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# Association between social determinants of health and olfactory dysfunction in older adults: A population-based analysis

Eli Stein, BS<sup>1</sup>, Alexander Chern, MD<sup>2</sup>, Honglei Chen, MD, PhD<sup>3</sup>, Eric J. Shiroma, ScD<sup>4</sup>, D. P. Devanand, MD<sup>5</sup>, David A. Gudis, MD<sup>2</sup>, Jonathan B. Overdevest, MD, PhD<sup>2</sup> <sup>1</sup>Northwestern University Feinberg School of Medicine, Chicago, Illinois, USA

<sup>2</sup>Department of Otolaryngology—Head and Neck Surgery, Columbia University Vagelos College of Physicians and Surgeons, New York, New York, USA

<sup>3</sup>Department of Epidemiology and Biostatistics, College of Human Medicine, Michigan State University, East Lansing, Michigan, USA

<sup>4</sup>Laboratory of Epidemiology and Population Science, Intramural Research Program of the National Institutes of Health, National Institute on Aging, Baltimore, Maryland, USA

<sup>5</sup>Department of Psychiatry and Neurology, Columbia University Irving Medical Center, New York, New York, USA

### Abstract

**Background:** Social determinants of health (SDoH) are environmental conditions that influence health outcomes. As olfactory dysfunction (OD) in older individuals is associated with increased morbidity and mortality, we sought to investigate the impact of specific SDoH on olfactory function.

**Methods:** A cross-sectional analysis of the Health, Aging and Body Composition Study, a US population-based epidemiologic cohort study, was performed. Olfactory function was assessed utilizing both a self-report and a psychophysical olfactory test (CC-SIT test). Multivariable logistic regressions were performed to examine associations between specific SDoH with self-reported anosmia (sOD) and objective anosmia (oOD) as assessed by psychophysical testing. Differences in sensitivity and specificity were evaluated with sample tests for equality of proportions.

**Results:** Of 2219 participants, 13% had oOD and 18% had objective hyposmia; only 10% had sOD. Individuals identifying as Black race had higher odds of oOD (odds ratio [OR]:1.41, 95% confidence interval [CI]:1.02–1.95), while females and those reporting family incomes \$50,000 had lower odds of oOD (OR: 0.46, CI:0.34–0.62; OR:0.52, CI:0.29–0.93), adjusting for covariates. No specific SDoH was significantly associated with sOD. The sensitivity and specificity of sOD

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**Correspondence:** Jonathan B. Overdevest, MD, PhD, Rhinology & Skull Base Surgery, Department of Otolaryngology - HNS, Columbia University Irving Medical Center, New York Presbyterian, 180 Fort Washington Avenue, HP8-873, New York, NY 10032, USA. jo2566@cumc.columbia.edu.

CONFLICT OF INTEREST

None to disclose.

SUPPORTING INFORMATION

for oOD was 23.1% and 92.0%, respectively. sOD had greater sensitivity in females than males (30.8% vs. 18.8%, p = 0.030), while specificity varied significantly depending on family income (range: 90.0%-94.8%, p = 0.033).

**Conclusions:** Utilizing a large population-based study, we find disparities in the prevalence and self-recognition of OD among individuals of different gender, race, and income levels. Further effort is needed to evaluate factors propagating these disparities and to raise awareness of OD across all patient populations.

#### Keywords

anosmia; disparities; hyposmia; olfaction; social determinants of health

# 1 | INTRODUCTION

Olfactory dysfunction (OD), and particularly complete olfactory loss, is an important public health concern that emerges disproportionately among aging individuals. Studies suggest that up to 27.5% of elderly adults in the United States have some form of OD, and up to one-third of this population reports dissatisfaction with their sense of smell.<sup>1–3</sup> The health ramifications of OD are numerous, contributing to diminished appetite and malnutrition, injury due to failed perception of environmental hazards, reduced quality of life (QoL), and major adverse health outcomes.<sup>4–6</sup> Indeed, recent studies provide mounting evidence that OD may be a harbinger of dementia and Parkinson's disease and predict higher long-term morbidity and mortality.<sup>7–10</sup>

The US Department of Health and Human Services defines social determinants of health (SDoH) as "the conditions in the environments where people are born, live, learn, work, play, worship, and age that affect a wide range of health, functioning, and QoL outcomes and risks."<sup>11</sup> Racial and socioeconomic disparities have been found in outcomes of cancer, diabetes, and heart disease.<sup>12,13</sup> In a recent study, we identified five themes which can be used to categorize specific SDoH that have been found to be associated with OD: socioeconomic status (SES), education status, occupational exposures, racial/ethnic disparities, and lifestyle/behavioral factors.<sup>14</sup>

The complexity and cost of performing broad population-based olfaction testing has led to a deficiency of knowledge about risk factors for age related, noninfectious OD. While several prior studies have looked at a limited set of available population-based datasets on OD, these investigations have shortcomings in not exploring both subjective olfactory dysfunction (sOD) and oOD in the same cohort or not fully accounting for facets of SDoH previously identified to be associated with OD.<sup>15–18</sup> In this study, we expand upon a previous investigation by Dong et al., which utilized a pooled analysis of two community-based studies, and found that Black individuals had markedly higher odds of anosmia compared with White individuals in age- and sex-adjusted analyses.<sup>16</sup> Here, we systematically investigate the association between the more comprehensive five independent categories of SDoH we previously identified, and both sOD and oOD, using a US population-based epidemiologic cohort study of older adults. We also investigate whether there are disparities in the prevalence of hyposmia or a reduced ability to detect odor that

does not meet the criteria for anosmia. In addition, as previous studies have found a lack of correlation between subjective and objective measures of olfaction, we assess whether specific SDoH may account for differences in olfactory function perception and objective measurement.<sup>2,19</sup>

### 2 | METHODS

#### 2.1 | Study population

Participants were drawn from the Health, Aging and Body Composition (HealthABC) study.<sup>20,21</sup> Briefly, the HealthABC is an interdisciplinary study focused on risk factors for the decline of function in healthier older persons, particularly change in body composition with age.<sup>20</sup> From 1997 to 1998, 3075 subjects were recruited from a random sample of Medicare beneficiaries and age-eligible participants in two cities. As part of the larger study, inperson interviews and examinations occurred at years 1, 2, and 3, with phone calls alternating every 6 months. A total of 2535 participants completed both the subjective and objective olfactory test, of whom 316 reported having a cold in the week prior to the exam and were excluded from further analysis. This secondary data analysis did not require institutional review board (IRB) review due to the utilization of a widely available, deidentified, national dataset.

#### 2.2 | Olfactory function testing

Olfactory function was tested as part of the year 3 clinical visit utilizing the 12-item cross-cultural smell identification test (CC-SIT), a psychophysical screening test commonly referred to as the B-SIT (Sensonics; Haddon Heights, NJ), which is widely used in clinical and epidemiological studies.<sup>22</sup> One point was given for each correct answer with a total score ranging from 0 to 12. Psychophysical testing offers a reproducibly quantifiable description of olfactory performance and allows for evaluation of treatment response. As in prior studies, we treat it as a surrogate of objective assessment because we lack truly quantitative measurement capabilities.<sup>2,15,23</sup> We defined objective olfactory dysfunction (oOD) as a CC-SIT score 6, a threshold previously utilized to define clinically confirmed *anosmia* among patients.<sup>16</sup> We also analyzed the association between SDoH and *hyposmia*, using a more sensitive cutoff of CC-SIT score 8.<sup>24</sup> sOD was also assessed as part of the year 3 clinical visit survey and was defined as an affirmative answer to the question: "Do you suffer from smell and/or taste problems?"<sup>16</sup>

#### 2.3 | Social determinates of health

Demographic and lifestyle information was obtained via interviews during the year 1 clinical visit, unless otherwise noted.<sup>16</sup> {Dong, 2017 #14} Gender and race were dichotomously coded as male/female and White/Black, respectively. Education was categorized as "less than high school," "high school," and "above high school." Measures of SES included: family income, which was categorized as "less than \$10,000," "\$10,000 to \$25,000," ">\$25,000 up to <\$50,000," " \$50,000"; whether they possessed supplemental insurance to Medicare; and whether they had enough money for food. Alcohol drinking behaviors and smoking, assessed in year 3, were defined as current, former, or never.<sup>16</sup>

#### 2.4 | Potential confounding variables

Statistical analyses were adjusted for the following comorbidities which have been shown to be associated with OD. These data were obtained via in-clinic interviews and examinations during year 3. Participants were asked how often they "feel excessively (overly) sleepy during the day" with a response of "often (5-15 times/month)" or "almost always (16-30 times/month)" considered positive for daytime sleepiness.<sup>16</sup> Prevalence of depressive symptoms was defined as 10 on the 15-item Center for Epidemiologic Studies depression (CES-D) scale.<sup>16</sup> Prevalence of anxiety symptoms, previously found to be associated with OD,<sup>18</sup> was defined as a positive response to "During the past week, have you felt nervous or shaky inside?" or "During the past week, have you felt tense or keyed up?" Head injury was defined as an affirmative answer to "Have you ever been hit in the head hard enough to make you faint?"<sup>16</sup> Body mass index (BMI) was categorized as normal (<25 kg/m<sup>2</sup>), overweight (25–29.9 kg/m<sup>2</sup>), and obese ( 30 kg/m<sup>2</sup>). Self-reported general health status was defined as "excellent or very good," "good," and "fair or poor." Cognition was assessed using the modified Mini-Mental State Examination (3MSE), a 100-point expansion of the 30-point Mini-Mental State Examination, with scores of <80 indicating dementia.<sup>20,25</sup> Comorbid conditions such as Parkinson's disease,<sup>26</sup> pneumonia in the preceding 6 months,<sup>27</sup> cardiovascular disease (myocardial infarction or congestive heart failure), hypertension, diastolic and systolic blood pressure,<sup>28</sup> cerebrovascular disease,<sup>29</sup> and diabetes mellitus<sup>30</sup> were also reported based on self-report.

#### 2.5 | Statistical analysis

Descriptive statistics were reported as median with interquartile range for continuous variables and as number and percent for categorical variables. Univariable logistic regressions were performed, separately for each SDoH, to identify factors associated with the presence of either sOD or oOD and were reported as an odds ratio (OR) with 95% confidence interval (CI) and p-value. All aforementioned SDoH independent variables, along with confounding variables that had a p < 0.2 on univariable regressions, were included in a stepwise forward multivariable logistic model. For ease of interpretation, diastolic and systolic blood pressure were converted to z-scores. Sensitivity, specificity, positive predictive value (PPV), and negative predictive value (NPV) of self-awareness of olfactory loss in self-reported sOD in evaluating for oOD were calculated both in the entire cohort and in subgroups of race, gender, education, and income. Differences in CC-SIT scores, sensitivity, specificity, PPV, and NPV by race, gender, education, and incomes were calculated and tested for statistical significance using sample tests without continuity correction, Wilcoxon rank sum tests, and Dunn's Kruskal-Wallis tests, as appropriate, and p-values were adjusted with the Benjamini-Hochberg method for multiple comparisons.<sup>31</sup> Missing values (<1.1% of total datapoints) were imputed utilizing multivariate imputation by chained equations.<sup>32</sup> Statistical analyses were performed using R, Version 4.0.5 (R Foundation for Statistical Computing, Vienna, Austria).

#### 3 | RESULTS

A total of 2219 individuals completed both the self-report olfactory awareness assessment and objective olfactory CC-SIT test and were included in our study. Fifty-two percent (n =

1164) of participants were female, 63% (n = 1,402) of participants self-described as White race, and the median age at the time of olfactory testing was 75 (interquartile range [IQR]: 73–78) (Table 1). Among all participants, the median CC-SIT score was 10 (IRQ: 8–11), 13% (n = 290) had anosmia or oOD with an additional 18% (n = 392) qualifying as having objective hyposmia, and 10% (n = 221) had sOD on questionnaire.

The factors associated with the prevalence of sOD differed from those associated with oOD. SDoH factors, including race, education, income, smoking or alcohol consumption, were not individually associated with sOD on unadjusted logistic analysis (Table 2). However, other participant factors, including anxiety symptoms (OR: 1.35, p = 0.038), a history of hitting one's head hard enough to faint (OR: 1.95, p < 0.001), and cerebrovascular disease (OR: 1.59, p = 0.033), were all associated with higher odds of sOD. In our complete multivariable model, none of the SDoH we investigated were associated with sOD, although geographic location of clinical testing site, history of hitting one's head hard enough to faint, and elevated systolic blood pressure were all significantly associated with sOD (Table 3).

When analyzing factors associated with oOD, female gender, increased educational attainment, and higher family income were all associated with lower odds of oOD on unadjusted analysis. Black race, status as a former drinker as opposed to never drinking alcohol, and being a current or former smoker as opposed to never smoking were all associated with higher odds of oOD on unadjusted analysis (Table 2). On stepwise multivariable logistic regression, we found that individuals identifying as Black race had higher odds of oOD (OR:1.41, 95% CI: 1.02-1.95), while females and those reporting family incomes \$50,000 had lower odds of oOD (OR:0.46, CI:0.34-0.62; OR:0.52, CI:0.29-0.93) even after considering all confounding variables (Table 4). Education, smoking status, and alcohol status did not maintain significance after accounting for all covariates. We also found that higher diastolic blood pressure (OR:1.19; CI:1.03-1.39) and cognitive impairment (OR:2.73; CI:1.86–3.98) were both associated with oOD. Liberalizing the definition of oOD to include both anosmia and hyposmia (CC-SIT score 8) expanded the prevalence of OD to 30.7% within this population. When comparing those with hyposmia with individuals with intact olfactory function, we found that female gender and higher income were associated with lower odds of hyposmia, while education and race did not maintain significance after accounting for all covariates (Table S1 and Table S2).

When comparing sOD with oOD within the entire cohort, we found a self-reported perception of sOD had a low sensitivity (23.1%) and PPV (30.3%) for positive oOD, while having a better specificity (92.0%) and NPV (88.8%) for negative oOD (Table 5). We explored whether specific SDoH may mediate the relationship between sOD and oOD, and if the association between the two would be strengthened within certain subgroups. We found that self-perception of olfactory function had greater sensitivity in females than males (30.8% vs. 18.8%, p = 0.030), while the specificity significantly varied depending on family income (p = 0.033). We also found that the NPV of sOD for oOD varied between subgroups of gender, race, education, and income, ranging from as low as 81.8% to as high as 93.1% depending on which population we were testing in (Table 5). Evaluating the entire cohort, we found that our objective measure of OD was associated with sOD, as participants who endorsed sOD performed worse on the CC-SIT test than those who

reported normal olfaction (CC-SIT score = 8 vs. 10, p < 0.001). We also found that CC-SIT scores were associated with sOD when divided into subgroups based on gender, race, and education; however, it was not related to sOD in those with family incomes below \$10k (Figure 1A–D). Lastly, when looking at only patients without sOD, we found that males, Black race, those with less than high school educational attainment, and those with family income <\$10,000 all had significantly lower CC-SIT scores than other subgroups within gender, race, education, and family income (Figure 2A–D).

### 4 | DISCUSSION

Utilizing data from 2219 participants in the HealthABC study, we found that 13% of individuals had objective absence of smell or anosmia (oOD) and 18% had objectively reduced sense of smell (hyposmia), compared with only 10%, who self-reported subjective olfactory impairment (sOD). When assessing risk factors for oOD, gender, race, and income remained associated with oOD even after accounting for comorbidities, while education, smoking status, and alcohol status were associated with oOD on univariable analysis but not after accounting for comorbidities. When looking at risk factors for sOD, we did not find that gender, race, education, income, smoking status, or alcohol status were associated with sOD on univariate analysis or on multivariate analysis after considering comorbidities. While participants who self-reported sOD performed worse on the CC-SIT than those who reported normal olfaction, self-reports showed a very poor sensitivity (23.1%) for identifying individuals with oOD. The NPV of sOD for oOD varied significantly between subgroups of gender, race, education, and family income. In addition, when looking at individuals who did not self-report OD, males, individuals of Black race, individuals who did not complete high school, and individuals with lower family income all had lower CC-SIT scores than females, White race, high school graduates, and higher income families, respectively. These results necessitate further efforts to evaluate factors propagating these disparities and compel physicians to consider differences in self-reporting when interpreting a negative response to a subjective olfactory screen.

Evaluating our results using the previously reported thematic construct for SDoH, such as racial/ethnic disparities, SES, education status, occupational exposures, and lifestyle/ behavioral factors, we found that race, similar to results from other studies, was an important predictor of OD. Individuals reporting a Black racial identity had greater odds of oOD than those reporting White race, even when accounting for comorbidities (OR: 1.41, p = 0.037).<sup>16–18</sup> The complexity of race as a construct challenges this interpretation, where rather than a true biologic or genetic relation to OD, race is more likely a proxy for different occupational exposures, life experience, and conditions that we are not capturing in our limited dataset. For example, prior studies have found that Black race, people at lower educational levels, and people at lower income levels were significantly more likely to live within a mile of a polluting facility—an exposure that can damage olfactory capabilities.<sup>33</sup> Unlike in prior studies, we did not find that race was associated with elevated rates of self-reported poor olfaction.<sup>15,17</sup>

We find mixed results when exploring the role of SES in OD. In multivariate analysis, those with family incomes >\$50,000 were approximately half as likely to have oOD than those

with families with family income <\$10,000 (OR = 0.52, p = 0.027). Similarly, in an analysis of the Beaver Dam Offspring Study, the authors found that an income of >\$50,000/year was associated with less OD (OR = 0.48).<sup>34</sup> We did not, however, find an association between SES and sOD. Patients with lower SES may be more at risk for true OD due to a myriad of factors, such as living in areas with greater air pollution or having reduced access to healthcare that may otherwise delay the deterioration of their condition.<sup>33,35</sup>

Similarly, we found the relationship between educational attainment and OD to be inconsistent. There was no apparent association between education and sOD in this population. After adjusting for covariates, those with less than a high school degree were not more likely to have oOD than those with a high school degree. Importantly, the HealthABC utilizes the CC-SIT, a psychophysical olfactory assessment using only odorant identification to stratify olfactory performance. This contrasts with other tests which assess olfactory threshold, the minimum concentration at which an odor can be detected by participants, and discrimination, the ability of a participant to discern between two odorants. Previous studies have found that odor identification test are more susceptible to cognitive and neurologic impairments, with patients with Alzheimer's disease and Parkinson's disease found to do worse on odor identification than odor threshold tests.<sup>36,37</sup> When interpreting the effects of race, education, and SES on oOD in our study, responses and perceptions of odorants used in the CC-SIT may be attributable to differences in cultural norms or life experience, thus leading to lower odor identification rather than worse underlying disease.<sup>38</sup> This highlights the importance of continuing to develop objective measures of disease that perform consistently across all populations.

Our findings that a large proportion of individuals with objective olfactory deficits did not self-report a diminished sense of smell is in-line with a recent review which found that OD prevalence was significantly greater when measured by objective olfactory assessments compared with subjective measures (28.8% vs. 9.5%, p < 0.001).<sup>2</sup> Other studies have found discordance between self-assessed measures of chemosensation and objective findings. In an analysis of the National Social Life, Health, and Aging Project, the authors found that 74.2% of older adults with measured OD did not recognize it.<sup>39</sup> In a recent analysis of the National Health and Nutrition Examination Survey, the authors found that increased age was associated with underreporting of olfactory impairment, though the analysis did not reveal how other demographic characteristics affect the sensitivity of self-report.<sup>40</sup> In another recent analysis of the Sister Study, a study of middle-aged and older women, the authors found that self-reports showed a low sensitivity (22.6%) for poor olfaction.<sup>15</sup> Here, we found that the sensitivity of sOD for oOD is 23.1%, with a sensitivity as low as 16.8% in those with less than a high school degree. Moreover, we found that the NPV of sOD for oOD—whether a patient reports no OD and truly has no olfactory issues—varied significantly based on gender, race, education, and income. In total, these results suggest that physicians should raise awareness of OD across all patient populations while strongly considering the use of objective olfactory testing to evaluate patients where there exists suspicion for OD.

In contrast to prior studies, in the HealthABC population we did not find that gender, race, education, SES, or behaviors such as smoking or alcohol consumption were associated with

differences in sOD, though we did find differences in self-awareness and reporting accuracy depending on gender, race, education, and SES.<sup>15,17</sup> Direct comparison across studies is difficult due to differences in study population and survey formulation; for example, the HealthABC asks participants whether they have smell or taste *problems*, and not if they have a *decrease* in smell as other surveys do.<sup>15</sup> Overall, we postulate that the accuracy of self-report of OD for objective disease is poor, as people interpret the question differently and disparities may exist in awareness and education of sensory deficits across various populations.

Our analysis has several limitations. Due to the study design of the dataset using an affirmative answer to the question: "Do you suffer from smell and/or taste problems?" to identify sOD, there remains ambiguity as to whether the participant's response is driven by olfactory or gustatory dysfunction. Our use of this question was based on prior analyses of this dataset,<sup>16</sup> and even when using this question encompassing both smell and taste, self-report of chemosensory dysfunction is lower than that uncovered by psychophysical testing. Moreover, prior studies have found that individuals often self-report codysfunction of these two senses, and individuals who do not report taste dysfunction rarely report OD.<sup>41</sup> Secondly, the variables provided within the HealthABC dataset, while comprehensive, may not fully account for all confounders. For example, the database lacks questions about vocation, thereby limiting our analysis into the association between OD and occupational exposure risks, a previously identified category of SDoH.<sup>14</sup> Additionally, the dataset does not include diseases, such as chronic rhinosinusitis (CRS), and environmental exposures, such as lead, that have previously been found to be associated with OD and might exist as underlying confounding factors; this is especially true considering previously published studies that have found disparities in care for CRS and that lower SES was associated with a higher CRS rate.<sup>42–44</sup> Lastly, the HealthABC cohort comprised only elderly adults above the age of 70(median age: 75, IQR:73–78) limiting its generalizability to younger patients. Nonetheless, the known association between OD and morbidity and mortality in elderly adults necessitates further investigation into a population with a disproportionate burden of noninfectious OD. A recent meta-analysis found that in studies with a mean age >55 years, the prevalence of OD was 34.5%, compared with an OD prevalence rate of 7.5% in studies with a mean age <55 years.<sup>2</sup> The association between specific SDoH and OD remains underexplored across all ages due to the hurdles to widespread olfactory testing in population studies, and we believe this study provides valuable insight into the disparities in olfactory function within older adults.

# 5 | CONCLUSION

While our study reports no association between SDoH and awareness of sOD, certain SDoH factors may impact olfactory function when measured using the CC-SIT psychophysical olfactory test (oOD). We also found that the accuracy of self-reports of sOD for oOD varies based on gender, race, education, and income. As OD is associated with morbidity and mortality, it is imperative that we make an effort to further evaluate factors propagating these disparities while providing more equitable care for our patients.

# **Supplementary Material**

Refer to Web version on PubMed Central for supplementary material.

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#### FIGURE 1.

Comparison of cross-cultural smell identification test (CC-SIT) scores between those with and without subjective olfactory dysfunction, within subgroups of gender (A), race (B), education (C), and family income (D). *p*-value from Wilcoxon rank sum test. ns = not significant (p = 0.05). \*< 0.05, \*\*< 0.01, \*\*\*< 0.001

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#### FIGURE 2.

Among those without subjective olfactory dysfunction, comparison of cross-cultural smell identification test (CC-SIT) scores within subgroups of gender (A), race (B), education (C), and family income (D). Red diamond represents the mean. Panels A and B: Wilcoxon p < 0.001 for both. Panels C and D: Kruskal–Wallis p < 0.001 for both

Patient characteristics and distribution of variables

Characteristic	<i>N</i> = 2,219
Site	
Memphis	1,082 (49%)
Pittsburgh	1,137 (51%)
Gender	
Male	1,055 (48%)
Female	1,164 (52%)
Race	
White	1,402 (63%)
Black	817 (37%)
Age at year 3 clinic visit	
70–74	921 (42%)
75–79	1,050 (47%)
80–84	248 (11%)
Education	
Less than high school	479 (22%)
High school graduate	744 (34%)
Post-secondary schooling	996 (45%)
Alcohol consumption	
Never	629 (28%)
Current	1,141 (51%)
Former	449 (20%)
Smoking status	
Never	1,019 (46%)
Former	1,044 (47%)
Current	156 (7.0%)
Family income	
<\$10,000	262 (12%)
\$10,000 to \$25,000	822 (37%)
> \$25,000 up to \$50,000	750 (34%)
\$50,000	385 (17%)
Supplemental insurance to Medicare	1,850 (83%)
Enough money for food	2,039 (92%)
Primary care received at	
Private doctor's office	1,891 (85%)
Public clinic	96 (4.3%)
Health maintenance organization	147 (6.6%)
Hospital outpatient clinic	85 (3.8%)
Daytime sleepiness	293 (13%)
Depressive symptoms	249 (11%)

Characteristic	<i>N</i> = 2,219
Anxiety symptoms	735 (33%)
Hit head hard enough to faint	211 (9.5%)
Parkinson's disease	12 (0.5%)
BMI	
<25	749 (34%)
25–29.9	934 (42%)
>30	536 (24%)
Avg sitting systolic BP, mm Hg	134.00 (120.00, 148.00)
Avg sitting diastolic BP, mm Hg	70.00 (64.00, 78.00)
Self-reported health	
Excellent/very good	1,003 (45%)
Good	840 (38%)
Fair/poor	376 (17%)
Cognitive impairment	235 (11%)
Pneumonia, preceding 6 months	15 (0.7%)
MI/heart disease	494 (22%)
Cerebrovascular disease	195 (8.8%)
Hypertension	1,251 (56%)
Diabetes mellitus	368 (17%)
CC-SIT score	10.00 (8.00, 11.00)
Objective olfactory dysfunction (anosmia)	290 (13%)
Subjective olfactory dysfunction	221 (10.0%)
Hyposmia or anosmia	682 (30.7%)

*Note*: *n* (%); Median (interquartile range [IQR]). Anosmia (objective olfactory dysfunction) defined as CC-SIT score 6. Hyposmia defined as CC-SIT score 8.

Abbreviations: BMI, body mass index; BP, blood pressure; CC-SIT, cross-cultural smell identification test; MI, myocardial infarction.

Univariable logistic regression analysis of variables associated with subjective olfactory dysfunction (left panel) and objective olfactory dysfunction (right panel)

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	Subjectiv	e olfactory dysfunc	ction (sOD), $n = 221$	Objective	olfactory dysfunct	tion (0OD) $n = 290$
Characteristic	OR	95% CI	<i>p</i> -value	OR	95% CI	<i>p</i> -Value
Gender						
Male	I					
Female	1.11	0.84, 1.47	0.472	0.46	0.35, 0.59	<0.001
Race						
White						
Black	0.99	0.74, 1.32	0.957	1.81	1.41, 2.32	<0.001
Age at year 3 clinic visit						
70–74	I					
75–79	1.14	0.85, 1.54	0.392	1.58	1.20, 2.09	0.001
80–84	1.2	0.75, 1.88	0.432	2.51	1.72, 3.65	<0.001
Education						
Less than high school						
High school graduate	0.99	0.67, 1.49	0.978	0.49	0.36, 0.68	<0.001
Post-secondary schooling	1.21	0.85, 1.77	0.3	0.52	0.39, 0.70	<0.001
Alcohol consumption						
Never						
Current	1.2	0.86, 1.69	0.279	1.07	0.79, 1.45	0.681
Former	1.25	0.83, 1.88	0.285	1.58	1.12, 2.23	0.01
Smoking status						
Never						
Former	1.17	0.88, 1.57	0.286	1.35	1.04, 1.75	0.026
Current	1.12	0.62, 1.92	0.681	1.68	1.04, 2.62	0.027
Family income						
<\$10,000						
\$10,000 to \$25,000	0.72	0.46, 1.13	0.141	0.62	0.43, 0.89	0.00
> \$25,000 up to \$50,000	1	0.66, 1.57	0.988	0.52	0.36, 0.76	<0.001

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	Subjective	olfactory dysfunc	tion (sOD), $n = 221$	Objective	olfactory dysfuncti	(00 (00D) $n = 290$
Characteristic	OR	95% CI	<i>p</i> -value	OR	95% CI	<i>p</i> -Value
\$50,000	0.61	0.35, 1.04	0.066	0.44	0.28, 0.69	<0.001
Spirituality/religion						
Not important						
Important	0.7	0.39, 1.37	0.26	1.2	0.64, 2.48	0.601
Site						
Memphis						
Pittsburgh	0.68	0.51, 0.90	0.007	0.98	0.76, 1.25	0.841
BMI						
<25						
25-29.9	0.8	0.58, 1.11	0.179	0.95	0.72, 1.25	0.706
>30	0.88	0.61, 1.26	0.493	0.77	0.55, 1.08	0.136
Avg sitting systolic BP, z-score	0.85	0.73, 0.98	0.026	0.96	0.85, 1.09	0.529
Avg sitting diastolic BP, z-score	0.97	0.85, 1.12	0.698	1.12	0.99, 1.26	0.076
Self-reported health						
Excellent/very good						
Good	1.23	0.90, 1.68	0.187	1.15	0.87, 1.52	0.332
Fair/poor	1.38	0.93, 2.01	0.101	1.8	1.30, 2.49	<0.001
Daytime sleepiness	1.22	0.82, 1.78	0.314	1.1	0.76, 1.55	0.615
Depressive symptoms	1.45	0.96, 2.12	0.067	1.18	0.80, 1.70	0.374
Anxiety symptoms	1.35	1.01, 1.80	0.038	0.91	0.70, 1.19	0.499
Hit head hard enough to faint	1.95	1.30, 2.85	<0.001	1.07	0.69, 1.59	0.76
Parkinson's disease	1.82	0.28, 6.94	0.443	3.36	0.89, 10.7	0.049
Cognitive impairment	0.83	0.50, 1.30	0.433	3.86	2.82, 5.24	<0.001
Pneumonia, preceding 6 months	1.39	0.22, 5.09	0.663	0.47	0.03, 2.37	0.47
MI/heart disease	0.94	0.66, 1.30	0.708	0.88	0.64, 1.18	0.4
Cerebrovascular disease	1.59	1.02, 2.40	0.033	1.03	0.65, 1.55	0.909
Hypertension	0.93	0.70, 1.23	0.608	0.82	0.64, 1.05	0.113
Diabetes mellitus	1.24	0.86, 1.76	0.227	1.18	0.85, 1.61	0.318
Abbreviations: BMI, body mass inc	lex; BP, bloo	d pressure; CI, conf	idence interval; MI, n	nyocardial in	farction OR, odds ra	ttio.

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Multivariable logistic regression models of variables associated with subjective olfactory dysfunction

0.99 (0.71, 1.37), 0.959 1.00 (0.71, 1.39), 0.994 1.13 (0.79, 1.61), 0.499 1.16 (0.85, 1.58), 0.360 1.29 (0.89, 1.88), 0.190 1.29 (0.83, 2.01), 0.258 1.14 (0.83, 1.59), 0.416 1.23 (0.89, 1.70), 0.207 1.34 (0.82, 2.13), 0.229 1.09 (0.70, 1.72), 0.706 1.33 (0.85, 2.11), 0.214 0.72 (0.45, 1.18), 0.184 1.06 (0.64, 1.79), 0.834 0.58 (0.31, 1.10), 0.096 0.99 (0.53, 1.74), 0.971 Model 5 1.00 (0.66, 1.53), 0.990 1.00 (0.66, 1.54), 0.988 1.15 (0.85, 1.56), 0.366 1.16 (0.85, 1.60), 0.346 1.09 (0.82, 1.46), 0.554 1.19 (0.87, 1.61), 0.276 0.70 (0.44, 1.13), 0.135 1.21 (0.84, 1.75), 0.318 1.24 (0.77, 1.94), 0.367 1.30 (0.86, 2.01), 0.222 0.93 (0.56, 1.54), 0.762 0.52 (0.28, 0.97), 0.041 1.25 (0.81, 1.92), 0.316 1.09 (0.60, 1.90), 0.761 Subjective olfactory dysfunction (n = 221) - odds ratio (95% confidence interval), p-value Model 4 1.14 (0.84, 1.54), 0.393 0.70 (0.44, 1.13), 0.136 1.31 (0.87, 2.01), 0.204 1.22 (0.76, 1.91), 0.401 0.93 (0.57, 1.55), 0.774 0.53 (0.28, 0.98), 0.042 Model 3 1.12 (0.84, 1.48), 0.443 1.14 (0.86, 1.51), 0.380 0.99 (0.74, 1.32), 0.939 1.04 (0.76, 1.42), 0.790 0.99 (0.66, 1.50), 0.956 1.15 (0.85, 1.55), 0.375 1.20 (0.75, 1.88), 0.430 1.23 (0.84, 1.84), 0.292 Model 2 1.14 (0.85, 1.55), 0.376 1.21 (0.75, 1.88), 0.424 Model 1 High school graduate Less than high school Alcohol consumption \$10,000 to \$25,000 \$25,000 to \$50,000 Characteristic Smoking status Family income Some college \$50,000 Education <\$10,000 Current Former Former Current Gender Female White 70-74 75-79 80 - 84Never Never Black Male Race Age

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Characteristic	Model 1	Model 2	Model 3	Model 4	Model 5
Site					
Memphis					Ι
Pittsburgh					0.66(0.49,0.89),0.006
BMI					
<b>25</b>					Ι
25-29.9					$0.82\ (0.59,1.15),0.25I$
>30					$0.88\ (0.59,1.30),0.526$
Systolic BP					$0.83\ (0.70,\ 0.99),\ 0.042$
Diastolic BP					$1.08\ (0.92,1.29),0.344$
Self-reported health					
Excellent/very good					
Good					1.21 (0.88, 1.68), 0.240
∃air/poor					1.32 (0.85, 2.03), 0.214
Depressive symptoms					$1.28\ (0.81,\ 1.98),\ 0.275$
Anxiety symptoms					$1.18\ (0.85,1.61),0.316$
Hit head hard					1.97 (1.29, 2.94), 0.001
Parkinson's disease					1.40(0.21, 5.67), 0.674
Cognitive impairment					0.73 (0.41, 1.25), 0.271
Hypertension					$0.92\ (0.68,1.25),0.602$
Cerebrovascular disease					1.54(0.97, 2.36), 0.057

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Note: Model 1: Gender + race + age; Model 2: Model 1 + education; Model 2 + socioeconomic status; Model 4: Model 3 + lifestyle/behavioral factors; Model 5: Model 4 + comorbidities. Abbreviations: BMI, body mass index; BP, blood pressure.

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Multivariable logistic regression models of variables associated with objective olfactory dysfunction

	Objective olfactory dysfu	nction $(n = 290)$ - odds rati	o (95% confidence interva	l), <i>p</i> -value	
Characteristic	Model 1	Model 2	Model 3	Model 4	Model 5
Gender					
Male					
Female	0.42 (0.32, 0.55), <0.001	0.43 (0.33, 0.56), <0.001	0.39 (0.30, 0.52), <0.001	0.41 (0.31, 0.55), <0.001	0.46 (0.34, 0.62), <0.001
Race					
White		I	I		I
Black	2.13 (1.65, 2.76), <0.001	1.85 (1.40, 2.44), <0.001	1.64 (1.22, 2.20), <0.001	1.61 (1.20, 2.17), <0.001	1.41 (1.02, 1.95), <0.001
Age					
70–74		I	I		I
75–79	1.59 (1.20, 2.11), 0.001	1.60 (1.20, 2.13), 0.001	1.56 (1.18, 2.09), 0.002	1.58 (1.19, 2.11), 0.002	1.54 (1.15, 2.07), 0.004
80-84	2.70 (1.83, 3.96), <0.001	2.75 (1.86, 4.03), <0.001	2.72 (1.84, 4.00), <0.001	2.77 (1.87, 4.09), <0.001	2.57 (1.71, 3.84), <0.001
Education					
Less than high school					
High school graduate		0.62 (0.44, 0.87), 0.006	0.68 (0.48, 0.97), 0.031	0.68 (0.48, 0.97), 0.032	0.90 (0.62, 1.33), 0.604
Some college		0.63 (0.46, 0.88), 0.006	0.75 (0.52, 1.07), 0.108	0.76 (0.53, 1.09), 0.132	1.03 (0.69, 1.54), 0.891
Family income					
<\$10,000			I		I
\$10,000 to \$25,000			0.64 (0.43, 0.95), 0.025	0.65 (0.44, 0.97), 0.032	$0.70\ (0.47,1.06),0.086$
\$25,000 to \$50,000			0.58 (0.37, 0.91), 0.018	0.59 (0.38, 0.93), 0.023	$0.64 \ (0.40, 1.03), 0.062$
\$50,000			0.47 (0.27, 0.81), 0.007	0.48 (0.28, 0.84), 0.011	0.52 ( $0.29$ , $0.93$ ), $0.027$
Alcohol consumption					
Never					
Current				$1.03\ (0.74, 1.46), 0.852$	1.05 (0.74, 1.50), 0.769
Former				$1.15\ (0.79, 1.68), 0.456$	$1.20\ (0.81,\ 1.77),\ 0.362$
Smoking status					
Never					
Former				1.08 (0.80, 1.44), 0.624	1.12 (0.83, 1.52), 0.445
Current				1.31 (0.79, 2.11), 0.279	1.13 (0.67, 1.87), 0.633

Characteristic	Model 1	Model 2	Model 3	Model 4	Model 5
Site					
Memphis					Ι
Pittsburgh					1.10(0.84,1.45),0.480
BMI					
<25					Ι
25–29.9					0.94 (0.70, 1.28), 0.701
>30					$0.79\ (0.54,1.14),0.205$
Systolic BP					0.87 (0.74, 1.02), 0.101
Diastolic BP					1.19(1.03, 1.39), 0.023
Self-reported health					
Excellent/very good					
Good					1.13(0.84, 1.53), 0.421
Fair/poor					1.42 (0.97, 2.07), 0.073
Depressive symptoms					$0.87\ (0.56,1.33),0.533$
Anxiety symptoms					0.94 (0.69, 1.27), 0.690
Hit head hard					1.01 (0.64, 1.55), 0.963
Parkinson's disease					3.38 (0.87, 11.4), 0.057
Cognitive impairment					2.73 (1.86, 3.98), <0.001
Hypertension					0.77 (0.58, 1.01), 0.063
Cerebrovascular disease					0 00 (0 52 1 45) 0 235

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Note: Model 1: Gender + race + age; Model 2: Model 1 + education; Model 2 + socioeconomic status; Model 4: Model 3 + lifestyle/behavioral factors; Model 5: Model 4 + comorbidities. Abbreviations: BMI, body mass index; BP, blood pressure.

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Sensitivity, specificity, positive predictive value (PPV), and negative predictive value of self-reported olfactory dysfunction (subjective OD) for objective OD

	Sensitivity	d	Specificity	d	γqq	d	NPV	d
Entire cohort	23.1%	Ι	92.0%	I	30.3%	I	88.8%	I
Gender		0.030		0.513		0.219		<0.001
Male	18.8%		92.5%		35.0%		84.2%	
Female	30.8%		91.6%		26.4%		93.1%	
Race		0.479		0.684		0.133		<0.001
White	25.2%		91.8%		26.4%		91.3%	
Black	21.0%		92.4%		37.0%		84.6%	
Education		0.115		0.635		0.539		<0.001
Less than high school	16.8%		92.7%		36.4%		81.8%	
High school graduate	22.2%		92.5%		26.5%		90.7%	
Post-secondary schooling	28.9%		91.4%		30.3%		90.9%	
Family income		0.429		0.033		0.723		<0.001
<\$10,000	18.9%		%0.0%		32.3%		81.4%	
\$10,000 to \$25,000	19.8%		93.0%		30.6%		88.1%	
\$25,000 to \$50,000	27.6%		90.2%		27.0%		90.5%	
\$50,000	28.2%		94.8%		37.9%		92.1%	

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Note: p-value from sample test of equality of proportions.