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Testosterone Use and Sexual Function among Transgender Men and Gender Diverse People Assigned Female at Birth

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- 1 Testosterone Use and Sexual Function among Transgender Men and Gender Diverse
- 2 People Assigned Female at Birth

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- **Tweetable Statement:** In a study of 1219 transgender men, testosterone use among transgender
- 37 men is associated with positive sexual function and vulvovaginal pain during sex.

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Short Title: Testosterone and Sexual Function

- 41 AJOG at a Glance:
- Why was this study conducted? The study aimed to improve our understanding of the
- impact of testosterone on the sexual health of transgender men and gender diverse people
- 44 assigned female at birth.
- **Key findings.** In our cross-sectional analysis of 1,219 adult transgender participants, 65%
- reported any genital pain/discomfort during sexual activity in the past 30 days. Current

- testosterone use was associated with a higher interest in sexual activity and ability to orgasm as well as vulvovaginal pain during sexual activity.
- What does this add to what is known? Although data from the extant literature are scare, available evidence demonstrates that >60% of transgender men experience vulvovaginal pain during sexual activity. Given this high burden, there is an urgent need to identify effective and acceptable interventions for this population.

53 ABSTRACT

54 **Background:** Testosterone use among transgender people likely impacts their experience of 55 sexual function and vulvovaginal pain via several complex pathways. Testosterone use is 56 associated with decreased estrogen in the vagina and atrophic vaginal tissue, which may be 57 associated with decreased vaginal lubrication and/or discomfort during sexual activity. At the 58 same time, increased gender affirmation through testosterone use may be associated with 59 improved sexual function. However, data on pelvic and vulvovaginal pain among transgender 60 men and nonbinary people assigned female at birth is scarce. 61 **Objective:** To assess the association between testosterone and sexual function, with a focus on 62 symptoms that are commonly associated with vaginal atrophy. 63 **Study Design:** We conducted a cross-sectional analysis of 1,219 participants ages 18-72 years 64 old using 2019-2021 data from an online, prospective, longitudinal, cohort study of sexual and/or 65 gender minority people in the US (The PRIDE Study). Our analysis included adult transgender 66 men and gender diverse participants assigned female at birth who were categorized as never, 67 current, and former testosterone users. Sexual function was measured across eight Patient-68 Reported Outcomes Measurement Information System (PROMIS) Sexual Function and 69 Satisfaction (SexFS) domains. 70 **Results:** Overall, 516 (42.3%) had never used testosterone and 602 (49.4%) currently used 71 testosterone. Median duration of use was 37.7 months (range 7 days to >27 years). Most 72 participants (64.6%) reported genital pain/discomfort during sexual activity in the past 30 days, 73 most commonly in the vagina/frontal genital opening (52.2%), followed by the clitoris (29.1%) 74 and labia (24.5%). Current testosterone use was associated with higher interest in sexual activity 75 $(\beta=6.32, 95\% \text{ CI: } 4.91-7.74)$ and more vaginal pain/discomfort during sexual activity $(\beta=1.80,$

76 95% CI: 0.61-3.00). No associations were observed between current testosterone use and 77 satisfaction with sex life, lubrication, labial pain/discomfort, or orgasm pleasure. 78 **Conclusions:** Testosterone use among transgender men and gender diverse people was 79 associated with a higher interest in sexual activity and ability to orgasm as well as vaginal 80 pain/discomfort during sexual activity. Notably, the available evidence demonstrates that >60% 81 of transgender men experience vulvovaginal pain during sexual activity. The causes of pelvic 82 and vulvovaginal pain are poorly understood but are likely multifactorial and include 83 physiological (e.g., testosterone-associated vaginal atrophy) and psychological factors (e.g., 84 gender affirmation). Given this high burden, there is an urgent need to identify effective and 85 acceptable interventions for this population.

87 **KEYWORDS:** Dyspareunia, Sexual Function, Testosterone, Transgender, Vulvovaginal Pain

INTRODUCTION

At least 1.6 million transgender adults and adolescents live in the United States, among whom an estimated 70% of transgender men have ever used testosterone as gender-affirming hormone therapy (GAHT). Vaginectomy is rare (<3%) in this population, and the majority of transgender men and gender diverse people retain their vagina. Testosterone GAHT likely impacts sexual function via several complex pathways: Testosterone GAHT is associated with vaginal atrophy, which may be associated with decreased lubrication and/or discomfort during sexual activity. At the same time, increased gender affirmation through testosterone use may be associated with improved sexual function.

There is limited research on the sexual function of transgender men and gender diverse people assigned female at birth (AFAB). What does exist suggests that, although testosterone GAHT is associated with increased desire and arousal,⁶ a high proportion of transgender men also reported dyspareunia (painful sex), a common symptom of vaginal atrophy. The prevalence of dyspareunia may be as high as 60-62% among transgender men,^{7,8} markedly higher than the prevalence reported among cisgender women (3-48%).⁹ Several studies also suggest that transgender men using testosterone may experience chronic genital pain and discomfort, with one study reporting that 10-16% of transgender men had been diagnosed with vulvodynia (defined as chronic burning, stinging, or irritating vulvovaginal pain for three consecutive months or longer).^{7,8} Only one prior small study directly assessed the impact of testosterone GAHT on genital pain during sexual activity. Although 30% of transgender men reported that testosterone had caused genital dryness and 14% experienced genital tearing since initiating testosterone, they did not observe an association between testosterone use and vulvodynia or dyspareunia symptoms.¹⁰

The present study aimed to improve our understanding of the impact of testosterone on the sexual health of transgender people, with a focus on symptoms that are commonly associated with vaginal atrophy, including decreased lubrication, and pain during sexual activity. Using data from a large national online sample of sexual and/or gender minority adults in the US, we examined the association between current testosterone use and self-reported measures of sexual function and satisfaction experienced by transgender men and gender diverse people.

MATERIALS AND METHODS

This analysis used data from The Population Research in Identity and Disparities for Equality (PRIDE) Study, an online, prospective, longitudinal cohort study of sexual and/or gender minority people in the US. We conducted a cross-sectional analysis using 2019-2021 data from a questionnaire administered annually to study participants among adult transgender men and gender diverse participants AFAB. We included participants who self-reported currently having a vagina or frontal genital opening (FGO) and who completed the Patient-Reported Outcomes Measurement Information System (PROMIS) Sexual Function and Satisfaction (SexFS) items. We excluded participants who did not self-report having a vagina/FGO or who reported having a phalloplasty or vaginectomy.

Measures

Demographic Characteristics. Participants self-reported data on race and ethnicity, current gender identity, current sexual orientation, and sex assigned at birth. Participants could select multiple response options for race, ethnicity, gender, and sexual orientation. Participants self-reported the gender(s) of people they had any sexual activity within the past year.

Testosterone Use. We categorized participants as never, current, and former testosterone users. Current testosterone use was assessed on the 2019-2021 annual questionnaires and included participants who, at the time, indicated they were currently taking testosterone (of any type in any formulation such as gel, injection or patch), testosterone cypionate, testosterone enanthate, or testosterone undecanoate for gender affirmation. To differentiate participants who had never used testosterone from participants who formerly used testosterone, we also incorporated participants' responses to a baseline survey that assessed participants' lifetime testosterone use. Duration of testosterone use was calculated based on participant's self-reported month and year of initiating testosterone and the date of survey completion. We did not collect information regarding the dose of testosterone.

Sexual Function. The PROMIS SexFS assesses self-reported sexual function and satisfaction over the past 30 days. This instrument was originally developed for cancer populations and was subsequently validated among a broad group of sexually active adults in the US who were not cancer survivors. Although sexual minority (e.g., bisexual, gay, and lesbian) individuals were involved in the development of the PROMIS SexFS, transgender people were not. The PRIDE Study implemented a modified version of the PROMIS SexFS that allowed participants to select their preferred anatomical language (i.e., "vagina" or "frontal genital opening"). In completing the survey, each participant's selection for their preferred terminology was propagated throughout subsequent survey items.

Our analysis included eight PROMIS SexFS domains: interest in sexual activity, satisfaction with sex life, vaginal/FGO lubrication, ability to orgasm, orgasm pleasure, as well as vaginal/FGO, labial, or clitoral pain/discomfort during sexual activity. Interest in sexual activity was assessed for all participants while all other domains were only assessed for participants who

reported any sexual activity in the past 30 days (which was broadly defined and included masturbation). For each domain, we calculated each participant's raw score and T-scores. A T-score of 50 represents the mean for the US population and has a standard deviation of 10. Higher T-scores indicate more of the construct measured by the domain; for example, more interest in sexual activity, increased vaginal/FGO lubrication, and more pain/discomfort relative to lower scores.

Covariates. We considered covariates that are associated with sexual function and pelvic pain based on prior literature. These included standardized clinical assessment tools and self-reported lifetime diagnoses. We broadly considered five categories of covariates that have been previously associated with sexual function among cisgender women and transgender men: structural changes to the pelvis, inflammatory conditions in the pelvis, hormonal influences, mental health, and substance use. 17–30

Structural changes to the pelvis included prior pregnancies, hysterectomy, and uterine fibroids. Inflammatory conditions in the pelvis included pelvic inflammatory disease, inflammatory bowel disease (including Crohn's disease and ulcerative colitis), irritable bowel syndrome, and bacterial sexually transmitted infections (STI; defined as chlamydia, gonorrhea, or syphilis diagnosis in the past year). Hormonal influences include oophorectomy, polycystic ovary syndrome, current hormonal contraceptive use (including oral contraceptives, transdermal patch, vaginal rings, medroxyprogesterone acetate injections, and etonogestrel implants), and intrauterine device use.

Mental health measures included lifetime diagnoses of depression and post-traumatic stress disorder (PTSD) as well as lifetime experiences of sexual abuse, rape, and sexual assault.

Depressive symptoms in the past two weeks were assessed using the Patient Health

Questionnaire-9 (PHQ-9; score range 0-27 with higher scores indicating more depressive symptoms)¹⁴ and PTSD symptoms in the last month were assessed with the brief 6-item version of the PTSD Check List (PCL-6; score range 6-30).¹⁵ Substance use variables included current smoking and current alcohol consumption behaviors; the latter was assessed using the Alcohol Use Disorders Identification Test (AUDIT; score range 0-40 with higher scores indicating more disordered alcohol use).¹⁶

Statistical Analysis

We assessed cross-sectional associations between current testosterone use and sexual function. For participants who completed more than one annual questionnaire, we restricted our analysis to only include their first year of responses. We calculated descriptive statistics stratified by testosterone use and conducted Chi-squared tests. We conducted one sample t-tests to test if the SexFS domain T-scores differed from the population mean of 50 and calculated the Pearson correlation coefficient to estimate the strength and direction of the relationship between PROMIS SexFS domains.

We then used multivariable linear regression to estimate the association between current testosterone use (relative to never testosterone use) and the T-scores for each sexual function domain. We used causal diagrams to select covariates to include in our model (Supplemental Figure 1). We estimated three models: an unadjusted model, a minimally adjusted model, and a robustly adjusted model. We chose our primary analysis to be a minimally adjusted model that included confounding variables (*i.e.*, covariates that were associated with both the exposure [testosterone use] and outcomes [sexual function] in our sample). The minimally adjusted model included age, current depression symptoms (PHQ-9 scores), current PTSD symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, and hormonal

contraception use. We conducted secondary analyses with a robustly adjusted model that included all covariates that are associated with sexual function and dyspareunia in prior literature, but were not associated with testosterone use in our sample, including history of sexual assault, prior pregnancy, inflammatory bowel disease, irritable bowel syndrome, uterine fibroids, pelvic inflammatory disease, polycystic ovary syndrome, and intrauterine devices.^{17–30}

Our primary analysis considers current *versus* never testosterone users. Results comparing former to current users are provided in the Supplementary Materials. We hypothesized that asexual identity may be a potential effect modifier, as asexual participants may differ from non-asexual participants with respect to interest in sexual activity. Therefore, we conducted sensitivity analyses stratified by self-reported asexual sexual orientation. All analyses were conducted in R version 4.2.1. This study received ethical approval from University of California, San Francisco, Stanford University School of Medicine, and WCG Institutional Review Boards.

RESULTS

Our analysis included 1,219 participants ages 18-72 years old (median age 27.1; Table 1). Most participants (61.1%) endorsed more than one gender identity: most commonly non-binary (54.2%), transgender man (46.0%), genderqueer (33.6%), and man (21.5%). Participants were diverse in sexual orientation, although most identified as queer (65.1%). The majority (80.6%) of participants only reported White race and/or ethnicity and 12.0% of participants selected more than one racial and/or ethnicity.

There were 516 (42.3%) participants who had never used testosterone, 602 (49.4%) who currently used testosterone, and 76 (6.2%) former testosterone users. Twenty-five participants

were missing information about GAHT use. The median duration of GAHT with testosterone use was 37.7 months (range 7 days to >27 years). Current testosterone users were significantly more likely to identify as a man or transgender man, and less likely to identify as agender, genderqueer, nonbinary, or questioning (Supplemental Table 1). There were no differences in testosterone use by age, race, or ethnicity.

Most participants reported having sex with another person in the past year (68.9%), were in a relationship (59.4%), and reported any sexual activity (including masturbation) in the past 30 days (89.5%; Table 2). Among participants who reported having sex with another person in the past year, 88.5% reported any receptive oral sex or receptive vaginal/FGO sex. Participants currently using testosterone were more likely to be sexually active in the past year (73.3% v. 63.4%, p=0.001) and in the past 30 days (93.9% v. 84.7%, p<0.001).

Table 2 reports participants' pelvic health histories. Current testosterone users were more likely to have a hysterectomy (17.9% v. 5.4%, p<0.001) and oophorectomy (14.5% v. 3.3%, p<0.001), and less likely to currently use hormonal contraceptives (5.8% v. 17.8%, p<0.001) compared to participants who never used testosterone. Current testosterone users were also more likely to have a past year bacterial STI diagnosis (3.0% v. 1.6%, p=0.001). Testosterone use was not associated with pelvic inflammatory disease, polycystic ovary syndrome, uterine fibroids, inflammatory bowel disease, irritable bowel syndrome, pregnancy history, or intrauterine device use.

Participants reported very high levels of lifetime experiences of sexual abuse, sexual assault, and ever having received a depression and PTSD diagnoses (Table 2). Testosterone use was associated with lower scores (*i.e.*, better mental health) for current depressive and PTSD

symptoms. Testosterone use was also associated was substance use, including current smoking
 (7.5% v. 4.3%, p=0.033) and higher AUDIT scores.

Among the 1,091 sexually active participants, most (n=693, 64.6%) reported genital pain/discomfort during sexual activity in the past 30 days, most commonly in the vagina/FGO (52.2%), followed by the clitoris (29.1%) and labia (24.5%; Table 3). There were 103 (9.6%) participants who reported pain at all three genital sites. T-scores for pain/discomfort were higher than the population mean (p-values<0.001) while T-scores for orgasm pleasure (mean T-score 45.0, p-value<0.001) and satisfaction with sex life (mean T-score 45.6, p-value<0.001) were lower than the population mean. Figure 1 shows the correlation between each sexual function T-score among participants. Interest in sexual activity, satisfaction, lubrication, orgasm ability and pleasure were positively correlated. Measures of pain/discomfort were negatively correlated with all other domains.

Compared to participants who never used testosterone, current testosterone users were less likely to report difficulty with lubrication (58.5% v. 66.7%, p=0.01), more like to report any vaginal pain/discomfort (56.0% v. 49.2%, p=0.04), and more likely to achieve orgasm (p=0.003; Table 3). In the minimally adjusted regression models (Table 4), current testosterone use was associated with higher interest in sexual activity (β =6.32, 95% CI: 4.91-7.74), higher ability to orgasm (β =1.50, 95% CI: 0.19-2.81), pain/discomfort during sexual activity in the vaginal/FGO (β =1.80, 95% CI: 0.61-3.00) and in the clitoris (β =1.20, 95% CI: 0.10-2.30). The association between testosterone use, ability to orgasm, and clitoral pain/discomfort were not statistically significant in the robustly adjusted model. No associations were observed between current testosterone use and satisfaction with sex life, lubrication, labial pain/discomfort, or orgasm pleasure.

Duration of testosterone use was associated with increased interested in sexual activity, but not with any other outcomes (Supplemental Table 4). In sensitivity analyses, we did not observe any evidence of effect modification by asexual sexual orientation (Supplemental Figure 2), although asexual identity was independently associated with most sexual function domains.

COMMENT

In our study, testosterone use among transgender men and gender diverse people AFAB was associated with some domains of positive sexual function (such as a higher interest in sexual activity and ability to orgasm) as well as pain/discomfort during sexual activity. Specifically, we observed a strong, consistent association between current testosterone use and higher interest in sex as well as vaginal/FGO pain during sexual activity.

Our findings are consistent with prior studies which found that testosterone use is associated with increased desire and interest in sex.⁶ GAHT is associated with significant improvements in overall mental health, quality of life, and body image,^{31–33} which in turn likely have positive impacts on other areas of wellbeing, including sexual function. For example, other studies have found that access to gender-affirming chest reconstruction surgery is associated with higher sexual function scores, while experiencing barriers to accessing gender-affirming care is associated with lower sexual function scores.²⁵ Although prior studies have not observed an association between testosterone use and orgasm,⁶ testosterone use was associated with a higher ability to orgasm in our study.

Notably, we observed that a majority of transgender and gender diverse people AFAB using testosterone (67%) experienced vulvovaginal pain during sexual activity. This prevalence is consistent with prior studies, which have reported that 60-62% of transgender men experience

dyspareunia.^{7,8} Although testosterone was associated with vaginal/FGO pain in all regression analyses, the prevalence of any genital pain among individuals who were testosterone naïve was also quite high (63%). The causes of genital pain during sex (including dyspareunia, vulvodynia, and vaginismus) are multifactorial.²⁴ Although physiological factors (such as vaginal atrophy, endometriosis, and pelvic floor injury) are associated with genital pain during sex,⁵ there are complex associations with psychological and social factors, including co-occurrence with other pain disorders, mental health, substance use, and sexual trauma.^{22,23} Our sample reported an alarmingly high level of sexual abuse (78%), sexual assault (50%), depression (81%) and PTSD diagnoses (40%)—factors that are correlated with chronic pelvic pain among presumably cisgender women.^{17–21} This may partially account for the high prevalence of pain in our study.

Genital pain, including dyspareunia and chronic vulvodynia, can have a significant impact on people's well-being and quality of life.^{34,35} Although one prior study reports that 1 in 5 transgender men reported that pain during sexual activity was causing significant problems in their life or relationship,⁷ there has been limited investigation into how this may impact the well-being and quality of life of transgender people.

Intravaginal estrogen (delivered via cream, tablets, or a ring) is recommended for transgender men experiencing testosterone-associated dyspareunia, vaginitis, and cervicitis.³⁶
Locally administered estrogen therapy has been demonstrated to be a safe and effective therapy in post-menopausal cisgender women, who also experience vaginal atrophy associated with estrogen deficiency.^{37–39} However, few existing reports likewise suggest that a minority of transgender men (<5%) have ever used intravaginal estrogen.⁸ Therefore, there may be barriers to the uptake of this intervention among transgender populations, including acceptability,

provider awareness, participant/patient knowledge, insurance coverage, and other structural barriers, in addition to the limited evidence base to inform clinical use.

Strengths and Limitations

To our knowledge, this is the first study to report PROMIS SexFS outcomes for transgender men, and one of few studies to examine the association between testosterone use and vulvovaginal pain. This study has several strengths, including a large, national sample of transgender participants who were diverse with respect to their age, gender identity, and sexual orientations.

Our results should be interpreted considering several limitations. The PRIDE Study is a convenience sample of majority White participants that relies on self-reported health outcomes and diagnoses and therefore may be subject to sampling, recall, and social desirability biases. Although our study demonstrates the feasibility of using the PROMIS SexFS with transgender men, the PROMIS SexFS has not been validated in transgender populations. This is notable, given that the item calibration and scoring is stratified by sex assigned at birth. Although scoring for interest in sexual activity, orgasm ability, and orgasm pleasure are identical for male and female populations, the items specific to vaginal/FGO anatomy were calibrated to a sample primarily composed of (presumably) cisgender women. Therefore, there remain opportunities to develop and validate sexual function measures for transgender and gender diverse people.

In addition, our survey did not collect data on several important variables. Although most participants reported any receptive oral or receptive vaginal/FGO sex, we did not assess other types of vaginal/FGO sexual activity (e.g., penetration with fingers or sex toys) and acknowledge that some transmasculine people do not use their vagina/FGO at all during sex. We also lacked detailed information of hormone doses and were unable to distinguish between low- and high-

dose testosterone. We did not collect data on several factors that also influence genital pain, including intravaginal estrogen use or the use of other medications used to manage symptoms of vaginal atrophy and painful sex (*e.g.*, topical lidocaine).

Lastly, although duration of testosterone use was associated with few outcomes in our study, this may in part be a limitation of our cross-sectional design. Future prospective longitudinal research is important for identifying changes to sexual function over time, accounting for cyclical variability in vaginal symptoms, variation in testosterone access and dosing, as well as for understanding how long-term testosterone use may impact sexual function and vulvovaginal pain.

Conclusions

Using data from a large national sample of transgender men and gender diverse people AFAB, we observed that testosterone use was both associated with positive sexual function and dyspareunia. The relationship between receipt of testosterone GAHT and sexual function is complex, and likely includes both physiological (*e.g.*, vaginal atrophy) and psychological factors (*e.g.*, affirmation). However, given the high burden (>60%) of dyspareunia observed among transgender people AFAB, there is a need to assess its impact on the overall quality of life, identify effective and acceptable interventions, and reduce barriers to accessing treatment for transgender people experiencing dyspareunia.

Author Contributions: DMT and JOM conceived of the study design and methodology and had access to all data. DMT performed and verified all analyses and wrote the original manuscript draft, with support from JOM. MRL, AF, ZD, MEL, MC, and JOM contributed to data collection. All authors provided input on data interpretation and provided manuscript edits. **Acknowledgements:** The PRIDE Study is a community-engaged research project that serves and is made possible by LGBTQ+ community involvement at multiple points in the research process, including the dissemination of findings. We acknowledge the courage and dedication of The PRIDE Study participants for sharing their stories; the careful attention of PRIDEnet Participant Advisory Committee (PAC) members for reviewing and improving every study application; and the enthusiastic engagement of PRIDEnet Ambassadors and Community Partners for bringing thoughtful perspectives as well as promoting enrollment and disseminating findings. For more information, please visit https://pridestudy.org/pridenet. **Data sharing:** We welcome the opportunity to facilitate high-quality, community-engaged, research collaborations that aim to improve the health and well-being of LGBTQ+ communities. Through The PRIDE Study's ancillary studies, a wide variety of investigators working on academic or community-based projects related to LGBTQ+ health can apply to work collaboratively with The PRIDE Study team and access data. For more information, please visit:

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https://pridestudy.org/collaborate

378 REFERENCES

- 379 1. Herman JL, Flores AR, O'neill KK. How Many Adults and Youth Identify as Transgender in
- the United States? The Williams Institute, UCLA School of Law; 2022. Accessed June 23,
- 381 2022. https://williamsinstitute.law.ucla.edu/publications/trans-adults-united-states/
- 382 2. James SE, Herman JL, Rankin S, Keisling M, Mottet L, Anafi M. The Report of the 2015
- 383 *U.S. Transgender Survey.* National Center for Transgender Equality; 2016.
- 384 3. Krakowsky Y, Potter E, Hallarn J, et al. The Effect of Gender-Affirming Medical Care on
- the Vaginal and Neovaginal Microbiomes of Transgender and Gender-Diverse People.
- 386 Front Cell Infect Microbiol. 2022;11. doi:10.3389/FCIMB.2021.769950
- 387 4. Baldassarre M, Giannone FA, Foschini MP, et al. Effects of long-term high dose
- testosterone administration on vaginal epithelium structure and estrogen receptor- α and - β
- 389 expression of young women. *Int J Impot Res.* 2013;25(5):172-177. doi:10.1038/ijir.2013.9
- 390 5. Kingsberg S, Kellogg S, Krychman M. Treating dyspareunia caused by vaginal atrophy: a
- review of treatment options using vaginal estrogen therapy. *Int J Womens Health*.
- 392 2010;1:105-111.
- 393 6. Mattawanon N, Charoenkwan K, Tangpricha V. Sexual Dysfunction in Transgender People:
- 394 A Systematic Review. *Urol Clin North Am.* 2021;48(4):437-460.
- 395 doi:10.1016/j.ucl.2021.06.004
- 396 7. Abern L, Maguire K, Cook J, Carugno J. Prevalence of Vulvar Pain and Dyspareunia in
- Trans Masculine Individuals. *LGBT Health*. Published online February 3, 2022.
- 398 doi:10.1089/LGBT.2020.0357
- 399 8. Zwickl S, Burchill L, Wong AFQ, et al. Pelvic Pain in Transgender People Using
- Testosterone Therapy. *LGBT Health*. Published online January 4, 2023.
- 401 doi:10.1089/lgbt.2022.0187
- 402 9. Weijmar Schultz W, Basson R, Binik Y, Eschenbach D, Wesselmann U, Van Lankveld J.
- Women's sexual pain and its management. J Sex Med. 2005;2(3):301-316.
- 404 doi:10.1111/j.1743-6109.2005.20347.x
- 405 10. Dadasovich R, Auerswald C, Minnis AM, Raymond HF, McFarland W, Wilson EC.
- Testosterone and sexual risk among transmen: a mixed methods exploratory study. *Cult*
- 407 *Health Sex.* 2017;19(2):256-266. doi:10.1080/13691058.2016.1216605
- 408 11. Weinfurt KP, Lin L, Bruner DW, et al. Development and Initial Validation of the
- PROMIS® Sexual Function and Satisfaction Measures Version 2.0. *J Sex Med.*
- 410 2015;12(9):1961-1974. doi:10.1111/jsm.12966
- 411 12. Moseson H, Lunn MR, Katz A, et al. Development of an affirming and customizable
- electronic survey of sexual and reproductive health experiences for transgender and gender

- 413 nonbinary people. *PLOS ONE*. 2020;15(5):e0232154.
- **414** doi:10.1371/JOURNAL.PONE.0232154
- 415 13. Klein A, Golub SA. Enhancing Gender-Affirming Provider Communication to Increase
- Health Care Access and Utilization Among Transgender Men and Trans-Masculine Non-
- 417 Binary Individuals. *LGBT Health*. 2020;7(6):292-304. doi:10.1089/lgbt.2019.0294
- 418 14. Levis B, Benedetti A, Thombs BD. Accuracy of Patient Health Questionnaire-9 (PHQ-9) for
- screening to detect major depression: individual participant data meta-analysis. *BMJ*.
- 420 2019;365. doi:10.1136/BMJ.L1476
- 421 15. Han B, Wong EC, Mao Z, Meredith LS, Cassells A, Tobin JN. Validation of a brief PTSD
- screener for underserved patients in federally qualified health centers. *Gen Hosp Psychiatry*.
- 423 2016;38:84-88. doi:10.1016/j.genhosppsych.2015.07.009
- 424 16. Bohn MJ, Babor TF, Kranzler HR. The Alcohol Use Disorders Identification Test (AUDIT):
- validation of a screening instrument for use in medical settings. *J Stud Alcohol*.
- 426 1995;56(4):423-432. doi:10.15288/jsa.1995.56.423
- 427 17. Reed BD, Legocki LJ, Plegue MA, Sen A, Haefner HK, Harlow SD. Factors associated with
- 428 vulvodynia incidence. *Obstet Gynecol*. 2014;123(2 Pt 1):225-231.
- 429 doi:10.1097/AOG.0000000000000066
- 430 18. Chisari C, Monajemi MB, Scott W, Moss-Morris R, McCracken LM. Psychosocial factors
- associated with pain and sexual function in women with Vulvodynia: A systematic review.
- 432 Eur J Pain. 2021;25(1):39-50. doi:10.1002/ejp.1668
- 433 19. Siqueira-Campos VME, Da Luz RA, de Deus JM, Martinez EZ, Conde DM. Anxiety and
- depression in women with and without chronic pelvic pain: prevalence and associated
- 435 factors. J Pain Res. 2019;12:1223-1233. doi:10.2147/JPR.S195317
- 436 20. Meltzer-Brody S, Leserman J, Zolnoun D, Steege J, Green E, Teich A. Trauma and
- posttraumatic stress disorder in women with chronic pelvic pain. *Obstet Gynecol*.
- 438 2007;109(4):902-908. doi:10.1097/01.AOG.0000258296.35538.88
- 439 21. Fishbain DA, Pulikal A, Lewis JE, Gao J. Chronic Pain Types Differ in Their Reported
- Prevalence of Post -Traumatic Stress Disorder (PTSD) and There Is Consistent Evidence
- That Chronic Pain Is Associated with PTSD: An Evidence-Based Structured Systematic
- 442 Review. *Pain Med Malden Mass*. 2017;18(4):711-735. doi:10.1093/pm/pnw065
- 443 22. Bornstein J, Goldstein AT, Stockdale CK, et al. 2015 ISSVD, ISSWSH and IPPS Consensus
- Terminology and Classification of Persistent Vulvar Pain and Vulvodynia. *Obstet Gynecol*.
- 445 2016;127(4):745-751. doi:10.1097/AOG.000000000001359
- 446 23. Lamvu G, Carrillo J, Ouyang C, Rapkin A. Chronic Pelvic Pain in Women: A Review.
- 447 *JAMA*. 2021;325(23):2381-2391. doi:10.1001/JAMA.2021.2631

- 448 24. Alimi Y, Iwanaga J, Oskouian RJ, Loukas M, Tubbs RS. The clinical anatomy of dyspareunia: A review. *Clin Anat N Y N*. 2018;31(7):1013-1017. doi:10.1002/ca.23250
- 450 25. Reisner SL, Pletta DR, Potter J, Deutsch MB. Initial Psychometric Evaluation of a Brief
- 451 Sexual Functioning Screening Tool for Transmasculine Adults: Transmasculine Sexual
- 452 Functioning Index. Sex Med. 2020;8(3):350-360. doi:10.1016/j.esxm.2020.05.006
- 453 26. McCool-Myers M, Theurich M, Zuelke A, Knuettel H, Apfelbacher C. Predictors of female
- sexual dysfunction: a systematic review and qualitative analysis through gender inequality
- 455 paradigms. *BMC Womens Health*. 2018;18:108. doi:10.1186/s12905-018-0602-4
- 456 27. Eftekhar T, Sohrabvand F, Zabandan N, Shariat M, Haghollahi F, Ghahghaei-Nezamabadi
- 457 A. Sexual dysfunction in patients with polycystic ovary syndrome and its affected domains.
- 458 *Iran J Reprod Med*. 2014;12(8):539-546.
- 459 28. O'Connor A, Gracie DJ, Hamlin PJ, Ford AC. Predictors of Dyspareunia Among Female
- Patients With Inflammatory Bowel Disease. Clin Gastroenterol Hepatol Off Clin Pract J
- 461 Am Gastroenterol Assoc. 2020;18(4):1000-1001. doi:10.1016/j.cgh.2019.07.065
- 462 29. Sakinci M, Ercan CM, Olgan S, Coksuer H, Karasahin KE, Kuru O. Comparative analysis
- of copper intrauterine device impact on female sexual dysfunction subtypes. *Taiwan J*
- 464 *Obstet Gynecol.* 2016;55(1):30-34. doi:10.1016/j.tjog.2014.12.011
- 465 30. Rosen NO, Dawson SJ, Binik YM, et al. Trajectories of Dyspareunia From Pregnancy to 24
- 466 Months Postpartum. *Obstet Gynecol*. 2022;139(3):391-399.
- 467 doi:10.1097/AOG.0000000000004662
- 468 31. Nguyen HB, Chavez AM, Lipner E, et al. Gender-Affirming Hormone Use in Transgender
- Individuals: Impact on Behavioral Health and Cognition. *Curr Psychiatry Rep.*
- 470 2018;20(12):110. doi:10.1007/s11920-018-0973-0
- 471 32. Foster Skewis L, Bretherton I, Leemaqz SY, Zajac JD, Cheung AS. Short-Term Effects of
- Gender-Affirming Hormone Therapy on Dysphoria and Quality of Life in Transgender
- Individuals: A Prospective Controlled Study. *Front Endocrinol*. 2021;12:717766.
- 474 doi:10.3389/fendo.2021.717766
- 475 33. Owen-Smith AA, Gerth J, Sineath RC, et al. Association Between Gender Confirmation
- 476 Treatments and Perceived Gender Congruence, Body Image Satisfaction, and Mental Health
- in a Cohort of Transgender Individuals. *J Sex Med.* 2018;15(4):591-600.
- 478 doi:10.1016/j.jsxm.2018.01.017
- 479 34. Xie Y, Shi L, Xiong X, Wu E, Veasley C, Dade C. Economic burden and quality of life of
- vulvodynia in the United States. Curr Med Res Opin. 2012;28(4):601-608.
- 481 doi:10.1185/03007995.2012.666963
- 482 35. Schneider MP, Vitonis AF, Fadayomi AB, Charlton BM, Missmer SA, DiVasta AD. Quality
- of Life in Adolescent and Young Adult Women With Dyspareunia and Endometriosis. J

| 484 485 | | Adolesc Health Off Publ Soc Adolesc Med. 2020;67(4):557-561. doi:10.1016/j.jadohealth.2020.02.024 |
|-------------------|-----|---|
| 486 487 488 | 36. | Obedin-Maliver J. Pelvic pain and persistent menses in transgender men. UCSF Transgender Care & Treatment Guidelines. Published 2016. Accessed December 7, 2022 https://transcare.ucsf.edu/guidelines/pain-transmen |
| 489 490 491 | 37. | Krause M, Wheeler TL, Snyder TE, Richter HE. Local Effects of Vaginally Administered Estrogen Therapy: A Review. <i>J Pelvic Med Surg</i> . 2009;15(3):105. doi:10.1097/SPV.0B013E3181AB4804 |
| 492 493 494 | 38. | Krause M, Wheeler TL, Richter HE, Snyder TE. Systemic Effects of Vaginally Administered Estrogen Therapy: A Review. <i>Female Pelvic Med Reconstr Surg</i> . 2010;16(3):188. doi:10.1097/SPV.0B013E3181D7E86E |
| 495 496 497 | 39. | Weber MA, Kleijn MH, Langendam M, Limpens J, Heineman MJ, Roovers JP. Local Oestrogen for Pelvic Floor Disorders: A Systematic Review. <i>PloS One</i> . 2015;10(9). doi:10.1371/JOURNAL.PONE.0136265 |
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500 Tables and Figures501

Table 1. Participant Characteristics

| Table 1. Participant Characteristics | | | | | |
|---|------------------|--|--|--|--|
| N | 1219 | | | | |
| Age, years (median, IQR) | 27.1 (22.6-33.0) | | | | |
| Gender Identity ¹ (n, %) | | | | | |
| Agender | 166 (13.6) | | | | |
| Genderqueer | 410 (33.6) | | | | |
| Man | 262 (21.5) | | | | |
| Non-binary | 661 (54.2) | | | | |
| Transgender man | 561 (46.0) | | | | |
| Two-spirit | 14 (1.1) | | | | |
| Questioning | 39 (3.2) | | | | |
| Another gender identity | 166 (13.6) | | | | |
| Sexual Orientation ¹ (n, %) | | | | | |
| Asexual | 262 (21.5) | | | | |
| Bisexual | 406 (33.3) | | | | |
| Gay | 227 (18.6) | | | | |
| Lesbian | 85 (7.0) | | | | |
| Pansexual | 245 (20.1) | | | | |
| Queer | 793 (65.1) | | | | |
| Same-gender loving | 60 (4.9) | | | | |
| Straight/heterosexual | 49 (4.0) | | | | |
| Two-spirit | 5 (0.4) | | | | |
| Questioning | 58 (4.8) | | | | |
| Another sexual orientation | 84 (6.9) | | | | |
| Race and Ethnicity ¹ (n, %) | | | | | |
| American Indian or Alaskan Native | 39 (3.2) | | | | |
| Asian | 59 (4.8) | | | | |
| Black, African American or African | 49 (4.0) | | | | |
| Hispanic, Latinx, or Spanish | 82 (6.7) | | | | |
| Middle Eastern or North African | 15 (1.2) | | | | |
| Native Hawaiian or Pacific Islander | 5 (0.4) | | | | |
| White | 1120 (91.9) | | | | |
| Another race/ethnicity | 16 (1.3) | | | | |
| Missing | 10 (0.8) | | | | |
| Derticipants were able to select more than one response therefore | | | | | |

¹Participants were able to select more than one response, therefore, proportions may sum to greater than 100%. 61.1% selected more than one gender identity, 53.2% selected more than one sexual orientation, and 12.0% selected more than one race and/or ethnicity.

Table 2. Sexual Behavior and Medical History of Participants, Stratified by Never or Current Testosterone Use

| | Overall | Never Testosterone Use | Current Testosterone Use | p- value |
|---|-------------|---------------------------|-----------------------------|-------------|
| N | 1219 | 516 | 602 | |
| Sex in the past year (n, %) | 750 (68.9) | 291 (63.4) | 400 (73.3) | 0.001 |
| Receptive oral sex ¹ | 598 (79.8) | 218 (74.9) | 333 (83.5) | 0.008 |
| Receptive vaginal/FGO sex ¹ | 400 (53.9) | 174 (59.8) | 193 (49.1) | 0.007 |
| Interest in sexual activity, past 30 days ² (mean, sd) | 46.6 (11.1) | 44.2 (11.3) | 49.0 (10.5) | < 0.001 |
| Any sexual activity, past 30 days ³ (n, %) | 1091 (89.5) | 437 (84.7) | 565 (93.9) | <0.001 |
| In a relationship (n, %) | 701 (59.4) | 291 (58.3) | 355 (60.5) | 0.509 |
| Gender(s) of past year sex partners (n, %) | | | | |
| Cisgender men | 354 (29.0) | 147 (28.5) | 182 (30.2) | 0.567 |
| Cisgender women | 286 (23.5) | 107 (20.7) | 160 (26.6) | 0.027 |
| Genderqueer, non-binary, or gender non-conforming people AFAB | 183 (15.0) | 75 (14.5) | 92 (15.3) | 0.791 |
| Genderqueer, non-binary, or gender non- conforming people AMAB | 125 (10.3) | 52 (10.1) | 60 (10.0) | >0.999 |
| Transgender Men | 112 (9.2) | 25 (4.8) | 81 (13.5) | <0.001 |
| Transgender Women | 103 (8.4) | 37 (7.2) | 57 (9.5) | 0.203 |
| Pelvic Health History (n, %) | | | | |
| Pelvic inflammatory disease | 27 (2.2) | 13 (2.5) | 11 (1.8) | 0.556 |
| Polycystic ovary syndrome | 120 (9.8) | 51 (9.9) | 55 (9.1) | 0.747 |
| Uterine Fibroids | 50 (4.1) | 24 (4.7) | 22 (3.7) | 0.493 |
| Inflammatory bowel disease ⁴ | 24 (2.0) | 11 (2.1) | 12 (2.0) | >0.999 |
| Irritable bowel syndrome | 218 (17.9) | 86 (16.7) | 112 (18.6) | 0.443 |
| Bacterial STI diagnosis ⁵ | 19 (1.6) | 1 (0.2) | 18 (3.0) | 0.001 |
| Ever pregnant | 130 (10.7) | 60 (11.6) | 56 (9.3) | 0.241 |
| Hysterectomy | 142 (11.6) | 28 (5.4) | 108 (17.9) | <0.001 |
| Oophorectomy | 108 (8.9) | 17 (3.3) | 87 (14.5) | <0.001 |
| Current hormonal contraceptive use ⁶ | 138 (11.3) | 92 (17.8) | 35 (5.8) | <0.001 |
| Current hormonal intrauterine device use ⁷ | 91 (8.4) | 49 (10.0) | 32 (6.5) | 0.056 |
| Current non-hormonal intrauterine device use ⁷ | 29 (2.7) | 11 (2.3) | 16 (3.2) | 0.454 |
| Mental Health and Substance Use (n, %) | | | | |
| History of sexual abuse | 686 (78.0) | 289 (76.3) | 340 (78.3) | 0.532 |
| Ever experienced rape or sexual assault | 437 (49.9) | 196 (51.9) | 206 (47.7) | 0.266 |
| Ever diagnosed with depression | 988 (81.1) | 399 (77.3) | 503 (83.6) | 0.011 |
| PHQ-9 score for depression8 (mean, sd) | 10.2 (6.4) | 10.7 (6.4) | 9.6 (6.2) | 0.005 |
| Ever diagnosed with PTSD | 484 (39.7) | 198 (38.4) | 238 (39.5) | 0.737 |
| PCL-6 score for PTSD ⁹ (mean, sd) | 15.6 (5.3) | 16.1 (5.2) | 15.0 (5.2) | 0.001 |
| Current smoker | 74 (6.1) | 22 (4.3) | 45 (7.5) | 0.033 |
| AUDIT score for alcohol use10 (mean, sd) | 3.6 (4.3) | 3.2 (3.8) | 3.9 (4.6) | 0.005 |
| Language Preferences (n %) | | | | |

Language Preferences (n, %)

| Vagina | 891 (73.1) | 428 (82.9) | 386 (64.1) | < 0.001 |
|-------------------------|------------|------------|------------|---------------|
| Frontal genital opening | 315 (25.8) | 86 (16.7) | 207 (34.4) | \0.001 |

AFAB, assigned female at birth; AMAB, assigned male at birth; AUDIT, alcohol use identification test; PHQ-9, 9-item patient health questionnaire; PTSD, post-traumatic stress disorder; sd, standard deviation

¹Among participants who reported having sex in the past year

²T-scores of 50 represents the population average for the US population, and 10 points represents one standard deviation from the population average

³Including masturbation as well as sexual activity with a partner

⁴Including Crohn's disease, ulcerative colitis, etc

⁵Diagnosis with chlamydia, gonorrhea, or syphilis in the past year

⁶Including oral contraceptives, transdermal patch, vaginal rings, medroxyprogesterone acetate injections, and etonogestrel implants ⁷Excluding participants who had a hysterectomy

⁸The PHQ-9 measures depressive symptoms in the past two weeks, where scores of 10 or higher are suggestive of moderate to severe depression

⁹The PCL-6 measures PTSD symptoms in the past month, where scores of 17 or higher are associated with probable PTSD

¹⁰The AUDIT measures current alcohol consumptions behaviors, where scores of 15 or higher are suggestive of alcohol use disorder 503

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Table 3. Sexual Function Stratified by Never and Current Testosterone Use

| Overall | Never Testosterone Use | Current Testosterone Use | p- value |
|------------|--|--|--|
| 128 | 79 | 37 | |
| | | | |
| 93 (72.7) | 67 (84.8) | 17 (45.9) | < 0.001 |
| 43 (33.6) | 26 (32.9) | 14 (37.8) | 0.756 |
| 32 (25.0) | 23 (29.1) | 4 (10.8) | 0.053 |
| 20 (15.6) | 10 (12.7) | 7 (18.9) | 0.544 |
| 15 (11.7) | 10 (12.7) | 5 (13.5) | >0.999 |
| 11 (8.6) | 4 (5.1) | 5 (13.5) | 0.225 |
| 9 (7.0) | 5 (6.3) | 2 (5.4) | >0.999 |
| 18 (14.1) | 12 (15.2) | 6 (16.2) | >0.999 |
| 1091 | 437 | 565 | |
| | | | |
| 670 (62.2) | 289 (66.7) | 326 (58.5) | 0.010 |
| 50.7 (8.5) | 50.2 (8.4) | 51.2 (8.4) | 0.045 |
| 693 (64.6) | 274 (63.3) | 368 (66.5) | 0.317 |
| | | | |
| 561 (52.2) | 213 (49.2) | 310 (56.0) | 0.040 |
| 51.6 (9.1) | 50.9 (8.9) | 52.3 (9.2) | 0.013 |
| | | | |
| 263 (24.5) | 118 (27.3) | 126 (22.8) | 0.124 |
| 51.2 (7.5) | 51.6 (7.7) | 51.0 (7.4) | 0.168 |
| | | | |
| 312 (29.1) | 117 (27.1) | 167 (30.2) | 0.317 |
| 53.1 (8.3) | 52.6 (7.7) | 53.4 (8.5) | 0.136 |
| | | | |
| 61 (5.6) | 34 (7.8) | 19 (3.4) | 0.003 |
| 559 (52.8) | 216 (51.2) | 289 (52.2) | 0.811 |
| 49.2 (9.8) | 48.4 (10.8) | 50.1 (9.0) | 0.010 |
| 45.0 (8.4) | 45.1 (8.3) | 45.2 (8.3) | 0.918 |
| | | | |
| 45.6 (7.7) | 45.7 (7.9) | 45.9 (7.6) | 0.670 |
| | 93 (72.7) 43 (33.6) 32 (25.0) 20 (15.6) 15 (11.7) 11 (8.6) 9 (7.0) 18 (14.1) 1091 670 (62.2) 50.7 (8.5) 693 (64.6) 561 (52.2) 51.6 (9.1) 263 (24.5) 51.2 (7.5) 312 (29.1) 53.1 (8.3) 61 (5.6) 559 (52.8) 49.2 (9.8) 45.0 (8.4) | Overall Testosterone Use 128 79 93 (72.7) 67 (84.8) 43 (33.6) 26 (32.9) 32 (25.0) 23 (29.1) 20 (15.6) 10 (12.7) 15 (11.7) 10 (12.7) 11 (8.6) 4 (5.1) 9 (7.0) 5 (6.3) 18 (14.1) 12 (15.2) 1091 437 670 (62.2) 289 (66.7) 50.7 (8.5) 50.2 (8.4) 693 (64.6) 274 (63.3) 561 (52.2) 213 (49.2) 51.6 (9.1) 50.9 (8.9) 263 (24.5) 118 (27.3) 51.2 (7.5) 51.6 (7.7) 312 (29.1) 117 (27.1) 53.1 (8.3) 52.6 (7.7) 61 (5.6) 34 (7.8) 559 (52.8) 216 (51.2) 49.2 (9.8) 48.4 (10.8) 45.0 (8.4) 45.1 (8.3) | Overall Testosterone Use Testosterone Use 128 79 37 93 (72.7) 67 (84.8) 17 (45.9) 43 (33.6) 26 (32.9) 14 (37.8) 32 (25.0) 23 (29.1) 4 (10.8) 20 (15.6) 10 (12.7) 7 (18.9) 15 (11.7) 10 (12.7) 5 (13.5) 11 (8.6) 4 (5.1) 5 (13.5) 9 (7.0) 5 (6.3) 2 (5.4) 18 (14.1) 12 (15.2) 6 (16.2) 1091 437 565 670 (62.2) 289 (66.7) 326 (58.5) 50.7 (8.5) 50.2 (8.4) 51.2 (8.4) 693 (64.6) 274 (63.3) 368 (66.5) 561 (52.2) 213 (49.2) 310 (56.0) 51.6 (9.1) 50.9 (8.9) 52.3 (9.2) 263 (24.5) 118 (27.3) 126 (22.8) 51.2 (7.5) 51.6 (7.7) 51.0 (7.4) 312 (29.1) 117 (27.1) 167 (30.2) 53.1 (8.3) 52.6 (7.7) 53.4 (8.5) 61 (5.6) 34 (7.8)< |

FGO, frontal genital opening

T-scores of 50 represents the population average for the US population, and 10 points represents one standard deviation from the population average

¹High scores indicate more lubrication, ability to achieve orgasm, pleasure from orgasms, and satisfaction with sex life

²Higher scores indicate more pain/discomfort

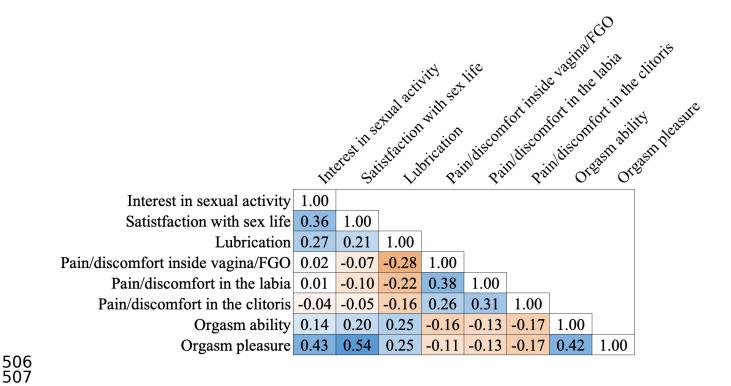


Figure 1. Correlation between sexual function T-scores.

The correlation coefficient measures the strength and directional of the linear association between two variables. It ranges from -1 to 1, with zero indicating no correlation. Blue indicates that variables are positive correlated with one another, and orange indicates variables are negatively correlated. FGO, frontal genital opening.

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Table 4. Association between Current Testosterone Use and Sexual Function

| | Current Testosterone Use vs Never Testosterone Use | | | | | |
|-----------------------------------|--|---------|---------------------------------|---------|-----------------------------|---------|
| | Unadjusted | | Minimally Adjusted ² | | Fully Adjusted ³ | |
| Outcomes ¹ | β (95% CI) | p-value | β (95% CI) | p-value | β (95% CI) | p-value |
| Interest in sexual activity | 6.44 (5.10, 7.78) | <0.001 | 6.32 (4.91, 7.74) | <0.001 | 6.02 (4.35, 7.69) | <0.001 |
| Satisfaction with sex life | 0.21 (-0.75, 1.17) | 0.669 | 0.16 (-0.85, 1.16) | 0.762 | 0.42 (-0.75, 1.58) | 0.486 |
| Lubrication | 1.08 (0.01, 2.15) | 0.048 | 0.61 (-0.52, 1.75) | 0.289 | 1.15 (-0.14, 2.44) | 0.081 |
| Pain/discomfort inside vagina/FGO | 1.45 (0.31, 2.59) | 0.013 | 1.80 (0.61, 3.00) | 0.003 | 1.95 (0.53, 3.36) | 0.007 |
| Pain/discomfort in the labia | -0.67 (-1.61, 0.28) | 0.168 | -0.38 (-1.38, 0.62) | 0.455 | -0.75 (-1.93, 0.44) | 0.218 |
| Pain/discomfort in the clitoris | 0.78 (-0.25, 1.82) | 0.139 | 1.20 (0.10, 2.30) | 0.033 | 1.21 (-0.09, 2.51) | 0.068 |
| Orgasm ability | 1.62 (0.39, 2.86) | 0.010 | 1.50 (0.19, 2.81) | 0.025 | 0.36 (-1.14, 1.86) | 0.636 |
| Orgasm pleasure | 0.06 (-1.02, 1.13) | 0.919 | -0.04 (-1.15, 1.08) | 0.951 | 0.46 (-0.81, 1.73) | 0.475 |

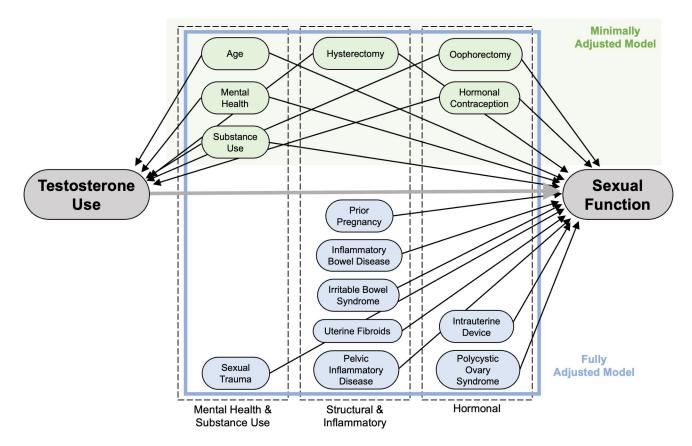
CI, confidence interval; FGO, frontal genital opening

¹T-scores of 50 represents the population average for the US population, and 10 points represents one standard deviation from the population average

²Adjusted for age, current depression symptoms (PHQ-9 scores), current post-traumatic stress symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, and hormonal contraception use

³Adjusted for age, current depression symptoms (PHQ-9 scores), current post-traumatic stress symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, intrauterine devices use, hormonal contraception use history of sexual assault, inflammatory bowel disease, irritable bowel syndrome, uterine fibroids, pelvic inflammatory disease, polycystic ovary syndrome, prior pregnancy, and intrauterine devices use

Supplemental Materials



Supplemental Figure 1. Directed acyclic graph (DAG) illustrating covariates included in regression models

Supplemental Table 1. Participant Characteristics Stratified by Never, Current, and Former Testosterone Use

| | Never Testosterone Use | Current Testosterone Use | p-value (Never v· Current) | Former Testosterone Use | p-value (Current v· Former) |
|--|------------------------------|--------------------------------|----------------------------------|-------------------------------|-----------------------------------|
| N | 516 | 602 | | 76 | |
| Age, years (mean, range) | 28.3 (8.7) | 29.9 (9.6) | 0.003 | 30.7 (9.9) | 0.505 |
| Gender Identity ¹ (n, %) | | | | | |
| Agender | 99 (19·2) | 50 (8.3) | <0.001 | 15 (19.7) | 0.003 |
| Genderqueer | 234 (45·3) | 135 (22.4) | <0.001 | 30 (39.5) | 0.002 |
| Man | 21 (4.1) | 231 (38.4) | <0.001 | 8 (10.5) | <0.001 |
| Non-binary | 379 (73.4) | 209 (34.7) | <0.001 | 54 (71·1) | <0.001 |
| Transgender man | 67 (13.0) | 458 (76·1) | <0.001 | 31 (40.8) | <0.001 |
| Two-spirit | 8 (1.6) | 5 (0.8) | 0.401 | 1 (1.3) | 1.000 |
| Questioning | 27 (5.2) | 10 (1.7) | 0.002 | 1 (1.3) | 1.000 |
| Another gender identity | 86 (16.7) | 59 (9.8) | 0.001 | 16 (21·1) | 0.006 |
| Sexual Orientation ¹ (n, %) | | | | | |
| Asexual | 145 (28·1) | 95 (15.8) | <0.001 | 17 (22.4) | 0.196 |
| Bisexual | 173 (33.5) | 203 (33.7) | 0.996 | 21 (27.6) | 0.350 |
| Gay | 56 (10.9) | 154 (25.6) | <0.001 | 16 (21·1) | 0.473 |
| Lesbian | 59 (11.4) | 14 (2.3) | <0.001 | 7 (9.2) | 0.004 |
| Pansexual | 115 (22.3) | 108 (17.9) | 0.082 | 14 (18.4) | 1.000 |
| Queer | 336 (65·1) | 384 (63.8) | 0.689 | 58 (76.3) | 0.042 |
| Same-gender loving | 17 (3.3) | 39 (6.5) | 0.022 | 3 (3.9) | 0.542 |
| Straight/heterosexual | 6 (1.2) | 42 (7.0) | <0.001 | 1 (1.3) | 0.097 |
| Two-spirit | 2 (0.4) | 2 (0.3) | 1.000 | 1 (1.3) | 0.764 |
| Questioning | 18 (3.5) | 34 (5.6) | 0.117 | 4 (5.3) | 1.000 |
| Another sexual orientation | 39 (7.6) | 34 (5.6) | 0.243 | 9 (11.8) | 0.066 |
| Race and Ethnicity ¹ (n, %) | | | | | |
| American Indian or Alaskan Native | 23 (4.5) | 15 (2.5) | 0.100 | 1 (1.3) | 0.814 |
| Asian | 25 (4.8) | 28 (4.7) | 0.991 | 6 (7.9) | 0.346 |
| Black, African American or African | 19 (3.7) | 29 (4.8) | 0.432 | 1 (1.3) | 0.270 |
| Hispanic, Latinx, or Spanish | 36 (7.0) | 39 (6.5) | 0.832 | 6 (7.9) | 0.824 |
| Middle Eastern or North African | 6 (1.2) | 7 (1.2) | 1.000 | 2 (2.6) | 0.601 |
| Native Hawaiian or Pacific Islander | 1 (0.2) | 3 (0.5) | 0.728 | 0 (0.0) | 1.000 |
| White | 480 (93.0) | 551 (91.5) | 0.413 | 72 (94.7) | 0.458 |
| Another race/ethnicity | 8 (1.6) | 7 (1.2) | 0.764 | 1 (1.3) | 1.000 |
| Missing | 0 (0.0) | 2 (0.3) | 0.548 | 0(0.0) | 1.000 |

¹Participants were able to select more than one response, therefore, proportions may sum to greater than 1

Supplemental Table 2. Sexual Behavior and Medical History of Participants, Stratified by Current and Former Testosterone Use

| | Current | Former | n valua |
|---|-------------------------|-------------------------|---------|
| | Testosterone Use | Testosterone Use | p-value |
| N | 602 | 76 | |
| Sex in the past year (n, %) | 400 (73.3) | 50 (75.8) | 0.774 |
| Interest in sexual activity, past 30 days ¹ (mean, sd) | 48.96 (10.54) | 41.99 (9.93) | <0.001 |
| Any sexual activity, past 30 days ² (n, %) | 565 (93.9) | 64 (84.2) | 0.005 |
| In a relationship (n, %) | 355 (60.5) | 45 (61.6) | 0.948 |
| Gender(s) of past year sex partners (n, %) | | | |
| Cisgender men | 182 (30.2) | 18 (23.7) | 0.296 |
| Cisgender women | 160 (26.6) | 16 (21·1) | 0.370 |
| Genderqueer, non-binary, or gender non- conforming people AFAB | 92 (15·3) | 14 (18·4) | 0.588 |
| Genderqueer, non-binary, or gender non- conforming people AMAB | 60 (10.0) | 13 (17·1) | 0.090 |
| Transgender Men | 81 (13.5) | 6 (7.9) | 0.237 |
| Transgender Women | 57 (9.5) | 8 (10.5) | 0.930 |
| Pelvic Health History (n, %) | | | |
| Pelvic inflammatory disease | 11 (1.8) | 3 (3.9) | 0.426 |
| Polycystic ovary syndrome | 55 (9·1) | 9 (11.8) | 0.581 |
| Uterine Fibroids | 22 (3.7) | 4 (5·3) | 0.711 |
| Inflammatory bowel disease ³ | 12 (2.0) | 1 (1.3) | 1.000 |
| Irritable bowel syndrome | 112 (18.6) | 15 (19.7) | 0.934 |
| Bacterial STI diagnosis ⁴ | 18 (3.0) | 0 (0.0) | 0.250 |
| Ever pregnant | 56 (100.0) | 12 (100.0) | 0.116 |
| Hysterectomy | 108 (100.0) | 6 (100.0) | 0.041 |
| Oophorectomy | 87 (14.5) | 4 (5·3) | 0.042 |
| Current hormonal contraceptive use ⁵ | 35 (5.8) | 8 (10.5) | 0.181 |
| Current hormonal intrauterine device use ⁶ | 32 (6.5) | 8 (11.4) | 0.207 |
| Current non-hormonal intrauterine device use ⁶ | 16 (3.2) | 2 (2.9) | 1.000 |
| Mental Health and Substance Use (n, %) | | | |
| History of sexual abuse | 340 (78.3) | 46 (88.5) | 0.127 |
| Ever experienced rape or sexual assault | 206 (47.7) | 29 (55.8) | 0.340 |
| Ever diagnosed with depression | 503 (83.6) | 70 (92·1) | 0.076 |
| PHQ-9 score for depression ⁷ (mean, sd) | 9.6 (6.2) | 11.08 (7.12) | 0.053 |
| Ever diagnosed with PTSD | 238 (39.5) | 42 (55.3) | 0.012 |
| PCL-6 score for PTSD ⁸ (mean, sd) | 15.1 (5.2) | 16.8 (5.8) | 0.007 |
| Current smoker | 45 (7.5) | 6 (7.9) | 1.000 |
| AUDIT score for alcohol use9 (mean, sd) | 3.9 (4.6) | 4.0 (4.7) | 0.886 |
| Language Preferences (n, %) | | | |
| Vagina | 386 (64·1) | 54 (71·1) | 0.309 |
| Front genital opening | 207 (34.4) | 20 (26.3) | 0.309 |

AFAB, assigned female at birth; AMAB, assigned male at birth; AUDIT, alcohol use identification test; PHQ-9, 9-item patient health questionnaire; PTSD, post-traumatic stress disorder

¹T-scores of 50 represents the population average for the US population, and 10 points represents one standard deviation from the population average

²Including masturbation as well as sexual activity with a partner

³Including Crohn's disease, ulcerative colitis, etc

⁴Diagnosis with chlamydia, gonorrhea, or syphilis in the past year

⁵Including oral contraceptives, transdermal patch, vaginal rings, medroxyprogesterone acetate injections, and etonogestrel implants

⁶Excluding participants who had a hysterectomy

⁷The PHQ-9 measures depressive symptoms in the past two weeks, where scores of 10 or higher are suggestive of moderate to severe depression

⁸The PCL-6 measures PTSD symptoms in the past month, where scores of 17 or higher are associated with probable PTSD
⁹The AUDIT measures current alcohol consumptions behaviors, where scores of 15 or higher are suggestive of alcohol use disorder

Supplemental Table 3. Sexual Function Stratified by Current and Former Testosterone Use

| | Current Testosterone Use | Former Testosterone Use | p-value |
|--|-----------------------------|----------------------------|---------|
| Participants who were NOT sexually active in the past 30 days, N | 37 | 12 | |
| Reasons for No Sexual Activity (n, %) | | | |
| Not interested | 17 (45.9) | 9 (75.0) | 0.156 |
| Dryness or pain in or around my vaginal/FGO | 5 (13.5) | 2 (16·7) | 1.000 |
| Difficulties with orgasm/climax | 5 (13.5) | 0(0.0) | 0.427 |
| Don't enjoy sexual activity | 4 (10.8) | 5 (41.7) | 0.049 |
| Health condition | 2 (5.4) | 2 (16·7) | 0.528 |
| No partners | 14 (37.8) | 3 (25.0) | 0.643 |
| Partner(s) away, not interested, or health condition | 7 (18.9) | 3 (25.0) | 0.966 |
| Another reason | 6 (16.2) | 0(0.0) | 0.326 |
| Participants who WERE sexually active in the past 30 days, N | 565 | 64 | |
| Lubrication | | | |
| Any difficulty achieving or maintaining lubrication (n, %) | 326 (58.5) | 41 (66·1) | 0.308 |
| T-score ¹ (mean, sd) | 51.2 (8.4) | 48.6 (10.2) | 0.020 |
| Pain/discomfort inside vagina/FGO | | | |
| Any pain or discomfort (n, %) | 310 (56.0) | 27 (43.5) | 0.084 |
| T-score ² (mean, sd) | 52.3 (9.2) | 50.7 (9.6) | 0.182 |
| Pain/discomfort in the labia | | | |
| Any pain or discomfort (n, %) | 126 (22.8) | 15 (24-2) | 0.928 |
| T-score ² (mean, sd) | 51.0 (7.4) | 51.4 (7.9) | 0.664 |
| Pain/discomfort in the clitoris | | | |
| Any pain or discomfort (n, %) | 167 (30.2) | 21 (34.4) | 0.594 |
| T-score ² (mean, sd) | 53.4 (8.5) | 54.3 (9.3) | 0.419 |
| Orgasm | | | |
| Did not have an orgasm (n, %) | 19 (3.4) | 6 (9.4) | 0.047 |
| Any difficulty achieving orgasm (n, %) | 289 (52.2) | 41 (67-2) | 0.036 |
| T-score for achieving orgasm¹ (mean, sd) | 50.1 (9.0) | 46.1 (10.7) | 0.001 |
| T-score for orgasm pleasure ¹ (mean, sd) | 45.2 (8.3) | 42.3 (9.0) | 0.013 |
| Satisfaction with sex life | | | |
| T-score ¹ (mean, sd) | 45.9 (7.6) | 42.9 (7.0) | 0.003 |
| FGO. frontal genital opening | 45.9 (7.6) | 42.9 (7.0) | 0.003 |

FGO, frontal genital opening

T-scores of 50 represents the population average for the US population, and 10 point represents one standard deviation from the population average

¹High scores indicate more lubrication, ability to achieve orgasm, pleasure from orgasms, and satisfaction with sex life

²Higher scores indicate more pain/discomfort

Supplemental Table 4. Association between Duration of Testosterone Use (Years) and Sexual Function

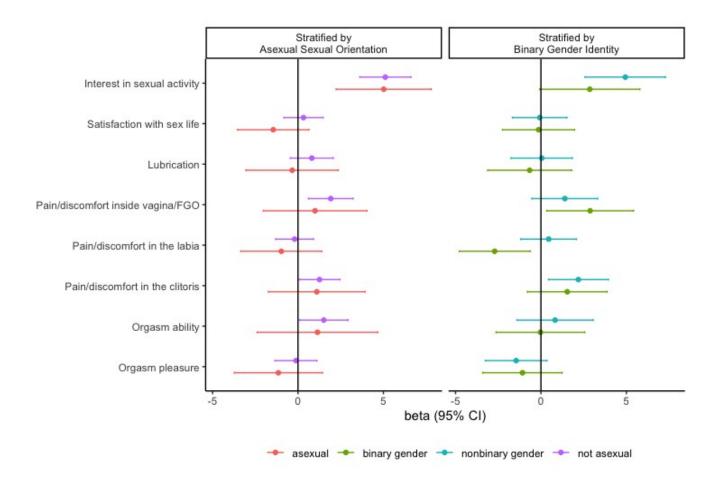
| | Unadjusted | | Minimally Adjusted ² | | Fully Adjusted ³ | |
|-----------------------------------|---------------------|---------|---------------------------------|---------|-----------------------------|---------|
| Outcomes ¹ | beta (95% CI) | p-value | beta (95% CI) | p-value | beta (95% CI) | p-value |
| Interest in sexual activity | 0.38 (0.20, 0.56) | <0.001 | 0.39 (0.18, 0.59) | <0.001 | 0.35 (0.11, 0.60) | 0.005 |
| Satisfaction with sex life | -0.01 (-0.13, 0.12) | 0.917 | 0.00 (-0.14, 0.14) | 0.969 | 0.01 (-0.16, 0.17) | 0.949 |
| Lubrication | 0.10 (-0.04, 0.23) | 0.165 | 0.11 (-0.04, 0.27) | 0.161 | 0.17 (-0.01, 0.35) | 0.062 |
| Pain/discomfort inside vagina/FGO | 0.01 (-0.14, 0.15) | 0.945 | 0.06 (-0.11, 0.23) | 0.500 | 0.05 (-0.14, 0.25) | 0.599 |
| Pain/discomfort in the labia | -0.10 (-0.22, 0.02) | 0.099 | -0.07 (-0.21, 0.07) | 0.325 | -0.08 (-0.24, 0.09) | 0.367 |
| Pain/discomfort in the clitoris | -0.17 (-0.3, -0.04) | 0.012 | -0.05 (-0.2, 0.11) | 0.557 | -0.06 (-0.24, 0.12) | 0.521 |
| Orgasm ability | 0.20 (0.04, 0.35) | 0.012 | 0.16 (-0.02, 0.34) | 0.078 | 0.03 (-0.17, 0.24) | 0.753 |
| Orgasm pleasure | 0.06 (-0.07, 0.2) | 0.335 | 0.03 (-0.12, 0.18) | 0.655 | 0.08 (-0.09, 0.25) | 0.358 |

CI, confidence interval; FGO, frontal genital opening

¹T-scores of 50 represents the population average for the US population, and 10 points represents one standard deviation from the population average

²Adjusted for age, current depression symptoms (PHQ-9 scores), current post-traumatic stress symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, and hormonal contraception use

³Adjusted for age, current depression symptoms (PHQ-9 scores), current post-traumatic stress symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, intrauterine devices use, hormonal contraception use, history of sexual assault, inflammatory bowel disease, irritable bowel syndrome, uterine fibroids, pelvic inflammatory disease, polycystic ovary syndrome, prior pregnancy, and intrauterine devices use



Supplemental Figure 2. Minimally Adjusted Regression Results Stratified by Binary *versus* **Nonbinary Gender Identity.** Adjusted for age, current depression symptoms (PHQ-9 scores), current post-traumatic stress symptoms (PCL-6 scores), alcohol use (AUDIT score), current smoker, hysterectomy, oophorectomy, and hormonal contraception use.