Effect of anogenital distance on stress urinary incontinence

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Abstract
Background: To clarify the effect of anogenital distance (AGD) on stress urinary incontinence (SUI) in female patients.
Methods: Charts of patients who admitted to urogynecology polyclinic between December 2020 and February 2021 were analyzed retrospectively. The AGD parameters including anogenital distance from the anus to the clitoris (AGDAC), anogenital distance from the anus to the fourchette (AGDAF) and genital hiatus (GH) were measured. To identify effect of AGD parameters on SUI, patients were divided into the two subgroups (patients with SUI and without SUI). Demographic characteristics of patients and perineal anatomy measurement were compared between groups.
Results: Totally, 256 female patients met study inclusion criteria. In comparison of patients with and without SUI, and demonstrated that age, weight, height, and BMI were similar between groups (p = 0.200, p = 0.455, p = 0.131, and p = 0.215, respectively). The AGDAF was measured 22.6 mm in patients with SUI and 25.5 mm in patients without SUI (p = 0.014). In contrast, AGDAC was significantly longer in patients with SUI (81.1 mm vs. 72.2 mm, p = 0.001). Also, GH was significantly longer in patients with SUI (p = 0.016). Multivariate logistic regression analysis revealed that one mm increment in AGDAC and GH, is associated with 1.108- and 1.038-fold time of SUI development risk, respectively (p = 0.001 and p = 0.004). In contrast, decrease in AGDAF is resulted with significantly higher risk for SUI (p = 0.001).
Conclusion: The present study demonstrated that shorter AGDAF, and longer AGDAC and GH were resulted with significantly higher incidence of SUI. Considering the ease and non-invasiveness of anogenital distance measurement, an important implication of the results of current study is that it is a measurement that can be easily performed during routine gynecological and obstetric examinations as a suitable tool for use in the prediction of women who will develop stress incontinence in the future.

Keywords: Anogenital distance, Genital hiatus, Stress urinary incontinence, Female, Reproductive age

1 Background
Stress urinary incontinence (SUI) is described as any involuntary loss of urine with increased abdominal pressure such as coughing, sneezing or physical exercise, according to the International Continence Society [1]. Incidence of SUI is very high in female population and Jha et al. demonstrated that quarter of women experienced with life quality deterioration due to SUI in their lifespan [2]. Factors affecting on SUI are still under investigation and previous reports tried to identify reasons for SUI including age, body mass index (BMI), number of births, previous gynecological surgery history, and perineal anatomic properties [3].

Effects of perineal anatomy characteristics on SUI is one of the hottest topic in urogynecology. Shin et al. investigated the relation between urethral length and SUI, and authors claimed that patients with SUI had a significantly shorter urethral length than in patients without SUI [4]. In another study, Athanasopoulos and colleagues evaluated the correlation between perineal measurement parameters (perineal body length, fourchette-coccyx distance, anal-coccyx length) and SUI in women; however, authors found perineal measurement parameters to be not significantly associated with SUI [5].
parameters had no effect on SUI [5]. Distance between the anus and external genital, also called anogenital distance (AGD), is dimorphic sexual characteristic, and AGD is accepted as a sign of androgen exposure during prenatal period [6]. Sanchez-Ferrer et al. analyzed the relation between AGD parameters and pelvic organ prolapse, and longer genital hiatus length and anoclitoral distance found as predictive factors for pelvic organ prolapse. However, Sanchez-Ferrer and colleagues did not focus on AGD parameters’ effects on SUI [7].

Although previous studies analyzed the effect of AGD on fertility, premature ejaculation, pelvic organ prolapses, and prostate cancer, none of study has investigated the effect of AGD on SUI. In present study, we aim to clarify the effect of AGD on SUI in female patients.

2 Methods
Charts of patients who admitted to urogynecology polyclinic between December 2020 and February 2021, were analyzed retrospectively. Ethical approval was achieved from local ethics committee, and informed consent for participate to study was signed for all patients. Detailed medical history was obtained and patients’ characteristics including age, weight, height, BMI, ASA (American society of anesthesiologists) score, and number of parity were recorded. Physical examination was performed for all patients, and AGD parameters including anogenital distance from the anus to the clitoris (AGDAC), anogenital distance from the anus to the fourchette (AGDAF) and genital hiatus (GH) were measured. Also, all patients were evaluated according to Pelvic Organ Prolapse Quantifications System (POP-Q). To diagnose SUI, all patients did Valsalva maneuver with empty bladder, then 300 cc of fluid is injected into the bladder and Valsalva maneuver performed again. Patients with urge and/or mixed incontinence type, severe mental inability, severe neurological disease, neurogenic bladder and refusal to content, were excluded from the study. Other exclusion criteria were being <18 years old, history of stress incontinence surgery, presence of gynecological tumor, rectal disease and active infection in anogenital area.

2.1 Anogenital distance measures

All AGD parameters were measured in lithotomy position with 45° angle of thighs. Stainless steel digital caliper (Supplier: VWR® International, LLC, West Chester, PA, USA) was used for all measurements. Distance between the upper edge of the anus and clitoris was accepted as AGDAC, and distance from upper edge of the anus to posterior fourchette was accepted as AGDAF, retrospectively. According to POP-Q, genital hiatus length defined from urethral meatus center to the edge of the perineum nucleus or posterior midline of hymen. To achieve more accurate outcomes, each AGD parameters were measured twice by two physician.

To identify effect of AGD parameters on SUI, patients were divided into the two subgroups (patients with SUI and without SUI). Demographic characteristics of patients and perineal anatomy measurement were compared between groups.

2.2 Data analysis

The Statistical Package for the Social Sciences version 25 (SPSS IBM Corp., Armonk, NY, USA) program was used. Normality of distribution of the variables was checked by Shapiro–Wilk test and Q–Q plots. Independent Student’s t test was used for comparison of the normally distributed variable between the groups, and Mann–Whitney U test was used for non-normally distributed data. Quantitative data are showed as mean ± standard deviation values. Categorical variables were grouped and compared using the χ² test or Fisher’s exact test. Binary logistic regression analysis was used for evaluate to parameters of anogenital distance in terms of incontinence risk. The data were analyzed at a 95% confidence level and P value of less than 0.05 was accepted as statistically significant.

3 Results

Totally, 256 female patients with mean age 36.9 years, met study inclusion criteria. The mean weight and height of patients were 74.9 kg and 162.2 cm, respectively. The mean BMI of study population was 28.5 kg/m². Only, 26 (10.2%) patients were nulliparous. The mean AGDAF and AGDAC were measured 24.9 mm and 73.8 mm, respectively. The distance of GH was 22.7 mm. According to POP-Q test, 143 patients were categorized as stage 0, and 77 patients and 36 patients were classified into stage 1 and stage 2, respectively. Demographic data of patients are summarized in (Table 1).

In comparison of patients with and without SUI, demonstrated that age, weight, height, BMI, and ASA score were similar between groups (p = 0.200, p = 0.455, p = 0.131, p = 0.215, and p = 0.867, respectively). Also, nulliparous rate was not significant different (10.9% vs. 10.0%, p = 0.964). The AGDAF was measured 22.6 mm in patients with SUI and 25.5 mm in patients without SUI (p = 0.014). In contrast, AGDAC was significantly longer in patients with SUI (81.1 mm vs. 72.2 mm, p = 0.001). Also, GH was significantly longer in patients with SUI in comparison of patients without SUI (P = 0.016). The POP-Q stages were comparable between groups (p = 0.478) (Table 2).

Multivariate logistic regression analysis revealed that one mm increment in AGDAC and GH, is associated with 1.108- and 1.038-fold time of SUI development risk, respectively (p = 0.001 and p = 0.004). In contrast,
ASA score 1.2
AGDAF (mm) 24.9
Parity
Nulliparous 26 (10.2%)
Parity n,(%) 0.964

ASA American society of anesthesiologists, AGDAF Anogenital distance from the anus to the clitoris, AGDAC Anogenital distance from the anus to the fourchette, BMI Body mass index, GH Genital hiatus, POP Q Pelvic Organ Prolapse Quantifications System

Table 1 Demographics data of all patients

|                               | n:256                |
|-------------------------------|----------------------|
| Age (years)*                 | 36.9±9.3             |
| Weight (kg)*                 | 74.9±15.9            |
| Height (cm)*                 | 162.2±5.4            |
| BMI (kg/m²)*                 | 28.5±5.8             |
| ASA score                    | 1.2±0.4              |
| Parity n,(%)                 | 0.964                |
| Nulliparous                  | 26 (10.2%)           |
| Parity ≥ 1                   | 230 (89.8%)          |
| AGDACp (mm)*                 | 73.8±12.0            |
| AGDAC (mm)*                  | 22.7±8.0             |
| POP Q stage                  | 0.478                |

| Presence of SUI (n:46)       | Absence of SUI (n:210) | p value |
|------------------------------|-------------------------|---------|
| Age (years)*                 | 38.4±9.0                | 36.6±9.3| 0.200               |
| Weight (kg)*                 | 78.1±16.7               | 74.2±15.6| 0.455               |
| Height (cm)*                 | 162.9±4.6               | 162.0±5.8| 0.131               |
| BMI (kg/m²)*                 | 29.4±6.3                | 28.3±5.6| 0.215               |
| ASA score                    | 1.2±0.4                 | 1.2±0.4| 0.867               |
| Parity n,(%)                 | 0.964                   |         |
| Nulliparous                  | 5 (10.9%)               | 21 (10.0%)| 0.656               |
| Parity ≥ 1                   | 41 (89.1%)              | 189 (90.0%)| 0.014               |
| AGDACp (mm)*                 | 22.6±7.1                | 25.5±5.5| 0.014               |
| AGDAC (mm)*                  | 81.1±9.5                | 72.2±11.9| 0.001               |
| GH (mm)*                     | 24.4±6.1                | 22.2±8.3| 0.016               |
| POP Q stage                  | 0.478                   |         |

Table 2 Comparison of patient data by stress urinary incontinence status

| Odds ratio | %95 CI | P value |
|------------|--------|---------|
| AGDAF (mm) | 0.833  | 0.772–0.898 | 0.001 |
| AGDAC (mm) | 1.108  | 1.069–1.148 | 0.001 |
| GH (mm)    | 1.038  | 1.008–1.045 | 0.004 |

AGDAC: Anogenital distance from the anus to the clitoris, AGDAF Anogenital distance from the anus to the fourchette, GH Genital hiatus

Table 3 Logistic regression analysis of anogenital distance parameters in terms of incontinence risk

A decrease in AGDAF is resulted with significantly higher risk for SUI (p=0.001) (Table 3). The ROC analyses showed that the AUC values of AGDAC, AGDAF and GH for predicting SUI were 0.706, 0.692, and 0.671, respectively. (p=0.001, p=0.003 and p=0.005, respectively) (Fig. 1).

4 Discussion

The AGD is simple and non-invasive anthropometric measurement, and the length of AGD is twice as longer in males than in females. Previous animal and human studies have demonstrated that AGD is an indirect sign of androgen exposure during prenatal period which affect the length of AGD in both sex. However, androgen exposure during prenatal period has different effects on both genders, resulted with shorter AGD in male and longer AGD in female [8, 9]. In male, shorter AGD length was found a predictive factor for atrophic testes, cryptorchidism, hypospadias, infertility, and lower testosterone hormone levels [10]. Similarly, longer AGD was detected in hormone related diseases such as endometriosis, congenital adrenal hyperplasia, and polycystic ovary syndrome [11]. First time, we analyze the AGD parameters effect on SUI, and we found longer AGDAC and GH, and shorter AGDAF was predictive factors for SUI (p = 0.001, p = 0.016 and p = 0.014, respectively).

The effect of AGDAF and AGDAC on clinic outcomes controversial issue and still under investigation. Bump et al. claimed subcutaneous structures including muscle and connective tissues, and position of hymen have an effect of AGDAF and AGDAC measurement [12]. In addition, Swan and colleagues stated uncertain anatomical landmarks make measure AGDAF more difficult [13]. Despite the negativities, Hernandez et al. found significantly correlation between longer AGDAC and polycystic ovary syndrome [14]. On the other hand, longer AGDAF was defined as a predictive factor for higher ovarian follicle number, presence of endometriomas, and deep endometriosis [15]. Moreover, longer AGDAC and shorter AGDAF were found in patients with pelvic organ prolapses [7]. In present study, we determined significantly longer AGDAC and shorter AGDAF distance in patients with SUI.

Previous reports investigated effect of GH length on the pelvic floor deficiency. Vakili and colleagues analyze the correlation between GH parameters and strength of
levator ani muscle contractions in patients with failed prolapse surgery, and authors concluded longer GH caused increment in surgical failure in early postoperative period of prolapse surgery [16]. In another study, Lowder et al. tried to identify predictive factors for pelvic organ prolapses, and authors stated that longer GH distance was related with vaginal support loss and higher pelvic organ prolapses rate [17]. To our knowledge, pelvic floor deficiency is common in patients with SUI. In accordance with this hypothesis, we found significantly longer GH distance in patients with SUI, in comparison of patients without SUI.

Although, this paper is the first study focused on relation between AGD parameters and SUI, we are aware of the limitations caused by the retrospective nature of study and limited patients’ number. In addition, we only investigated the AGD—SUI relationship, we did not evaluate the effect of AGD on SUI medical and/or surgical treatments’ outcomes, which may be subject of another study. Body structure including AGD parameters may be affected by genetic differences and races, and our study was performed in one tertiary academic center. We believe further studies in different races contribute to understand the relationship between AGD parameters and SUI.

5 Study limitations
Some limitations of this study were a small sample size as well as the fact that it was a single center study. Larger multicenter studies are required to establish the true effects of AGD on SUI.

6 Conclusion
Present study demonstrated that shorter AGDAF and longer AGDAC distances were resulted with significantly higher incidence of SUI. Additionally, our study found that one mm increment in GH length was associated with 1.038-fold time of SUI development risk. This study is the first to analyze relation between AGD parameters and SUI, our study findings should be supported by prospective studies with more sample size. An important aspect of the results of this study is that, considering the ease and non-invasiveness of anogenital distance measurement, it is a measurement that can be easily performed during routine gynecological and obstetric examinations, and that it is suitable for use in the prediction of women who will develop stress incontinence in the future. Although it is obvious that the introduction of anogenital distance measurement into routine clinical practice will be beneficial, it should be supported by the data of studies with long-term follow-up in order to form a general consensus and to be beneficial for women.
Availability of data and materials
The datasets used and/or analyzed during the current study are available from funding agencies.

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Authors’ contributions
M.E. and F.E. involved in surgical and medical practices; M.E. and F.E. participated in concept; M.E. and F.E. participated in design; M.E. and F.E. involved in surgical and medical practices; M.E. and F.E. participated in analysis or interpretation; M.E. and F.E. involved in literature search; M.E. and F.E. involved in writing. All authors read and approved the final manuscript.

Declarations

Ethics approval and consent to participate
This study was approved by the Haseki Training and Research Hospital (protocol number:17/2020). Informed written consent was obtained from all patients.

Consent for publication
Not Applicable.

Competing interests
The authors declare that they have no known competing interests.

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