Buccal mucosa graft in surgical management of Peyronie’s disease: Ultrasound features and clinical outcomes

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Summary
Introduction: Plaque incision and grafting represent the best surgical approach to the Peyronie’s Disease (PD). The grafting procedures must be restricted to patients with normal preoperative status, excessive curvature and/or deformities. However, the ideal graft has not been identified yet. Buccal mucosa grafts (BMG) provided excellent short-term results, ensuring the fast return of spontaneous erections and preventing shrinkage, which is the main cause of graft failure. Another fearsome surgical complication is de novo erectile dysfunction (ED). We report our results with BMG focusing on the analysis of ultrasonographic and clinical data demonstrating buccal mucosa as determinant factor that allow to avoid complications.

Materials and methods: From 2013 to 2019 we performed at our Urology Unit 27 corporoplasties with BMG to correct complex penile curvature due to PD. Clinical, post-surgical and ultrasound follow up data were evaluated. All patients were no responders to medical treatment or previous surgical procedures. The evaluation period was 72 months. Data regarding pre-operative work-up, including IIEF (IIEF-5) questionnaire administration, detailed clinical history and penile dynamic ultrasound (PGEl-induced erection) were collected. The time of spontaneous erection resumption was recorded for each patient. To improve blood supply to the graft, a low-dose PDES-i was prescribed for all patients for a period of two months, starting immediately after discharge. Check-ups were scheduled every 3 months, starting from 1 month after surgery. In each visit, patients underwent a penile ultrasound evaluation of graft features. After 6 and 12 months, all patients underwent a penile dynamic ultrasound for Erection Hardness Score determination, then standard ultrasound and clinical evaluation yearly. Our analyses were focused on BMG as a major determinant of the surgical success.

Results: Mean age of 27 patients was 57 years (42-71) with a maximum follow up time of 72 months and minimum of 3. Site of penile curvature was dorsal in 18 (67%) patients, ventral in 2 (7%), complex in 7 (26%). The degree of the curvature was < 60° in 11 (41%) patients, > 60° in 16 (59%). Straightening of penis was reached in 100% of cases. Penile shortening resulted in 7.4% (2/27). De novo ED appeared in 2/27 cases with a post-operative rate of PDESi users increasing from 12 to 14 patients (45% vs 52%). Ultrasound aspects of BMG, recorded at every follow up visit, results in a hypoechoic plaque with an iperechoic rim that become isoechoic over the time in all cases. No case of scars or seroma was registered. Small intra-graft cystic lesions were highlighted in 3 cases (11%).

Conclusions: BMG may represent a good choice in grafts procedures for PD surgical management. The functional results obtained by BMG procedures were related to the good anatomical characteristics of the patch and were highlighted in our series by use of penile ultrasound, during the follow up period.

Key words: Peyronie’s disease; Buccal mucosa; Ultrasound; Graft; Erectile dysfunction.

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INTRODUCTION
Peyronie’s disease (PD) is a benign condition causing penile deformities, shortening, loss of penile flexibility and sexual dysfunctions (1). The diagnoses are increasing in number, even in the younger population, because of the increased knowledge about the topic (2). Surgical treatment remains the gold standard and it includes penis plication, grafting, and placement of inflatable prosthesis with the aim to restore coital functions (3). Buccal mucosa grafts (BMG) provided excellent short-term results, suggested by the early recovery of spontaneous erections and the prevention of late shrinkage, which is the main cause of graft failure. It also seems to be safe and reproducible, thus representing a valuable treatment option for PD. Clinical series are limited yet but functional and cosmetics results are very promising (4). We analyzed our surgical results with the BMG technique applied to 27 patients focusing on the ultrasonographic follow-up and clinical data.

METHODS
From our surgical data base, we evaluated 27 consecutive cases treated with a plaque incision and BMG to correct the secondary penile curvature due to PD. Clinical, post-surgical and ultrasound follow up data were evaluated. All patients were no responders to medical treatment or previous surgical procedures. The evaluation period was 72 months. Data regarding pre-operative work-up, including IIEF (IIEF-5) questionnaire administration, detailed clinical history and penile dynamic ultrasound (PGEl-induced erection) features were collected (5, 6). The surgical procedures were done previous penile degloving and using a relaxing incision (double Y or H shaped) at the point of maximum curvature on the penis. The albuginea defects were covered using BMG grafting...
technique (7, 8). The buccal mucosa grafts were harvested as described by Epbley et al. (9). After “defatting”, BMG was apposite to cover the albuginea defect with the submucosa surface in contact with the cavernous tissue in order to obtain a quick blood supply and sutured with a 3/0 absorbable running suture (Maxon™) in each side (Figure 1). Artificial erection was repeated to evaluate the curvature and deformity correction, defining the need of complementary tunica albuginea plications (10). Buck’s fascia was accurately closed with interrupted absorbable suture, especially in correspondence with the patch site to avoid patch enlargement. In cases of ventral curvatures, urethra was fixed to corpora cavernosa with a tension-free stitches. A small drainage was placed between the Buck’s fascia and the dartos. Circumcisional incision was closed and dressed. Patients were discharged at third post-operative day after catheter, drainage and dressing removal. The time of spontaneous erection resumption was recorded for each patient. To improve blood supply to the graft, a low-dose PDE-5 inhibitor (PDE-5i) was prescribed for all patients for a period of two months, starting immediately after discharge. Check-ups were scheduled every 3 months, starting from 1 month after surgery. In each visit, they underwent a penile ultrasonic evaluation of graft features. After 6 and 12 months, all patients underwent a penile dynamic ultrasound for Erection Hardness Score determination, then standard ultrasound and clinical evaluation yearly. All consecutive cases who underwent BMG in a 72-month period of time were analyzed. We studied the ultrasonographic appearance of the BMG at 1, 3, 6 and 12 months after surgery and yearly cosmetic outcomes and functional outcomes.

Table 1. Salient pre-operative clinical data.

| Number of patients | 27 (100%) |
|--------------------|-----------|
| Age (years)        | 57 (42-71) |
| Penile curvature degree pre-op | 61.9° (45-90) |
| Duration of PD (months) | 16.5 (6-48) |
| IIEF score pre-operative | 22.5 (18-24) |
| Curvature side     | Dorsal 18 (67%) |
|                    | Ventral 2 (7%) |
|                    | Complex 7 (26%) |
| PDE5 use pre-op    | 12 (45%) |
| Diabetes mellitus  | 4 (14.8%) |
| Previous PD surgery| 1 (3.8%) |
| Previous radical prostatectomy | 4 (14.8%) |
| Duplexgyn disease associated | 1 (3.7%) |
| Pioniasis           | 1 (3.7%) |
| Prepatial Lichen Sclerosis | 2 (7.4%) |
| Cardiovascular disease | 2 (7.4%) |

Table 2. Results and complications.

**RESULTS**

Mean follow-up time (months) 26.3 (3-72)
Time of spontaneous erection resumption (days) 32 (1-7)
IIEF score post-operative at last follow-up 23.1 (15-24)
PDE5 post-operative use 14 (62%)
Functional penile straightening 27 (100%)
Penile shortening 2 (7.4%)

**COMPlications**

Removal at the donor site 1 (4.7%)
Penile shaft hematoma 1 (4.7%)
Glandular erection pain 1 (3.7%)
Prosthetic edema 4 (14.8%)
Skin necrosis 1 (3.7%)
Cyclical intra-graft lesions 3 (11.5%)
Penile abscesses 1 (3.7%)
Resnaturum raphae 2 (7.4%)
De novo erectile dysfunction 2 (7.4%)

Figure 1. Surgical steps for BMG procedures in a case of ventral curvature.
A. Buccal mucosa graft harvesting; B. Intracavernosal injection; C. Urethral Isolation;
D. Marked transversal incision (double Y) on the plaque; E. Buccal mucosa grafting; F. Final result.

**Results**

We performed 27 procedures in men with mean penile curvature of degree 61.9° (range 45°-90°), mean age 57 years (42-71) and mean follow-up of 28.3 months (min 3 max 72). The degree of the curvature was <60° in 11 (41%) patients, >60° in 16 (59%). Site of penile curvature (degree 61.9° in mean; range 45°-90°), was dorsal in 18 (67%) patients, ventral in 2 (7%), complex in 7 (26%). Pre-operative data and results are summarized in Tables 1, 2. The patient medical history was characterized from different comorbidities recognized as erectile dysfunction (ED) risk factors. Diabetes mellitus was reported in 4 (14.8%) patients; cardiovascular disease in 2 (7.4%), previous radical prostatectomy in 4 (14.8%), preputial lichen sclerosis in 2 (7.4%) and psoriasis in 1. One patient had a failed plication procedure. Pre-operative use of PDE-5i was referred by 12 patients (45%). Pre-operative IIEF score was 22.5 in mean (11-24). After a follow up period of 28.3 months (3-72), we reported a functional straightening of penis reached in 100% of cases. Penile shortening resulted in 7.4% (2/27). All patients had complete spontaneous erections in a period of 1 to 7 days after surgery, however paraphimosis occurred in three cases (14.2%), requiring surgical revision, and one patient (4.7%) develop skin preputial necrosis managed by topical therapy. Post-operative mean IIEF score, at the last follow up was 23.1 (15-24). De novo ED appeared in 2/27 cases. Post-operative amount of PDE5i users increasing from 12 to 14 patients (45% vs 52%). Ultrasound was applied at each follow-up visit. Ultrasound features during follow up are showed in
Figure 2.  
A. Ultrasound patch (white arrow) appearance 3 months after surgery.  
B. Ultrasound patch (green arrow) appearance after surgery (6 months).

Figure 2a, b and Figure 3. In all cases, post-operative images consisted in a hypoechoic plaque with hyperechoic rim, observed in the graft area during the first month after surgery. Progressively, we described the ultrasound disappearance of graft, which has become isoechoic respect the tunica albuginea and not distinguishable from albuginea rim. No case of scars or seroma was registered. Small intra-graft cystic lesions highlighted in 3 cases (11%).

DISCUSSION

Severe PD (> 60°) is still treated by surgical approach that leads an efficient solution in a relatively short time (11). In spite of this, results are not devoid of complications, the most serious is the development of de-novo ED, which seems to be caused by complete surgical excision of the plaque (12). Nowadays, plaque incision surgery with grafting is the preferred therapeutic approach (3, 8). In 2005, buccal mucosa was introduced as free autograft in the surgical treatment of PD (13) and then, during the years, evaluated in several others series (14-19). Promising results were reported by Shioshivi et al. (13) who observed the complete straightening of penis in 92.3% of 26 patients treated with a residual curvature (< 10°) in 7.7%. The shortening of penis was observed in four patients (15.4%) and in two patients (7.7%) a partial reduction of erectile function. They concluded that buccal mucosa showed high properties of adaptation and revascularization, good anatomical and functional clinical results, demonstrating a stable elasticity without late shrinkage. Similarly, Liu et al. (14) showed that buccal mucosa remained stable with good elasticity over time. They performed replacing the plaque by free autograft of buccal mucosa on 24 patients and obtained satisfactory results: complete penile straightening was achieved in 21/24, slight residual curvature (< 15 degrees) noted in 3 (12.5%), a little shortening of the penis (< 1 cm) in 2 (8.3%). As advancement in term of prevention of de novo ED, Cormio and co-workers (15) introduced important technical modifications consisting in the plaque incision instead of excision. BMG was performed similar than others series. In 15 consecutive surgical procedures they reported a 100% of penile straightening, no curvature recurrence and any de novo ED. These results were confirmed by Zucchi et al. (16) in 32 patients treated with plaque incision and BMG between 2006 and 2013: in 28 cases, they reported one curvature relapse after 1 year (3.5%) and an immediate significant improvement of IIEF scores in half of cases, although after 2 years the trend was in reduction. They concluded that corporoplasty with buccal mucosa represent a good treatment choice for most forms of PD. The complete penis straightening was achieved in 73% of 33 patients by Gvasalia et al. (17), however a residual curvature (< 20°) was observed in 27% and de novo ED in 18%, commonly in elderly patients, but with a good response to PDE5i. Molina-Escudero et al. (18) reported the short-term results (penile length, angle of curvature and erectile function) of 10 patients treated through corporoplasty with oral mucosa graft. The straightening of penis was reached in 100% with a shortening rate of 80% and 10 % of de novo ED. In our previous experience with 17 cases (19), we reported favorable results. In this series, that take account of 10 more patients passing from 17 to 27 cases, we have similar results with a longer follow-up. All the procedures were performed as the main indication the failure or inability to sexual intercourse and psychological impairment due to the penile curvature. Patients are variably aging, with a medical general history characterized from different comorbidities as, in example, diabetes mellitus, cardiovascular disease, previous pelvic surgical procedure for prostate cancer (Table 1). One patient underwent a previous failed plication procedure. Pre-operative use of PDE5 inhibitors was referred by 12 patients (45%). Recognized that several patients had more risk factors of post-operative ED (baseline sexual function, age, graft size or position) (12), the probability of de novo ED should be higher, than that observed. We obtained a very good functional results with 7.4% of de novo ED and a post-operative increase of PDE5i users to 52% from 45%. In literature the rate of de novo ED ranges between 0% to 18%. These results may be explained by the nature of buccal mucosa that is a living

Figure 3. Complete patch disappearance in ultrasound evaluation (star; 6 years after surgery).

Figure 4. Small intra-patch cystic lesion.
tissue with an elevated binding capacity and revascularization which is immediately supplied with blood from the cavernous tissue. In this way, buccal mucosa tends to heal rapidly, immediately integrating with the surrounding albuginea tissue. This translates into a more rapid resumption of spontaneous erections (mean of 3.26 days in our experience) and sexual activity and into a reduced risk of curvature relapse (20). Penile ultrasound validated these results showing the process of mucosal patch integration with tunica albuginea. Ultrasound has been showed to be a method of choice because his cost effectiveness, repeatability and low invasiveness, if compared to others methods (ie. Magnetic Resonance) (21). US is mainly used in diagnosis and staging of PD. There are only few reports regarding the ultrasound usefulness in the follow up after treatment. In relapses or in dilations patch evaluation, ultrasound is still decisive (22). In our series, US showed that BMG appears initially as hypoechoic plaques with hyper-echoic rim (Figure 2a, b) and then evolved in isoechoic over the time. After six years from surgical procedure, we reported a complete disappearance of the patch aspect, confounded with the albuginea (Figure 3). No cases of scars, and consequently, of curvature relapse, or seroma were registered. Small intra-graft cystic lesions highlighted in 3 cases (11%) (Figure 4). Any patch bulging has been reported yet. The perfect seal of BMG and the accurate closure of Buck’s fascia are important key-factors to obtain a successful result. The biological characteristics of buccal mucosa reduced the risk of curvature relapse as demonstrated by the high rate of penile straightening reported in 100% in our series while is 87.5-100% in literature. Patch ultrasound evaluation confirm these aspects and it is needed in the surgical management of PD.

CONCLUSIONS

The functional results obtained by BMG procedures in PD surgical treatment were related to the great anatomical characteristics of the patch and highlighted in our series by use of penile ultrasound during the follow up period. Ultrasound features confirmed the characteristics of buccal mucosa as perfect seal in the tunica albuginea defect, created after fibrotic plaque incision, allowing to reduce the risk of de novo ED.

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