

Obesity Facts

Obes Facts , DOI: 10.1159/000533301

Received: November 18, 2021

Accepted: July 19, 2023

Published online: August 8, 2023

How could different obesity scenarios alter the burden of type 2 diabetes and liver disease in Saudi Arabia?

Coker T, Saxton J, Retat L, Guzek J, Card-Gowers J, Bindhim NF, Althumiri NA, Aldubayan K, Razack HIA, Webber L, Alqahtani SA

ISSN: 1662-4025 (Print), eISSN: 1662-4033 (Online)

<https://www.karger.com/OFA>

Obesity Facts

Disclaimer:

Accepted, unedited article not yet assigned to an issue. The statements, opinions and data contained in this publication are solely those of the individual authors and contributors and not of the publisher and the editor(s). The publisher and the editor(s) disclaim responsibility for any injury to persons or property resulting from any ideas, methods, instructions or products referred to the content.

Copyright:

This article is licensed under the Creative Commons Attribution-NonCommercial 4.0 International License (CC BY-NC) (<http://www.karger.com/Services/OpenAccessLicense>). Usage and distribution for commercial purposes requires written permission.

© 2023 The Author(s). Published by S. Karger AG, Basel

Research Article

How could different obesity scenarios alter the burden of type 2 diabetes and liver disease in Saudi Arabia?

Timothy Coker^a, Jennifer Saxton^a, Lise Retat^a, John Guzek^a, Joshua Card-Gowers^a, Nasser F. BinDhim^{b,c,d}, Nora A. Althumiri^b, Khalid Aldubayan^e, Habeeb I. A. Razack^f, Laura Webber^a, Saleh A. Alqahtani^{g,h*}

^a HealthLumen, 35 Ballards Lane, London, United Kingdom

^b Sharik Association for Health Research, Riyadh, Saudi Arabia

^c College of Medicine, Alfaisal University, Riyadh, Saudi Arabia

^d Saudi Food and Drug Authority, Riyadh, Saudi Arabia

^e Department of Community Health Sciences, College of Applied Medical Sciences, King Saud University, Riyadh, Saudi Arabia

^f College of Medicine, King Saud University, Riyadh, Saudi Arabia

^g Liver Transplant Centre, King Faisal Specialist Hospital & Research Centre, Riyadh, Saudi Arabia

^h Division of Gastroenterology & Hepatology, Johns Hopkins University, Baltimore, MD, United States of America

Short Title: Obesity policy and non-communicable diseases in Saudi Arabia

*Corresponding Author:

Dr. Saleh A. Alqahtani, MD

Liver Transplant Centre,

King Faisal Specialist Hospital & Research Centre,

Riyadh, Saudi Arabia

Tel: +966 539393286

E-mail: salalqahtani@kfshrc.edu.sa

Number of Tables: 0

Number of Figures: 2

Number of Online Supplementary Material: 1 (containing Tables S1-S5, Figure S1 and Detailed Methodology)

Word count: 286 (abstract), 2999 (full text)

Accepted Manuscript

Abstract

Introduction

Obesity is a major risk factor for type 2 diabetes (T2DM) and liver disease, and obesity-attributable liver disease is a common indication for liver transplant. Obesity prevalence in Saudi Arabia (SA) has increased in recent decades. SA has committed to the WHO 'halt obesity' target to shift prevalence to 2010 levels by 2025. We estimated the future benefits of reducing obesity in SA on incidence and costs of T2DM and liver disease under two policy scenarios: 1) SA meets the 'halt obesity' target; 2) population body mass index (BMI) is reduced by 1% annually from 2020 to 2040.

Methods

We developed a dynamic microsimulation of working-age people (20-59 years) in SA between 2010 and 2040. Model inputs included population demographic, disease and healthcare cost data, and relative risks of diseases associated with obesity. In our two policy scenarios, we manipulated population BMI and compared predicted disease incidence and associated healthcare costs to a baseline 'no change' scenario.

Results

Adults <35 years are expected to meet the 'halt obesity' target, but those ≥ 35 years are not. Obesity is set to decline for females, but to increase amongst males 35-59 years. If SA's working-age population achieved either scenario, >1.15 million combined cases of T2DM, liver disease and liver cancer could be avoided by 2040. Healthcare cost savings for the 'halt obesity' and 1% reduction scenarios are 46.7 and 32.8 billion USD, respectively.

Discussion/Conclusion

SA's younger working-age population is set to meet the 'halt obesity' target, but those aged 35-59 are off-track. Even a modest annual 1% BMI reduction could result in substantial future health and economic benefits. Our findings strongly support universal initiatives to reduce population-level obesity, with targeted initiatives for working-age people ≥ 35 years of age.

Keywords: Obesity, type 2 diabetes, liver disease, liver cancer, dynamic microsimulation modelling, non-communicable diseases

Introduction

In 2013, The World Health Organization (WHO) set an ambitious global target for countries to ‘halt’ obesity at 2010 levels by 2025, through measures such as fiscal policies to reduce consumption of sweetened beverages, supporting breastfeeding, and promoting healthier food in schools and other public institutions [1]. This target arose from a broader agreement to reduce by one-quarter the number of premature deaths from major non-communicable diseases (NCDs) – cardiovascular disease, diabetes and cancer – by 2025, for which obesity is a key risk factor [2]. A 2020 report from the World Obesity Federation is critical of global efforts towards the ‘halt obesity’ target [3]. Their projections suggest most countries have a less than 10% chance of meeting their goal, and they have put out a call to action to redress shortcomings in country responses to the obesity epidemic [3].

Developing a clearer understanding of the health and economic benefits of reaching the 2025 obesity target could renew political will for improved prevention and treatment in advance of the deadline. Beyond 2025, policy-makers could benefit from a snapshot of the longer-term future health of the population and the likely demands on the health system resulting from obesity. From a policy and planning perspective, it is useful for these snapshots to include the potential impact of particular policies on obesity, obesity-attributable diseases and associated healthcare costs [4].

Mathematical modelling is one approach for projecting future epidemiological and cost-related health outcomes in a population, which can be modified according to different policy scenarios [5]. Dynamic microsimulation models simulate the behaviour of hypothetical individuals over time [6], and are described by the Organisation for Economic Co-operation and Development (OECD) as the most appropriate method for risk factor and chronic disease modelling [7]. Key strengths include 1) large sample sizes of hypothetical individuals, replicating real, heterogeneous populations; 2) flexibility to account for the influence of pre-existing risk factors on an individual’s disease status, which may progress and regress in line with a disease’s natural history; 3) the ability to introduce hypothetical interventions to assess their potential influence on population-level outcomes; and 4) scope to estimate economic costs of diseases and potential cost savings associated with different policies [8-10]. Findings from microsimulation models are widely used to inform policy decisions, particularly in relation to budget allocation and interventions [8], a notable example being Minimum Unit Pricing in Scotland in 2018 [11].

Saudi Arabia (SA) is a co-signatory on the WHO ‘halt obesity’ pledge and is a pertinent case-study country for quantifying the potential benefits of different obesity policies. SA national estimates show that 24.1% of men and 33.5% of women were obese in 2013 [12], much higher than the 2016 global average of 11% for men and 15% for women [13]. SA has correspondingly high and increasing levels of type 2 diabetes (T2DM) [14], for which obesity increases the risk more than seven-fold [15]. Obesity is also implicated in the rising prevalence of non-alcoholic fatty liver disease (NAFLD), which affects more than 30% of the Middle East population [16] and could affect 48% of SA’s population by 2030 [17]. A more serious NAFLD variant – non-alcoholic steatohepatitis (NASH) – is linked to advanced liver fibrosis, cirrhosis and liver cancer [18]. NASH has overtaken hepatitis C virus as the most common indication for liver transplants in SA [19]. The health and economic burden of T2DM, liver disease and liver cancer could be dramatically reduced with effective obesity policies and lifestyle interventions [20-22].

We used a well validated dynamic microsimulation model [22-27] to quantify the future health and economic benefits of two obesity policy scenarios compared to a baseline ‘no change’ scenario in SA from 2010 to 2040:

- Scenario 1: attainment of the ‘halt obesity’ target (i.e. no movement of individuals into the obese category between 2010 and 2025)
- Scenario 2: a reduction in population-level BMI by 1%, annually from the year 2020 to 2040

We estimated cumulative incidence avoided and direct healthcare costs avoided for obesity-attributable T2DM, liver disease and liver cancer between 2010 and 2040.

Materials and Methods

Dynamic microsimulation model

Our model included a hypothetical population of 100 million individuals, developed from the most recent SA age and sex population distribution available from the UN [28]. We implemented the model from 2010 to enable assessment of the ‘halt obesity’ scenario, which relates to the period 2010 to 2025. We modelled outcomes to 2040. Our model has been previously used to predict UK obesity trends and other NCDs [22, 25], and further adapted and implemented across 70 countries, including Saudi Arabia [24] and the wider Middle East [26]. The method has also previously been

employed to quantify the impact of different policy scenarios on burden and direct healthcare costs of related diseases [29]. The full method is described elsewhere [21] and in the Online Supplementary Material.

Focused literature reviews to source model input data

We searched for prevalence estimates for different categories of BMI (our risk factor) by age and sex. Incidence, prevalence, and mortality estimates for T2DM, liver disease, and liver cancer (our disease outcomes) were also collected, as well as papers reporting relative risks linking BMI categories to our diseases. Our BMI categories were: healthy weight (BMI < 25.00 kg/m²), pre-obese (25.00-29.99 kg/m²), and obese (≥ 30.00 kg/m²), following WHO definitions [30]. We also searched for direct healthcare cost estimates for our focal diseases.

Inclusion/exclusion criteria

We included data for all adults in SA, but restricted our reporting to people of working age (20-59 years) given their greater influence on the economy. We excluded data older than 15 years to ensure our predictions were not unduly influenced by historical data no longer reflective of SA's population characteristics. We prioritized data firstly from SA, then the Middle East region, then Asia, and finally to other countries outside of Asia until we found suitable data sources.

Final data sources

Cross-sectional BMI data were drawn from multiple sources and years (see Table S1, Online Supplementary Material). Data on T2DM in SA were drawn from the 2017 Global Burden of Disease study [31], with relative risks drawn from a global study, reporting estimates for Asian males and females [32]. Liver disease data were drawn from the Global Burden of Disease study [31], which provides incidence and mortality estimates from 2016, for which 'Cirrhosis and other chronic liver diseases' was the most appropriate category and could be linked to the limited available relative risk data [33]. Liver cancer incidence was drawn from the 2014 Saudi Cancer Registry [34], and mortality data were from the 2018 registry [35]. Relative risks were drawn from a 2012 global meta-analysis, which included an estimate for Asian populations [36].

Due to limited cost data availability for Saudi Arabia, for all diseases, we used cost data from the United Kingdom [37, 38], converted to US dollars (2010 value) using a purchasing power parity approach [39] (Table S2, Online Supplementary Material).

Implementing policy scenarios

The simulated population was exposed to three scenarios: 1) no additional policy (baseline); 2) obesity halted at 2010 levels: no-one in the population could move from the healthy-weight or pre-obese groups into the obese group between 2010 and 2025; 3) an annual 1% reduction in population-level BMI was implemented from the year 2020 until 2040.

Microsimulation outputs

Microsimulation outputs were generated for ~100 million hypothetical individuals representing the entire SA population. Here, we report only results for the working age (20-59 years) population, representing ~60 million trials.

Results

Confidence limits (+/-) for all results refer to Monte Carlo error of trials in the microsimulation.

Estimated obesity prevalence under current trends (baseline scenario 2020-2040):

Obesity prevalence estimates for the period 2020-2040 are presented in Table S3 (Online Supplementary Material) by five-year age groups and sex. Our model suggests strong sex and generational effects on obesity. This modeling approach estimated that 33.7% of all working-age males (20-59 years) were obese in 2020, with a maximum obesity prevalence of 34.8% is predicted in 2030 and declining to 33.3% in 2040. Trends for the youngest males (20-34 years) are optimistic, showing a steady decline from 19.8% in 2020 to 15.5% in 2040. Obesity trends for 35-59 years-old males are more concerning, predicted to increase from 43.7% in 2020 to 56.3% in 2040 in the 35-44 age group, and from 45.5% to 59.9% in the 45-54 age group. Obesity amongst the oldest males (55-59 years) is expected to increase from 38.9% to 46.4%.

For working-age females, although declines in obesity prevalence are predicted, the prevalence is still high: 34.5% in 2020 to 27.7% in 2040. The magnitude of decline varies considerably by age group, as does the starting prevalence. Females < 35 years of age have the lowest starting and finishing prevalence: 20.0% in 2020 to 10.7% in 2040. In contrast, nearly 50% of females 35-54 years were estimated to be obese in 2020, dropping to 40% by 2040. Females

55-60 years have the highest starting obesity prevalence (54.3% in 2020) and the smallest reduction over time (50.2% in 2040).

Estimated cumulative incidence avoided for T2DM, liver disease and liver cancer (2010-2040), by obesity policy scenario relative to baseline

Cumulative incidence avoided, for each disease, for each year of the microsimulation are shown in Figure 1, for both scenarios compared to baseline. T2DM was predicted to show the greatest reduction of all diseases between 2010 and 2040, at 994,453 [\pm 2,569] for scenario 1 (halt obesity) and 1,019,715 [\pm 2,560] for scenario 2 (1% BMI reduction). For liver disease, 156,095 [\pm 1,165] cases could be avoided in scenario 1, and 147,041 [\pm 1,170] in scenario 2, by 2040. Reductions for liver cancer were predicted as 324 [\pm 172] and 534 [\pm 171] for scenarios 1 and 2, respectively, by 2040. In total, this represents a reduction of 1,150,871 [\pm 2,826] disease cases in scenario 1, and 1,167,290 [\pm 2,820] cases in scenario 2 (see Table S4 [Online Supplementary Material] for full results by scenario, sex and age-group). In both scenarios, the majority of incidence avoided was in males (~68-73% of cases avoided). By age, the greatest reductions were seen in adults aged 35-49 (~51-63% of cases avoided).

Estimated direct healthcare costs avoided for T2DM, liver disease and liver cancer (2010-2040), by obesity policy scenario

Cumulative healthcare costs avoided for each disease, through each year of the microsimulation, are shown in Figure 2, for both scenarios compared to baseline. Figure S1 (Online Supplementary Material) shows the schematic of risk factor model and microsimulation model. Costs avoided are cumulative from 2010. The greatest savings were predicted for T2DM, at 31,606,477,828 [\pm 32,682,902] USD for scenario 1 (halt obesity), and 22,245,802,810 [\pm 33,149,239] USD for scenario 2 (1% BMI reduction), by 2040. Costs avoided for liver disease were 15,124,470,064 [\pm 56,156,104] USD and 10,557,215,034 [\pm 56,655,461] USD for scenarios 1 and 2, respectively. Costs avoided for liver cancer were 7,805,755 [\pm 4,037,343] and 11,462,574 [\pm 4,028,680] respectively. In total, costs avoided for the three diseases were predicted to be 46,738,753,648 [\pm 65,099,771] USD for scenario 1, and 32,814,480,419 [\pm 65,764,303] USD for scenario 2 (see Table S5 [Online Supplementary Material] for full results by scenario, sex and age group).

Discussion/Conclusion

Future trends in obesity prevalence

Our results show a mixed picture of future obesity trends if there is no change in current policy, with large variation by age and sex. The 40.5% 'halt obesity' target for women [1] is set to be reached overall, but not by women 35-59 years who will remain considerably off-target. Whilst it is encouraging that obesity amongst working-age women is set to decline over time, for women 35-59 years, prevalence starts high and remains high in 2040. Men overall are not set to meet the 'halt obesity' target of 27.2% [3], though men <35 years may reach it. Men 35-59 years are further away from the 2025 target than women, with increasing obesity predicted over time. Almost 60% of men 45-54 years could be obese by 2040.

One other study has modelled SA obesity trends in relation to the halt obesity target [3]. The World Obesity Federation (WOF) report estimates that neither males nor females are on track to meet the target. The study authors used ordinary least squares (OLS) regression to derive 2025 obesity estimates of 40.0% for adult males and 49.1% for adult females in 2025. Our results are more optimistic for women overall, and more pessimistic for men, though our estimates overlap for women 35-54 years of age. There are several reasons why our results differ, including our focus on the working-age population (rather than all adults), and methodological differences (e.g. the use of a dynamic microsimulation with the flexibility to move between granular states versus a deterministic approach). Another key difference is that we excluded BMI data sources older than 15 years, and the WOF report included data from the 1980s, which could affect the gradient. Our results complement the WOF report findings because they provide additional information about obesity by age group, highlighting which segments of the population are off track.

Quantifying future benefits of achieving obesity scenarios

Our findings suggest that reducing obesity would be particularly beneficial for preventing T2DM, for which around a million cases would be avoided for both scenarios. The management of T2DM is extremely costly, and the achievement of either scenario represents T2DM cost savings of over 22.2 billion USD. For liver disease, each scenario could avoid around 150,000 cases by 2040 – amounting to savings of over 10.5 billion USD. Liver cancer, despite being the least common of the diseases evaluated, would also lead to substantial cost savings of around 7.8 to 11.5 million

USD. We are not aware of other studies quantifying the potential benefits of meeting the 'halt obesity' target, or other policy scenarios (such as achieving an annual 1% reduction in BMI).

Implications for SA policy

The need for targeted weight reduction interventions

Our results strongly suggest the need for targeted weight reduction interventions, particularly for people 35 years and older. For women in this age group, obesity trends are decreasing, but not rapidly, so interventions that hasten the reduction would be worthwhile. For men, of concern is the increasing obesity trend predicted for those 35 years and older. Interventions will be needed to address the factors underlying this increase. A number of effective obesity prevention and treatment interventions exist, which can have meaningful individual- and population-level impacts [20-22].

Include working-age people ≥ 35 years as a target group in SA's obesity strategy

SA's 2019 obesity strategy explicitly refers to the commitment to reduce key NCDs by one-quarter by 2025 [40]. The strategy is being delivered through a framework that combines a life-course approach with a socio-ecological outlook to recognize the multi-factorial nature of obesity. However, the target groups for obesity monitoring do not explicitly include middle- or older-aged people; and instead, focus mostly on those engaged with antenatal services. This omission could work against the overarching life-course approach upon which the strategy is based. Our data show that people 35-59 years of age are at high risk of developing obesity and obesity-attributable diseases, many of whom would not be captured by antenatal services. Targeted interventions to these groups, and the use of general population surveys to monitor obesity and obesity-related diseases will be paramount to achieving NCD targets, preventing harm and reducing healthcare costs attributable to obesity.

Saudi Vision 2030

SA is planning to restructure the country's vital service sectors through ambitious Saudi Vision 2030 objectives. The Health Sector Transformation Program is one of the eight themes that Saudi Vision 2030 covers [41] and aims to facilitate access to healthcare services, improve the quality and efficiency of health services, and promote prevention of health risks [41, 42]. Community-based health programs, healthy food and eating behavior promotion, and multilevel health education initiatives are expected to be a vital part of this program. In particular, one of the current initiative strategic pillars focuses on reducing the prevalence of risk factors (including obesity and diabetes) for NCDs. This vision is supported through various reform initiatives, which include but are not limited to health sector reform, the introduction of a modern care model (i.e., change in the design of care delivery), private sector participation and investment, and e-health initiatives to improve access to care and care management; and is further supported by the new agreements signed with multinational pharmaceutical companies to commence medicine manufacturing within SA and reinforcing medical research [43]. A recent Gulf Cooperation Council report mentions that the SA's health investment will increase to 66.67 billion USD (cumulative) by 2030 [44]. These healthcare reforms provide promise to address the current public health challenges in SA, notably to support reducing obesity prevalence in the Kingdom [45].

Strengths, limitations & future work

We have developed a novel SA microsimulation study designed to inform policy decisions to promote health in SA. Our study provides timely obesity estimates in the lead up to the 2025 'halt obesity' deadline. Our findings highlight the urgency of addressing obesity, particularly amongst middle-to-older age working people. We demonstrate that a relatively small 1% annual reduction in BMI could bring substantial health and economic benefits. Though we indicate optimistic projections for younger groups, changing environmental factors, such as COVID-19 restrictions that can affect physical activity and diet, will need monitoring across age groups [46, 47].

Our study was limited by the availability of data, and we used data from outside of SA for certain microsimulation modules. We identified a need for more up-to-date, national and regionally representative data on obesity incidence and prevalence, as well as specific liver diseases such as NAFLD, which are on the rise in SA [19]. More frequent, high-quality data will enable more detailed modelling of the associations between obesity and NCDs in SA, and more accurate projections of the economic and health burden from NCDs. We also restricted our reporting to people of working age, who have the greatest economic influence, but this meant we could not see the true extent of obesity's influence on slower progressing diseases such as NAFLD/NASH-related cirrhosis and liver cancer, more commonly seen in older age [48, 49].

Acknowledgement

None

Statement of Ethics

Our study used open-access secondary data and did not require ethical approval.

Conflict of Interest Statement

The authors have no conflicts of interest to declare.

Funding Sources

This research received no specific grant from any funding agency in the public, commercial, or not-for-profit sectors.

Data Availability Statement

The data supporting the findings of this study are taken from publicly available sources (as mentioned in the Methods section and Online Supplementary Material). Further enquiries can be directed to the corresponding author.

Author Contributions

SA conceived the study, with support from LW. JS wrote the manuscript, with redrafts supported by LW, LR and TC. JG, JC-G and HI conducted literature reviews and sourced data for the microsimulation. TC, LR and LW adapted the model and ran the microsimulation. JS, TC and LW interpreted the data. SA, NB, NA, KA and HI reviewed and substantially revised the manuscript content. All authors reviewed and approved the final version of the manuscript.

References

1. World Health Organization. *Noncommunicable diseases and mental health: Target 7: Halt the rise in obesity*. 2013 [cited 2021 21 June]; Available from: <https://www.who.int/news/item/19-01-2015-noncommunicable-diseases-prematurely-take-16-million-lives-annually-who-urges-more-action>.
2. Sixty-Sixth World Health Assembly. *Follow-up to the Political Declaration of the High-level Meeting of the General Assembly on the Prevention and Control of Non-communicable Diseases*. 2013 [cited 2021 30 June]; Available from: https://apps.who.int/gb/ebwha/pdf_files/WHA66/A66_R10-en.pdf?ua=1.
3. World Obesity Federation. *Obesity: missing the 2025 global targets*. . 2020 [cited 30 June 2021]; Available from: <https://www.worldobesity.org/resources/resource-library/world-obesity-day-missing-the-targets-report>.
4. Xue, H., et al., *Applications of systems modelling in obesity research*. *Obes Rev*, 2018. **19**(9): p. 1293-1308.
5. Kretzschmar, M., *Disease modeling for public health: added value, challenges, and institutional constraints*. *J Public Health Policy*, 2020. **41**(1): p. 39-51.
6. Li, J. and C. O'Donoghue, *A survey of dynamic microsimulation models: uses, model structure and methodology*. *International Journal of Microsimulation*, 2013. **6**(2): p. 3-55.
7. Oderkirk J, S.F., Cecchini M, Astolfi R; OECD Health Division. *Toward a New Comprehensive International Health and Health Care Policy Decision Support Tool*. *OECD Directorate of Employment, Labour and Social Affairs*. 2012 [cited 2021 30 June]; Available from: <https://www.oecd.org/els/health-systems/TowardANewComprehensiveInternationalHealthAndHealthCarePolicyDecisionSupportTool.pdf>
8. Rutter, C.M., A.M. Zaslavsky, and E.J. Feuer, *Dynamic microsimulation models for health outcomes: a review*. *Med Decis Making*, 2011. **31**(1): p. 10-8.
9. Zucchelli, E., A. Jones, and N. Rice, *The evaluation of health policies through dynamic microsimulation methods*. *International Journal of Microsimulation*, 2012. **5**: p. 2-20.
10. Odonoghue, C. and G. Dekkers, *Increasing the Impact of Dynamic Microsimulation Modelling*. *International Journal of Microsimulation*, 2018. **11**: p. 61-96.
11. The Scottish Government. *Alcohol and drugs*. [cited 2020 25 November]; Available from: <https://www.gov.scot/policies/alcohol-and-drugs/minimum-unit-pricing/>
12. Memish, Z.A., et al., *Obesity and associated factors--Kingdom of Saudi Arabia, 2013*. *Prev Chronic Dis*, 2014. **11**: p. E174.
13. World Health Organization. *Global Health Observatory (GHO) data*. 2016 [cited 2021 30 June]; Available from: <https://apps.who.int/gho/data/node.main.A896?lang=en>
14. Alotaibi, A., et al., *Incidence and prevalence rates of diabetes mellitus in Saudi Arabia: An overview*. *J Epidemiol Glob Health*, 2017. **7**(4): p. 211-218.
15. Abdullah, A., et al., *The magnitude of association between overweight and obesity and the risk of diabetes: a meta-analysis of prospective cohort studies*. *Diabetes Res Clin Pract*, 2010. **89**(3): p. 309-19.
16. Younossi, Z.M., et al., *Global epidemiology of nonalcoholic fatty liver disease-Meta-analytic assessment of prevalence, incidence, and outcomes*. *Hepatology*, 2016. **64**(1): p. 73-84.
17. Alswat, K., et al., *Nonalcoholic fatty liver disease burden - Saudi Arabia and United Arab Emirates, 2017-2030*. *Saudi J Gastroenterol*, 2018. **24**(4): p. 211-219.
18. Alqahtani, S.A., et al., *Saudi Association for the Study of Liver diseases and Transplantation practice guidelines on the diagnosis and management of hepatocellular carcinoma*. *Saudi J Gastroenterol*, 2020. **26**(Suppl 1): p. S1-s40.
19. Alqahtani, S.A., et al., *Changing trends in liver transplantation indications in Saudi Arabia: from hepatitis C virus infection to nonalcoholic fatty liver disease*. *BMC Gastroenterol*, 2021. **21**(1): p. 245.
20. Arena, R., et al., *Healthy lifestyle interventions to combat noncommunicable disease-a novel nonhierarchical connectivity model for key stakeholders: a policy statement from the American Heart Association, European Society of Cardiology, European Association for Cardiovascular Prevention and Rehabilitation, and American College of Preventive Medicine*. *Eur Heart J*, 2015. **36**(31): p. 2097-2109.
21. Webber, L., et al., *The future burden of obesity-related diseases in the 53 WHO European-Region countries and the impact of effective interventions: a modelling study*. *BMJ Open*, 2014. **4**(7): p. e004787.

22. Retat, L., et al., *Screening and brief intervention for obesity in primary care: cost-effectiveness analysis in the BWeL trial*. *Int J Obes (Lond)*, 2019. **43**(10): p. 2066-2075.
23. Pineda, E., et al., *Forecasting Future Trends in Obesity across Europe: The Value of Improving Surveillance*. *Obesity Facts*, 2018. **11**(5): p. 360-371.
24. Coker, T., et al., *The future health and economic burden of obesity-attributable type 2 diabetes and liver disease among the working-age population in Saudi Arabia*. *PLoS One*, 2022. **17**(7): p. e0271108.
25. Wang, Y.C., et al., *Health and economic burden of the projected obesity trends in the USA and the UK*. *Lancet*, 2011. **378**(9793): p. 815-25.
26. Kilpi, F., et al., *Alarming predictions for obesity and non-communicable diseases in the Middle East*. *Public Health Nutr*, 2014. **17**(5): p. 1078-86.
27. Pimpin, L., et al., *Estimating the costs of air pollution to the National Health Service and social care: An assessment and forecast up to 2035*. *PLOS Medicine*, 2018. **15**(7): p. e1002602.
28. United Nations. *World Population Prospects 2019* [cited 2021 6 Junr]; Available from: <https://population.un.org/wpp/>
29. Knuchel-Takano, A., et al., *Modelling the implications of reducing smoking prevalence: the benefits of increasing the UK tobacco duty escalator to public health and economic outcomes*. *Tob Control*, 2018. **27**(e2): p. e124-e129.
30. World Health Organization. *Obesity and overweight*. 2021 [cited 2021 30 June]; Available from: <https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight>
31. Institute for Health Metrics and Evaluation. *Global Burden of Disease Project*. 2017 [cited 2021 6 June]; Available from: <http://ghdx.healthdata.org/>
32. Vazquez, G., et al., *Comparison of body mass index, waist circumference, and waist/hip ratio in predicting incident diabetes: a meta-analysis*. *Epidemiol Rev*, 2007. **29**: p. 115-28.
33. Harris, R., et al., *Obesity Is the Most Common Risk Factor for Chronic Liver Disease: Results From a Risk Stratification Pathway Using Transient Elastography*. *Am J Gastroenterol*, 2019. **114**(11): p. 1744-1752.
34. International Association of Cancer Registries. *Saudi Cancer Registry*. 2018 [cited 2021 6 June]; Available from: http://www.iacr.com.fr/index.php?option=com_comprofiler&task=userprofile&user=1215&Itemid=498
35. Global Cancer Observatory. *No Title*. 2018 [cited 2020 1 September]; Available from: <https://gco.iarc.fr/>
36. Chen, Y., et al., *Excess body weight and the risk of primary liver cancer: an updated meta-analysis of prospective studies*. *Eur J Cancer*, 2012. **48**(14): p. 2137-45.
37. Public Health England. *The health and social care costs of a selection of health conditions and multimorbidities*. 2020 [cited 2021 30 June]; Available from: <https://www.gov.uk/government/publications/health-and-social-care-costs-of-a-selection-of-health-conditions>
38. Georghiou, T., et al. *Understanding patterns of health and social care at the end of life*. 2021 [cited 2021 30 June]; Available from: <https://www.nuffieldtrust.org.uk/research/understanding-patterns-of-health-and-social-care-at-the-end-of-life>
39. CCEMG - EPPI-Centre Cost Converter. 2019 [cited 2021 3 June]; Available from: <https://eppi.ioe.ac.uk/costconversion/default.aspx>
40. Saudi Center for Disease Prevention and Control, *Obesity Prevention & Control Strategy 2020-2030*. 2019. p. 1-34.
41. *Saudi Vision 2030: Health Sector Transformation Program*. [cited 2021 22 September]; Available from: <https://www.vision2030.gov.sa/v2030/vrps/hstp/>.
42. News, A. *Saudi Arabia signs deals with AstraZeneca, Pfizer for local vaccines production, medical research*. 2021 [cited 2021 22 September]; Available from: <https://www.arabnews.com/node/1928396/business-economy>.
43. Chowdhury, S., D. Mok, and L. Leenen, *Transformation of health care and the new model of care in Saudi Arabia: Kingdom's Vision 2030*. *J Med Life*, 2021. **14**(3): p. 347-354.
44. News, A. *Saudi Arabia to invest \$66 billion in healthcare infrastructure by 2030*. 2021 [cited 2021 22 September]; Available from: <https://www.arabnews.com/node/1837271/business-economy>.
45. Althumiri, N.A., et al., *Obesity in Saudi Arabia in 2020: Prevalence, Distribution, and Its Current Association with Various Health Conditions*. *Healthcare (Basel)*, 2021. **9**(3).

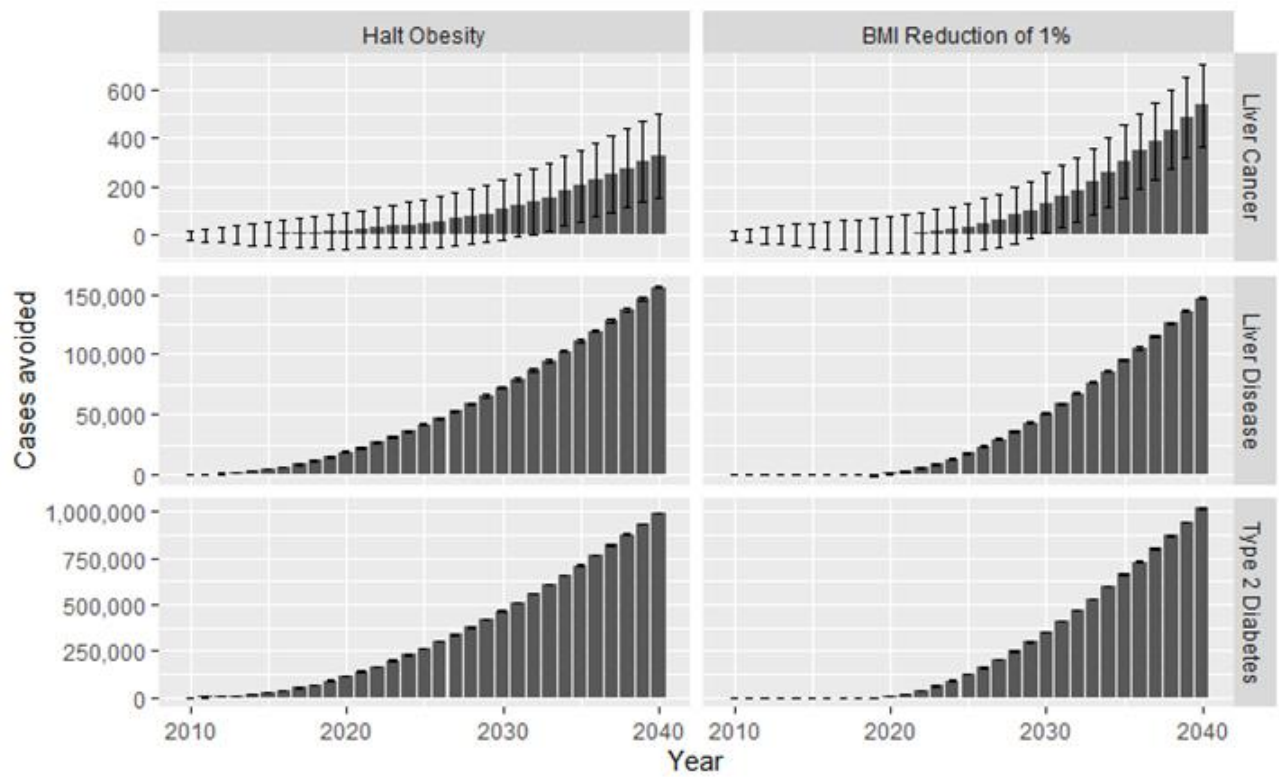
46. Flanagan, E.W., et al., *The Impact of COVID-19 Stay-At-Home Orders on Health Behaviors in Adults*. Obesity (Silver Spring), 2021. **29**(2): p. 438-445.
47. Ammar, A., et al., *Effects of COVID-19 Home Confinement on Eating Behaviour and Physical Activity: Results of the ECLB-COVID19 International Online Survey*. Nutrients, 2020. **12**(6).
48. Alexander, M., et al., *Real-world data reveal a diagnostic gap in non-alcoholic fatty liver disease*. BMC Med, 2018. **16**(1): p. 130.
49. Abdo, A.A., et al., *Saudi guidelines for the diagnosis and management of hepatocellular carcinoma: technical review and practice guidelines*. Ann Saudi Med, 2012. **32**(2): p. 174-99.

Accepted Manuscript

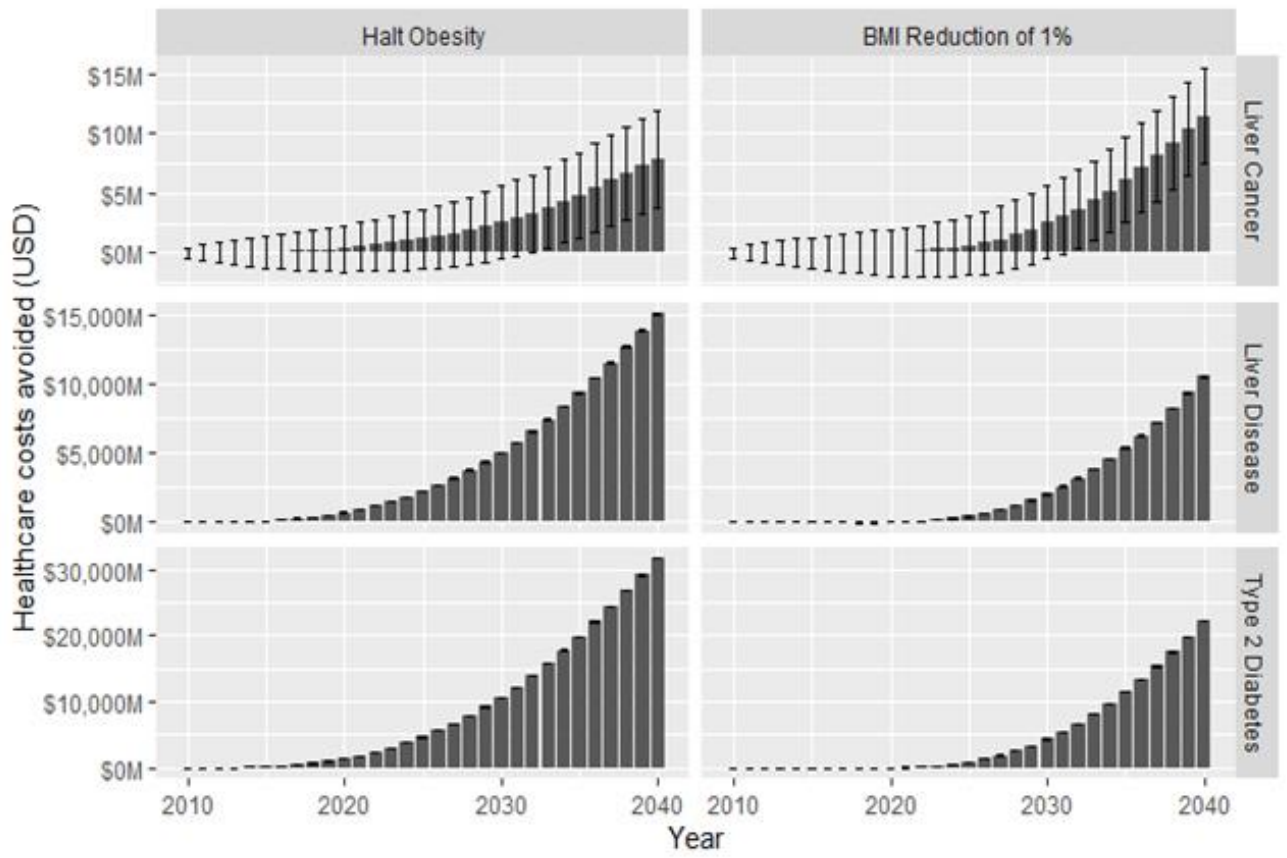
Fig. 1. Cumulative incidence avoided for type 2 diabetes, liver disease and liver cancer, for each obesity policy scenario compared to baseline, amongst people 20-59 years, 2010-2040.

Fig. 2. Cumulative direct healthcare costs avoided (USD) for type 2 diabetes, liver disease and liver cancer, for each obesity policy scenario compared to baseline amongst people 20-59 years, 2010-2040.

Accepted Manuscript



Accepted Manuscript



Accepted