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Abstract- Priapism is defined as a penile erection that persists more 4 hours or longer and is unrelated to sexual activity. The three main subtypes are low flow (ischemic), high flow (non-ischemic) and stuttering (intermittent or recurrent) priapism. Alpha-1 antagonistic activities of some antipsychotics, especially chlorpromazine, have been reported to be responsible for development of low-flow priapism. Chlorpromazine has the greatest alpha-adrenergic affinity among the conventional antipsychotic agents and the most frequently reported to be associated with priapism. We report a case of priapism after a first single dose of chlorpromazine (Neurazine ® 100 mg tablet, Misr Co. For Phramind-Egypt). The case was presented first by neglected low-flow priapism that had been converted to high-flow priapism due to injury of a branch of the cavernosal artery, during aspiration and irrigation treatment procedure, proved by blood gas testing of the aspirated penile blood, penile color duplex Doppler ultrasonography and selective left internal pudendal arteriogram. This conversion is an uncommon complication of such procedure.

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High-Flow Priapism following Chlorpromazine Induced Low-Flow Priapism

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Abstract- Priapism is defined as a penile erection that persists more 4 hours or longer and is unrelated to sexual activity. The three main subtypes are low flow (ischemic), high flow (nonischemic) and stuttering (intermittent or recurrent) priapism. Alpha-1 antagonistic activities of some antipsychotics, especially chlorpromazine, have been reported to be development of responsible for low-flow priapism. Chlorpromazine has the greatest alpha-adrenergic affinity among the conventional antipsychotic agents and the most frequently reported to be associated with priapism. We report a case of priapism after a first single dose of chlorpromazine (Neurazine ® 100 mg tablet, Misr Co. For Phramind-Egypt). The case was presented first by neglected low-flow priapism that had been converted to high-flow priapism due to injury of a branch of the cavernosal artery, during aspiration and irrigation treatment procedure, proved by blood gas testing of the aspirated penile blood, penile color duplex Doppler ultrasonography and selective left internal pudendal arteriogram. This conversion is an uncommon complication of such procedure.

I. CASE REPORT

A 24-year-old male patient presented with priapism to our Andrology Department, Kobry El-Koba Military Armed Forces Medical Complex, Cairo-Egypt. He was fully awake, sitting calm and alert. On, Inspection, the penis was fully erected and has a dorsoventral oscillatory movement (atrial pulse). Although the shaft was very rigid on palpation, the glans penis was semi rigid with no tenderness. Patient is not married, not known to have hypertension, sickle cell trait, history of malignancy. He denied any previous episodes of priapism, recent sexual activity or arousal, use of phosphodiesterase inhibitors or perineal trauma.

Three weeks before the patient came to us, he experienced agitation and insomnia for which he took a first single dose of chlorpromazine (Neurazine ® 100 mg tablet, Misr Co. For Phramind.-Egypt). He had observed a prolonged penile erection 5 hours after taking chlorpromazine. The patient did not give a care for his priapism for the first 18 hours. However, as his penile erection became painful and did not resolve spontaneously and after repeated ejaculation trials, he went to ER of Embaba General Hospital in next 24

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hours. He was in obvious physical pain and reported that the erection had remained rigid throughout the entire 36hrs since becoming tumescent. Urologists had performed two sessions of intra-corporeal injection of 1 ma ephedrine, drainage, and irrigation with sterile saline that were not sufficient to obtain de-tumescence and they transferred the patient to Andrology Specialty Center and Department, Kasr El-Ainy Faculty of Medicine, Cairo University in the next day. Andrologists admitted the patient and diagnosed him to have a lowflow priapism by penile duplex Doppler ultrasonography with multiple intra-cavernosal fibrosis and echogenic blood clots. Corporal aspiration revealed dark, red blood with a pH of 7.24, pO2 of 20, and pCO2 of 65, consistent with low-flow, ischemic priapism. Since then, intra-corporeal injection of 1 mg ephedrine, drainage, and irrigation with sterile saline were handled at a rate of 3 cycles per session for 2 sessions over the first 24hrs and de-tumescence was achieved with some residual corporal edema. Unfortunately, the corporal bodies were again fully erect, but surprisingly not painful. Many sessions of intra-corporeal injection of alpha-adrenergic receptor agonists with aspiration and ice-pack compression in between were done for the next consecutive 6 dayswithout improvement. They discharged the patient thereafter as the patient refused surgical intervention.

As long as erection was inevitable yet; patient came to us on day 23rd since becoming tumescent and he was admitted in our andrology in-patient unit, where basic laboratory investigations including: complete blood counts, coagulation profile, electrolytes, hepatic and renal function and sickle cell tests were done. All laboratory investigations were normal except for mild leukocytosis (TLC=14.2). Penile arterial blood gas (ABG) from the lateral base of the corporal body was obtained. The latter was consistent with high-flow priapism (bright red blood with a pH of 7.45, pO2 of 90, and pCO2 35). Penile duplex of Doppler ultrasonography revealed bilateral echogenic, distorted cavernosa reflecting significant bilateral corporal fibrosis and the sinusoids are typically not compressible with probe pressure and engorged with sinusoidal thrombosis. Doppler assessment of the both left and right cavernosal artery, revealed significantly reduced arterial flow (<10 cm/s) giving picture of low-flow priapism except for a turbulent arteriocavernosal fistula which was determined with high-velocity and low-

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resistance flow (>90 cm/s) within the left proximal corpus cavernosus (peri-lacunar) (fig.1) giving a picture of high-flow priapism. Selective left internal pudendal arteriogram confirmed a large fistula arising from the peri-lacunar area of left corpus cavernosus (fig. 2-10).Medical interference was postponed (no embolization) according to the patient's decision.Treatment non-surgical strategy for neglected priapism was recommended in order to reduce the ongoing fibrosis process and risk of infections in the form of the following:

- Acetylsalicylic Acid (Jusprin 81 mg tab / 24 hr, Julphar, Gulf Pharmaceutical Industries, Ras Al Khaimah, U.A.E.).
- 2. Pentoxifylline (Trental SR 400 mg tab / 8 hr, sanofi-aventis, Egypt).

- 3. Vitamin E (Vitamin E 1000 mg cap / 12 hr, Pharco Pharmaceuticals Alexandria, Egypt).
- Cefotaxime (Cefotax 1 gm vial I.V / 12 hr, Egyptian Int. Pharmaceutical Industries Co. (E.I.P.I.CO.), Egypt).

This treatment strategy is continued, except for the antibiotic (only 7 days), and priapism is not resolved yet, the rigidity is decreased by 20%, which is retained whenever sexual arousal is present and decreased back by 20% after masturbation. Follow up was handled monthly for 18 months without any other complications except for the persistent penile erection, which is still surprisingly tumescent. The case will be presented later on to report the efficacy of the new non-surgical treatment strategy we followed.



Figure 1



Figure 2







Figure 4







Figure 6





Figure 8





Figure 10

II. Discussion

Priapism is defined as a penile erection that persists for 4 hours or longer and is unrelated to sexual activity that was first described in medical literature in 1845¹. The corpus cavernosa consist of vascular spaces called sinusoids, which include smooth muscles on their walls. Arterioles in the penis that supply blood to the corpus cavernosa are in a tonic state during flaccid periods². The three main subtypes are low flow (ischemic), high flow (non-ischemic) and stuttering (intermittent or recurrent) priapism³. The identification of priapism subtype is important as delayed emergent medical interference (particularly in low flow ischemic subgroup) can result in persisting erectile dysfunction^{1,3,4}.

Patients with low-flow priapism will often present with rigid corpus cavernosa and pain, whereas those with high-flow priapism will often present with tumescent but not rigid corpus cavernosa and seldom report pain. The most considerable modalities to distinguish lowand high-flow priapism are blood gas testing and color Doppler ultrasonography⁵. Low flow priapism is characterized by a reduced venous outflow, hypoxia, rising PaCO2 and acidosis that can cause progressive ischemia within the cavernosal tissue with timedependent changes in the corporal metabolic environment. This lead to smooth muscle irreversible injury and necrosis to the erectile tissue with consequent fibrosis of the corporas^{3,6}.

Alpha-1 antagonistic activities of some antipsychotics, especially chlorpromazine, have been thought to be responsible for development of priapism. Based on their ability to block the α -adrenergic receptors of sinusoidal smooth muscles, they lead to a reduction in resistance and uncontrolled blood flow into the corpora cavernosa^{2,7,8}. Chlorpromazine has the alpha-adrenergic affinity among greatest the conventional antipsychotic agents and the most frequently reported to be associated with priapism^{7,8}. It had been reported that, priapism occurred in a patient after he had taken one 25-mg tablet of chlorpromazine (the lowest dose to produce this complication). On contrary, another patient developed priapism after he had been using chlorpromazine for 3 years. In our case report, the patient developed low-flow priapism after first single 100-mg takina his tablet of chlorpromazine. This shows that priapism can occur after short-term or long-term and even after high or low dose therapy with chlorpromazine⁹. So, it is believed that low-flow priapism is not a dose-or duration-specific complication to chlorpromazine as has been reported in different studies^{2,7,8,10,11}.

High-flow priapism is a persistent erection caused by unregulated cavernous arterial inflow⁴. The usual cause of high-flow priapism is blunt perineal trauma. However, it may be associated with metastatic

malignancy to the penis, with acute spinal cord injury and occasionally may complicate low-flow priapism after shunt procedures^{4,12}. Moreover, high-flow priapism caused by iatrogenic needle trauma during pharmacological agent injections in the cavernosal body, as a treatment of early cases of low-flow priapism, is considered as an uncommon complication of such procedure. This may occur due to a lacerated cavernous artery or one of its branches leading to a high-flow fistula between the artery and the lacunar spaces of the sinusoidal tissue^{13,14}. During priapism, the edematous corporal state, septal and/or intra-corporal fibrosis may anchor the cavernous arteries to a more eccentric place immediately adjacent to the intercorporal septum. These arteries are therefore become relatively fixed and are possibly more susceptible to needle injury during corporal aspiration as happened in our case report¹³.

III. Conclusion

It seems hard to predict the dose and/or duration that may cause priapism with antipsychotic medications, especially chlorpromazine, as discussed before. Among patients treated for low-flow priapism, who retain penile tumescent, a penile ABG determination and penile ultrasound are essential for early detection of converted priapism from low to highflow priapism. Selective internal pudendal arteriogram is the golden standard technique to diagnose and treat (arterial embolization) high-flow priapism due to arterial fistulae. However, in our case, non-surgical treatment strategy is continued for about 2 years. This acquires a promising strategy as it has reduced the risk of corporal fibrosis and its complications such as irresponsive penile erection on sexual excitation.

IV. Recommendations

Clinicians should be familiar with infrequent serious adverse events of antipsychotic medications especially chlorpromazine. Performing the corporal aspiration, irrigation and injections of alpha-adrenergic receptor agonists in low-flow priapism, under the guidance of penile duplex Doppler ultrasonography to prevent injury of corporal vessels during procedure and thus preventing conversion of low-flow priapism into high-flow one.Further prospective studies on larger number of cases are recommended in order to evaluate the effectiveness of non-surgical systemic oral medications versus the invasive surgical procedures especially in neglected cases of priapism.

References Références Referencias

 Halls J. E., Patel D. V., Walkden M., Patel U. Priapism: Pathophysiology and the role of the radiologist. Br J Radiol. 2012: 85 (Spec. Issue 1): 79-85. doi:10.1259/bjr/62360925.

- 2. Baytunca M. B., Kose S., Ozbaran B., Erermis S. Risperidone, quetiapine and chlorpromazine may have induced priapism in an adolescent. Pediatr Int. 2016: 58 (1): 61-63. doi:10.1111/ped.12741.
- Muneer A., Ralph D. Guideline of guidelines: priapism. BJU Int. 2017: 119 (2): 204-208. doi:10. 1111/bju.13717.
- 4. Salonia A., Eardley I., Giuliano F., et al. European association of urology guidelines on priapism. Eur Urol. 2014: 65 (2): 480-489. doi:10.1016/j.eururo. 2013.11.008.
- Wallis C. J. D., Hoag N., Pommerville P. J., Huk M. E. Recurrent idiopathic high-flow priapism treated with selective arterial embolization after repeated initial treatments for low-flow priapism. J Can Urol Assoc. 2009: 3 (1): 60-63.
- Habous M., Elkhouly M., Abdelwahab O., et al. Noninvasive treatments for iatrogenic priapism: Do they really work? A prospective multicenter study. Urol Ann. 2016: 8 (2): 193-196. doi:10.4103/0974-7796.176874.
- Shahrbabaki M. E., Sabzevari L. Early onset priapism under chlorpromazine and risperidone therapy. Iran J Psychiatry Behav Sci. 2011: 5 (2): 139-142.
- Turkan S., Kalkan M., Şahin C. Two Priapism Cases Following Short-Term Use of Chlorpromazine Kısa Dönem Klorpromazin Kullanımına Bağlı İki Priapizm Olgusu. J Clin Anal Med. 2014: 4 (Suppl 1): 47-48. doi:10.4328/JCAM.2710.
- Kilciler M., Bedir S., Sümer F., Dayanç M., Peker A.
 F. Priapism in a Patient Receiving Long-Term Chlorpromazine Therapy. Urol Int. 2003: 71 (1): 127-128. doi:10.1159/000071112.
- Andersohn F., Schmedt N., Weinmann S., Willich S. N., Garbe E. Priapism Associated With Antipsychotics. J Clin Psychopharmacol. 2010: 30 (1): 68-71. doi:10.1097/JCP.0b013e3181c8273d.
- Marrag I., Soussia R. Ben, Hajji K., Zarrouk L., Binous Y., Nasr M. Priapisme induit par la chlorpromazine: A propos de deux cas. African J Urol. 2016: 22 (2): 131-135. doi:10.1016/j.afju. 2015.05.006.
- 12. Lutz A., Lacour S., Hellstrom W. Conversion of Low-Flow to High-Flow Priapism: A Case Report and Review (CME). J Sex Med. 2012: 9 (4): 951-954. doi:10.1111/j.1743-6109.2012.02692.x.
- McMahon C. G. High flow priapism due to an arterial-lacunar fistula complicating initial venoocclusive priapism. Int J Impot Res. 2002: 14 (3): 195-196. doi:10.1038/sj.ijir.3900828.
- Park J. K., Jeong Y. B., Han Y. M., Kim H. J. Highflow priapism caused by injury of the bilateral cavernosal artery after needle trauma in a patient with low-flow priapism. BJU Int. 2003: 92 (Suppl. 3): 1-2. doi:10.1111/j.1464-410X.2003.02945.x.