





Frailty Among Sexual and Gender Minority Older Adults: The All of Us Database

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Abstract

Background: Despite known disparities in health status among older sexual and gender minority adults (OSGM), the prevalence of frailty is unknown. The aim of this study was to develop and validate a deficit-accumulation frailty index (AoU-FI) for the All of Us database to describe and compare frailty between OSGM and non-OSGM participants.

Methods: Developed using a standardized approach, the AoU-FI consists of 33 deficits from baseline survey responses of adults aged 50+. OSGM were self-reported as “not straight” or as having discordant gender and sex assigned at birth. Descriptive statistics characterized the AoU-FI. Regression was used to assess the association between frailty, age, and gender. Validation of the AoU-FI used Cox proportional hazard models to test the association between frailty categories (robust <0.15, 0.15 ≤ pre-frail ≤ 0.25, frail >0.25) and mortality.

Results: There were 9 110 OSGM and 67 420 non-OSGM with sufficient data to calculate AoU-FI; 41% OSGM versus 50% non-OSGM were robust, whereas 34% versus 32% were pre-frail, and 26% versus 19% were frail. Mean AoU-FI was 0.19 (95% confidence interval [CI]: 0.187, 0.191) for OSGM and 0.168 (95% CI: 0.167, 0.169) for non-OSGM. Compared to robust, odds of mortality were higher among frail OSGM (odds ratio [OR] 6.40; 95% CI: 1.84, 22.23) and non-OSGM (OR 3.96; 95% CI: 2.96, 5.29).

Conclusions: The AoU-FI identified a higher burden of frailty, increased risk of mortality, and an attenuated impact of age on frailty among OSGM compared to non-OSGM. Future work is needed to understand how frailty affects the OSGM population.

Keywords: Diversity in Aging, Frailty, LGBTQIA+

In the United States, there are approximately 3 million older sexual and gender minority adults (OSGM), and these numbers are projected to grow to over 5 million by 2030 (1). OSGM face a higher burden of chronic conditions (2–4), mental health issues (2,3,5), subjective cognitive decline (6,7), and health risk behaviors (3,4). These disparities are understood through the minority stress theory that describes that external stressors (eg, stigma and discrimination related to sexual orientation) become internalized, resulting in negative feelings about identity, a need to conceal one's identity, and expectations of rejection in future interactions. These specific stressors accumulate over everyday stress (8). The minority stress theory has been extended to include the unique experiences of people who are members of gender-minoritized populations (9), individuals with multiple minoritized identities (eg, sexual, gender, and racial) (10), and aging (11). The

current generation of OSGM experienced significant events specific to their minority identity, including homosexuality being labeled as a mental illness; the Lavender Scare where there was mass dismissal of SGM workers from U.S. government employment; and the human immunodeficiency virus infection and acquired immunodeficiency syndrome epidemic. Minority stress has been linked to negative health consequences including changes in inflammation, immune, and endocrine function, which contribute to poor cardiovascular, metabolic, and immunologic clinical outcomes (8,12,13) and potentially frailty.

The deficit-accumulation frailty index (FI) is one of 2 prevailing theories of frailty measurement (14,15). FI captures the aggregate burden of age-related health deficits and represents a multidimensional risk state (15). Moreover, the FI is a comprehensive measure that integrates physical and mental

health conditions more common among OSGM. However, a lack of available data precludes any systematic characterization of frailty in an OSGM population. The NIH-funded All of Us (AoU) Research Program is well positioned to address this critical need by using multimodal outreach to recruit a diverse cohort of participants who have been historically underrepresented in biomedical research, including OSGM (16). In the present study, we used the AoU database to study frailty in OSGM with 2 goals: First, to develop and validate an FI that can readily be applied to AoU; and second, to estimate the prevalence of frailty among OSGM and compare the burden of frailty in OSGM to a non-SGM older adult (non-OSGM) population. We hypothesized that frailty would be higher in OSGM compared to non-OSGM individuals.

Method

Population

The AoU Program goals, scientific rationale, recruitment methods, and sites have been described previously (16). In brief, eligible participants included U.S. residents over the age of 18, not incarcerated at the time of enrollment, and able to provide informed consent. More than 75% of participants are from populations underrepresented in biomedical research, including 50% from racial and ethnic minority groups. Participants volunteer to complete health surveys, authorize sharing of electronic health records (EHR), share mobile health data (eg Fitbit, Apple HealthKit), and are invited to an in-person visit to contribute biospecimen samples and undergo physical assessment. Structured EHR data are transferred from enrolling sites at least once per quarter (17). All experimental protocols and data collection involving human participants were approved by the Ethics Committee/Institutional Review Board of the AoU Institutional Review Board. The current study was reviewed and deemed exempt by the Northeastern University Institutional Review Board.

We used AoU Version 6 Curated Data Repositories (CDR) of the Registered Tier data set (updated on June 23, 2022, CDR Hotfix for v6 CDRs released January 13, 2023), which has undergone a series of data transformations to protect participant privacy and has been mapped to the Observational Medical Outcomes Partnership common data model (18). At the time of analysis, over 370 000 community-dwelling people in the United States had participated in AoU.

We restricted our older adult cohort to those aged 50 years or older, as conditions of aging have been shown to have higher prevalence at relatively younger ages among other stigmatized populations, such as people living with HIV (19). Data from 5 baseline surveys were used to develop the FI: “The Basics” (participant demographics), “Overall Health” (levels of individual health), “Lifestyle” (health risk behaviors), “Personal Medical History” (medical conditions), and “Health Care Access and Utilization” (access and use of health care).

Identifying Older Sexual and Gender Minority Participants

Self-reported sex assigned at birth was assessed in “The Basics” survey with the question “What was your biological sex assigned at birth?” (responses “Male | Female | Intersex | None of these describe me | Prefer not to answer”), and gender was assessed with “What terms best express how you describe your gender identity?” (responses “Man |

Woman | Non-binary | Transgender | None of these describe me and I've like to consider additional options | Prefer not to answer”). In the Registered Tier Data Set, responses for both sex and gender were aggregated by AoU administrators for participant privacy into “Male | Female | Not male/Not female/prefer not to answer/skip.” Sexual orientation was assessed by the question “Which of the following best represents how you think of yourself?” responses “Gay | Lesbian | Straight | Bisexual | None of these describe me and I'd like to see additional options | Prefer not to answer” were aggregated by AoU administrators into “Straight” or “Not Straight.” We divided participants into 2 mutually exclusive groups: OSGM and non-OSGM, based on concordance between sex assigned at birth, gender, and “straight” status (Table 1). For example, anyone reporting male sex assigned at birth, male gender, and “straight” was considered non-OSGM. However, an individual reporting male sex assigned at birth, male gender, and “not straight” was considered an OSGM man. Anyone reporting male or female sex assigned at birth and gender as not male/not female/prefer not to answer/skip was considered OSGM gender-diverse, regardless of “straight” status. Consistent with culturally appropriate gender identity language (20), we use the terms “Man,” “Woman,” and “Gender-diverse,” to refer to AoU gender responses “Male,” “Female,” and “Not male/Not female/prefer not to answer/skip.” As such, our sample consisted of the following subgroups: non-OSGM men, non-OSGM women, OSGM men, OSGM women, and OSGM gender-diverse. We compared the baseline demographic characteristics of OSGM and non-OSGM people using means and proportions.

Measures

We developed an AoU deficit-accumulation frailty index (AoU-FI) using methods outlined by Searle et al. (21). We matched AoU survey items with deficits used in other validated frailty indices (21,22). Each deficit was given a value between 0 and 1. Any item with more than 2 answer options had equally distributed graded values between 0 and 1. We assessed 33 deficits across 7 domains of health (Supplementary Table 1): cognition (concentration and dementia), morbidity (cancer, hypertension, peripheral vascular disease, stroke/transient ischemic attack, atrial fibrillation, heart failure, coronary artery disease, diabetes, kidney disease, asthma, and chronic lung disease), physical function (transportation, bathing, running errands alone, walking/climbing stairs, average pain in past 7 days, and everyday activities), geriatric syndromes (fractured bone, osteoporosis, and arthritis), general health status (general health, general social health, health literacy, general social satisfaction, and average fatigue in past 7 days), mental health (anxiety, depression, emotional problems in past 7 days, and general mental health), and sensory impairment (hearing and blindness). Principal components analysis was used to estimate the independent contributions of the 33 deficits. The AoU-FI is calculated as the sum of the conditions present for each participant divided by the number of deficits evaluated for an individual. Participants were excluded if they were missing more than 20% of the AoU-FI deficits or if the available data for the AoU-FI was >70% comorbidities (21). The second criterion ensured a more accurate estimation of frailty rather than a comorbidity index.

Table 1. Sex Assigned at Birth and Gender Categories by Sexual and Gender Minority Status

Population	Sex Assigned at Birth	Gender*	n (%)	Final Population	n (%)
Non-OSGM (n = 67 420)	Male	Male	25 572 (38%)	Non-OSGM, Man	25 572 (38%)
	Female	Female	41 850 (62%)	Non-OSGM, Woman	41 850 (62%)
OSGM (n = 9 110)	Male	Male	2 712 (29.8%)	OSGM, Man	2 712 (29.8%)
	Female	Female	2 480 (27.2%)	OSGM, Woman	2 480 (27.2%)
	Male	Not man only, not woman only, prefer not to answer, or skipped	305 (3.3%)	OSGM, gender-diverse	3 918 (43%)
	Female		386 (4.2%)		
	No matching concept	Male or Female	<30 (~0.3%)		
		Not man only, not woman only, prefer not to answer, or skipped	2 639 (29%)		
	Not male, not female, prefer not to answer, or skipped	Male	185 (2%)		
	Female	294 (3.2%)			
		Not man only, not woman only, prefer not to answer, or skipped	<100 (~1%)		

Notes: OSGM = older sexual and gender minority.
 *Gender was recoded from male and female to man and woman in the analyses.

Describing and Validating the AoU-FI

We used histograms and summary statistics to characterize the continuous AoU-FI for the OSGM and non-OSGM groups. Consistent with Searle et al. (21), we assessed the skew and density of the AoU-FI distribution, as well as the association between AoU-FI and age. To do this, we used linear regression to evaluate the association between log-transformed AoU-FI and age for OSGM and non-OSGM, as well as stratified by gender within each group. We conducted an unequal variance *t*-test to compare the AoU-FI to the nationally representative U.S. community using the National Health and Nutritional Examination Survey (NHANES) FI from Pridham et al. (23). We also categorized the AoU-FI using the following cut-points: robust <0.15, ≥0.15 pre-frail, and >0.25 as frail (24).

For validation analyses, we restricted our sample to participants who consented to share EHR data and used data on death from any cause recorded in the EHR. We validated the AoU-FI association against risk of mortality using Cox proportional hazards models. We included an a priori interaction term between groups (OSGM and non-OSGM) and categorical frailty (robust, pre-frail, frail) while adjusting for the following baseline characteristics as covariates: age group, race/ethnicity, alcohol consumption, smoking status, income, marital status, and HIV status. The prevalence of alcohol use, smoking, and HIV tend to be higher in OSGM (4) and have previously been shown to affect frailty (25). Analyses were done using R version 4.2.2 (26) in the AoU Researcher Workbench cloud-based platform.

Results

Population Characteristics

Of the 200 793 participants aged 50 and older, 124 263 were excluded for missing more than 20% of AoU-FI deficits. All remaining participants had <70% comorbidity-related deficits. This resulted in a final sample of 76 530 participants consisting of 67 420 non-OSGM and 9 110 OSGM. The OSGM had a mean age of 64.8 years (standard deviation [SD] 8.3), 2 480 (27%) were women, 4 896 (54%) identified

as White, 3 195 (35%) are married or living with a partner, and 355 (4%) report with living with HIV. The non-OSGM had a mean age (SD) of 65.7 (8.2) years, 41 850 (62%) were women, 54 244 (80%) identified as White, 43 963 (65%) are married or living with a partner, and 237 (0.4%) report with living with HIV (Table 2).

AoU Deficit-Accumulation Frailty Index

Of the 33 deficits included, principal components analysis demonstrated that each deficit independently contributed to the variance of the AoU-FI (Supplementary Figure 1). The AoU-FI distribution for both OSGM and non-OSGM groups was right-skewed, resembling a gamma distribution, as with previous FIs (27,28) (Figure 1). Comparing the AoU-FI to the NHANES FI, both had similar means (SD), AoU-FI 0.170 (0.10), and NHANES 0.176 (0.073), yet there was a significant difference between the means with a *t*(2 111) = -3.32, 95% confidence interval (CI; -0.009, -0.002), and *p* < .001, likely due to large sample sizes.

Comparing Frailty Between OSGM and Non-OSGM

Among OSGM there were 3 711 (41%) robust, 3 064 (34%) pre-frail, and 2 335 (26%) frail, and among non-OSGM, there were 33 401 (50%) robust, 21 277 (32%) pre-frail, and 12 742 (19%) frail (Table 2). The median, mean (95% CI), and 99% of the AoU-FI for OSGM compared to non-OSGM were 0.17, 0.19 (0.187, 0.191), and 0.47 and 0.15, 0.168 (0.167, 0.169), and 0.49 respectively (Table 2 and Figure 1).

There was a significant association between frailty and age (β = 0.005, *p* < .001). Further, the age-by-OSGM status interaction term was significant (β = -0.005, *p* < .001), suggesting that the association with age is stronger for non-OSGM compared to OSGM (Figure 2). When stratified by gender, non-OSGM women had higher frailty than non-OSGM men (β = 0.30, *p* < .001) and a significant age-by-gender interaction (β = -0.003, *p* < .001; Figure 2). For OSGM, OSGM women had higher frailty than OSGM men (β = 0.41, *p* = .007) and frailty for OSGM gender-diverse and OSGM men were similar (β = 0.19, *p* = .16). Further,

Table 2. Characteristics of All of Us Participants by Sexual and Gender Minority Status

Characteristics	Non-OSGM, <i>n</i> (%)	OSGM, <i>n</i> (%)
Sample size	67 420	9 110
Age		
Mean (<i>SD</i>)	65.7 (8.2)	64.8 (8.3)
Age group		
50–59	18 901 (28%)	2 992 (33%)
60–69	27 079 (40%)	3 540 (39%)
70–79	18 705 (28%)	2 230 (24%)
80+	2 735 (4.1%)	348 (3.8%)
Race/ethnicity		
Hispanic or Latino	4 132 (6.1%)	521 (5.7%)
Black or African American, not Hispanic or Latino	5 363 (8.0%)	543 (6.0%)
White, not Hispanic or Latino	54 244 (80%)	4 896 (54%)
Other	2 664 (4.0%)	274 (3.0%)
Choose not to answer	1 017 (1.5%)	2 876 (32%)
Education		
College graduate or advanced degree	42 910 (64%)	4 090 (45%)
Highest grade: college one to three	16 141 (24%)	1 477 (16%)
Highest grade: twelve or GED	6 118 (9.1%)	529 (5.8%)
Less than a high school degree or equivalent	1 766 (2.6%)	219 (2.4%)
Choose not to answer	485 (0.7%)	2 795 (31%)
Annual income		
<50K	17 562 (26%)	2 354 (26%)
>100K	23 931 (35%)	1 851 (20%)
Between 50 and 100K	18 452 (27%)	1 565 (17%)
Choose not to answer	7 475 (11%)	3 340 (37%)
Marital status		
Divorced or separated	12 199 (18%)	1 092 (12%)
Married or living with partner	43 963 (65%)	3 195 (35%)
Never married	5 708 (8.5%)	1 593 (17%)
Widowed	5 052 (7.5%)	415 (4.6%)
Choose not to answer	498 (0.7%)	2 815 (31%)
Smoked 100 cigarettes (lifetime)		
100 cigarette lifetime: No	39 281 (58%)	4 766 (52%)
100 cigarette lifetime: Yes	27 206 (40%)	4 170 (46%)
Skip choose not to answer	933 (1.4%)	174 (1.9%)
Alcohol use (past year)		
Drink frequency past year: 2 or more per week	22 154 (33%)	2 841 (31%)
Drink frequency past year: 4 or less per month	30 510 (45%)	4 027 (44%)
Ever drinker	11 381 (17%)	1 744 (19%)
Nondrinkers	3 130 (4.6%)	429 (4.7%)
Choose not to answer	245 (0.4%)	69 (0.8%)
Concerned about stable housing		
No	62 895 (93%)	5 598 (61%)
Yes	4 202 (6.2%)	773 (8.5%)
Choose not to answer	323 (0.5%)	2 739 (30%)
HIV status		
HIV/AIDS recorded	237 (0.4%)	355 (3.9%)
No HIV/AIDS recorded	67 183 (99.6%)	8 755 (96%)
Frailty index		
Mean (<i>SD</i>)	0.17 (0.10)	0.19 (0.11)
Median (IQR)	0.15 (0.09, 0.22)	0.17 (0.11, 0.25)
99%	0.47	0.49

Notes: AIDS = acquired immunodeficiency syndrome; GED = General Education Development; HIV = human immunodeficiency virus; IQR = interquartile range; OSGM = older sexual and gender minority; SD = standard deviation.

the interaction of age-by-gender among OSGM was significant for OSGM women compared to OSGM men ($\beta = -0.005, p < .001$), but not for OSGM gender-diverse compared to OSGM men ($\beta = -0.003, p = .14$; Figure 2). The 3 OSGM gender categories had similar median and mean (SD) frailty, OSGM men 0.17, 0.19 (0.1), OSGM women 0.18, 0.2 (0.11), and OSGM gender-diverse 0.17, 0.19 (0.11) (Supplementary Table 2).

Mortality

There were 378 deaths during follow-up: 346 among non-OSGM and 32 among OSGM. In the adjusted analysis, the adjusted hazard ratio (HR; 95% CI) for mortality among non-OSGM was 3.99 (2.99, 5.33), and among OSGM was 6.34 (1.83, 21.98) for frail compared to robust. However, a likelihood-ratio test comparing nested models with and

without the a priori interaction between SGM status and FI category was not statistically significant ($p = .4$; Table 3).

Discussion

Using the AoU survey data to construct an FI, we found that OSGM compared to non-OSGM have a higher burden of frailty maintained across the life span and a higher magnitude of mortality risk. To our knowledge, this is the first study to examine frailty among OSGM.

Overall, OSGM had higher rates of frailty, at younger ages and continuing into older age, compared to non-OSGM in our study, made evident by the significant association between frailty and age-by-OSGM status. It may be that among frail OSGM, those in the oldest age groups (70–79 and 80+) are less likely to participate or died earlier than frail non-OSGM. Additionally, this finding is analogous to the literature regarding people living with HIV, another vulnerable minority population who have been shown to have higher levels of frailty at relatively younger ages and among whom more than 50% also identify as SGM (29,30). This may be due to the minority stress experience of chronic additive stress associated with social stigmatization and discrimination which may lead to systemic inflammation (13) and contribute to the development of frailty at relatively younger ages. Considering this, HIV status was included as a covariate in the adjusted analysis, and the association between frailty and mortality remained significant. Among people living with HIV, 79% experienced stigma when receiving medical care (31), and stigma was associated with reduced access to care, no regular source of HIV care, and suboptimal adherence to life-critical medication (32). Similarly for OSGM, those with internalized stigma or discrimination were less likely to have a routine physical exam in the past year (33), screening pap smear (34), sexual minority women are less likely to have a usual place of care, and both sexual minority men and women report difficulties with affording care (35). Despite the consistent association

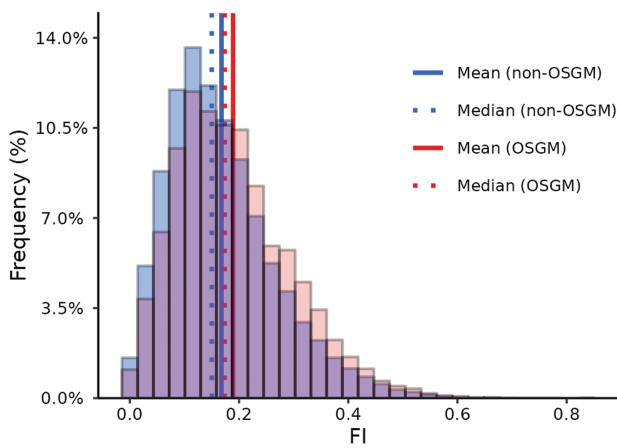


Figure 1. Distribution of AoU-FI for OSGM, $n = 9\ 110$ and non-OSGM, $n = 67\ 420$. AoU-FI = All of Us frailty index; OSGM = older sexual and gender minority. The AoU-FI distribution for both populations with mean (solid line) and median (dashed line).

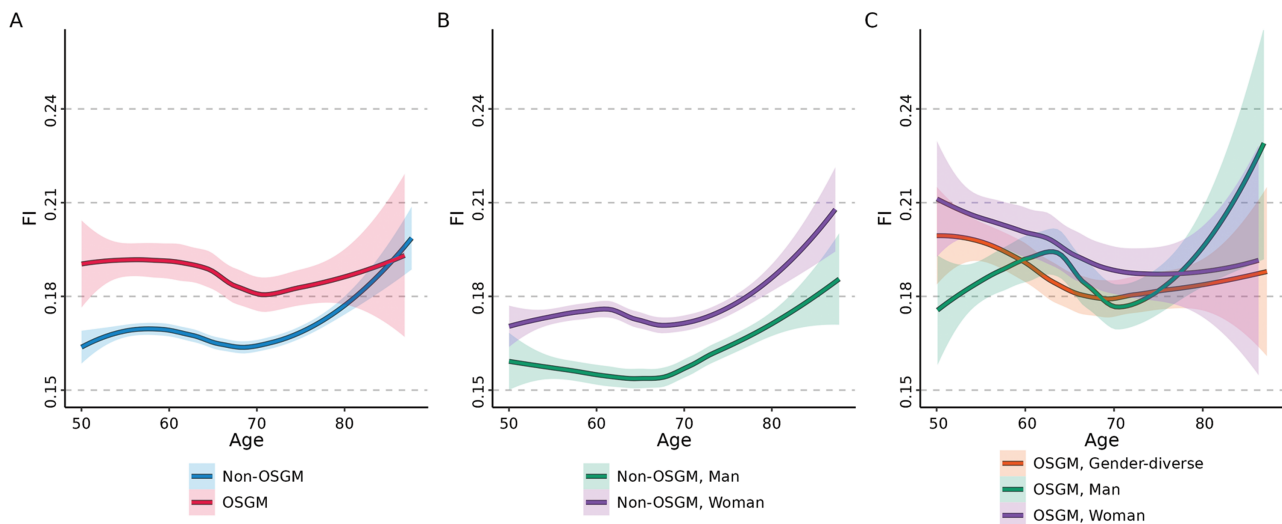


Figure 2. Association of AoU-FI values by age and gender for non-OSGM and OSGM. AoU-FI = All of Us frailty index; OSGM = older sexual and gender minority adults. (A) The association between AoU-FI by age ($\beta = 0.005, p < .001$), with separate lines for OSGM $n = 9\ 110$ and non-OSGM $n = 67\ 420$. (B) Non-OSGM association between AoU-FI by age and gender (age \times gender: $\beta = -0.003, p < .001$), non-OSGM men, $n = 25\ 572$ (ref) and non-OSGM women $n = 41\ 850$. (C) OSGM association between AoU-FI by age and gender, OSGM gender-diverse, $n = 3\ 918$ (age \times compared to OSGM men: $\beta = -0.003, p = .14$), OSGM men $n = 2\ 712$ (ref), and OSGM women $n = 2\ 480$ (age \times compared to OSGM men: $\beta = -0.005, p < .001$). For all panels, a trend line was added using the generalized additive models.

Table 3. Association of Frailty Category and Mortality by Sexual and Gender Minority Status

		Sample Size	Crude Analysis		Adjusted Analysis [†]	
		<i>n</i> (%)	HR	95% CI	HR	95% CI
Non-OSGM	Robust	33 401 (49.5%)	Reference		Reference	
	Pre-frail	21 277 (31.6%)	2.47*	1.87, 3.27	2.18*	1.64, 2.89
	Frail	12 742 (18.9%)	4.71*	3.58, 6.19	3.99*	2.99, 5.33
OSGM	Robust	3 711 (40.7%)	Reference		Reference	
	Pre-frail	3 064 (33.6%)	5.68*	1.63, 19.78	4.81*	1.38, 16.76
	Frail	2 335 (25.6%)	7.94*	2.30, 27.42	6.34*	1.83, 21.98

Notes: AoU-FI = All of Us Frailty Index; CI = confidence interval; HR = hazard ratio; OSGM = older sexual and gender minority.

[†]Adjusted for age, race/ethnicity, alcohol use, smoking status, HIV status, and marital status.

**p* value < .05. The *p* value for the OSGM-frailty interaction was .4.

between minority stress and health care utilization, little is known to date about how utilization affects health outcomes (36) or frailty among OSGM.

The magnitude of association between frailty and mortality was higher for OSGM compared to non-OSGM, potentially reflecting health disparities and may also contribute to the age difference in frailty between OSGM and non-OSGM. OSGM mortality rates are understudied and not well defined. For women, despite no differences in all-cause mortality, sexual minority women had a higher mortality risk due to self-harm (37) and breast cancer (38) compared to non-sexual minority women. In contrast, research on mortality for sexual minority men has shown both higher (39) and no difference (37) in HIV-related mortality risk. Beyond sexual minorities, there is growing interest in understanding transgender mortality rates given exaggerated disparities in morbidity, mental health issues, and discrimination compared to non-transgender individuals (40–43). Hughes and colleagues found the mortality rate of privately insured transgender people was almost double compared to non-transgender people and highest in transfeminine and unclassified transgender individuals (42). Together, these findings highlight that OSGM heterogeneity may give rise to differences in frailty and mortality of OSGM subpopulations, emphasizing that OSGM research will require discerning subpopulations to elucidate disparities in frailty.

As expected, we demonstrated differences in the association between frailty and age across genders among non-OSGM (44–46). By extending this to the OSGM population, our results demonstrated that the association between frailty and age was reduced for all OSGM genders compared to non-OSGM genders. Consistent with the minority stress theory, our results suggest that OSGM status may have a larger affect on frailty development compared to age. Clinically, this finding emphasizes the need for early frailty assessment among OSGM.

Strengths and Limitations

The major strength of our study is the implementation of a novel AoU-FI on a large OSGM population with detailed health information available. Despite this, there are limitations to our study. We used Searle et al. methods to develop the AoU-FI, but not all the original FI items were available and substitutions were made. Even still, the resulting AoU-FI had the expected distribution (21), and association between age (47,48), gender (48–50), and mortality (21,27) reinforcing our confidence in its effectiveness as an FI. Additionally, the

AoU database is a convenience sample of healthy volunteers, which may limit generalizability and introduce sampling bias. However, the AoU research program was designed to focus on those underrepresented in biomedical research, including sexual and gender minorities, enabling studies like ours. In this study, we identified distributions of OSGM baseline characteristics including age, race/ethnicity, annual income, and marital status that were consistent with the literature describing the OSGM population (2,3,6,49) despite a higher proportion of OSGM who chose not to answer. OSGM reported lower proportions of college or advanced degrees, some college, and high school/GED compared to non-OSGM. Studies have reported both higher (3,5) and lower (2) levels of education for OSGM participants. The similarity in the characteristics of the AoU OSGM with previously published OSGM populations is encouraging for external validity, despite being a convenience sample. Given these limitations, future work should also assess selection bias and the impact that missing data have on measures such as the AoU-FI.

Although we were able to develop the AoU-FI, we were unable to assess a frailty phenotype using the AoU survey data; however, given the extent of OSGM health disparities, FI provides a multidimensional assessment of aging. Further, we validated the AoU-FI with mortality with death from any cause recorded in the EHR, notably not all deaths are captured by the EHR, and deaths are not currently confirmed by next of kin contact or linked to the National Death Index. Despite this limitation, we were able to establish an association between AoU-FI and mortality to support that AoU-FI is a new representation of the frailty concept within the common data model that can be used by other investigators and can expand our understanding of frailty by applying it to underresearched and underrepresented populations.

Finally, the OSGM population is heterogeneous and consists of but is not limited to those who identify as gay, lesbian, bisexual, transgender, and nonbinary. Subpopulations of OSGM (eg bisexual or transgender individuals) have differences in health disparities (41,50), mortality (42), and likely frailty. Further, sexual minority and gender minority identities are not mutually exclusive so it is possible that individuals who belong to both gender and sexual minority groups (eg, transgender woman and lesbian) may be more frail than their cisgender, sexual minority counterparts as a result of stigma and discrimination from multiple minoritized identities. The AoU Registered Tier Data lacked the granularity to study the individual sexual orientation subpopulations, but we did assess gender subpopulations of OSGM. Future work should

evaluate frailty among subpopulations of OSGM as these differences may have significant implications for OSGM care strategies. Despite these limitations, this study was a necessary first step to understanding frailty among OSGM.

Conclusion

We successfully developed and validated an AoU-FI and describe frailty among OSGM for the first time. Our results support the hypothesis that OSGM have a higher burden of frailty, particularly at younger ages, and have a higher risk of mortality compared to non-OSGM. This highlights the need to consider aging physiology for OSGM populations, even at younger ages. Future work should assess how and when frailty develops among OSGM to identify potential targets for intervention.

Supplementary Material

Supplementary data are available at *The Journals of Gerontology, Series A: Biological Sciences and Medical Sciences* online.

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Conflict of Interest

None.

Author Contributions

Study design: All authors. Analysis: M.P.W., L.H.S., R.C., C.N.W., B.O.M. Data interpretation: All authors. Manuscript drafting: C.N.W., B.O.M. Manuscript editing: All authors.

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