184</sup>, including cardiovascular diseases<sup>185</sup>, type 2 diabetes<sup>186</sup>, several cancers<sup>187</sup>, including those of the breast and colon, bone and joint diseases (osteoporosis and osteoarthritis)<sup>188</sup>, and depression<sup>189</sup>. Exercise results in a decrease in morbidity from most major non-communicable diseases, including CVDs<sup>190</sup>, T2DM<sup>191</sup>, and cancer<sup>192</sup>. Further, it helps reduce premature mortality by at least 20%–30% in most major chronic medical conditions<sup>193</sup>. Most professional organizations recommend that adults aged 18 to 65 years perform moderate-intensity aerobic physical activity for a minimum of 30 min on five days each week or vigorous-intensity aerobic physical activity for a minimum of 20 min on three days each week. Muscle

and bone-strengthening exercises are recommended for a minimum of two days each week. Balance exercises are also important in the elderly <sup>194,195</sup>.

Lonely people are less likely to partake in regular physical exercise 196.197. In a study of older English individuals, Kobayashi and Steptoe found that socially isolated participants were less likely than non-isolated participants to partake in weekly moderate-to-vigorous physical activity 198. They found that socially isolated individuals spent more time spent in sedentary behavior and less time in light and moderate/vigorous physical activity, during the day. In another study of 267 community-based men and women aged 50-81 years taking part in the English Longitudinal Study of Ageing, Schrempft et al. also found that lonely individuals spent less time in moderate/vigorous physical activity and more time in sedentary behavior 199. These studies were done in high-income countries 198,199. Similar findings were noted in other parts of the world. In a study of 34,129 individuals aged 50 and older in low- and middle-income countries, loneliness was also associated with low physical activity<sup>200</sup>. There are confounding factors that can decrease physical activity in lonely individuals, such as chronically poor health and a low socioeconomic status<sup>201</sup>. Exercise has a multitude of benefits, and it also helps decrease loneliness. In one study involving 382 participants, there was a 6.9% decrease in loneliness and a 3.3% improvement in social connectedness, with exercise<sup>202</sup>. Unfortunately, the COVID-19 lockdown has not helped. A longitudinal study showed that a lockdown period due to COVID-19 decreased physical activity levels in a group of physically active Spanish adults<sup>203</sup>.

## **Diet**

Eating too many calories can result in excess body weight and obesity<sup>204</sup>. The harmful effects of obesity have been discussed earlier in this manuscript. A poor-quality diet is also detrimental to health - it increases the risk for CVDs<sup>205</sup>, cancer<sup>206</sup>, T2DM<sup>207</sup>, obesity<sup>208</sup>, OSA<sup>209</sup>, CKD<sup>210</sup>, GI diseases<sup>211</sup>, and many others<sup>212,213</sup>. The western diet is typical of an unhealthy diet<sup>214</sup>. It is usually rich in refined sugars, high in salt, white flour, processed meats, purified animal fats, and food additives<sup>215</sup>. It tends to be low in fruits, vegetables, and whole grains<sup>216</sup>. It is also heavy in processed foods, "fast foods", snacks, and sugary soft drinks<sup>217</sup>. The result is more 'empty calories' that are lacking in the beneficial phytochemicals, fiber, vitamins, and minerals. They are also usually energy-dense and exhibit harmful high glycemic indexes<sup>218</sup>. Besides the marked increase in morbidity <sup>219-224</sup>, an unhealthy diet also increases mortlity <sup>225</sup>. Suboptimal diets are estimated to have caused 1 in 5 premature deaths globally from 1990–2016<sup>226</sup>. A healthy diet on the other hand is rich in non-starchy vegetables, fruits, whole grains, and legumes, moderate in consumption of nuts, seafood, lean meats, low-fat dairy products, unsaturated vegetable oil, and limited in or void of trans-fats, saturated fats, sodium, red meat, refined carbohydrates, and sugar-sweetened beverages<sup>227</sup>. It is low in salt (<2,300 mg per day) and limits alcohol to 2 drinks or less in a day for men or 1 drink or less in a day for women. Mediterranean diets and DASH diets are healthy diets. Mediterranean diets, especially with extra virgin olive oil are extremely healthy<sup>228</sup>. Healthy diets are associated with a major reduction in morbidity and mortality<sup>229,230</sup>.

There is a link between loneliness higher consumption of unhealthy foods, both in caloric intake (causing obesity) and poor quality<sup>231,232</sup>. As mentioned before, lonely individuals are more likely to be obese. Social isolation and its link with low fruit or vegetable intake have been noted in

studies<sup>233</sup>. In a group of older English individuals, Kobayashi and Steptoe found that socially isolated participants were less likely to eat five daily fruit and vegetable servings<sup>234</sup>. Loneliness is also associated with stress and negative emotions, and this may lead to or aggravate several eating disorders-including anorexia nervosa<sup>235</sup> and bulimia nervosa<sup>236</sup>. In the elderly, loneliness is often associated with dietary inadequacies and undernutrition<sup>237</sup>. As noted with other lifestyles, the impact of COVID-19 has not been good on lifestyles. In an Italian study, 67% of people increased consumption of foods containing added sugars, when pre-COVID-19 eating patterns were compared to COVID-19 lockdown eating patterns<sup>238</sup>.

#### **Conclusion**

Loneliness is detrimentally associated with a wide array of health disorders. It results in frequent emergency room visits and higher hospitalizations. It imparts a poorer prognosis in several chronic diseases, lowers the quality of life, and increases premature mortality. It is associated with equal or more mortality than that due to obesity and smoking. Healthy lifestyles include not smoking, not drinking alcohol, or consuming it in moderation, maintaining a normal BMI, avoiding a sedentary lifestyle, exercising regularly, and following a prudent dietary pattern. Practicing healthy lifestyles not only increases disease-free life but also increases longevity. The actual lifespan at age 50 years, in those compliant with all healthy lifestyles, has been estimated to increase by14.1 years in women and 12.1 years in men. Unfortunately, loneliness has been aggravated by the COVID-19 pandemic. Overall, loneliness discourages healthy lifestyles. The repercussions of this negative association is highlighted in this manuscript.

#### References

- 1. Wenger GC, Davies R, Shahtahmasebi S, Scott A. Social isolation and loneliness in old age: review and model refinement. Ageing Soc. 1996;16(3):333–358. doi:10.1017/S0144686X00003457.
- 2. Petersen N., König H.-H., Hajek A. The link between falls, social isolation and loneliness: A systematic review. Arch. Gerontol. Geriatr. 2020;88:104020. doi: 10.1016/j.archger.2020.104020.
- 3. Bude H, Lantermann E-D. Soziale exklusion und exklusionsempfinden. KZfSS Kölner Zeitschrift für Soziologie und Sozialpsychologie. 2006;58(2):233–252. doi:10.1007/s11575-006-0054-1.
- 4. Petersen N., König H.-H., Hajek A. The link between falls, social isolation and loneliness: A systematic review. Arch. Gerontol. Geriatr. 2020;88:104020. doi: 10.1016/j.archger.2020.104020.
- 5. Rico-Uribe LA, Caballero FF, Olaya B, et al. Loneliness, Social Networks, and Health: A Cross-Sectional Study in Three Countries. PloS one. 2016;11(1):e0145264 doi: 10.1371/journal.pone.0145264.

- 6. Madsen KR, Holstein BE, Damsgaard MT, Rayce SB, Jespersen LN, Due P. Trends in social inequality in loneliness among adolescents 1991-2014. J Public Health (Oxf). 2019 Jun 1;41(2):e133-e140. doi: 10.1093/pubmed/fdy133.
- 7. Cassidy J, Asher SR. Loneliness and peer relations in young children. Child Development. 1992;63(2):350–365.; Asher S, Hymel S, Renshaw PD. Loneliness in children. Child Development. 1984;55:1456–1464.
- 8. De Jong Gierveld J, van Tilburg T. Living arrangements of older adults in the Netherlands and Italy: Coresidence values and behaviour and their consequences for loneliness. Journal of Cross-Cultural Gerontology. 1999;14(1):1–24.
- 9. Victor CD, Scambler SJ, Bowling A, Bondt J. The prevalence of and risk factors for loneliness in later life: A survey of older people in Great Britain. Ageing & Society. 2005:25:357–375.
- 10. F. Landeiro, P. Barrows, E.N. Musson, A.M. Gray, J. Leal. Reducing social isolation and loneliness in older people: A systematic review protocol. BMJ Open, 7 (2017), Article e013778).
- 11. N.K. Valtorta, D.C. Moore, L. Barron, D. Stow, B. Hanratty. Older adults' social relationships and health care utilization: A systematic review. American Journal of Public Health, 108 (2018), pp. e1-e10.
- 12. Cigna/Ipsos. 2018. Cigna U.S. loneliness index. https://www.cigna.com/about-us/newsroom/studies-and-reports/combatting-loneliness/ (accessed July 28, 2020).
- 13. Leigh-Hunt N, Bagguley D, Bash K, Turner V, Turnbull S, Valtorta N, et al. An overview of systematic reviews on the public health consequences of social isolation and loneliness. Public Health. 2017. November;152:157–71. 10.1016/j.puhe.2017.07.035.
- 14. Richard A, Rohrmann S, Vandeleur CL, Schmid M, Barth J, Eichholzer M. Loneliness is adversely associated with physical and mental health and lifestyle factors: Results from a Swiss national survey. PLoS One. 2017;12(7):e0181442. Published 2017 Jul 17. doi:10.1371/journal.pone.0181442.
- 15. Bu F, Zaninotto P, Fancourt D. Longitudinal associations between loneliness, social isolation and cardiovascular events. Heart. 2020 Sep;106(18):1394-1399. doi: 10.1136/heartjnl-2020-316614.
- 16. https://www.who.int/news-room/fact-sheets/detail/cardiovascular-diseases-(cvds) accessed March 11, 2022.
- 17. Valtorta NK, Kanaan M, Gilbody S, Ronzi S, Hanratty B. Loneliness and social isolation as risk factors for coronary heart disease and stroke: systematic review and meta-analysis of longitudinal observational studies. Heart. 2016;102:1009–1016.
- 18. Lauder W, Mummery K, Jones M, Caperchione C. A comparison of health behaviours in lonely and non-lonely populations. Psychol Health Med. 2006;11:233–245. doi: 10.1080/13548500500266607.
- 19. Fox CM, Harper AP, Hyner GC, Lyle RM. Loneliness, emotional repression, marital quality, and major life events in women who develop breast cancer. J Community Health. 1994;19(6):467–482. doi: 10.1007/BF02260327.
- 20. Drageset J, Eide GE, Kirkevold M, Ranhoff AH. Emotional loneliness is associated with mortality among mentally intact nursing home residents with and without cancer: A five-

- year follow-up study. Journal of Clinical Nursing. 2013;22(1–2):106–114. doi: 10.1111/j.1365-2702.2012.04209.x.
- 21. Marty PK, Novotny P, Benzo RP. Loneliness and ED Visits in Chronic Obstructive Pulmonary Disease. Mayo Clin Proc Innov Qual Outcomes. 2019 Aug 23;3(3):350-357. doi: 10.1016/j.mayocpiqo.2019.05.002.
- 22. Tański W. The Role of Clinical Activity, Loneliness, and Satisfaction with e-Health Services as Factors Affecting Quality of Life in Patients with Rheumatoid Arthritis During the SARS-CoV-2 Pandemic. Psychol Res Behav Manag. 2021 Oct 5;14:1581-1590. doi: 10.2147/PRBM.S332141.
- 23. Holwerda TJ. van Tilburg TG. Deeg DJ. Schutter N. Van R. Dekker J. Stek ML. Beekman AT. Schoevers RA. Impact of loneliness and depression on mortality: results from the Longitudinal Ageing StudyAmsterdam. Br J Psychiatry. 2016;209(2):127–134.
- 24. R. Dröes, R. Chattat, A. Diaz, D. Gove, M. Graff, K. Murphy, et al. Social health and dementia: A European consensus on the operationalization of the concept and directions for research and practice. Aging & Mental Health, 21 (2017), pp. 4-17.
- 25. Cohen S. Doyle WJ. Skoner DP. Rabin BS. Gwaltney JM., Jr Social ties and susceptibility to the common cold. JAMA. 1997;277(24):1940–1944.
- 26. Pressman SD, Cohen S, Miller GE, Barkin A, Rabin BS, Treanor JJ. Loneliness, social network size, and immune response to influenza vaccination in college freshmen. Health Psychol. 2005;24(3):297. doi: 10.1037/0278-6133.24.3.297.
- 27. Leigh-Hunt N, Bagguley D, Bash K, Turner V, Turnbull S, Valtorta N, Caan W. An overview of systematic reviews on the public health consequences of social isolation and loneliness. Public Health. 2017;152:157–171. doi: 10.1016/j.puhe.2017.07.035.
- 28. Chung, J., Kulkarni, G.S., Bender, J., Breau, R.H., Guttman, D., Maganti, M., Matthew, A., Morash, R., Papadakos, J. and Jones, J.M. (2020), Modifiable lifestyle behaviours impact the health-related quality of life of bladder cancer survivors. BJU Int, 125: 836-842. https://doi.org/10.1111/bju.15007.
- 29. M.T. Teguo, N. Simo-Tabue, R. Stoykova, C. Meillon, M. Cogne, H. Amiéva, et al. Feelings of loneliness and living alone as predictors of mortality in the elderly: The PAQUID study. Psychosomatic Medicine, 78 (2016), pp. 904-909,
- 30. Rico-Uribe LA, Caballero FF, Martín-María N, Cabello M, Ayuso-Mateos JL, Miret M. Association of loneliness with all-cause mortality: a meta-analysis. PLoS One. 2018;13(1):e0190033. doi: 10.1371/journal.pone.0190033.
- 31. Holt-Lunstad J, Smith TB, Layton B. Social Relationships and Mortality Risk: A Meta-analytic Review. PLoS Med. 2010;7(7):e1000316.
- 32. Freedman A, Nicolle J. Social isolation and loneliness: the new geriatric giants: Approach for primary care. Can Fam Physician. 2020;66(3):176-182.
- 33. National Academies of Sciences, Engineering, and Medicine. 2020. Social Isolation and Loneliness in Older Adults: Opportunities for the Health Care System. Washington, DC: The National Academies Press. https://doi.org/10.17226/25663.
- 34. Cacioppo JT, Hawkley LC, Thisted RA. Perceived social isolation makes me sad: 5-year cross-lagged analyses of loneliness and depressive symptomatology in the Chicago Health,

- Aging, and Social Relations Study. Psychol Aging. 2010;25(2):453–463. doi: 10.1037/a0017216.
- 35. Anderson CA, Harvey RJ. Brief report: discriminating between problems in living: an examination of measures of depression, loneliness, shyness, and social anxiety. J Soc Clin Psychol. 1988;6(3–4):482–491. doi: 10.1521/jscp.1988.6.3-4.482.
- 36. Gutkind S, Gorfinkel LR, Hasin DS. Prospective effects of loneliness on frequency of alcohol and marijuana use. Addict Behav. 2022 Jan;124:107115. doi: 10.1016/j.addbeh.2021.107115.
- 37. Rudatsikira E, Muula AS, Siziya S, Twa-Twa J. Suicidal ideation and associated factors among school-going adolescents in rural Uganda. BMC Psychiatry. 2007;7:67.
- 38. Povedano A, Cava MJ, Monreal MC, Varela R, Musitu G. Victimization, loneliness, overt and relational violence at the school from a gender perspective. Int J Clin Health Psychol. 2015 Jan-Apr;15(1):44-51. doi: 10.1016/j.ijchp.2014.09.001.
- 39. Hughes ME, Waite LJ, Hawkley LC, et al. A short scale for measuring loneliness in large surveys: results from two population-based studies. Res Aging 2004;26:655–72.
- 40. Jylhà M. Old age and loneliness: cross-sectional and longitudinal analyses in the Tampere Longitudinal Study on Aging. Can J Aging. 2004;23(2):157–168.
- 41. https://www.monalisa-asso.fr.
- 42. https://rise-cisa.ca.
- 43. https://www.endloneliness.com.au.
- 44. https://connect2affect.org/ accessed March 7, 2022.
- 45. M. Vozikaki, M. Linardakis, K. Micheli, A. Philalithis. Activity participation and well-being among European adults aged 65 years and older. Social Indicators Research, 131 (2017), pp. 769-795.
- 46. Li Y, Pan A, Wang DD, Liu X, Dhana K, Franco OH, et al. Impact of Healthy Lifestyle Factors on Life Expectancies in the US Population. Circulation. 2018 Jul 24;138(4):345-355. doi: 10.1161/CIRCULATIONAHA.117.032047.
- 47. Reed DM, Foley DJ, White LR, Heimovitz H, Burchfiel CM, Masaki K, 1998. Predictors of healthy aging in men with high life expectancies. Am J Public Health88:1463-1468.
- 48. Hawkley LC, Burleson MH, Berntson GG, Cacioppo JT. Loneliness in everyday life: cardiovascular activity, psychosocial context, and health behaviors. J Pers Soc Psychol. 2003;85(1):105. doi: 10.1037/0022-3514.85.1.105.
- 49. Helen B. Hubert, Daniel A. Bloch, John W. Oehlert, James F. Fries, Lifestyle Habits and Compression of Morbidity, The Journals of Gerontology: Series A, Volume 57, Issue 6, 1 June 2002, Pages M347–M351, <a href="https://doi.org/10.1093/gerona/57.6.M347">https://doi.org/10.1093/gerona/57.6.M347</a>.
- 50. Stenholm S, Head J, Kivimäki M, et al. Smoking, physical inactivity and obesity as predictors of healthy and disease-free life expectancy between ages 50 and 75: a multicohort study. Int J Epidemiol 2016;45:1260-70. 10.1093/ije/dyw126.
- 51. O'Doherty MG, Cairns K, O'Neill V, et al. Effect of major lifestyle risk factors, independent and jointly, on life expectancy with and without cardiovascular disease: results from the Consortium on Health and Ageing Network of Cohorts in Europe and the United States (CHANCES). Eur J Epidemiol 2016;31:455-68. 10.1007/s10654-015-0112-8.

- 52. Li Y, Schoufour J, Wang DD, et al. Healthy lifestyle and life expectancy free of cancer, cardiovascular disease, and type 2 diabetes: prospective cohort study. BMJ. 2020;368:16669. Published 2020 Jan 8. doi:10.1136/bmj.16669.
- 53. Mokdad AH, Forouzanfar MH, Daoud F, et al. Global burden of diseases, injuries, and risk factors for young people's health during 1990-2013: a systematic analysis for the Global Burden of Disease Study 2013. Lancet 2016;387:2383-401.10.1016/S0140-6736(16)00648-6.
- 54. Tamakoshi A, Tamakoshi K, Lin Y, Yagyu K, Kikuchi S, JACC Study Group Healthy lifestyle and preventable death: findings from the Japan Collaborative Cohort (JACC) Study. Prev Med 2009;48:486-92. 10.1016/j.ypmed.2009.02.017.
- 55. Khaw KT, Wareham N, Bingham S, Welch A, Luben R, Day N. Combined impact of health behaviours and mortality in men and women: the EPIC-Norfolk prospective population study. PLoS Med 2008;5:e12. 10.1371/journal.pmed.0050012.
- 56. Manuel DG, Perez R, Sanmartin C, et al. Measuring Burden of Unhealthy Behaviours Using a Multivariable Predictive Approach: Life Expectancy Lost in Canada Attributable to Smoking, Alcohol, Physical Inactivity, and Diet. PLoS Med 2016;13:e1002082. 10.1371/journal.pmed.1002082.
- 57. Li K, Hüsing A, Kaaks R. Lifestyle risk factors and residual life expectancy at age 40: a German cohort study. BMC Med 2014;12:59. 10.1186/1741-7015-12-59.
- 58. O'Doherty MG, Cairns K, O'Neill V, et al. Effect of major lifestyle risk factors, independent and jointly, on life expectancy with and without cardiovascular disease: results from the Consortium on Health and Ageing Network of Cohorts in Europe and the United States (CHANCES). Eur J Epidemiol 2016;31:455-68. 10.1007/s10654-015-0112-8.
- 59. Li Y, Schoufour J, Wang DD, et al. Healthy lifestyle and life expectancy free of cancer, cardiovascular disease, and type 2 diabetes: prospective cohort study. BMJ. 2020;368:16669. Published 2020 Jan 8. doi:10.1136/bmj.16669.
- 60. Hodgson S, Watts I, Fraser S, Roderick P, Dambha-Miller H. Loneliness, social isolation, cardiovascular disease and mortality: a synthesis of the literature and conceptual framework. J R Soc Med. 2020;113:185–192.
- 61. Xia N, Li H. Loneliness, social isolation, and cardiovascular health. Antioxid Redox Signal. 2018;28:837–851.
- 62. Brown EG, Gallagher S, Creaven AM. Loneliness and acute stress reactivity: a systematic review of psychophysiological studies. Psychophysiology. 2018;55:e13031.
- 63. Lieberz, J., et al. (2022) Behavioral and neural dissociation of social anxiety and loneliness. JNeurosci. doi.org/10.1523/JNEUROSCI.2029-21.2022.
- 64. Hawkley L.C., Cole S.W., Capitanio J.P., Norman G.J., Cacioppo J.T. Effects of social isolation on glucocorticoid regulation in social mammals. Horm. Behav. 2012;62(3):314–323.
- 65. Cacioppo J.T., Cacioppo S., Capitanio J.P., Cole S.W. The neuroendocrinology of social isolation. Annu. Rev. Psychol. 2015;66:733–767.
- 66. Wilson SJ, Woody A, Padin AC, Lin J, Malarkey WB, Kiecolt-Glaser JK. Loneliness and Telomere Length: Immune and Parasympathetic Function in Associations With Accelerated Aging. Ann Behav Med. 2019 May 3;53(6):541-550. doi: 10.1093/abm/kay064.

- 67. Lutgendorf SK. DeGeest K. Dahmoush L. Farley D. Penedo F, et al. Social isolation is associated with elevated tumor norepinephrine in ovarian carcinoma patients. Brain Behav Immun. 2011;25:250–255.
- 68. Edwards KM. Bosch JA. Engeland CG. Cacioppo JT. Marucha PT. Elevated macrophage migration inhibitory factor (MIF) is associated with depressive symptoms, blunted cortisol reactivity to acute stress, and lowered morning cortisol. Brain Behav Immun. 2010;24:1202–1208.
- 69. Li H, Xia N. The role of oxidative stress in cardiovascular disease caused by social isolation and loneliness. Redox Biol. 2020;37:101585.
- 70. Eisenberger NI, Moieni M, Inagaki TK, Muscatell KA, Irwin MR. In sickness and in health: the co-regulation of inflammation and social behavior. Neuropsychopharmacology. 2017:42:242–253.
- 71. Spithoven W, Cacioppo S, Goossens L, Cacioppo JT. Genetic contributions to loneliness and their relevance to the evolutionary theory of loneliness. Perspect Psychol Sci. 2019;14:3.
- 72. Baumeister RF, et al. Social exclusion impairs self-regulation. J Pers Soc Psychol. 2005;88:589–604.
- 73. Yarcheski A, et al. A meta-analysis of predictors of positive health practices. J Nurs Scholarsh. 2004;36(2):102–108.
- 74. Lauder W, Mummery K, Jones M, Caperchione C. A comparison of health behaviours in lonely and non-lonely populations. Psychol Health Med. 2006;11(2):233-245. doi:10.1080/13548500500266607.
- 75. Henriksen J, Larsen ER, Mattisson C, Andersson NW. Loneliness, health and mortality. Epidemiol Psychiatr Sci. 2019;28(2):234-239. doi:10.1017/S2045796017000580.
- 76. Hawkley LC, Thisted RA, Cacioppo JT. Loneliness predicts reduced physical activity: cross-sectional & longitudinal analyses. Health Psychol. 2009;28(3):354. doi: 10.1037/a0014400.
- 77. Gerst-Emerson K, Jayawardhana J. Loneliness as a public health issue: the impact of loneliness on health care utilization among older adults. Am J Public Health. 2015;105(5):1013-1019. doi:10.2105/AJPH.2014.302427
- 78. Kusaslan Avci D. Evaluation of the relationship between loneliness and medication adherence in patients with diabetes mellitus: A cross-sectional study. J Int Med Res. 2018 Aug;46(8):3149-3161. doi: 10.1177/0300060518773223.
- 79. Stimpson J.P., Ju H., Raji M.A., Eschbach K. Neighborhood deprivation and health risk behaviors in NHANES III. American Journal of Health Behavior. 2007;31(2):215–222.
- 80. Christakis N.A., Fowler J.H. The collective dynamics of smoking in a large social network. New England Journal of Medicine. 2008;358(21):2249–2258.
- 81. Rosenquist J.N., Murabito J., Fowler J.H., Christakis N.A. The spread of alcohol consumption behavior in a large social network. Annals of Internal Medicine. 2010;152(7):426–433. w141.
- 82. Müller F, Röhr S, Reininghaus U, Riedel-Heller SG. Social Isolation and Loneliness during COVID-19 Lockdown: Associations with Depressive Symptoms in the German Old-Age Population. Int J Environ Res Public Health. 2021;18(7):3615. Published 2021 Mar 31.

- 83. Shihadeh A, Schubert J, Klaiany J, El Sabban M, Luch A, Saliba NA. Toxicant content, physical properties and biological activity of waterpipe tobacco smoke and its tobacco-free alternatives. BMJ. 2015;24(1):22–30.
- 84. Smith P (2019) E-cigarettes-Tobacco Prevention and Control-Minnesota Department of Health. Retrieved from https://www.health.state.mn.us/ecigarettes.
- 85. Mishra S, Joseph RA, Gupta PC, Pezzack B, Ram F, Sinha DN, et al. Trends in bidi and cigarette smoking in India from 1998 to 2015, by age, gender and education. BMJ Glob Health. 2016;1(1):e000005.
- 86. World Health Organization . WHO Report on the Global Tobacco Epidemic, 2017: Monitoring Tobacco Use and Prevention Policies. WHO; Geneva, Switzerland: 2017. Bloomberg Philanthropies.
- 87. Salehi N, Janjani P, Tadbiri H, Rozbahani M, Jalilian M. Effect of cigarette smoking on coronary arteries and pattern and severity of coronary artery disease: a review. J Int Med Res. 2021 Dec;49(12):3000605211059893. doi: 10.1177/03000605211059893.
- 88. Li B, Li D, Liu JF, Wang L, Li BZ, Yan XJ, Liu W, Wu K, Xiang RL. "Smoking paradox" is not true in patients with ischemic stroke: a systematic review and meta-analysis. J Neurol. 2021 Jun;268(6):2042-2054. doi: 10.1007/s00415-019-09596-3.
- 89. Maddatu J, Anderson-Baucum E, Evans-Molina C. Smoking and the risk of type 2 diabetes. Transl Res 2017; 184: 101–107.
- 90. Carbone D. Smoking and cancer. Am J Med. 1992 Jul 15;93(1A):13S-17S. doi: 10.1016/0002-9343(92)90621-h.
- 91. Provenzano M, Serra R, Michael A, Bolignano D, Coppolino G, Ielapi N, Serraino GF, Mastroroberto P, Locatelli F, De Nicola L, Andreucci M. Smoking habit as a risk amplifier in chronic kidney disease patients. Sci Rep. 2021 Jul 20;11(1):14778. doi: 10.1038/s41598-021-94270-w.
- 92. Forey BA, Thornton AJ, Lee PN. Systematic review with meta-analysis of the epidemiological evidence relating smoking to COPD, chronic bronchitis and emphysema. BMC Pulm Med. 2011;11:36. doi: 10.1186/1471-2466-11-36.
- 93. Amiri S, Behnezhad S. Smoking and disability pension: a systematic review and meta-analysis. Public Health. 2020 Sep;186:297-303. doi: 10.1016/j.puhe.2020.04.013.
- 94. Qin W, Magnussen CG, Li S, Steffen LM, Xi B, Zhao M. Light Cigarette Smoking Increases Risk of All-Cause and Cause-Specific Mortality: Findings from the NHIS Cohort Study. Int J Environ Res Public Health. 2020 Jul 15;17(14):5122. doi: 10.3390/ijerph17145122. Erratum in: Int J Environ Res Public Health. 2020 Aug 25;17(17).
- 95. Doll R, Peto R, Boreham J, Sutherland I. Mortality in relation to smoking: 50 years' observations on male British doctors. BMJ 2004;328:1519.
- 96. https://www.fda.gov/tobacco-products/rules-regulations-and-guidance/harmful-and-potentially-harmful-constituents-tobacco-products-and-tobacco-smoke-established-list (accessed March 10, 2022)
- 97. Yalcin E, de la Monte S. Tobacco nitrosamines as culprits in disease: mechanisms reviewed. J Physiol Biochem. 2016 Mar;72(1):107-20. doi: 10.1007/s13105-016-0465-9.

- 98. Loeb LA, Ernster VL, Warner KE, Abbotts J, Laszlo J. Smoking and lung cancer: an overview. Cancer Res. 1984 Dec;44(12 Pt 1):5940-58. Erratum in: Cancer Res 1986 Oct;46(10):5453
- 99. World No Tobacco Day 2017. Tobacco threatens us all: protect health, reduce poverty and promote development. Geneva: World Health Organization; 2017.
- 100. Lauder W., Mummery K., Jones M., Caperchione C. A comparison of health behaviours in lonely and non-lonely populations. Psychology Health & Medicine. 2006;11(2):233–245.
- 101. Shankar A, McMunn A, Banks J, Steptoe A. Loneliness, social isolation, and behavioral and biological health indicators in older adults. Health Psychol. 2011;30(4):377. doi: 10.1037/a0022826.
- 102. Kobayashi LC, Steptoe A. Social Isolation, Loneliness, and Health Behaviors at Older Ages: Longitudinal Cohort Study. Ann Behav Med. 2018 May 31;52(7):582-593. doi: 10.1093/abm/kax033.
- 103. Dyal SR, Valente TW. A Systematic Review of Loneliness and Smoking: Small Effects, Big Implications. Subst Use Misuse. 2015;50(13):1697-1716. doi:10.3109/10826084.2015.1027933.
- 104. Habibi M, Hosseini F, Darharaj M, et al. Attachment Style, Perceived Loneliness, and Psychological Well-Being in Smoking and Non-Smoking University Students. J Psychol 2018;152:226-36. 10.1080/00223980.2018.1446894.
- 105. Zhang CL, Xu YM, Zhong BL. The association between smoking and loneliness among Chinese university freshmen. Ann Transl Med. 2020 May;8(10):649. doi: 10.21037/atm-20-3523.
- 106. Carreras G, Lugo A, Stival C, Amerio A, Odone A, Pacifici R, et al. Impact of COVID-19 lockdown on smoking consumption in a large representative sample of Italian adults. Tob Control. 2021 Mar 29:tobaccocontrol-2020-056440. doi: 10.1136/tobaccocontrol-2020-056440.
- 107. https://www.nhlbi.nih.gov/health/educational/lose\_wt/BMI/bmicalc.htm accessed October 27, 2021.
- 108. Jensen MD, Ryan DH, Apovian CM, Ard JD, Comuzzie AG, Donato KA, et al. 2013 AHA/ACC/TOS guideline for the management of overweight and obesity in adults: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines and The Obesity Society. Circulation. 2014;129(suppl 2):S102–S138. doi: 10.1161/01.cir.0000437739.71477.ee.
- 109. https://www.cdc.gov/obesity/adult/defining.html. Accessed March 10, 2022.
- 110. https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight. Accessed March 1, 2022.
- 111. Staynor JMD, Smith MK, Donnelly CJ, Sallam AE, Ackland TR. DXA reference values and anthropometric screening for visceral obesity in Western Australian adults. Sci Rep. 2020 Oct 30;10(1):18731. doi: 10.1038/s41598-020-73631-x.
- 112. https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight. Accessed March 5, 2022.
- 113. Powell-Wiley TM, Poirier P, Burke LE, Després JP, Gordon-Larsen P, Lavie CJ, Lear SA, Ndumele CE, Neeland IJ, Sanders P, St-Onge MP; American Heart Association Council

- on Lifestyle and Cardiometabolic Health; Council on Cardiovascular and Stroke Nursing; Council on Clinical Cardiology; Council on Epidemiology and Prevention; and Stroke Council. Obesity and Cardiovascular Disease: A Scientific Statement From the American Heart Association. Circulation. 2021 May 25;143(21):e984-e1010. doi: 10.1161/CIR.000000000000000073.
- 114. Saklayen MG. The Global Epidemic of the Metabolic Syndrome. Curr Hypertens Rep. 2018 Feb 26;20(2):12. doi: 10.1007/s11906-018-0812-z.),
- 115. Pillon NJ, Loos RJF, Marshall SM, Zierath JR. Metabolic consequences of obesity and type 2 diabetes: Balancing genes and environment for personalized care. Cell. 2021 Mar 18;184(6):1530-1544. doi: 10.1016/j.cell.2021.02.012.
- 116. Tsuboi N. Obesity Indices and the Risk of CKD. Intern Med. 2021 Jul 1;60(13):1987-1988. doi: 10.2169/internalmedicine.6921-20. Epub 2021 Feb 8.
- 117. Bence KK, Birnbaum MJ. Metabolic drivers of non-alcoholic fatty liver disease. Mol Metab. 2021 Aug;50:101143. doi: 10.1016/j.molmet.2020.101143.
- 118. Obesity and its relationship with cancer Part II: Agarwal, S. K. (2021). Obesity And Its Relationship with Cancer: Part Ii Cancer. American International Journal of Cancer Studies, 3(1), 14–24. https://doi.org/10.46545/aijcs.v3i1.217.
- 119. Erridge S, Moussa O, McIntyre C, Hariri A, Tolley N, Kotecha B, Purkayastha S. Obstructive Sleep Apnea in Obese Patients: a UK Population Analysis. Obes Surg. 2021 May;31(5):1986-1993. doi: 10.1007/s11695-020-05196-7.
- 120. Dai S, Zhang J, Zhu X, et al. Identification of key genes underlying the effects of obesity on knee osteoarthritis. Chin Med J (Engl). 2021;135(4):474-476. Published 2021 Sep 21. doi:10.1097/CM9.000000000001670.
- 121. National Academies of Sciences, Engineering, and Medicine. 2020. Social Isolation and Loneliness in Older Adults: Opportunities for the Health Care System. Washington, DC: The National Academies Press. https://doi.org/10.17226/25663.
- 122. Fulton S, Décarie-Spain L, Fioramonti X, Guiard B, Nakajima S. The menace of obesity to depression and anxiety prevalence. Trends Endocrinol Metab. 2022 Jan;33(1):18-35. doi: 10.1016/j.tem.2021.10.005.
- 123. Reilly JJ, Kelly J. Long-term impact of overweight and obesity in childhood and adolescence on morbidity and premature mortality in adulthood: systematic review. Int J Obes (Lond). 2011 Jul;35(7):891-8. doi: 10.1038/ijo.2010.222.
- 124. National Academies of Sciences, Engineering, and Medicine. 2020. Social Isolation and Loneliness in Older Adults: Opportunities for the Health Care System. Washington, DC: The National Academies Press. https://doi.org/10.17226/25663external.icon.
- 125. Macmillan M. The future of television and medicine. Proc R Soc Med. 1969 Apr;62((4)):401–2.
- 126. WHO Consultation on Obesity (1999: Geneva, Switzerland) & World Health Organization . Obesity: Preventing and Managing the Global Epidemic. Report of a WHO Consultation. World Health Organ Tech Rep Ser (2000) 894:i–xii, 1-253.
- 127. Hajek A, Kretzler B, König HH. The Association Between Obesity and Social Isolation as Well as Loneliness in the Adult Population: A Systematic Review. Diabetes Metab Syndr Obes. 2021;14:2765-2773. Published 2021 Jun 17. doi:10.2147/DMSO.S313873.

- 128. Petitte T, Mallow J, Barnes E, Petrone A, Barr T, Theeke L. A systematic review of loneliness and common chronic physical conditions in adults. Open Psychol J. 2015;8(Suppl 2):113–32. doi: 10.2174/1874350101508010113.
- 129. Hawkley LC, Thisted RA, Cacioppo JT. Loneliness predicts reduced physical activity: cross-sectional & longitudinal analyses. Health Psychol. 2009 May;28((3)):354–63.
- 130. Hawkley LC, Preacher KJ, Cacioppo JT. Loneliness impairs daytime functioning but not sleep duration. Health Psychol. 2010 Mar;29((2)):124–9.
- 131. Jacobs JM, Cohen A, Hammerman-Rozenberg R, Stessman J. Global sleep satisfaction of older people: the Jerusalem Cohort Study. J Am Geriatr Soc. 2006;54:325–9. doi: 10.1111/j.1532-5415.2005.00579.x.
- 132. Morse SA, Ciechanowski PS, Katon WJ, Hirsch IB. Isn't this just bedtime snacking? The potential adverse effects of night-eating symptoms on treatment adherence and outcomes in patients with diabetes. Diabetes Care. 2006;29:1800–4. doi: 10.2337/dc06-0315.
- 133. Whisman MA. Loneliness and the metabolic syndrome in a population-based sample of middle-aged and older adults. Health Psychol. 2010;29:550–4. doi: 10.1037/a0020760.
- 134. Lauder W, Mummery K, Jones M, Caperchione C. A comparison of health behaviours in lonely and non-lonely populations. Psychol Health Med. 2006 May;11((2)):233–45.
- 135. Jung FU, Luck-Sikorski C. Overweight and lonely? A representative study on loneliness in obese people and its determinants. Obes Facts. 2019;12:440–7. doi: 10.1159/000500095.
- 136. Obesity jobs
- 137. Puhl RM, Himmelstein MS, Quinn DM. Internalizing Weight Stigma: Prevalence and Sociodemographic Considerations in US Adults. Obesity (Silver Spring) 2018 Jan;26((1)):167–75.
- 138. Schumaker JF, Krejci RC, Small L, Sargent RG. Experience of loneliness by obese individuals. Psychol Rep. 1985;57:1147–54. doi: 10.2466/pr0.1985.57.3f.1147.
- 139. Phelan SM, Burgess DJ, Puhl R, Dyrbye LN, Dovidio JF, Yeazel M, et al. The Adverse Effect of Weight Stigma on the Well-Being of Medical Students with Overweight or Obesity: Findings from a National Survey. J Gen Intern Med. 2015 Sep;30(9):1251-8. doi: 10.1007/s11606-015-3266-x.
- 140. Heinberg LJ, Steffen K. Social Isolation and Loneliness During the COVID-19 Pandemic: Impact on Weight. Curr Obes Rep. 2021 Sep;10(3):365-370. doi: 10.1007/s13679-021-00447-9.
- 141. Nathan PE, Conrad M, Skinstad AH. History of the Concept of Addiction. Annu Rev Clin Psychol. 2016;12:29–51.
- 142. https://www.who.int/healthinfo/indicators/2015/chi\_2015\_64\_alcohol\_consumption.pdf? ua=1
- 143. Kalinowski A., Humphreys K. Governmental standard drink definitions and low-risk alcohol consumption guidelines in 37 countries. Addiction 2016; 111: 1293–1298.
- 144. Maisch B. Alcoholic cardiomyopathy: The result of dosage and individual predisposition. Herz. 2016 Sep;41(6):484-93. doi: 10.1007/s00059-016-4469-6.
- 145. Ajani UA, Hennekens CH, Spelsberg A, Manson JE. Alcohol consumption and risk of type 2 diabetes mellitus among US male physicians. Arch Intern Med. 2000;160(7):1025-1030.

- 146. 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 (Clinical research ed). 1999;319(7224):1523-1528.
- 147. Berger K, Ajani UA, Kase CS, et al. Light-to-moderate alcohol consumption and the risk of stroke among U.S. male physicians. N Engl J Med. 1999;341(21):1557-1564.
- 148. Buja A, Scafato E, Baggio B, et al. Renal impairment and moderate alcohol consumption in the elderly. Results from the Italian Longitudinal Study on Aging (ILSA). Public health nutrition. 2011;14(11):1907-1918.
- 149. Gaziano JM, Gaziano TA, Glynn RJ, et al. Light-to-moderate alcohol consumption and mortality in the Physicians' Health Study enrollment cohort. J Am Coll Cardiol. 2000;35(1):96-105.
- 150. Griswold M.G., Fullman N., Hawley C., et al. Alcohol use and burden for 195 countries and territories, 1990-2016: a systematic analysis for the Global Burden of Disease Study 2016. Lancet. 2018;392(10152):1015–1035. doi: 10.1016/S0140-6736(18)31310-2.
- 151. Huang C., Zhan J., Liu Y.J., et al. Association between alcohol consumption and risk of cardiovascular disease and all-cause mortality in patients with hypertension: a meta-analysis of prospective cohort studies. Mayo Clin. Proc. 2014;89(9):1201–1210. doi: 10.1016/j.mayocp.2014.05.014.
- 152. Butler L, Popkin BM, Poti JM. Associations of alcoholic beverage consumption with dietary intake, waist circumference, and body mass index in US adults: national health and nutrition examination survey 2003-2012. J Acad Nutr Diet. 2018;118(3):409–420.e3. doi: 10.1016/j.jand.2017.09.030.
- 153. Rumgay H, Shield K, Charvat H, Ferrari P, Sornpaisarn B, Obot I, Islami F, Lemmens VEPP, Rehm J, Soerjomataram I. Global burden of cancer in 2020 attributable to alcohol consumption: a population-based study. Lancet Oncol. 2021 Aug;22(8):1071-1080. doi: 10.1016/S1470-2045(21)00279-5.
- 154. Wu X, Liu X, Liao W, Kang N, Dong X, Abdulai T, Zhai Z, Wang C, Wang X, Li Y. Prevalence and characteristics of alcohol consumption and risk of type 2 diabetes mellitus in rural China. BMC Public Health. 2021 Sep 9;21(1):1644. doi: 10.1186/s12889-021-11681-0.
- 155. Krumpe PE, Cummiskey JM, Lillington GA. Alcohol and the respiratory tract. Med Clin North Am. 1984 Jan;68(1):201-19. doi: 10.1016/s0025-7125(16)31250-0.
- 156. Perneger TV, Whelton PK, Puddey IB, Klag MJ. Risk of end-stage renal disease associated with alcohol consumption. Am J Epidemiol. 1999;150(12):1275-1281.
- 157. Chow MSC, Poon SHL, Lui KL, Chan CCY, Lam WWT. Alcohol Consumption and Depression Among University Students and Their Perception of Alcohol Use. East Asian Arch Psychiatry. 2021 Dec;31(4):87-96. doi: 10.12809/eaap20108.
- 158. Morys F, Dadar M, Dagher A. Association Between Midlife Obesity and Its Metabolic Consequences, Cerebrovascular Disease, and Cognitive Decline. J Clin Endocrinol Metab. 2021 Sep 27;106(10):e4260-e4274. doi: 10.1210/clinem/dgab135.
- 159. Stermer E. Alcohol consumption and the gastrointestinal tract. Isr Med Assoc J. 2002 Mar;4(3):200-2.
- 160. Sun FR, Wang BY. Alcohol and Metabolic-associated Fatty Liver Disease. J Clin Transl Hepatol. 2021 Oct 28;9(5):719-730. doi: 10.14218/JCTH.2021.00173.

- 161. Thakkar MM, Sharma R, Sahota P. Alcohol disrupts sleep homeostasis. Alcohol. 2015 Jun;49(4):299-310. doi: 10.1016/j.alcohol.2014.07.019.
- 162. Chen Z, Li S, Guo L, Peng X, Liu Y. Prenatal alcohol exposure induced congenital heart diseases: From bench to bedside. Birth Defects Res. 2021 Apr 15;113(7):521-534. doi: 10.1002/bdr2.1743.
- 163. George WH. Alcohol and Sexual Health Behavior: "What We Know and How We Know It". J Sex Res. 2019 May-Jun;56(4-5):409-424. doi: 10.1080/00224499.2019.1588213.
- 164. Cherpitel CJ. Alcohol and violence-related injuries in the emergency room. Recent Dev Alcohol. 1997;13:105-18. doi: 10.1007/0-306-47141-8\_6.
- 165. Kalichman SC, Simbayi LC, Kaufman M, Cain D, Jooste S. Alcohol use and sexual risks for HIV/AIDS in sub-Saharan Africa: systematic review of empirical findings. Prev Sci. 2007 Jun;8(2):141-51. doi: 10.1007/s11121-006-0061-2.
- 166. https://www.iarc.who.int/wp-content/uploads/2018/07/WCR\_2014\_Chapter\_2-3.pdf accessed March 10, 2022.
- 167. https://world-heart-federation.org/news/no-amount-of-alcohol-is-good-for-the-heart-says-world-heart-federation/ accessed March 11, 2022..
- 168. Akerlind I, Hornquist JO. Loneliness and alcohol abuse: A review of evidences of an interplay. Social Science & Medicine. 1992;34(4):405–414.
- 169. McKay MT, Konowalczyk S, Andretta JR, Cole JC. The direct and indirect effect of loneliness on the development of adolescent alcohol use in the United Kingdom. Addict Behav Rep. 2017;6:65-70. Published 2017 Aug 1. doi:10.1016/j.abrep.2017.07.003.
- 170. Barbosa Filho V.C., Campos W., Lopes Ada S. Prevalence of alcohol and tobacco use among Brazilian adolescents: A systematic review, Revista de Saúde Pública. 2012;46:901–917.
- 171. Pedersen W., von Soest T. Adolescent alcohol use and binge drinking: An 18-year trend study of prevalence and correlates. Alcohol and Alcoholism. 2015;50:219–225.
- 172. Gonzalez V.M., Skewes M.C. Solitary heavy drinking, social relationships, and negative mood regulation in college drinkers. Addiction Research and Theory. 2013;21:285–294.
- 173. Akerlind I, Hörnquist JO. Loneliness and alcohol abuse: a review of evidences of an interplay. Soc Sci Med. 1992 Feb;34(4):405-14. doi: 10.1016/0277-9536(92)90300-f.
- 174. Hwang TJ, Rabheru K, Peisah C, Reichman W, Ikeda M. Loneliness and social isolation during the COVID-19 pandemic. Int Psychogeriatr. 2020;32(10):1217-1220. doi:10.1017/S1041610220000988.
- 175. Grossman ER, Benjamin-Neelon SE, Sonnenschein S. (2020) Alcohol consumption during the COVID-19 pandemic: a cross-sectional survey of US adults. Int J Environ Res Public Health 17:9189.
- 176. Pollard MS, Tucker JS, Green HD. (2020) Changes in adult alcohol use and consequences during the COVID-19 pandemic in the US. JAMA Netw Open 3:e2022942.
- 177. Yanguas J, Pinazo-Henandis S, Tarazona-Santabalbina FJ. The complexity of loneliness. Acta Biomed. 2018;89(2):302-314. Published 2018 Jun 7. doi:10.23750/abm.v89i2.7404.
- 178. Koivumaa-Honkanen H, Kaprio J, Korhonen T, Honkanen RJ, Heikkilä K, Koskenvuo M. Self-reported life satisfaction and alcohol use: a 15-year follow-up of healthy adult twins. Alcohol Alcohol. 2012 Mar-Apr;47(2):160-8. doi: 10.1093/alcalc/agr151.

- 179. Jetté M, Sidney K, Blümchen G. Metabolic equivalents (METS) in exercise testing, exercise prescription, and evaluation of functional capacity. Clin Cardiol. 1990 Aug;13(8):555-65. doi: 10.1002/clc.4960130809.
- 180. Sedentary Behaviour Research Network . Ottawa: Sedentary Behaviour Research Network; 2020. SBRN Terminology Consensus Project: 2017-2020 [Internet] [cited 2020 Nov 6]. Available from: https://www.sedentarybehaviour.org/sbrn-terminology-consensus-project/.
- 181. Global Action Plan on Physical Activity 2018–2030: More Active People for a Healthier World. World Health Organization; Geneva: 2018.
- 182. Ainsworth BE, Haskell WL, Herrmann SD, Meckes N, Bassett DR, Jr, Tudor-Locke C, et al. 2011 Compendium of physical activities: a second update of codes and MET values. Med Sci Sports Exerc. 2011;43:1575–81.
- 183. Owen N, Healy GN, Matthews CE, Dunstan DW. Too much sitting: the population health science of sedentary behavior. Exerc Sport Sci Rev. 2010; 38:105–113. doi: 10.1097/JES.0b013e3181e373a2.
- 184. Gibbs BB, Hergenroeder AL, Katzmarzyk PT, Lee IM, Jakicic JM. Definition, measurement, and health risks associated with sedentary behavior. Med Sci Sports Exerc. 2015;47(6):1295–1300. doi: 10.1249/MSS.000000000000517.
- 185. Bailey DP, Hewson DJ, Champion RB, Sayegh SM. Sitting time and risk of cardiovascular disease and diabetes: a systematic review and meta-analysis. Am J Prev Med. 2019;57(3):408–416. doi: 10.1016/j.amepre.2019.04.015.
- 186. Biswas A, Oh PI, Faulkner GE, Bajaj RR, Silver MA, Mitchell MS, Alter DA. Sedentary time and its association with risk for disease incidence, mortality, and hospitalization in adults: a systematic review and meta-analysis. Ann Intern Med. 2015 Jan 20;162(2):123-32. doi: 10.7326/M14-1651.
- 187. Patel AV, Friedenreich CM, Moore SC, Hayes SC, Silver JK, Campbell KL, et al. American College of Sports Medicine Roundtable Report on Physical Activity, Sedentary Behavior, and Cancer Prevention and Control. Med Sci Sports Exerc. 2019 Nov;51(11):2391-2402. doi: 10.1249/MSS.0000000000002117.
- 188. Grady CL, Muirhead F, Skelton DA, Mavroeidi A. Exploring osteoporosis sufferers knowledge on sedentary behaviour in the management of their disease. J Frailty Sarcopenia Falls. 2021;6(2):36-42. Published 2021 Jun 1. doi:10.22540/JFSF-06-036.
- 189. Huang Y, Li L, Gan Y, Wang C, Jiang H, Cao S, Lu Z. Sedentary behaviors and risk of depression: a meta-analysis of prospective studies. Transl Psychiatry. 2020 Jan 22;10(1):26. doi: 10.1038/s41398-020-0715-z.
- 190. Fiuza-Luces C, Santos-Lozano A, Joyner M, Carrera-Bastos P, Picazo O, Zugaza JL, Izquierdo M, Ruilope LM, Lucia A. Exercise benefits in cardiovascular disease: beyond attenuation of traditional risk factors. Nat Rev Cardiol. 2018 Dec;15(12):731-743. doi: 10.1038/s41569-018-0065-1.
- 191. Sampath Kumar A, Maiya AG, Shastry BA, Vaishali K, Ravishankar N, Hazari A, Gundmi S, Jadhav R. Exercise and insulin resistance in type 2 diabetes mellitus: A systematic review and meta-analysis. Ann Phys Rehabil Med. 2019 Mar;62(2):98-103. doi: 10.1016/j.rehab.2018.11.001.

- 192. Idorn M, Thor Straten P. Exercise and cancer: from "healthy" to "therapeutic"? Cancer Immunol Immunother. 2017 May;66(5):667-671. doi: 10.1007/s00262-017-1985-z.
- 193. Warburton DE, Nicol CW, Bredin SS. Health benefits of physical activity: the evidence. CMAJ. 2006;174(6):801-809. doi:10.1503/cmaj.051351.
- 194. Katrina L. Piercy and Richard P. Troiano. Physical Activity Guidelines for Americans From the US Department of Health and Human Services. Circulation: Cardiovascular Quality and Outcomes. 2018;11:e005263.https://doi.org/10.1161/CIRCOUTCOMES.118.005263.
- 195. https://www.cdc.gov/physicalactivity/basics/age-chart.html accessed March 12, 2022.
- 196. Hawkley LC, Thisted RA, Cacioppo JT. Loneliness predicts reduced physical activity: cross-sectional & longitudinal analyses. Health Psychol. 2009;28(3):354. doi: 10.1037/a0014400.
- 197. Steptoe A, Shankar A, Demakakos P, Wardle J. Social isolation, loneliness, and all-cause mortality in older men and women. Proc Natl Acad Sci U S A 2013; 110: 5797–5801.
- 198. Kobayashi LC, Steptoe A. Social Isolation, Loneliness, and Health Behaviors at Older Ages: Longitudinal Cohort Study. Ann Behav Med. 2018 May 31;52(7):582-593. doi: 10.1093/abm/kax033.
- 199. Schrempft S, Jackowska M, Hamer M, Steptoe A. Associations between social isolation, loneliness, and objective physical activity in older men and women. BMC Public Health. 2019 Jan 16;19(1):74. doi: 10.1186/s12889-019-6424-y.
- 200. Vancampfort D, Lara E, Smith L, Rosenbaum S, Firth J, Stubbs B, Hallgren M, Koyanagi A. Physical activity and loneliness among adults aged 50 years or older in six low- and middle-income countries. Int J Geriatr Psychiatry. 2019 Dec;34(12):1855-1864. doi: 10.1002/gps.5202.
- 201. Allen L, Williams J, Townsend N, et al. Socioeconomic status and non-communicable disease behavioural risk factors in low-income and lower-middle-income countries: a systematic review. The Lancet Global Health. 2017;5(3):e277–e289.
- 202. Allison Moser Mays, Sungjin Kim, Katrina Rosales, Tam Au, Sonja Rosen. The Leveraging Exercise to Age in Place (LEAP) Study: Engaging Older Adults in Community-Based Exercise Classes to Impact Loneliness and Social Isolation. The American Journal of Geriatric Psychiatry, 2020; DOI: 10.1016/j.jagp.2020.10.006.
- 203. Martínez-de-Quel Ó, Suárez-Iglesias D, López-Flores M, Pérez CA. Physical activity, dietary habits and sleep quality before and during COVID-19 lockdown: A longitudinal study. Appetite. 2021 Mar 1;158:105019. doi: 10.1016/j.appet.2020.105019.
- 204. Ryan DH, Kahan S. Guideline recommendations for obesity management. Med Clin North Am. 2018;102:49–63. doi: 10.1016/j.mcna.2017.08.006.
- 205. Yu E, Malik VS, Hu FB. Cardiovascular Disease Prevention by Diet Modification: JACC Health Promotion Series. J Am Coll Cardiol. 2018;72(8):914-926. doi:10.1016/j.jacc.2018.02.085.
- 206. Schwingshackl L, Schwedhelm C, Galbete C, Hoffmann G. Adherence to Mediterranean Diet and Risk of Cancer: An Updated Systematic Review and Meta-Analysis. Nutrients. 2017 Sep 26;9(10):1063. doi: 10.3390/nu9101063

- 207. Martín-Peláez S, Fito M, Castaner O. Mediterranean Diet Effects on Type 2 Diabetes Prevention, Disease Progression, and Related Mechanisms. A Review. Nutrients. 2020 Jul 27;12(8):2236. doi: 10.3390/nu12082236.
- 208. Bales CW, Porter Starr KN. Obesity Interventions for Older Adults: Diet as a Determinant of Physical Function. Adv Nutr. 2018 Mar 1;9(2):151-159. doi: 10.1093/advances/nmx016.
- 209. Reid M, Maras JE, Shea S, Wood AC, Castro-Diehl C, Johnson DA, Huang T, Jacobs DR Jr, Crawford A, St-Onge MP, Redline S. Association between diet quality and sleep apnea in the Multi-Ethnic Study of Atherosclerosis. Sleep. 2019 Jan 1;42(1):zsy194. doi: 10.1093/sleep/zsy194.
- 210. Molina P, Gavela E, Vizcaíno B, Huarte E, Carrero JJ. Optimizing Diet to Slow CKD Progression. Front Med (Lausanne). 2021 Jun 25;8:654250. doi: 10.3389/fmed.2021.654250.
- 211. Khalili H, Chan SSM, Lochhead P, Ananthakrishnan AN, Hart AR, Chan AT. The role of diet in the aetiopathogenesis of inflammatory bowel disease. Nat Rev Gastroenterol Hepatol. 2018 Sep;15(9):525-535. doi: 10.1038/s41575-018-0022-9.
- 212. Gioia C, Lucchino B, Tarsitano MG, Iannuccelli C, Di Franco M. Dietary Habits and Nutrition in Rheumatoid Arthritis: Can Diet Influence Disease Development and Clinical Manifestations? Nutrients. 2020 May 18;12(5):1456. doi: 10.3390/nu12051456.
- 213. Kepka A, Ochocinska A, Borzym-Kluczyk M, Skorupa E, Stasiewicz-Jarocka B, Chojnowska S, Waszkiewicz N. Preventive Role of L-Carnitine and Balanced Diet in Alzheimer's Disease. Nutrients. 2020 Jul 3;12(7):1987. doi: 10.3390/nu12071987.
- 214. Odermatt A. The Western-style diet: a major risk factor for impaired kidney function and chronic kidney disease. Am J Physiol Renal Physiol. 2011 Nov;301(5):F919-31. doi: 10.1152/ajprenal.00068.2011.
- 215. Vossen E, Goethals S, De Vrieze J, Boon N, Van Hecke T, De Smet S. Red and processed meat consumption within two different dietary patterns: Effect on the colon microbial community and volatile metabolites in pigs. Food Res Int. 2020 Mar;129:108793. doi: 10.1016/j.foodres.2019.108793.
- 216. Weaver CM. Potassium and health. Adv Nutr. 2013;4(3):368S-77S. Published 2013 May 1. doi:10.3945/an.112.003533.
- 217. Monteiro C.A. Moubarac J.C. Cannon G. Ng S.W. Popkin B. Ultra-processed products are becoming dominant in the global food system. Obes. Rev. 2013; 14: 21-28.
- 218. Mozaffarian D.Dietary and Policy Priorities for Cardiovascular Disease, Diabetes, and Obesity: A Comprehensive Review. Circulation. 2016; 133: 187-225.
- 219. Yu E, Malik VS, Hu FB. Cardiovascular Disease Prevention by Diet Modification: JACC Health Promotion Series. J Am Coll Cardiol. 2018;72(8):914-926. doi:10.1016/j.jacc.2018.02.085
- 220. https://www.diabetes.co.uk/news/2005/dec/western-diets-diabetes-link.html. Accessed March 12, 2022.
- 221. http://nrs.harvard.edu/urn-3:HUL.InstRepos:11940214.
- 222. Stoll BA. Western diet, early puberty, and breast cancer risk. Breast Cancer Res Treat. 1998 Jun;49(3):187-93. doi: 10.1023/a:1006003110909.

- 223. Norat T, Lukanova A, Ferrari P, Riboli E. Meat consumption and colorectal cancer risk: dose–response meta-analysis of epidemiological studies. Int J Cancer. 2002;98:241–256.
- 224. Więckowska-Gacek A, Mietelska-Porowska A, Wydrych M, Wojda U. Western diet as a trigger of Alzheimer's disease: From metabolic syndrome and systemic inflammation to neuroinflammation and neurodegeneration. Ageing Res Rev. 2021 Sep;70:101397. doi: 10.1016/j.arr.2021.101397.
- 225. Fadnes, L.T., et al. (2022) Estimating impact of food choices on life expectancy: A modeling study. PLOS Medicine. doi.org/10.1371/journal.pmed.1003889.
- 226. Abajobir AA, Abate KH, Abbafati C et al. Global, regional, and national comparative risk assessment of 84 behavioural, environmental and occupational, and metabolic risks or clusters of risks, 1990–2016: a systematic analysis for the Global Burden of Disease Study 2016. Lancet 2017;390:1345–1422.
- 227. Slavin J. Structure, Nomenclature, and Properties of Carbohydrates. In: Stipanuk M.H., Caudill M.A., editors. Biochemistry, Physiology and Molecular Aspects of Human Nutrition. 3rd ed. Elsevier Saunders; St. Louis, MO, USA: 2013. pp. 50–68.
- 228. Nocella C, Cammisotto V, Fianchini L, D'Amico A, Novo M, Castellani V, Stefanini L, Violi F, Carnevale R. Extra Virgin Olive Oil and Cardiovascular Diseases: Benefits for Human Health. Endocr Metab Immune Disord Drug Targets. 2018;18(1):4-13. doi: 10.2174/1871530317666171114121533.
- 229. Satija A, Hu FB. Plant-based diets and cardiovascular health. Trends Cardiovasc Med. 2018 Oct;28(7):437-441. doi: 10.1016/j.tcm.2018.02.004.
- 230. Kim H, Caulfield LE, Rebholz CM. Healthy Plant-Based Diets Are Associated with Lower Risk of All-Cause Mortality in US Adults. J Nutr. 2018 Apr 1;148(4):624-631.
- 231. Lauder W, Mummery K, Jones M, Caperchione C. A comparison of health behaviours in lonely and non-lonely populations. Psychology Health & Medicine. 2006;11:233–245.
- 232. Rotenberg KJ, Flood D. Loneliness, dysphoria, dietary restraint, and eating behavior. Int J Eat Disord. 1999;25(1):55–64.
- 233. Weyers S., Dragano N., Mobus S., Beck E.M., Stang A., Mohlenkamp S....Siegrist J. Poor social relations and adverse health behaviour: Stronger associations in low socioeconomic groups? International Journal of Public Health. 2010;55(1):17–23.
- 234. Kobayashi LC, Steptoe A. Social Isolation, Loneliness, and Health Behaviors at Older Ages: Longitudinal Cohort Study. Ann Behav Med. 2018 May 31;52(7):582-593. doi: 10.1093/abm/kax033.
- 235. Ramic E, Pranjic N, Batic-Mujanovic O, Karic E, Alibasic E, Alic A. The effect of loneliness on malnutrition in elderly population. Med Arh. 2011;65(2):92-5.
- 236. Levine MP. Loneliness and eating disorders. J Psychol. 2012 Jan-Apr;146(1-2):243-57. doi: 10.1080/00223980.2011.606435.
- 237. Ferry M, Sidobre B, Lambertin A, Barberger-Gateau P. The SOLINUT study: analysis of the interaction between nutrition and loneliness in persons aged over 70 years. J Nutr Health Aging. 2005 Jul-Aug;9(4):261-8.
- 238. Prete M, Luzzetti A, Augustin LSA, Porciello G, Montagnese C, Calabrese I, et al. Changes in Lifestyle and Dietary Habits during COVID-19 Lockdown in Italy: Results of an Online Survey. Nutrients. 2021 Jun 3;13(6):1923. doi: 10.3390/nu13061923.

