HIV/AIDS mortality attributable to alcohol use in South Africa: a comparative risk assessment by socioeconomic status

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ABSTRACT

Objectives To quantify HIV/AIDS mortality attributable to alcohol use in the adult general population of South Africa in 2012 by socioeconomic status (SES).

Design Comparative risk assessment based on secondary individual data, aggregate data and risk relations reported in the literature.

Setting South African adult general population.

Participants For metrics of alcohol use by SES, sex and age: 27 070 adults that participated in a nationally representative survey in 2012. For HRs of dying from HIV/AIDS by SES: 87 029 adults that participated in a cohort study (years 2000 to 2014) based out of the Umkhanyakude district, KwaZulu-Natal.

Main outcome measures Alcohol-attributable fractions for HIV/AIDS mortality by SES, age and sex were calculated based on the risk of engaging in condom-unprotected sex under the influence of alcohol and interactions between SES and alcohol use. Age-standardised HIV/AIDS mortality rates attributable to alcohol by SES and sex were estimated using alcohol-attributable fractions and SES-specific and sex-specific death counts. Rate ratios were calculated comparing age-standardised rates in low versus high SES by sex.

Results The age-standardised HIV/AIDS mortality rate attributable to alcohol was 31.0 (95% uncertainty interval (UI) 21.6 to 41.3) and 229.6 (95% UI 108.8 to 351.6) deaths per 100 000 adults for men of high and low SES, respectively. For women the respective rates were 10.8 (95% UI 5.5 to 16.1) and 75.5 (95% UI 31.2 to 144.9). The rate ratio was 7.4 (95% UI 3.4 to 13.2) for men and 7.0 (95% UI 2.8 to 18.2) for women. Sensitivity analyses corroborated marked differences in alcohol-attributable HIV/AIDS mortality, with rate ratios between 2.7 (95% UI 0.8 to 7.6; women) and 15.1 (95% UI 6.8 to 27.7; men).

Conclusions The present study showed that alcohol use contributed considerably to the socioeconomic differences in HIV/AIDS mortality. Targeting HIV infection under the influence of alcohol is a promising strategy for interventions to reduce the HIV/AIDS burden and related socioeconomic differences in South Africa.

INTRODUCTION

HIV/AIDS was the disease with the highest health burden in South Africa in 2016, accounting for over 30% of the total burden of disease.1 Research showed that the HIV/AIDS burden in South Africa was unequally distributed across different levels of socioeconomic status (SES). Persons of low SES were shown to have a higher HIV prevalence,2 and an at least 1.5 times higher risk of dying from HIV/AIDS compared with persons of high SES.3 With a Gini index over 60, South Africa is among the countries worldwide where income is most unequally distributed.4 Not least due to the history of colonialism and apartheid, socioeconomic differences in South Africa are heavily intertwined with race and despite decreasing trends, the average income of a white person is still over 10 times that of a black African person.5

One factor possibly contributing to the socioeconomic differences in HIV/AIDS burden is alcohol use. While in South Africa and elsewhere, current drinkers of low SES were shown to drink higher quantities and in overall riskier patterns, they were—partly due to lower affordability of alcohol—also more likely to abstain from alcohol completely.6–9 A study based on nationally representative data from South Africa recently found an
interaction between SES and alcohol use on HIV infection risk, indicating that persons of low SES increase their risk of contracting HIV significantly more when drinking alcohol compared with their counterparts of high SES.10

Alcohol is a psychotropic depressant of the central nervous system that likely impairs decision-making by diminishing the perception of risk and reducing inhibitions.11 Particularly occasional heavy drinking has been linked to sexual risk behaviour and an increased HIV infection risk in cross-sectional and longitudinal studies from Sub-Saharan Africa as well as other countries.12–16 A series of randomised experiments corroborated the findings by showing that (HIV-positive) participants were significantly more likely to consider condom-unprotected sex when under the influence of alcohol.17 Overall, the evidence supports causal effects of alcohol use on HIV infection risk.18 Interactive effects of alcohol use and SES could be the result of a number of contextual factors such as the drinking environment, malnutrition and poverty that work to the disadvantage of persons of low SES.10 As a consequence, alcohol use is likely to contribute to socioeconomic differences in HIV/AIDS mortality in South Africa above and beyond potential differences in the drinking patterns.

Figure 1 shows a conceptual framework of the associations between alcohol use, SES and HIV/AIDS mortality. The focus of the current study was on alcohol use as a risk factor for HIV infection via condom-unprotected sex.19–22 Interactions between alcohol use and a low SES,10 likely due to interactions between alcohol use and malnutrition,23 24 were taken into account. Effects of alcohol use on the mortality risk after HIV infection (eg, through delayed treatment initiation, reduced treatment adherence and increased risk of opportunistic infections)25 26 were not modelled in the current study. Previous research showed that less than 0.5% of HIV/AIDS deaths could be explained through non-adherence to antiretroviral therapy due to alcohol use in Sub-Saharan Africa.

Even though there is worldwide a large body of evidence showing increased risk of all-cause and alcohol-attributable mortality for persons of low SES, there have been few attempts to incorporate SES in burden of disease estimations.27 28 The objective of the current study was to quantify the burden of HIV/AIDS mortality attributable to alcohol use by SES, age and sex in South Africa in 2012.

**METHODS**

A comparative risk assessment was performed to estimate alcohol-attributable HIV/AIDS mortality for different subgroups defined by SES, age and sex. The methodology was based on previous burden of disease and mortality analyses.29–31 Figure 2 gives a schematic overview of the procedures and data sources. In step 1, alcohol-attributable fractions (AAF) were calculated, representing the proportion of HIV/AIDS deaths that would not have occurred under the counter-factual scenario that everyone would have been a lifetime abstainer. To calculate the AAF, information on exposure to alcohol in each subgroup was combined with the risk associated with the level of exposure in the particular subgroup (see formula 1 in the online supplementary material).

In the second step, the AAF of each subgroup was applied to the HIV/AIDS deaths in the subgroup to calculate the alcohol-attributable deaths. This required knowledge on the HIV/AIDS death counts in each subgroup (see section ‘Mortality’). In the third step, the alcohol-attributable deaths were combined with population counts to calculate HIV/AIDS mortality rates per 100 000 adults. The mortality rates were age-standardised using the WHO reference population.32 Mortality rate ratios were obtained by dividing the age-standardised mortality rate in the low SES group by the one in the high SES group.

The following paragraph will define the core measures used in the comparative risk assessment and provide details on the four areas shown in figure 2, namely alcohol exposure, risk, mortality and population. If not indicated differently, all analyses were performed in R V.3.2.3. The analyses conformed to the principles embodied in the Declaration of Helsinki, and ethics approval was obtained from the Centre for Addiction and Mental Health Research Ethics Board.
Definition of core measures

The analyses refer to the adult general population defined as 15 years of age or older. A subgroup was defined by SES (high or low), sex (male or female) and age category (15 to 34, 35 to 54 and 55 years and older). SES was operationalised as an asset score splitting the population at the median into two equal halves. Alcohol use was categorised into current abstainers (no alcohol use in the past 12 months) and current drinkers. Average grams of pure alcohol per day were calculated among current drinkers.

Exposure to alcohol

The prevalence of current abstinence and current alcohol use, as well as the average grams of pure alcohol per day in each subgroup, were calculated based on nationally representative survey data. Secondary data from the nationally representative South African National HIV Prevalence, Incidence and Behaviour Survey (SABSSM) from the year 2012 were used. In SABSSM, households were sampled based on a multistage sampling frame, excluding homeless and institutionalised persons. A total of 27,070 adults were interviewed by trained field workers. Household assets such as television ownership or type of water supply were recorded. The frequency of alcohol use was assessed with the question ‘How often did you have a drink containing alcohol in the past 12 months?’, with response categories ranging from ‘Not in the past 12 months’ to ‘4 or more times a week’. The quantity was assessed as ‘How many drinks containing alcohol do you have on a typical day when you are drinking?’, with response categories ranging from ‘1 or 2’ to ‘10 or more’. The frequency of irregular heavy drinking occasions with four/five (for women/for men) drinks or more was assessed in categories ranging from ‘never’ to ‘daily or almost daily’. Based on these variables, the grams of pure alcohol per day among current drinkers in each subgroup were calculated. A standard drink was defined as 12 g of pure alcohol. All calculations were performed using survey weights. More details on SABSSM are provided in the online supplementary data.

Population

Prevalence of alcohol use and grams per day by subgroup in 2012 [individual data: SABSSM]

Exposure to alcohol by subgroup corrected for underestimation [method: Kehoe et al., 2012]

Total adult per capita alcohol consumption in to total population in 2012 [aggregate data: GISAH]

Interaction between SES and alcohol use for HIV risk [evidence: Probst et al., 2017]

Relative risk of acquiring HIV through unprotected sex under influence of alcohol vs. abstinence [evidence: Scott-Sheldon et al., 2016]

Number of HIV/AIDS deaths in South Africa in 2012 by sex and age [aggregate data: WHO]

Relative risk to split HIV/AIDS deaths by SES [evidence: Probst et al. 2016; data: DSA]

Population counts in each subgroup in 2012 [individual data: SABSSM][aggregate data: Statistics South Africa]

Figure 2  Schematic overview of the data sources, relative risks and core data processing steps used to quantify alcohol-attributable HIV/AIDS mortality by SES in South Africa for the year 2012. A subgroup was defined by SES, age and sex. HIV risk is referring to HIV acquired through condom-unprotected sex under the influence of alcohol. Sources: Probst et al, Probst et al, Scott-Sheldon et al, Ahmad et al, Shisana et al, Kehoe et al, Statistics South Africa, AAF, alcohol-attributable fraction; GISHA, Global Information System on Alcohol and Health; SABSSM, South African National HIV Prevalence, Incidence and Behaviour Survey; SES, socioeconomic status.
alcohol use, leading to an underestimation of the ‘true’ amount of alcohol consumed. The standard triangulation technique of the Global Burden of Disease study and WHO Global Status Reports was applied to correct for underestimation. The consumption in the subgroups was thereby shifted up to match in sum 80% of the total per capita alcohol consumption in 2012. The aggregate total adult per capita consumption in litres of pure alcohol was obtained from the WHO Global Information System on Alcohol and Health. A total of 10.39 L of pure alcohol were consumed per capita in South Africa in 2012.

Alcohol-related risk

The risk relating different levels of alcohol use to HIV/AIDS was derived from two main sources. An OR from a meta-analysis (including 28 experimental studies predominantly from North America and the UK) performed by Scott-Sheldon et al was used. The OR of 1.54 (95% CI 1.31 to 1.78) referred to intentions to engage in condom-unprotected sex under the influence of alcohol use compared with current abstinence. Specifically, the OR referred to a minimum of 61 and 49 g per drinking occasion among men and women, respectively (see also online supplementary material). A detailed discussion on the use of the above OR in comparative risk assessments of HIV/AIDS mortality can be found elsewhere. Since this OR was lower than those reported in observational cohort studies from Sub-Saharan African countries, it can be considered as conservative; in addition, it captured only the causal effect of alcohol as it was based exclusively on experimental studies.

Furthermore, a recent study using three cross-sectional representative surveys from South Africa in the years 2005, 2008 and 2012 showed that alcohol use and SES interact to increase the risk of testing HIV positive among current drinkers of low SES. The OR (interaction term) of 1.94 (95% CI 1.29 to 2.93) from the year 2012 was used to account for the increased risk of current drinkers of low SES.

Mortality

Aggregate HIV/AIDS death counts in 2012 by age group and sex were obtained from WHO. To obtain death counts in each subgroup, that is, split by SES, a relative risk of dying from HIV/AIDS for people of low compared with high SES was used.

South African cohort studies that reported a relative risk of dying from HIV/AIDS by SES, based on an asset score, were identified through a recently published systematic review and recalculated to comply with the operationalisation of SES used in the current study. In the case of the Agincourt health and sociodemographic surveillance system, Houle et al were asked to recalculate the results accordingly. In the case of the Demographic Surveillance Area (DSA), original data were used to recalculate the respective HR, as described below. Both measures of relative risk were combined in a random-effects meta-analysis, performed in Stata V.13 (see online supplementary material).

The secondary cohort data were obtained from the DSA of the Africa Centre for Health and Population Studies. The study was based in a predominantly rural area in the Umkhanyakude district of KwaZulu-Natal, comprising also periurban informal settlements and a township. An open cohort of about 11 000 households was surveyed between 2000 and 2014. Each household was visited at least two to three times a year. Trained field workers assessed household demographic and socioeconomic information in face-to-face interviews. Cause of death was determined using verbal autopsy interviews with the next of kin. The sample used in this study comprised 87 029 adults, 757 404 person-years (average...
follow-up time was 8.7 years) and 5212 deaths from HIV/AIDS. Cox proportional hazards survival analyses were performed using the DSA data. The HR of dying from HIV/AIDS was calculated for the median split and for a mapped asset score, adjusting for age and sex. Kaplan-Meier Survival Curves are shown in online supplementary material.

Population
Aggregate population counts were based on 2011 census information. To calculate counts in each subgroup, proportions observed in SABSSM data were applied. Population counts were used as denominators to calculate mortality rates per 100,000 adults.

Calculate asset scores
An asset score was calculated for SABSSM as well as DSA data using Multiple Correspondence Analysis (MCA; eg, from South Africa see Wabiri and Taffa). In a first step, all variables for which 95% of all participants gave the same answer were excluded to assure stability of the model. Then missing values in individual asset variables were imputed based on an iterative MCA algorithm, using the imputeMCA command from the missMCA package in R. After fitting the MCA on the imputed data, variables for which all categories contributed less than 1.5% to the first dimension were excluded. The asset score was calculated as a median split of the first dimension resulting from the MCA. This resulted in two socioeconomic groups representing the lower and the upper 50% of the socioeconomic distribution in the adult population. A second version of the asset score of the DSA was calculated to correct for the predominantly rural and lower SES population of the study area. The continuous asset score (first dimension) was mapped onto the continuous asset score of the nationally representative SABSSM data and split at the percentile corresponding to the median of the national distribution.

Race groups were highly unequally distributed across the two levels of SES with 98% of all whites being in the upper SES group and 97% of all persons in the low SES group being black African (SABSSM data). Detailed information on the asset scores and associations with race are shown in the online supplementary material.

Uncertainty estimation
Uncertainty of all calculated metrics (ie, AAFs, mortality rates and mortality rate ratio) was determined using Monte Carlo simulations, sampling from the distribution of each parameter 100,000 times. The 95% uncertainty interval (UI) was estimated using the 2.5th and 97.5th percentile of the distribution of the sampled metric.

Sensitivity analyses
Two sensitivity analyses were performed to examine the stability of the results under different assumptions. Sensitivity analysis I used the mapped asset score instead of the median split asset score. Sensitivity analysis II was performed applying the interaction term exclusively for current alcohol use above the critical

Figure 4 One minus the cumulative density of the gamma distribution of alcohol use for high (dashed) and low SES (dotted) and by age and sex in the adult (15+) South African general population in 2012. The graph indicates the proportion of current drinkers (y-axis) with an average level of drinking of alcohol use equal to or above the grams of pure alcohol per day indicated on the x-axis.
Table 1  Total adult population, HIV/AIDS deaths, alcohol-attributable fractions in per cent, HIV/AIDS deaths attributable to alcohol and alcohol-attributable HIV/AIDS mortality rates (per 100 000) by sex, socioeconomic status and age in South Africa in 2012

<table>
<thead>
<tr>
<th>Sex</th>
<th>SES</th>
<th>Age</th>
<th>Total adult population</th>
<th>Total HIV/AIDS deaths*</th>
<th>AAF (%)†</th>
<th>HIV/AIDS deaths attributable to alcohol‡</th>
<th>Mortality rate per 100 000 **</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td></td>
<td>%§  95% CI   N¶</td>
<td>PE  95% UI</td>
<td>PE  95% UI</td>
<td>PE  95% UI</td>
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</tr>
<tr>
<td>Male</td>
<td>High</td>
<td>15–34</td>
<td>12.2  11.2 to 13.3  4475846</td>
<td>9618  9325 to 9912</td>
<td>8.8  5.3 to 12.6</td>
<td>846  508 to 1210</td>
<td>18.9  11.4 to 27.0</td>
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<tr>
<td></td>
<td></td>
<td>35–54</td>
<td>8.4  7.6 to 9.2   3059895</td>
<td>26517  25799 to 27232</td>
<td>7.1  4.2 to 10.4</td>
<td>1895  1115 to 2768</td>
<td>61.9  36.4 to 90.5</td>
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<td></td>
<td>55+</td>
<td>3.6  3.2 to 4.0   1325354</td>
<td>1384  1347 to 1420</td>
<td>7.0  4.1 to 10.3</td>
<td>97  57 to 142</td>
<td>7.3  4.3 to 10.7</td>
</tr>
<tr>
<td>Low</td>
<td>15–34</td>
<td>14.0  13.0 to 15.0  5115000</td>
<td>17763  17468 to 18056</td>
<td>31.6  9.1 to 55.3</td>
<td>5611  1608 to 9826</td>
<td>109.7  31.4 to 192.1</td>
<td></td>
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<tr>
<td></td>
<td></td>
<td>35–54</td>
<td>7.0  6.3 to 7.7   2550831</td>
<td>35724  35009 to 36442</td>
<td>35.4  10.7 to 59.2</td>
<td>12640  3815 to 21170</td>
<td>495.5  149.6 to 829.9</td>
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<tr>
<td></td>
<td></td>
<td>55+</td>
<td>2.8  2.5 to 3.2   1036844</td>
<td>1749  1713 to 1786</td>
<td>34.6  11.0 to 58.1</td>
<td>605  192 to 1016</td>
<td>58.3  18.5 to 98.0</td>
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<td>High</td>
<td>15–34</td>
<td>11.6  10.7 to 12.7  4259681</td>
<td>14885  14421 to 15352</td>
<td>3.3  1.1 to 5.6</td>
<td>499  168 to 840</td>
<td>11.7  3.9 to 19.7</td>
</tr>
<tr>
<td></td>
<td></td>
<td>35–54</td>
<td>9.0  8.3 to 9.8   3305373</td>
<td>19933  19385 to 20480</td>
<td>2.6  0.7 to 4.6</td>
<td>515  144 to 926</td>
<td>15.6  4.4 to 28.0</td>
</tr>
<tr>
<td></td>
<td></td>
<td>55+</td>
<td>5.0  4.5 to 5.6   1830590</td>
<td>1017  989 to 1045</td>
<td>2.8  0.9 to 4.8</td>
<td>29  9 to 49</td>
<td>1.6  0.5 to 2.7</td>
</tr>
<tr>
<td>Low</td>
<td>15–34</td>
<td>14.2  13.0 to 15.5  5213940</td>
<td>29444  28978 to 29909</td>
<td>11.7  2.2 to 27.3</td>
<td>3457  637 to 8033</td>
<td>66.3  12.2 to 154.1</td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td>35–54</td>
<td>7.8  7.2 to 8.6   2869320</td>
<td>27963  27416 to 28512</td>
<td>13.4  2.2 to 30.7</td>
<td>3737  622 to 8592</td>
<td>130.2  21.7 to 299.4</td>
</tr>
<tr>
<td></td>
<td></td>
<td>55+</td>
<td>4.4  3.9 to 4.9   1594161</td>
<td>1431  1403 to 1459</td>
<td>11.1  2.4 to 25.4</td>
<td>159  35 to 363</td>
<td>10.0  2.2 to 22.8</td>
</tr>
</tbody>
</table>

*Calculated from aggregate WHO data and relative risk to split HIV/AIDS deaths by SES (see mortality panel in figure 2).
†Calculated from exposure to alcohol and related risk in each subgroup (see step one in figure 2).
‡Calculated from AAFs and estimated HIV/AIDS deaths (see step 2 in figure 2).
¶Calculated from the proportion and Census 2011 information (see population panel in figure 2).
**Calculated from alcohol-attributable HIV/AIDS deaths and population counts (see step 3 in figure 2).
AAF, alcohol-attributable fraction; PE, point estimate; SES, socioeconomic status; UI, uncertainty interval from Monte Carlo simulation.
thresholds described in the ‘Alcohol-related risk’ section.

RESULTS

Alcohol use

Results on alcohol use based on SABSSM data are shown in figure 3. Across age groups and sex, the prevalence of current drinking was higher at high compared with low SES.

Figure 4 shows the distribution (one minus the cumulative gamma density distribution) of alcohol use in each subgroup that has been corrected for under-reporting. At each level of alcohol use measured in average grams of pure alcohol per day, the graph indicates the proportion of current drinkers that drank at this level or above it. With the only exception of young men, the proportion of current drinkers above the threshold of 49/61 g of pure alcohol per day was greater in the low than in the high SES group.

Mortality

The survival analysis using DSA data resulted in an HR of 1.60 (95% CI 1.51 to 1.70) for persons of low compared with high SES, adjusting for sex and age at baseline. The recalculated relative risk from Houle et al was 1.65 (95% CI 1.52 to 1.78) for persons of low compared with high SES, adjusting for sex and age at baseline. Combining both using a random-effects meta-analysis resulted in a relative risk of 1.62 (95% CI 1.54 to 1.96) for persons of low compared with high SES of 1.62 (95% CI 1.54 to 1.96). HIV/AIDS deaths in each subgroup are shown in table 1. The HR based on the mapped asset score, used in sensitivity analysis I, was 3.30 (95% CI 2.82 to 3.87).

Comparative risk assessment

AAFs showed that while in women of high SES, roughly 3% of all HIV/AIDS deaths were attributable to alcohol; among women of low SES, this was the case for just over 10% (table 1). Similarly, among men of high SES, 7% to 9% of all HIV/AIDS deaths were alcohol attributable, while over 30% of the HIV/AIDS death in men of low SES was alcohol attributable. In 2012, approximately 30 000 HIV/AIDS deaths among adults were attributable to alcohol in South Africa, about 26 000 of which appeared in persons of low SES.

Among women, the age-standardised mortality rates for HIV/AIDS mortality attributable to alcohol were 10.8 (95% UI 5.5 to 16.1) per 100 000 and 75.5 (95% UI 31.2 to 144.9) per 100 000 at high and low SES, respectively. The age-standardised mortality rates among men were 31.0 (95% UI 21.6 to 41.3) per 100 000 in men of high and 229.6 (95% UI 108.8 to 351.6) per 100 000 in men of low SES.

The ratio comparing age-standardised rates of alcohol-attributable HIV/AIDS mortality in high versus low SES was 7.4 (95% CI 3.4 to 13.2) in men and 7.0 (95% CI 2.8 to 18.1) in women, indicating a seven times elevated risk of an alcohol-attributable HIV/AIDS death in persons of low SES compared with high SES.

Sensitivity analyses

The results of both sensitivity analyses are shown in the online supplementary material. Sensitivity analysis I used an asset score that was mapped onto the distribution in the nationally representative data to calculate the HR used to split deaths by SES. This led to increased age-standardised mortality rates for HIV/AIDS mortality attributable to alcohol among men and women of low SES resulting in a mortality rate ratio of 14.3 (95% UI 5.6 to 37.9) among women and 15.1 (95% UI 6.8 to 27.7) among men. Sensitivity analysis II applied the interaction term between SES and alcohol use exclusively for current alcohol use above the critical thresholds of 61 (men) and 49 g (women) of pure alcohol per day. This led to lower AAFs for persons of low SES, resulting in a mortality rate ratio of 2.7 (95% UI 0.8 to 7.6) among women and 4.0 (95% UI 1.4 to 8.3) among men.

DISCUSSION

This study was to our knowledge the first to quantify alcohol-attributable HIV/AIDS mortality burden for different socioeconomic strata. The study showed that age-standardised HIV/AIDS mortality rates attributable to alcohol are much higher for persons of low SES compared with high SES. Based on the vast socioeconomic inequalities between racial groups, the excess burden was to bear predominantly by the black African population. According to our findings, about 10% to 30% of HIV/AIDS deaths in persons of low SES could be prevented when (heavy) alcohol use would be prevented. Among persons of high SES, the estimated proportion of deaths that would have been prevented was about 3% to 9%.

A number of factors contributed to the findings including the higher average alcohol use in grams per day among current drinkers of low SES, the interaction expressing a higher increase in the risk for alcohol users of low SES as well as the higher overall risk of dying from HIV/AIDS at low SES. The findings on drinking patterns by SES were in line with other research from South Africa including the most recent demographic and health survey. Regarding alcohol use as a contributor to socioeconomic differences in mortality, the findings match those of a recent study that investigated the contribution of alcohol use to socioeconomic differences in self-reported diagnoses in South Africa, concluding that alcohol use accounted for about a quarter of the socioeconomic inequalities in self-reported health.

Overall, the findings of the current study suggest that alcohol use contributes to socioeconomic differences above and beyond socioeconomic differences in drinking patterns. Factors such as malnutrition, contextual and behavioural factors should be considered as potential contributors to interaction effects between alcohol use
and SES to increase the HIV infection risk for persons of low SES in future research. Studies from Sub-Saharan Africa support the relevance of the above factors in the associations between HIV/AIDS and sexual risk taking under the influence of alcohol, particularly in a low SES context.\textsuperscript{30–32} However, a systematic analysis of the underlying causal pathways is missing as of now.

Limitations

In two regards, the national representativeness of the individual data used in the analyses was limited. With respect to data on exposure to alcohol, the SABSSM sampling frame excluded institutionalised and homeless populations, whereby alcohol use of these potentially low SES populations was not accounted for. Furthermore, to this date, there are no nationally representative mortality statistics on HIV/AIDS mortality and SES for South Africa. In the latest release on vital statistics from Statistics South Africa, education and occupation were unknown or not specified for 50\% and 70\% of the deaths, respectively.\textsuperscript{53} Consequently, SES-specific death counts had to be estimated using the relative risk of dying for people with low compared with high SES. A systematic literature review identified no nationally representative cohort studies on cause-specific mortality.\textsuperscript{5} However, the current study included estimates of relative risk from both major South African cohort studies on mortality from Umkhanyakude\textsuperscript{54} and Agincourt.\textsuperscript{55} The overall lower SES of the South African cohort studies on mortality from Umkhanyakude\textsuperscript{54} and Agincourt.\textsuperscript{55} The overall lower SES of the populations in both cohorts could have led to an underestimation of the socioeconomic differences in HIV/AIDS mortality. Sensitivity analysis I using a mapped asset score indicated that the mortality gap might be much larger than assumed based on the simple median split without mapping.

The division of the total population in two halves was rather crude. However, given the available data and relative risks, the categorisation was chosen to achieve stable and consistent estimates. Furthermore, the representation of the whole of the population was preferred over an extreme group comparison.

A recent study investigating the coverage (ie, the proportion of the total alcohol consumed per capita that gets reported in representative surveys) of five nationally representative South African surveys (including SABSSM) found coverage below 20\% in all of the surveys investigated.\textsuperscript{35} The authors concluded that an over-reporting of abstention from alcohol could have caused the low coverage additional to reporting lower quantities of alcohol consumed. The triangulation technique used to estimate the ‘true’ consumption among current drinkers relies on the accuracy of the reported proportion of current abstainers. It is possible that the prevalence of current alcohol use was underestimated based on the survey data, while the average grams per day among current drinkers and thereby the respective AAF were overestimated in the current study.

To this day, it is the common practice in comparative risk assessments to use identical relative risks across countries.\textsuperscript{56,57} As the required data are often not available for the specific country under investigation, the risk relations are mostly derived from pooled cohort studies or meta-analyses. However, this practice might introduce bias as the same relative risks might not apply to all countries, especially if the underlying causal pathways are not mainly biological. In the present study, the OR relating alcohol use to HIV/AIDS was based on a meta-analysis of experimental studies that were to the large part performed in high-income countries.\textsuperscript{17} However, all other model inputs were based on recent data from South Africa, specifically.

Possible implications for policies and interventions

Targeting interventions on high-risk groups such as 35 to 54-year-old male current drinkers of low SES could be a potentially cost-effective strategy to reduce the HIV/AIDS burden in South Africa. Interventions addressing alcohol-attributable HIV/AIDS burden should target contextual and behavioural factors associated with alcohol use and the drinking environment of persons of low SES instead of only addressing alcohol use directly. Such factors include malnutrition which is likely to increase the infection risk under the influence of alcohol by exacerbating the immune suppressive effects of alcohol, as well as availability of condoms at drinking venues or alternatives for leisure activities besides drinking venues in low SES housing areas.\textsuperscript{58–60} Of course, improving the wealth inequality between strata in South Africa by reducing poverty would also contribute to an improved situation, but such general measures seem to be more elusive than more concrete actions.

CONCLUSION

Overall, the study proposed and carried out a methodology to quantify socioeconomic differences in alcohol-attributable mortality, specifically for HIV/AIDS which can be adapted for other contexts. The metrics and data included were based almost exclusively on the most up-to-date data from large, original studies from South Africa—in most cases from the year 2012. High consistency was ensured by using the same measurements of alcohol use, SES and age groups throughout the calculations. The uncertainty around all reported estimations was assessed, reflecting the uncertainty in all metrics that went into the calculation. Despite limitations with regard to the assessment of exposure and the quantification of deaths in each SES group irrespective of alcohol use, the reported estimates are the most adequate given the available data. The study showed that alcohol use contributed to the socioeconomic differences in HIV/AIDS mortality with an about sevenfold mortality rate among persons of low SES compared with high SES in South Africa in 2012. As alcohol-attributable burden is in theory fully preventable, research and policy-makers should work together to address the high burden of alcohol-attributable HIV/AIDS mortality in persons of low SES in South Africa.
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Contributors CP had the overall responsibility of all steps, performed the statistical analyses and wrote the first draft. JR and CDHP supervised the whole working process and added specific knowledge on comparative risk assessments for alcohol use and on the literature about alcohol use and HIV in the South African context. JR contributed to the conceptualisation and to the methodology of the modelling process and the statistical analysis. All authors refined various drafts of the manuscript and approved the final version.

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