# Disability and costs of ischemic heart disease attributable to the consumption of trans fatty acids in Brazil

Magda do Carmo Parajára<sup>1,\*</sup>, Aline Siqueira Fogal Vegi<sup>1</sup>, Ísis Eloah Machado<sup>1,2</sup>, Mariana Carvalho de Menezes<sup>1,3</sup>, Eliseu Verly-Jr<sup>4</sup> and Adriana Lúcia Meireles<sup>1,3</sup>

<sup>1</sup>Programa de Pós-Graduação em Saúde e Nutrição, Escola de Nutrição, Universidade Federal de Ouro Preto, Ouro Preto, Minas Gerais, Brazil:

<sup>2</sup>Departamento de Medicina de Família, Saúde Mental e Coletiva, Escola de Medicina, Universidade Federal de Ouro Preto, Ouro Preto, Minas Gerais, Brazil:

<sup>3</sup>Departamento de Nutrição Clínica e Social, Escola de Nutrição, Universidade Federal de Ouro Preto, Ouro Preto, Minas Gerais, Brazil:

<sup>4</sup>Departamento de Epidemiologia, Instituto de Medicina Social, Universidade do Estado do Rio de Janeiro, Rio de Janeiro, Rio de Janeiro, Brazil

\*Corresponding author: Magda do Carmo Parajára, Programa de Pós-Graduação em Saúde e Nutrição, Escola de Nutrição, Universidade Federal de Ouro Preto, Rua Dois, 607, Campus Morro do Cruzeiro, Bauxita, Ouro Preto, Minas Gerais, Brazil. CEP: 35400-000; magda.parajara@ufop.ufop.edu.br; +55 (31) 3559-1838.

Short title: Disability and costs of trans-fatty acids



This is an Accepted Manuscript for Public Health Nutrition. This peer-reviewed article has been accepted for publication but not yet copyedited or typeset, and so may be subject to change during the production process. The article is considered published and may be cited using its DOI 10.1017/S1368980024001101

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[Disclosure statements, as outlined below. These must be included on the title page and not in the manuscript file, to enable double-blind reviewing; if the paper is accepted, they will be inserted into the manuscript during production. If any are not applicable, please state this.]

**Acknowledgements:** We are grateful to the Universidade Federal de Ouro Preto (UFOP) for their support and to the Grupo de Pesquisa e Ensino em Nutrição e Saúde Coletiva (GPENSC) for their invaluable encouragement.

Financial Support: This study was supported by the Conselho Nacional de Desenvolvimento Científico e Tecnológico (CNPq/Brazil) (Í.E.M., grant number 442636/2019-9); received a research fellowship from the Fundação de Amparo à Pesquisa do Estado de Minas Gerais (FAPEMIG/Brazil) (M.C.P.); received a Research Productivity fellowship from the CNPq (Í.E.M.); was financed in part by the Coordenação de Aperfeiçoamento de Pessoal de Nível Superior - Brazil (CAPES); and used data from the Institute for Health Metrics and Evaluation (IHME), funded by Bill & Melinda Gates Foundation. None of the funders had a role in this article's design, analysis, or writing.

#### **Conflict of Interest: None.**

**Authorship:** M.C.P. Conceptualization, methodology, formal analysis, writing – original draft. A.S.F.V. Methodology, writing – review and editing. Í.E.M. and M.C.M. Methodology, writing – review and editing, supervision, project administration, funding acquisition. E.V.J. Writing – review and editing. A.L.M. Conceptualization, methodology, writing – review and editing, supervision, project administration, funding acquisition. All authors contributed to data interpretation and approved the final version of the submitted manuscript, accepting full responsibility for all aspects of this work.

**Ethical Standards Disclosure:** This study was exempted from approval by the Research Ethics Committee at the Universidade Federal de Ouro Preto (UFOP) in Brazil since was explored secondary data which does not allow identification of the individual level (CAAE no. 33396720.7.0000.5150).

**Abstract** 

**Objective:** To estimate the disability and costs of the Brazilian Unified Health System (SUS)

for ischemic heart disease (IHD) attributable to trans fatty acid (TFA) consumption in 2019.

Design: This ecological study used secondary data from the Global Burden of Disease

(GBD) Study 2019 to estimate the years lived with disability (YLDs) from IHD attributable

to TFA in Brazil in 2019. Data on direct costs (purchasing power parity: 1 Int\$ = R\$ 2.280)

were obtained from the Hospital and Ambulatory Information Systems of the SUS. Moreover,

the total costs in each state were divided by the resident population in 2019 and multiplied by

10,000 inhabitants. The relationship between the socio-demographic index (SDI), disease,

and economic burden was investigated.

**Setting:** Brazil and its 27 states.

**Participants:** Adults aged  $\geq 25$  years of both sexes.

Results: IHD attributable to TFA consumption resulted in 11,165 YLDs (95% uncertainty

interval [UI]: 932-18,462) in 2019 in Brazil. A total of Int\$ 54,546,227 (95% UI: 4,505,792-

85,561,810) was spent in the SUS in 2019 due to IHD attributable to TFA, with the highest

costs of hospitalizations, for males and individuals aged  $\geq 50$  years or over. The highest costs

were observed in Sergipe (Int\$ 6,508/10,000; 95% UI: 576-10,265), followed by the two

states from the South. Overall, as the SDI increases, expenditures increase.

**Conclusions:** TFA consumption results in a high disease and economic IHD burden in Brazil,

reinforcing the need for more effective health policies, such as industrial TFA elimination,

following the international agenda.

Keywords: Global burden of disease; Noncommunicable diseases; Trans fatty acids; Health

care costs; Costs and cost analysis; Cost savings.

#### Introduction

Over the last three decades, ischemic heart disease (IHD) has consistently been ranked as one of the leading causes of death and loss of health in Brazil. Among noncommunicable diseases (NCDs), cardiovascular diseases (CVDs) are recognized as a significant global public health issue because they cause premature mortality, have poor survival, impact health and quality of life, reduce workforce productivity, threaten economic prosperity due to enhanced healthcare costs, and create enormous disparities in opportunity. In the face of demographic, epidemiological, and nutritional transition, there has been an increase in life expectancy and NCDs (including CVDs), and consequently, in the impact on morbidity and disabilities caused by them. It is perceived that, in isolation, mortality measures would not be most adequate to describe the overall population health status since NCDs often have nonfatal impacts on health; then, individuals live with the disease and its consequences for many years. Also, the poorest and most vulnerable individuals regionally and at a sub-national level are at the highest risk of developing CVDs and are least likely to have access to detection and control, even with the rising healthcare expenditure caused by them. The CVD burden in low- and middle-income countries (LMICs) is challenging.

The huge CVD burden is attributable to environmental, metabolic, and behavioural risk factors, including unhealthy diet. For example, trans fatty acids (TFA), a preventable dietary risk factor, have a good level of evidence as a critical risk factor for CVD. TFA are defined as unsaturated fats that have at least one double bond in the trans configuration, and are divided into two groups: TFA naturally produced by ruminants (rTFA), found in meat and dairy, and industrial TFA (iTFA), found mainly in partially hydrogenated vegetable oils (PHO).

Given the alarming burden of CVD, the World Health Organization (WHO) has been implementing an action package called REPLACE, which aims to reduce and eliminate iTFA globally by 2023. Following this global agenda, the Member States of the Pan American Health Organization (PAHO), which Brazil participates, approved the "Plan of Action for the Elimination of Industrially Produced Trans-Fatty Acids 2020-2025". Consequently, in 2019, it was published in Brazil a resolution to limit TFA to 2% of total fats in all foods between July 1, 2021, and January 1, 2023, and to ban the production and use of PHO from January 2023.

In Brazil, although a temporal trend of IHD attributable to TFA from 1990 to 2019 revealed a decrease of approximately 60% in mortality and disability-adjusted life years (DALYs) age-standardized rates, this burden is still relevant to the country and more effective policies, such as TFA ban, only implemented in 2021, could have more marked effects on this burden. A modelling study showed that banning PHO could prevent or postpone approximately 10,500 deaths (95% uncertainty interval [95% UI]: 9,963–10,909) in the Brazilian population in 2018.

A systematic analysis of the health and economic burden of TFA consumption may reinforce the importance of reducing iTFA levels in Brazil and provide benchmarks for policy and decision-makers. Moreover, expenses arising from premature deaths and disabilities caused by NCDs threaten the efficiency and sustainability of health systems. To the best of our knowledge, the disability and cost of the disease attributable to TFA in the Brazilian Unified Health System (SUS) have not yet been investigated. Therefore, this study aimed to estimate the years lived with disability (YLDs) and the direct costs to the SUS from IHD attributable to TFA consumption in Brazil and its states in 2019.

# **Methods**

# Study design, sources of data, and population

This descriptive ecological study used secondary data from the Global Burden of Disease (GBD) Study 2019, publicly available at <a href="https://ghdx.healthdata.org/">https://ghdx.healthdata.org/</a> and retrieved in March 2023, to measure disease burden. GBD 2019, led by the Institute for Health Metrics and Evaluation (IHME), aims to quantify health loss in populations worldwide, ensuring comparative, detailed, and current results for evidence-based policymaking. Further details of the GBD data, methods, and results have been previously reported. (1,6)

Economic burden estimation of the Brazilian public health system (SUS) was based on publicly available information from the Department of Informatics of the Unified Health System (DATASUS), Health Ministry, at <a href="https://datasus.saude.gov.br/">https://datasus.saude.gov.br/</a>, retrieved in May 2022. DATASUS allows access to transparent information on procedures and service provider payments. Specifically, the Outpatient Information System (SIA/SUS) and Hospital Information System (SIH/SUS) databases were used to estimate costs. Both systems provide values practised in the Brazilian public health system and those transferred to health institutions that carry out health actions and services for the SUS. (14)

The disease and economic burden of IHD attributable to TFA were estimated for the Brazilian population aged  $\geq$  25 years in 2019.

# Disease burden

Disability was estimated using YLDs (obtained from GBD 2019), which are understood as the years of healthy life lost. YLDs were calculated by multiplying the prevalence of a sequela by the disability weights of diseases and injuries for that sequela. (1) Exposure, the consumption of TFA, is defined in the GBD 2019 as any intake (in % daily energy) of TFA from all sources. (6) The GBD 2019 sourced the information about TFA intake from national sales data provided by Euromonitor Passport. (6) To split the data into standard age groups, GBD defined the global age and patterns of the dietary factor using data obtained from 24-hour dietary recall (24HR). Afterwards, GBD utilized the recognized age patterns to divide the sales data into standard age categories. (6,15) For continuous data not originating from the 24HR, considered the gold standard by the GBD, such as TFA consumption data, various adjustments are applied to render them more consistent and suitable for modelling. (6) Additionally, the spatiotemporal Gaussian process regression (ST-GPR) framework still allows other types of information that have plausible relationships with dietary intakes, such as country-level covariates, to control and adjust for data biases. However, for TFA, the GBD 2019 did not include any adjusting covariates in the model. (6)

The attributable burden for the risk-outcome pair is measured by GBD using a comparative risk assessment, also called the population attributable fraction (PAF). The PAFs were obtained from GBD 2019 and correspond to the proportion of YLDs that could be avoided if the population achieved counterfactual exposure in the past (i.e., the theoretical minimum risk exposure level, TMREL). For TFA, TMREL represents no TFA intake. In addition to the TMREL, the PAF includes two other inputs: the average daily TFA intake and the relative risks (RRs) to the risk-outcome pair (**Supplementary Tables S1 and S2**). Then, the attributable YLDs were calculated by GBD multiplying PAFs for each age-sex-location-year by the outcome quantity, enabling stratified analyses. The PAFs of IHD attributable to TFAs are presented in **Supplementary Table S3**.

GBD select the dietary risk factors based on some criteria: the importance of the risk factor to disease burden or policy; sufficient data to estimate risk factor exposure; the strength of the epidemiological evidence supporting a causal relationship between risk factor exposure and disease, along with the accessibility of data to measure the extent of this relationship for each

unit of exposure change; and substantial evidence of the applicability of the effects to diverse populations. The World Cancer Research Fund criteria was used by GBD for grading convincing or probable evidence of risk-outcome pairs. Then, based on published systematic reviews, GBD identified IHD as the only outcome attributed to the TFA consumption.

IHD is a disease of the coronary arteries, mainly from atherosclerosis, leading to myocardial infarction or ischemia and stable angina, <sup>(1)</sup> mapped to the GBD 2019 using the International Classification of Diseases 10th Revision (ICD-10) codes: I20–I25.9, Z82.4–Z82.49. <sup>(16)</sup> Furthermore, GBD 2019 considers two metabolic mediators in the physiological pathway between TFA consumption and IHD: high low-density lipoprotein cholesterol (LDL-c) and systolic blood pressure (ICD-10 codes E78.0 and I10, respectively). <sup>(6)</sup>

The number, crude, and age-standardized rate of YLDs from IHD attributable to TFA in Brazil and its 27 federative units, referred to here as states and divided into five regions (**Supplementary Figure S1**), in 2019 were described. The YLD attributable burden estimates in states were expressed in quartiles. GBD uses a standard GBD world population to calculate age-standardized rates. The rates were expressed per 100,000 inhabitants in this study.

# Economic burden

Direct costs in the public health system related to outpatient care and hospitalizations were evaluated in this study, including the following expenditures: specialized medical consultations, hospital admissions, medications administered in hospital outpatient settings, orthoses and prostheses, and complementary procedures for secondary and tertiary care. The expenses did not include primary healthcare or medications beyond the scope of secondary and tertiary healthcare services.

SIA/SUS and SIH/SUS identify the cost per procedure related to IHD using the ICD-10 codes. These codes were used to link the costs and PAFs obtained from GBD 2019 by sex, age, location, and year. The Individualized Outpatient Production Bulletin (BPA-I) and the Authorizations for High Complexity Procedures (APAC) were used to obtain data from SIA/SUS in 2019, 2020, and 2021, while the reduced Hospital Admission Authorization (AIH) was used for SIH/SUS for the same period. The years 2020 and 2021 were included to consider information from 2019, which was corrected in subsequent years. Subsequently,

only the 2019 data were retained. In addition, to avoid age typographical errors, individuals aged > 110 years were excluded.

Cost data were extracted and processed using the R Microdatasus package. The Stata ICD-10 package version 13 was used to group the procedures in SIA/SUS and SIH/SUS, and the IHD attributable to TFA consumption from GBD 2019 through the ICD-10 codes. Subsequently, the IHD costs attributable to TFA consumption in 2019 were obtained by multiplying the total cost estimated for IHD by the respective PAF for each sex, age group, and location. In addition, to remove the effect of population size, the total costs per state were divided by the estimated resident population in the respective state in 2019<sup>(19)</sup> and multiplied by 10,000 inhabitants.

All costs were estimated in Brazilian Reais (R\$) and then converted into International Dollars (Int\$), a hypothetical unit of currency equivalent to the purchasing power of one US Dollar (US\$), considering the 2019 purchasing power parity (PPP) (Int\$ 1= R\$ 2.280). (20)

# Disease and economic burden and their relationship with the socio-demographic index

The correlations of the age-standardized YLD rate and the economic burden of IHD attributable to TFA consumption with the socio-demographic index (SDI) in 2019 were assessed. SDI is a composite metric developed by the GBD related to health outcomes. <sup>(6)</sup> Briefly, SDI ranges from 0 (less developed) to 1 (most developed) and comprises income per capita, mean years of schooling for those aged  $\geq$  15 years, and fertility rate in females under 25 years. <sup>(1,6)</sup>

#### Statistical analysis

The estimates were presented for both sexes, all ages, and the country pooled and stratified by type of healthcare (outpatient care and hospitalization), sex, age group, and state, with 95% UIs. The 95% UIs consist of a range of values probable to include the correct estimate of health loss for a specific cause. They are calculated by GBD 2019 to incorporate the uncertainty of parameters (such as the exposure, RR, ideal level of intake, and mortality) through Monte Carlo simulation iterations, wherein the UIs are the 2.5th and 97.5th values of the ordered 1,000 values and are chosen after repeating all calculations 1,000 times using one draw of each parameter. These percentiles represent the lower and upper bounds of the interval, respectively.

QGIS version 3.22.3 was used to create maps of the disease burden. Additionally, costs were analysed using STATA version 13.0 (*College Station*, Texas, USA).

#### **Results**

In 2019, IHD attributable to TFA consumption caused 11,165 YLDs (95% UI: 932–18,462) in the Brazilian population. The crude YLD rate was 5.15/100,000 (95% UI: 0.43–8.52. The results are shown in **Supplementary Table S4**.

Figure 1 (Supplementary Table S4) shows the total number, crude rates, and age-standardized rates of YLDs from IHD attributable to TFA consumption for the 27 Brazilian states. In general, the highest YLDs number was in states from the Northeast, South and Southeast states, especially in São Paulo, state with the highest value: 2,654.05 YLDs (95% UI: 221.07–4,399.80) (Figure 1A). The crude rates revealed almost two more YLDs in South and Southeast states when compared with those from the North (Figure 1B). Although the age-standardized rates were quite similar across the country, the Federal District, in the Central-West region, presented the lowest rates of YLDs: 3.92/100,000 (95% UI: 0.33–6.44) (Figure 1C).

The resulting total direct cost to the SUS in Brazil in 2019 with IHD attributable to TFA consumption was Int\$ 54,546,227 (95% UI: 4,505,792–85,561,810), as shown in **Table 1**. The higher cost share (93.72%) was for the hospitalizations (Int\$ 51,121,821; 95% UI: 4,195,620–80,168,034), whereas the costs of outpatient care were estimated at Int\$ 3,430,406 (95% UI: 310,172–5,393,776). Comparing the costs by sex, male individuals had higher costs (Int\$ 35,006,662; 95% UI: 2,708,429–55,099,866) than females (Int\$ 19,539,565; 95% UI: 1,797,363–30,461,944), even when compared by age group and type of healthcare. As expected, there was a general trend of cost increase with age, with the highest expenditure concentrated in individuals aged  $\geq$  50 years.

Including direct costs related to the mediators between TFA consumption and IHD, LDL-c and systolic blood pressure would increase by more than Int\$ 2,033,644 (95% UI: 165,656–3,176,781), totalling almost Int\$ 57 million in expenditure on the Brazilian health system (**Supplementary Table S5**).

As shown in **Figure 2** (**Supplementary Table S6**), from the top ten higher expenditures with IHD attributable to TFA to the SUS at the state level in 2019, the first six states were from

the South and Southeast. São Paulo, Paraná, and Minas Gerais had the highest costs. The states from the North, such as Acre, Roraima, and Amapá, spent less.

Additionally, to better understand and compare the states that contributed the highest cost to the SUS, we considered the size of their resident population and these costs for every 10,000 inhabitants (**Figure 3, Supplementary Table S7**). The three highest costs were observed in Sergipe (Int\$ 6,508/10,000; 95% UI: 576–10,265), from the Northeast region, followed by Paraná (Int\$ 6,296/10,000; 95% UI:520–9,883) and Santa Catarina (Int\$ 4,490/10,000; 95% UI: 369–7,111), both from the South. On the contrary, Maranhão (Int\$ 581/10,000; 95% UI: 48–920), from the Northeast region, followed by Acre (Int\$ 655/10,000; 95% UI: 51–1,038) and Pará (Int\$ 840/10,000; 95% UI: 71–1,322), both from the North, spent less with IHD attributable to TFA.

Figure 4 (Supplementary Tables S4, S7, and S8) shows the relationship between disease burden and economic costs to the SUS of IHD attributable to TFA consumption and the SDI by state. First, it is noteworthy to highlight that the states in the North and Northeast have the lowest SDI compared to their counterparts in the South, Southeast and Central-West. Overall, populations from the Northeast and North regions, with the lowest SDI, had fewer YLDs and spent less, except for Sergipe and Rio Grande do Norte, which presented higher expenditures. In contrast, Maranhão had low expenditure and high YLDs. Of note is the trend of increasing costs as the SDI increases, as shown in the states from the South, Southeast, and Central-Oest regions, except for the Federal District.

The results should be interpreted with attention because the 95% UI range incorporates the uncertainty of the parameters. The broad UIs do not allow the identification of differences between the states.

#### **Discussion**

This study revealed that IHD attributable to TFA consumption in Brazil contributed to many years of healthy life loss and represented an enormous direct cost to the country's public health system. Adhering to the lack of TFA intake (as recommended by GBD) would have avoided approximately 11,166 YLDs and saved Int\$ 54.5 million (R\$ 124.4 million) for the country in 2019. The highest costs were for hospitalizations, males, and the population aged ≥ 50 years. Our results also provide insights into these burdens across the country, highlighting the highest crude YLD rate in states from the South and Southeast and a similar pattern in

age-standardized YLD rates over the country with some heterogeneity regarding the costs with states from the Northeast and South counting with the bulk of these costs. However, it seems that as the SDI increases, expenditures increase.

The disease burden of IHD attributable to TFA is high in the Brazilian population. Of the total YLDs due to IHD in Brazil, the consumption of TFA contributes significantly to this burden (7.6%, the PAF for both sexes, all ages, in 2019). Deaths and DALYs attributable to TFA decreased in the country between 1990 and 2019, but diets high in TFA rose some positions in the ranking in this period. In addition, age-standardized YLD rates of CVD attributable to dietary risks have increased over the last 30 years. These estimates, in part, reflect improvements in CVD diagnosis and control. They can also be explained by the higher TFA intake in Brazil, 1.1% and 1.4% of the total daily energy intake in years before 2019, not complying with the WHO recommendations of a maximum of 1% total daily energy intake.

More recent research shows a consumption of TFA between 0.70% and 0.75% of the total daily energy for the Brazilian population, with stabilization of values between 2008 and 2009 and 2017–2018. Despite this, a scenario considering the elimination of PHO in Brazil, the largest source of TFA in the diet, would contribute to three to five times fewer deaths and costs because of premature deaths compared with limiting the TFA content in foods. These estimates and our results align with the current policies regarding TFA in Brazil (following an international agenda), which aimed for a 2% reduction in TFA from total fats by 2021 with complete elimination starting in January 2023. Besides that, it is worth mentioning that Brazil has been implementing policies to reduce TFA intake over time. In 2003, the requirement for trans-fat disclosure on nutritional labels was established as mandatory, succeeded by the zero TFA claims. Subsequently, in 2008, the national food industries voluntarily committed to the Ministry of Health's action plan for reducing iTFA through the Declaration of Rio de Janeiro.

The disability caused by CVD contributes to a substantial financial cost to the Brazilian health system<sup>(28)</sup>. However, the impact of dietary risk factors such as sodium,<sup>(29)</sup> sugar-sweetened beverages,<sup>(30)</sup> and processed meat<sup>(31)</sup> has recently been highlighted in Brazil. To the best of our knowledge, our study provides the first cost analysis of the disease attributable to TFA in SUS. These estimates regarding diets high in TFA are insightful for policymakers when considering the potential of health policies and the savings in the direct costs for the

health system, as already shown for Australia, <sup>(32)</sup> or even when considering the savings associated with the years of productivity lost in Brazil (US\$ 166.7 million). <sup>(13)</sup>

Although our study included TFA from all sources (ruminant and industrial), and it would be difficult to exclude total TFA from the diet, it is known that food and diet contain more iTFA.<sup>(8,33)</sup> While beef, lamb, and dairy products comprise 2–9% of the total fatty acids as TFA, PHO comprises up to 60% TFA. The reduction in iTFA could have a substantial impact on decreasing the health burden in Brazil.<sup>(13)</sup>

The cost analysis by sex and age revealed a higher burden of IHD attributable to TFA in men and individuals aged  $\geq 50$  years. Higher exposure to risk factors, such as an unhealthy diet, and reduced concern towards disease prevention and use of services for the detection and control may promote higher disability in male individuals. (5,21,25) Additionally, the fact that may explain the economic burden of IHD related to TFA consumption in the adult and elderly populations is the suboptimal diet during childhood and adolescence, (25) which may suggest that the promotion of a healthy diet should begin in the early stages of life. Besides the lifestyle changes brought on by urbanization and globalization, which affect diet and physical activity patterns, the rapid population growth due to the increase in life expectancy can help explain the significant impact of health problems at the oldest ages. (34) Thus, although NCDs may appear early in life, they progress commonly slowly, starting at younger ages but manifesting during adulthood, having a cumulative effect with advancing age. (35) Then, even though TFA consumption may decrease with age, (25) the effects of TFA consumption may accumulate over a lifetime, and the burden of IHD disease increases with age. Furthermore, we hypothesize that although ultra-processed foods are sources of iTFA, some products, such as soft drinks, certain types of bread, and cookies, may contain little or no TFA. Additionally, we propose that other factors, such as age itself, may have more impact on the disease burden among older people than iTFA (and consequently, ultra-processed foods). Perhaps these factors could explain why the burden of IHD does not follow the intake of ultra-processed foods with increasing age in Brazil.

Another important finding of our study is the higher costs for hospitalizations compared to outpatient care. Hospitalizations due to CVD resulted in the highest expenditure related to hospital admissions in Brazil. These results suggest that disease prevention contributes to the reduction of more complex and specialized treatments. However, new efforts are still

needed to improve CVD prevention and control in the country with the aim of reducing risk factors. (2,3)

In terms of differences in YLDs by state, this may reflect the complex relationship among diet, sociodemographic characteristics, interventions, and health services targeting CVD, which reflects regional inequalities and inequities in health. Moreover, this study advances the current understanding of the distribution of the direct costs of TFA on IHD, providing information stratified by states. Evaluating the absolute number, we found that states from the South and Southeast regions, with the highest SDI, presented the highest costs, mainly São Paulo, possibly because of their large population size and aging in these states. Therefore, the increasing absolute number of incidents and prevalent cases of IHD means that national health systems need to address more IHD-related procedures to detect and control diseases as the trend continues. The rising costs can result in substantial increases in the costs of public healthcare services that could be avoided by food and regulatory policies focused on interventions to reduce TFA consumption and then reduce IHD, save related expenditures, and promote the population's well-being.

After population size was considered in our analysis, the states with the highest direct costs to the SUS per 10,000 inhabitants were from the Northeast and South. Notably, these findings need to be better understood, but states with the lowest and highest SDI are experiencing increased costs of IHD attributable to TFA. Nonetheless, it seems that there is a general pattern of increasing expenditures as the SDI increases. The states with the highest SDI also presented high YLD rates attributable to TFA. Populations with better socioeconomic conditions (such as those from the South and Southeast in Brazil) consume more fat and ultra-processed foods (23,25,37) and have a broader network of secondary and tertiary care services<sup>(38)</sup> (accounted for in the study and which have a high cost). Therefore, if the population accesses these services more, they can contribute to an increase in expenditure. In contrast, although the Brazilian population in states with low SDI has been increasing ultraprocessed food consumption over time, (37) studies have already shown-individuals in low SDI regions in Brazil and worldwide presenting pronounced poor eating habits and fewer investments in medical care. (38-40). In addition, the Brazilian states with the lowest private health insurance coverage have the lowest cost, further revealing the inequalities in the country. (41)

Reflecting this hypothesis regarding the regional inequalities and the public financing of hospitalizations related to CVD in Brazil, a study showed that in terms of per capita for people aged  $\geq$  40 years, the expenditures were considerably lower in states from the North (US\$ 6.07) and Northeast (US\$ 10.28) than in the southern states (US\$ 20.32), even with CVD mortality varying little across the regions. Additional potential explanations for the observed lower costs in the North and Northeast states could be attributed to their younger populations and lower YLDs. Another possible explanation is the high coverage of primary health care in these regions, mainly in the Northeast, perhaps contributing to more access to preventive care instead of curative care.

The Federal District, with a high SDI and small disease and economic burden, seems closer to a more economically developed state with controlled outcome and risk factor actions, different from Maranhão, with a high disease burden but low expenditures to control them. Regarding social conditions, our results align with the nationwide findings in 2019, wherein the Federal District exhibited more favourable social indicators, while Maranhão demonstrated less favourable ones. (44) In Brazil, many NCDs have social gradients towards the most socially vulnerable populations. (21) Despite the poorest population experiencing worse health outcomes and having more difficult access, especially concerning secondary and tertiary care, progress made by the SUS over the past 30 years has led to improved outcomes and reduced health inequities. (45)

The SUS is the Brazilian public health system responsible for providing free access to health care at all levels for the entire Brazilian population. In 2019, approximately 76% of the population exclusively depended on the SUS to access medical services. The SUS plays a crucial role as the primary healthcare provider for the poorest population and those with limited access to private health insurance. Knowledge of the economic burden possibility the prioritizing policies, interventions and allocation of health resources according to budgetary constraints of this system. Therefore, in addition to highlighting the disease burden on the population and the potential cost savings for the SUS if iTFA consumption is zero, our findings also contribute to optimizing healthcare investments and providing guidance on resource allocation at the subnational level. Moreover, it should be mentioned that considering the health promotion model, the best quality service would prevent illness by promoting access to a healthy diet, a more cost-effective action. Notably, our results enable better management of healthcare and related costs across the country according to the TFA-attributable disease burden and sociodemographic conditions, but most importantly, can

contribute to the decisions of public policies to target iTFA to reduce disease and economic burden more effectively.

This study had some limitations which are worth noting. Although secondary data sources are essential for public health, the disease may be misclassified. In addition, disease classification and data depend on access to the diagnosis and awareness of the correct completion of information systems used for epidemiological surveillance. For example, subnational data can have different characteristics. Also, data related to states can result in imprecise estimates because most epidemiological data sources, such as surveys, are disaggregated to regions and not to states. In GBD, the impact of dietary risk factors on disease outcomes primarily stems from meta-analyses of prospective observational studies. While adjustments for confounding variables such as age, sex, smoking, and physical activity have been made in many cases, residual confounding remains a potential concern. (15) GBD uses many strategies to improve and compare data; however, biases are inevitable because many databases are used. (15)

Another limitation is that the most recent Brazilian consumption surveys<sup>(25)</sup> were not considered in GBD 2019. On the contrary, the data was obtained from a global market information database on sales. For example, this could weaken our conclusions on differences according to age since IHD increases with age, whereas ultra-processed foods consumption (the dietary sources of iTFA) decreases with age in Brazil.<sup>(25)</sup> However, as mentioned before, ultra-processed consumption has cumulative effects on NCDs; some ultra-processed foods consumed by this population may have little or no iTFA or may even have a lesser impact on the disease burden during ageing. Although our study included TFA from all sources (natural and industrial), which would make it difficult to exclude total TFA from the diet, it is known that food and diet contain more industrial TFA.<sup>(8,33)</sup>

Regarding the costs of IHD attributable to TFA consumption, it is worth mentioning that our findings can be underestimated because we only considered public healthcare expenditure and direct costs. The Brazilian private publicly available data does not include individualized data and ICD codes—essential in our study to determine the TFA attributable burden. Moreover, we do not include the costs of primary health care and other direct costs, such as medications to use at home provided by the SUS, rehabilitation, payment for caregivers, transportation of the patient to the health facility, etc. Finally, using PAF may not represent the picture of disease and economic burden because of limitations. For example, limited or inaccurate data, complexity of calculations and assumption of homogeneity of effects among

different populations. These uncertainties can propagate through the PAF calculation, affecting the reliability of the results. However, this is a well-established approach to quantify the contribution of specific risk factors to the disease and economic burden in a population, inform public health priorities, and have been used in other similar studies. (30,31,48–50)

Our study has several strengths. The originality and strength of this work lie in the disease and economic burden assessment related to the consumption of TFA in Brazil, especially in its 27 states. To the best of our knowledge, this study is the first to provide evidence on disability and direct costs to the Brazilian public health system attributable to TFA at the subnational level. We revealed the potential health benefits to the Brazilian population and cost savings to the SUS if TFA consumption was reduced. Moreover, we further investigated sex, age groups, and SDI, wherein this reduction could result in the highest cost savings. As a perspective, we also encourage future analyses incorporating other variables not included in this study, such as differences between the area of residence (urban or rural), access to health services, income, education, race, etc. Another strength is that we considered the riskoutcome pair, RRs, PAF, and uncertainty estimated by GBD, which has a robust, standardized, and updated methodology. (1,6) Lastly, this study strengthens the awareness of policymakers and other stakeholders towards TFA's global public health agenda<sup>(9)</sup> based on evidence found in the country. These estimates help draw attention to the international agenda accepted by Brazil, aiming to reduce and eliminate iTFA globally by 2023. (9,10) Therefore, these estimates reinforce that the country should prioritize the elimination of iTFA as already in progress, prioritizing the population's health in the face of long-term policies. (13)

#### **Conclusions**

This study showed that IHD attributable to TFA consumption contributed to high disability in the population and costs to the Brazilian health system in 2019. Overall, heterogeneity in the economic burden across Brazilian states is observed, which reveals inequalities regarding disease expenditures over the country; however, there is an indication that as their SDI increases, direct costs also increase. Thus, our findings reinforce that more stringent policies, such as iTFA elimination, as suggested by the international agenda, can contribute to health gains and economic savings, in addition to reducing subnational inequalities by prioritizing the allocation of resources and sustainability of the SUS.

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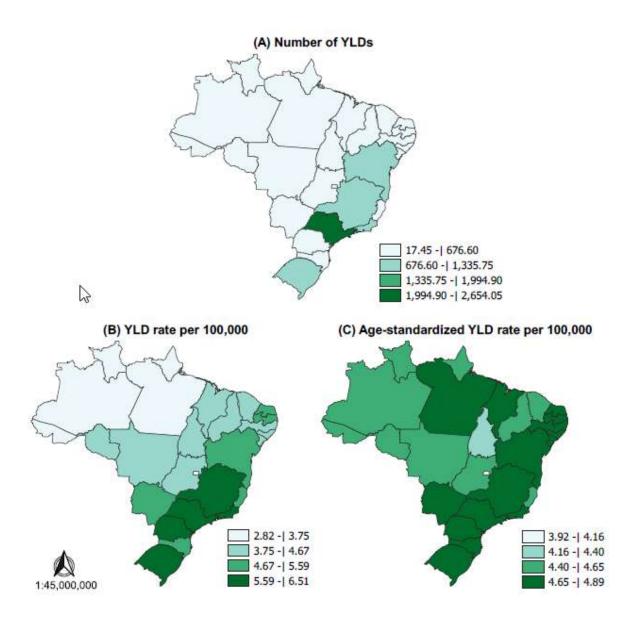
**Table 1.** The direct cost of ischemic heart disease attributable to the trans fatty acids consumption to the Unified Health System in Brazil by type of procedure, sex, and age group, 2019.

Age group		SIA	SIH	Total
(years)	Sex	Int\$ (95% UI)	Int\$ (95% UI)	Int\$ (95% UI)
	Male	1,681	106,957	108,638
25–29		(107–2,629)	(7,072–166,688)	(7,179–169,317)
	Female	1,408	45,603	47,011
		(102-2,215)	(3,277-70,721)	(3,379–72,936)
	Both	3,089	152,560	155,649
		(209–4,844)	(10,349–237,409)	(10,558–242,253)
30–34	Male	1,935	249,725	251,660
30-34		(128-3,104)	(16,253–399,280)	(16,381–402,384)
	Female	1,817	113,808	115,625
		(139–2,834)	(8,251-177,642)	(8,390–180,476)
	Both	3,752	363,533	367,285
		_(267–5,938)	(24,504–576,922)	(24,771–582,860)
35–39	Male	3,693	654,847	658,540
33 37		(235–5,818)	(42,309-1,026,718)	(42,544–1,032,536)
	Female	2,976	278,141	281,117
		(232–4,647)	(21,925–429,445)	(22,157–434,092)
	Both	6,669	932,988	939,657
		(467-10,465)	(64,234–1,456,163)	(64,701–1,466,628)
40–44	Male	5,363	1,243,121	1,248,484
10 11		(371–8,506)	(85,953–1,963,679)	(86,324–1,972,185)
	Female	3,829	611,640	615,469
		(298–6,045)	(47,309–953,233)	(47,607–959,278)
	Both	9,192	1,854,761	1,863,953
		(669–14,551)	(133,262–2,916,912)	(133,931–2,931,463)
45–49	Male	8,306	2,552,043	2,560,349
		(570–13,218)	(171,300–4,027,756)	(171,870–4,040,974)
	Female	6,387	1,218,778	1,219,165
	D 4	(544–9,898)	(103,794–1,872,099)	(104,338–1,881,997)
	Both	14,693	3,770,821	3,779,514
	N.C. 1.	(1,114–23,116)	(275,094–5,899,855)	(276,208–5,922,971)
50-54	Male	15,518	4,292,398	4,307,916
	Eamala	(1,081–24,821)	(295,557–6,769,326) 1,919,303	(296,638–6,794,147) 1.931,821
	Female	12,518 (1,030–19,624)	(158,943–2,986,870)	(159,973–3,006,494)
	Both	28,036	6,211,701	6,239,737
	Dom	(2,111–44,445)	(454,500–9,756,196)	(456,611–9,800,641)
	Male	32,988	5,581,717	5,614,705
55–59	Maic	(2,449–52,546)	(417,261–8,801,608)	(419,710–8,854,154)
	Female	29,057	2,621,758	2,650,815
	1 Ciliaic	(2,588–46,058)	(235,613–4,140,142)	(238,201–4,186,200)
	Both	62,045	8,203,475	8,265,520
	Dom	(5,037–98,604)	(652,874–12,941,750)	(657,911–13,040,354)
	Male	63,644	6,051,446	6,115,090
60–64	1,1410	(4,895–100,637)	(475,073–9,469,378)	(479,968–9,570,015)
	Female	62,337	3,002,874	3,065,211
	1 0111410	(5,726–97,715)	(274,796–4,671,548)	(280,522–4,769,263)
	Both	125,981	9,054,320	9,180,301
	2011	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	, , , , , , , , , , , , , , , , , , ,	-,,

			(10,621–198,352)	(749,869–14,140,926)	(760,490–14,339,278)
6 <b>7</b> 60		Male	114,065	5,138,951	5,253,016
65–69			(9,327–179,963)	(437,843–8,119,338)	(447,170–8,299,301)
		Female	114,745	2,894,115	3,008,860
			(10,622–179,849)	(271,233–4,520,254)	(281,855–4,700,103)
		Both	228,810	8,033,066	8,261,876
			(19,949–359,812)	(709,076–12,639,592)	(729,025–12,999,404)
70.74		Male	188,959	3,632,432	3,821,391
70–74			(15,772–298,638)	(294,388–5,701,393)	(310,160–6,000,031)
		Female	183,065	2,299,603	2,482,668
			(18,060–285,645)	(227,580–3,597,230)	(245,640–3,882,875)
		Both	372,024	5,932,035	6,304,059
			(33,832–584,283)	(521,968–9,298,623)	(555,800–9,882,906)
75 70		Male	255,868	2,213,135	2,469,003
75–79			(21,639–400,325)	(184,971-3,451,561)	(206,610–3,851,886)
		Female	233,892	1,575,579	1,809,471
			(23,205–363,804)	(154,471–2,435,812)	(177,676–2,799,616)
		Both	489,760	3,788,714	4,278,474
			(44,844–764,129)	(339,442–5,887,373)	(384,286–6,651,502)
80–84		Male	384,848	1,045,347	1,430,195
0U-04			(32,977–614,853)	(89,863–1,648,643)	(122,840-2,263,496)
		Female	349,888	837,058	1,186,946
			(36,108–544,269)	(86,011-1,295,943)	(122,119-1,840,212)
		Both	734,736	1,882,405	2,617,141
			(69,085-1,159,122)	(175,874–2,944,586)	(244,959–4,103,708)
85–89		Male	317,984	382,434	700,418
03-07			(27,068–506,128)	(33,212–603,473)	(60,280-1,109,601)
		Female	296,830	349,324	646,154
			(27,266–462,947)	(32,186–540,692)	(59,452–1,003,639)
		Both	614,814	731,758	1,346,572
			(54,334–969,075)	(65,398–1,144,165)	(119,732–2,113,240)
90–94		Male	231,142	81,001	312,143
, , , ,			(20,266–364,774)	(7,147–127,384)	(27,413–492,158)
		Female	221,348	99,461	320,809
		D 4	(21,598–344,163)	(9,323–154,565)	(30,921–498,728)
		Both	452,490	180,462	632,952
		3.7.1	(41,864–708,937)	(16,470–281,949)	(58,334–990,886)
+95		Male	143,281	11,833	155,114
		E1.	(12,341–228,931)	(1,001–18,750)	(13,342–247,681)
		Female	141,034	17,389	158,423
		Dot1-	(13,428–219,172)	(1,705–26,863)	(15,133–246,035)
		Both	284,315	29,222	313,537 (28,475–493,716)
		Male	(25,769–448,103) 1,769,275	(2,706–45,613) 33,237,387	35,006,662
Total		iviale	(149,226–2,804,891)	(2,559,203–52,294,975)	(2,708,429–55,099,866)
Int\$	(95%	Female	1,661,131	17,884,434	19,539,565
UI)	(3370	Temale	(160,946–2,588,885)	(1,636,417–27,873,059)	(1,797,363–30,461,944)
01)		Both	3,430,406	51,121,821	54,546,227
		Doni	(310,172–5,393,776)	(4,195,620–80,168,034	(4,505,792–85,561,810)
			(310,174-3,373,770)	(7,193,020-00,100,034	(7,505,194-05,501,010)

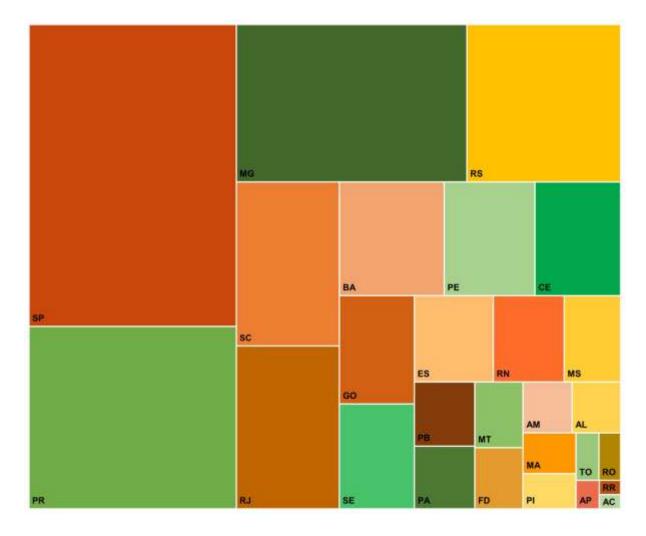
95% UI: 95% uncertainty interval; Int\$: International dollar, Int\$ 1 = US\$ 1; SIA: Outpatient

Information System; SIH: Hospital Information System.



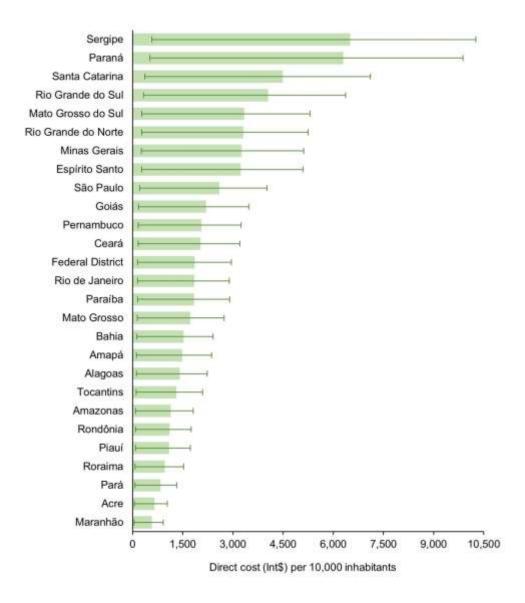
**Figure 1** Number and rates of years lived with disability, per 100,000, for ischemic heart disease attributable to trans fatty acids consumption in Brazil, 2019.

95% UI: 95% uncertainty interval; Int\$: International dollar, Int\$ 1 = US\$ 1; YLD: years lived with disability.



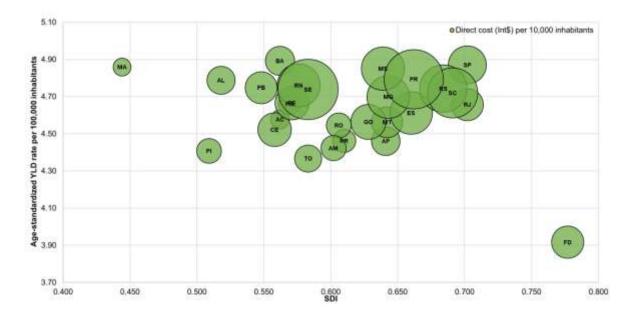
**Figure 2** The direct cost (Int\$) of ischemic heart disease attributable to the trans fatty acids consumption to the Unified Health System in Brazil by states, 2019.

AC: Acre; AP: Amapá; AM: Amazonas; PA: Pará; RO: Rondônia; RR: Roraima; TO: Tocantins; AL: Alagoas; BA: Bahia; CE: Ceará; MA: Maranhão; PB: Paraíba; PE: Pernambuco; PI: Piauí; RN: Rio Grande do Norte; SE: Sergipe; FD: Federal District; GO: Goiás; MT: Mato Grosso; MS: Mato Grosso do Sul; ES: Espírito Santo; MG: Minas Gerais; RJ: Rio de Janeiro; SP: São Paulo; PR: Paraná; RS: Rio Grande do Sul; SC: Santa Catarina; Int\$: International dollar, Int\$ 1 = US\$ 1.



**Figure 3** The direct cost (Int\$) per 10,000 inhabitants of ischemic heart disease attributable to the trans fatty acids consumption to the Unified Health System in Brazil by states, 2019.

Int\$: International dollar, Int\$ 1 = US\$ 1.



**Figure 4** Relationship between the disease burden and economic costs (Int\$) to the Unified Health System in Brazil of ischemic heart disease attributable to trans fatty acids consumption and the SDI by state, 2019.

AC: Acre; AP: Amapá; AM: Amazonas; PA: Pará; RO: Rondônia; RR: Roraima; TO: Tocantins; AL: Alagoas; BA: Bahia; CE: Ceará; MA: Maranhão; PB: Paraíba; PE: Pernambuco; PI: Piauí; RN: Rio Grande do Norte; SE: Sergipe; FD: Federal District; GO: Goiás; MT: Mato Grosso; MS: Mato Grosso do Sul; ES: Espírito Santo; MG: Minas Gerais; RJ: Rio de Janeiro; SP: São Paulo; PR: Paraná; RS: Rio Grande do Sul; SC: Santa Catarina; SDI: Socio-demographic index. Int\$: International dollar, Int\$ 1 = US\$ 1.