Indirect inguinal hernia with uterine tissue in a male: A case of persistent Mullerian duct syndrome and literature review

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Abstract

A 32-year-old male presenting with a direct inguinal hernia was found to have uterine tissue extending through the inguinal canal, warranting a diagnosis of persistent Mullerian duct syndrome (PMDS). PMDS is an extremely rare form of internal male pseudo-hermaphroditism in which female internal sex organs, including the uterus, cervix and proximal vagina, persist in a 46XY male with normal external genitalia. The condition results from a congenital insensitivity to anti-Mullerian hormone, or lack of anti-Mullerian hormone, leading to persistence of the female internal sex organs in a male. Clinically, this condition is associated with cryptorchidism. Controversy persists regarding the appropriate treatment of PMDS, since resection of the remnant structures is associated with potential morbidity, but retention risks development of occasional malignancies. We review the literature and discuss various aspects of pathophysiology, diagnosis, and management of PMDS.

Case presentation

A 32-year-old gentleman was referred to the general surgery clinic for a painful right groin mass. The patient described the pain as dull and persistent with no clear exacerbating factors. As an infant, he had a history of a right inguinal hernia and a left undescended testicle. At 1 year of age, he underwent right inguinal hernia repair and left orchiectomy. There was no additional surgical or medical history. Physical examination demonstrated a well-appearing male with a right inguinal hernia and a left undescended testicle. At 1 year of age, he underwent right inguinal hernia repair and left orchiectomy. There was no additional surgical or medical history. Physical examination demonstrated a well-appearing male with a well-healed right groin scar overlying a reducible groin mass with palpable cough impulse. The right testis was of normal size, with no palpable masses, and normally located within the right hemiscrotum. The left testicle was absent and with no evidence of a detectable hernia. The abdominal examination was entirely unremarkable. The patient was diagnosed with a recurrent right-sided inguinal hernia, and open mesh repair of the hernia was planned.

Following informed consent, the patient had surgery. A 4-cm surgical incision was made at the right groin crease. Contents of the inguinal canal, in particular, the right spermatic cord, were identified as emerging from the right internal inguinal ring. Adjacent to the spermatic cord, however, an unidentified tubulo-muscular structure passing from the internal inguinal ring could be seen (Fig. 1). A small incision was made within the structure and flexible endoscopy demonstrated a hollow tubular structure that eventually led to a blind end (Fig. 2). Contrast was passed into the tubular structure and intra-operative radiographs revealed that the structure continued into a blind end within the pelvis. Communication within the prostatic urethra, but not the bladder, was evident. At this point, a decision was made to excise the structure at its point of exit from the inguinal canal. It measured about 10 cm in length and 2 cm × 2 cm × 3 cm at its proximal end (Fig. 3). It was seen to contain mucus only. The indirect hernia was subsequently repaired using a standard Lichtenstein repair with a plug and patch mesh. There were no perioperative complications.

Subsequent pathological evaluation of the excised structure identified poorly developed endometrium and myometrium, consistent with uterine tissue (Fig. 4, Fig. 5). The patient underwent genotypic analysis, which revealed a normal 46XY karyotype, as well as computed-tomography (CT) scanning of the abdomen and pelvis, which was unremarkable. He had an uneventful follow-up visit in the general surgery clinic 1 month later, and was discharged from surgical care without further routine follow-up. The diagnosis was persistent Mullerian duct syndrome (PMDS).
PMDS is an extremely rare form of internal male pseudo-hermaphroditism in which female internal sex organs, including the uterus, cervix and proximal vagina (terminating in the prostatic utricle), persist in a 46XY male with normal external genitalia. The persistent Mullerian structures often tether the testes and impede their descent, causing unilateral or bilateral undescended testes. The condition was first described in 1895, with about 200 cases described in the literature since.1

In normal fetal sex organ development, the Wolffian and Mullerian ducts give rise to male and female internal sex organs, respectively. During the eighth and ninth weeks of male fetal development, Sertoli cells produce an anti-Mullerian hormone (AMH), also known as Mullerian inhibiting factor or substance (MIF/MIS). This hormone induces regression of the Mullerian ducts, thereby inhibiting the development of the uterus, cervix, proximal vagina and fallopian tubes. Testosterone produced by Leydig cells subsequently induces virilization of the Wolffian ducts into epididymides, vasa deferentia and seminal vesicles. External male genitalia development is largely under the control of dihydrotestosterone (DHT).2 In males, failure to produce functional AMH or a defective type II AMH receptor (AMHR-II) leads to failure of Mullerian duct regression.3 The Mullerian structures thus develop into uterus, upper vagina, and fallopian tubes in a 46XY male with normal male external genitalia, leading to PMDS.4 In 15% of patients, the cause of PMDS is unknown. However, in the remaining 85% of cases, autosomal recessive inheritance of defects in AMH (chromosome 19) and its receptor AMHR-II (chromosome 12) account for development of PMDS.5 In males, virilization generally remains unaffected, but infertility is common secondary to azoospermia.6 Homozygous mutations in AMH or AMHR-II have been linked with premature loss of germ cells in female mouse models. However, homozygous human females are normal and fertile at least in early reproductive life.2

Clinically, PMDS is most commonly diagnosed as an incidental finding during an operation, but the presence of unilateral or bilateral undescended testes may increase clinical suspicion.7-11 Imaging modalities for the diagnosis of PMDS have thus far proven unreliable.12 Most (60%-70%)
of patients with PMDS have bilateral undescended testes located in an area analogous to the ovaries in women. This specific form of PMDS is occasionally referred to as class A, or the “female” form, of PMDS. A second 20% to 30% of patients present with a unilateral testis contained within a hernia sac or hemiscrotum, with a contralateral undescended testis and inguinal hernia, this being known as class B PMDS (Fig. 1). Many class B PMDS patients, such as the patient reported in this case, have a fallopian tube and uterus herniating through the inguinal canal, a condition classically described as “hernia uteri inguinale.” In class C PMDS, seen in the remaining 10% of patients, both testes (along with the fallopian tubes and uterus) herniate into the same hemiscrotum, a condition known as “transverse testicular ectopia.” About 30% to 50% of all cases of transverse testicular ectopia are thought to be secondary to PMDS. Rarely, the second testis will become entrapped in the inguinal canal ipsilateral to the normally descended testis.

Management of PMDS is controversial. An element of deciding on intervention is largely based on reducing the risk of malignancy, while maintaining maximal reproductive function. In PMDS patients with undescended testes, the rate of testicular cancer is about 12%, which is comparable to the rate seen in undescended testes not associated with PMDS. To detect testicular disease, as with any case of undescended testis, orchiopexy is indicated after 12 months of age. In addition, orchiopexy may be attempted for cases of bilateral cryptorchidism, with hopes of preserving fertility and testosterone production. Unilateral or bilateral cryptorchidism in adults should be treated with orchiectomy, as orchiopexy of the testes is unlikely to preserve fertility or lead to sufficient testosterone production to forego hormonal replacement.

Beyond orchiopexy, the decision to resect the Mullerian structures depends largely on maximizing testicular function.

**Table 1. Key points**

- Persistent Mullerian duct syndrome (PMDS) is typically a rare autosomal recessive disorder caused by non-functional mutations in anti-Mullerian hormone or its type II receptor.
- Clinically, PMDS is defined as the presence of Mullerian-derived structures (uterus, fallopian tubes, and proximal vaginal) inserting at the prostatic utricle in a normal 46XY male with normal appearing external genitalia. This condition is often associated with unilateral or bilateral undescended testicles.
- Undescended testes in PMDS patients have a similar risk of developing testicular cancers as patients with non-PMDS intra-abdominal testes, quoted at about 12%.
- Surgical management of PMDS is controversial, and focuses on management of undescended testes with possible removal of Mullerian structures, but only if resection of the structures does not impair testicular function, or damage the vas deferens, through damage to the blood supply during dissection.
and minimizing morbidity. Several authors have asserted that the Mullerian structures should not be removed, as there is minimal risk associated with their retention, and excision of the Mullerian structures risks damaging primary blood supply to the PMDS testis via the internal spermatic and the deferential arteries. However, more recent reports have demonstrated rare malignancies, usually adenocarcinomas of the Mullerian duct, associated with retained Mullerian structures. If resection is attempted, great care must be taken to avoid damage to the vascular supply to the testis, or vas deferens, which run in close proximity to the remnant uterus and proximal vagina. Interestingly, laparoscopic management of PMDS has now also been successfully reported. See Table 1 for our key points.

Competing interests: None declared.

This paper has been peer-reviewed.

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