Age at Surgery and Outcomes of an Undescended Testis

Francisco Javier Schneuer, PhD, Andrew J.A. Holland, MB BS, PhD, Gavin Pereira, PhD, Sarra Jamieson, PhD, Carol Bower, MB BS, PhD, Natasha Nassar, PhD

BACKGROUND: Undescended testis (UDT) is the most common genital anomaly in boys. Current guidelines recommend surgery before 12 months of age to maximize fertility and potentially reduce the risk of future malignancy. We investigated the prevalence of UDT and examined rates of surgery and age at surgery in an Australian population.

METHODS: UDT was identified from all live-born infants in New South Wales, Australia, from 2001 to 2011 using routinely collected record-linked birth and hospital data. The prevalence of UDT, surgery rates, age at surgery, postsurgical outcomes, and risk factors for surgery performed later than the recommended age were evaluated.

RESULTS: There were 10,875 (2.1%) boys with a recorded diagnosis of UDT. Corrective surgery was performed in 4,980 (45.8%), representing a cumulative prevalence of 9.6 per 1,000 male births. Five percent of surgeries were orchidectomies, and 9% of boys had revision surgery. Median age at surgery was 16.6 months (interquartile range 11.8 to 31.0 months), decreasing from 21 months for boys born in 2001 to 13 months for boys born in 2010. Among those boys having surgery before 36 months (n = 3,897), 67% had corrective surgery after the recommended 12 months of age; socioeconomic disadvantage, regional/remote area of residence, and lack of private health insurance were risk factors for having corrective surgery after 12 months.

CONCLUSIONS: One in 50 boys born are diagnosed with UDT; two-thirds had no report of corrective surgery. The age at surgery is decreasing; however, two-thirds of surgeries are performed after 12 months of age.

WHAT’S KNOWN ON THIS SUBJECT: Recent reports have revealed that a majority of undescended testis are surgically treated after the recommended age of 12 months, but most include acquired cases that have later presentation. There is a lack of population-based information about outcomes after surgery.

WHAT THIS STUDY ADDS: Two-thirds of undescended testis cases are treated later than the recommended age, with the age at surgery decreasing in the past decade; however, diagnosis at birth remains inadequate. One in 10 boys requires revision surgery.

Drs Schneuer and Nassar conceived and designed the study, conducted the statistical analysis, and drafted the initial manuscript; Dr Holland provided clinical advice; Drs Holland, Pereira, Jamieson, and Bower critically reviewed and revised the manuscript; Drs Pereira, Jamieson, and Bower provided epidemiological advice; and all authors approved the final manuscript as submitted and agree to be accountable for all aspects of the work.

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Undescended testis (UDT), or cryptorchidism, is the failure of 1 or both testes to descend to a normal scrotal position and is the most common genital anomaly in boys. The prevalence at birth, otherwise known as congenital UDT, ranges from 2% to 9%.1 Although a large proportion of cases descend spontaneously to a normal position by 3 months of age, nearly 1% remain undescended and require surgery.2 UDT can also develop in a previously descended testis, considered an acquired UDT, although these tend to present later in childhood and have increased prevalence in prepubertal boys.3

Approximately one-third of UDTs are persistent and need to be treated with orchidopexy, surgical repositioning of the testis within the scrotum.4 Although most surgical procedures are successful, ≤33% have been reported to experience failures, depending on the original location of the testis, including severe complications such as testicular atrophy. These complications may ultimately require removal of the testis (orchiectomy) or revision orchidopexy. These findings may be overestimated, as many previous studies have tended to be from single pediatric referral centers, which include a disproportionate number of complex cases more likely to be associated with surgical failure.5

To date, there is a lack of data on outcomes of UDT surgery from large multicenter studies. The importance of surgery for UDT is underpinned by the fact that boys with UDT have an increased risk of testicular cancer later in life.6 In addition, it has been shown that UDT inhibits the differentiation of primitive germ cells, starting at 4 to 12 months, which is crucial for the production of germ cells that subsequently enable spermatogenesis.7

Delayed repositioning of an undescended testis may result in a reduction in germ cell development and low testicular volume, potentially diminishing subsequent fertility.8 Given these findings, there has been a general consensus that early orchidopexy improves results associated with markers of fertility and testicular malignancy.8 Recent international guidelines have recommended orchidopexy to be performed before 12 months of age to preserve fertility potential and avoid risk of malignancy.9,10 Despite guideline recommendations, a recent systematic review reported that >75% of orchidopexies were performed later than the recommended age,11 with most studies combining congenital and acquired UDT cases. Although they may have similar etiology, combining the 2 types of UDT may be misleading, as they have different ages at presentation and outcomes, and results may be skewed. Persistent congenital cases are of particular interest, as research suggests that longer testicular exposure to higher temperatures leads to increased risks of subfertility and cancer.12 Hence, population-based monitoring of cases from birth to distinguish congenital from acquired cases, current practices, and identification of potential factors contributing to delays in orchidopexy are important to ensure future reproductive health of affected boys.

We hypothesize that a majority of boys with congenital UDT in Australia undergo corrective surgery later than the recommended age. The aims of this population-based study were to investigate the prevalence of UDT and examine the rates of corrective surgery, success or failure of surgery, current trends in age at surgery, and factors influencing early or late surgery in an Australian population.

METHODS

Study Population and Data Sources

The study population included all live-born males diagnosed with UDT in New South Wales (NSW), Australia, from January 1, 2001, to December 31, 2011. NSW is the most populated Australian state, with one-third of all births in the nation. Health care is based on a combination of a national publicly funded (60%) and private (40%) health care system.13

Two main data sources were used: the NSW Perinatal Data Collection, which is a statutory surveillance system covering all live births and stillbirths in NSW, and the NSW Admitted Patient Data Collection, which is a census of all inpatient hospital admissions that collects demographic and clinical information. All diagnoses and procedures for each admission are coded according to the 10th revision of the International Classification of Diseases, 10th Revision, Australian Modification (ICD10-AM) and the Australian Classification of Health Interventions (ACHI), respectively. Longitudinal record linkage of individual birth and hospital data were conducted to provide information on each boy’s birth and any subsequent hospital admissions until March 2014. Record linkage was conducted by the NSW Centre for Health Record Linkage independent of the research. Ethics approval for access and linkage of data were obtained from the NSW Population and Health Services Research Ethics Committee.

Boys with UDT were identified from the NSW Admitted Patient Data Collection if they had a relevant ICD-10-AM code (Q53) (Table 1) recorded in any birth or hospital admission. These were classified as unilateral, bilateral, unspecified, and ectopic. Isolated UDT was defined in cases for which no other congenital anomaly diagnosis was recorded. For those diagnosed with other anomalies, other testicular anomalies, hypospadias, and other genitourinary anomalies were each differentiated from all other types. All
TABLE 1 Definitions and Diagnostic and Procedures Codes Related to UDT

<table>
<thead>
<tr>
<th>Definition</th>
<th>Code</th>
</tr>
</thead>
<tbody>
<tr>
<td>Undescended testicle, unilateral</td>
<td>Q53.1, Q53.10, Q53.11, Q53.12, Q53.13, and Q53.19</td>
</tr>
<tr>
<td>Undescended testicle, bilateral</td>
<td>Q53.2, Q53.20, Q53.21, Q53.22, Q53.23, and Q53.28</td>
</tr>
<tr>
<td>Undescended testicle, unspecified</td>
<td>Q53.9, Q53.90, Q53.91, Q53.92, Q53.93, and Q53.99</td>
</tr>
<tr>
<td>Ectopic testis</td>
<td>Q53.0, Q53.01, Q53.00, Q53.02, Q53.03, and Q53.09</td>
</tr>
<tr>
<td>Bifid scrotum, absence of scrotum, polyorchism, and other unspecified congenital malformations of testis or scrotum</td>
<td>Q55.2, Q55.8, Q55.9, Q55.3, and Q55.4</td>
</tr>
<tr>
<td>Hypospadias</td>
<td>Q54.0, Q54.1 Q54.2, Q54.3, Q54.8, and Q54.9</td>
</tr>
<tr>
<td>Other congenital malformations of male genital organs, lower urinary system, and indeterminate gender</td>
<td>Q56 and Q64</td>
</tr>
<tr>
<td>Other congenital malformations, deformations, and chromosomal abnormalitiesa</td>
<td>Q00–Q99 (excluding the above codes)</td>
</tr>
<tr>
<td>Atrophy of testis</td>
<td>N50</td>
</tr>
<tr>
<td>Torsion of testis</td>
<td>N44</td>
</tr>
<tr>
<td>Complications resulting from surgical procedure including wound infection, sepsis, hemorrhage, and hematoma</td>
<td>T81, T88, Z98, N99</td>
</tr>
<tr>
<td>ACHI procedure code</td>
<td></td>
</tr>
<tr>
<td>Primary procedures</td>
<td></td>
</tr>
<tr>
<td>Orchidopexy for undescended testis, unilateral</td>
<td>37803-00, 37806-00, and 37806-02</td>
</tr>
<tr>
<td>Orchidopexy for undescended testis, bilateral</td>
<td>37803-01, 37806-01, and 37806-03</td>
</tr>
<tr>
<td>Orchitectomy, unilateral</td>
<td>30641-00, and 30641-02</td>
</tr>
<tr>
<td>Orchitectomy, bilateral</td>
<td>30641-01, and 30641-03</td>
</tr>
<tr>
<td>Exploration of scrotal contents with fixation of testis, unilateral and bilateral</td>
<td>37604-04, and 37604-05</td>
</tr>
<tr>
<td>Exploration of groin for impalpable testis</td>
<td>37812-00</td>
</tr>
<tr>
<td>Laparoscopy</td>
<td>30390-00, and 30391-00</td>
</tr>
<tr>
<td>Secondary procedures</td>
<td></td>
</tr>
<tr>
<td>Revision orchidopexy for undescended testis, unilateral and bilateral</td>
<td>37808-00, and 37808-01</td>
</tr>
<tr>
<td>Refixation of testis, unilateral and bilateral</td>
<td>37604-07, and 37604-08</td>
</tr>
</tbody>
</table>

a Excluding minor congenital anomalies, ie, tongue-tie, nevus, skin tags, and unstable hip and feet defects.

related ICD10-AM–coded diagnoses are presented in Table 1.

Study Outcomes and Data

The main study outcomes were primary surgical procedures, testicular atrophy, testicular torsion, procedures after failed orchidopexy, and readmissions for postsurgical wound-related complications. Primary surgical procedures were orchidopexy (including codes for fixation of testis) and orchitectomy. All surgical procedures were identified using ACHI procedure codes and are listed in Table 1. Procedures after failed orchidopexy included revisions of orchidopexy, refixation of testis, and repeat orchidopexy. We excluded repeat orchidopexy performed within 12 months after primary orchidopexy for nonpalpable testis to avoid including routine second-stage orchidopexy.14 Postsurgical complications were defined as hospital readmissions occurring within 28 days if caused by surgery-related wound infections or hematomas.

Study factors included timing of diagnosis, coexistent congenital anomalies, preterm birth (<37 weeks’ gestation), age at primary surgery, public or private health insurance status, area of residence, and socioeconomic disadvantage. Age at primary surgery and time at first recorded diagnosis were categorized into 6 groups: birth, <12, 12 to <18, 18 to <24, 24 to <36, and ≥36 months. The area of residence was dichotomized into major cities versus regional/remote areas using the Accessibility/Remoteness Index of Australia,15 and socioeconomic disadvantage was determined using the Socioeconomic Indexes for Areas relative disadvantage scores developed by the Australian Bureau of Statistics16 and classified as disadvantaged (<20th centile) versus nondisadvantaged (≥20th percentile) (Table 2).

Statistical Analysis

The proportion of UDTs diagnosed and the prevalence of surgery was examined as total number of reported cases per 1000 male births in NSW, 2001 to 2011. The trend in annual prevalence was evaluated including only boys with similar follow-up time, ie, born up to 2010 and with corrective surgery performed before 36 months of age. Descriptive statistics were calculated to assess characteristics of boys undergoing surgery, and association between study factors, procedures, and age at surgery were assessed by using χ² tests and Cochran–Armitage test for trend. Spearman coefficient was used to determine correlation between date of birth and date at surgery, and multivariate logistic regression was applied to examine the association between risk factors for primary corrective surgery performed before or later than the recommended 12 months of age.9,10 Characteristics of boys and age of subsequent surgery by timing of first recorded diagnosis (birth or later) was also investigated. A P value <.05 was considered statistically significant, and all analyses were performed by using SAS version 9.3 (SAS Institute, Cary, NC).

RESULTS

There were 518 846 boys born live in NSW from 2001 to 2011; of these, 10 875 (2.1%) had a recorded diagnosis of UDT. Overall, 4980 (45.7%) boys with a recorded diagnosis of UDT had
corrective surgery (in 65 hospitals), representing a prevalence of 9.6 per 1000 male births. Of these, 80% (n = 3987) had surgery before 36 months of age, with the prevalence of these cases decreasing from 11.6 in 2001 to 7.8 per 1000 male births in 2010. Of boys undergoing surgery, more than three-quarters (78%) had unilateral and 17% bilateral UDT, whereas 5% were either ectopic or unspecified. Most cases were isolated (72%), with 8.7% having other genitourinary anomalies and 14.7% anomalies of other organs. Although the majority of boys had an orchidopexy (97.8%), 589 (11.8%) required surgery for impalpable testis and 223 (4.5%) had an orchietomy. As such, a small proportion of boys (n = 92, 1.8%) were diagnosed with testicular atrophy, and 58 (1.2%) had an associated diagnosis of testicular torsion. Overall, 26 (0.5%) boys were readmitted to the hospital within 28 days after surgery with postprocedural complications. Almost 10% (n = 446) required revision procedures after 1 year for failed primary surgery, and of these, 16 (3.6%) had a recorded diagnosis of testicular atrophy.

Among boys undergoing surgery, 78% were aged <36 months and 22% were >36 months (Table 3), with an overall median age at primary surgery of 16.6 months (interquartile range 11.8 to 31.0 months). As age of surgery increased, there was a decreasing trend in rates of orchietomy and surgery for impalpable testis, unilateral UDT, and testicular atrophy (P < .01) but an increasing trend in surgery for bilateral UDT (P < .01) (Table 3). For those with complete follow-up to 36 months (n = 3897), one-third had surgery at age <1 year and 54% at 12 to 24 months. The trend in the median age at surgery for these boys decreased from 14.7 months in 2001 to 12.7 months in 2010 (r = −0.05; P < .01). Primary surgery was more likely to be performed after 12 months of age for boys from socioeconomically disadvantaged backgrounds (adjusted odds ratio [aOR] 1.37; 95% confidence interval [CI] 1.13–1.67) and living in regional/remote areas (aOR 1.27; 95% CI 1.08–1.49), and less likely among boys from families with private health insurance (aOR 0.61; 95% CI 0.53–0.71). There was no association between preterm birth or coexistent congenital anomalies and surgery after 12 months of age.

When age at surgery was examined by timing of diagnosis, only one-third of boys (35%) had a recorded diagnosis of UDT at birth, with the remaining diagnosed in later health checks but first identified in this study at the time of surgery. Even for boys undergoing surgery at <12 months, less than half (44%) had UDT recorded at birth (Fig 1). However, there was no difference in the characteristics of boys and timing of reported diagnosis by socioeconomic or health insurance status, area of residence, preterm birth status, or type of surgery; those with other congenital anomalies were more likely to have UDT recorded at birth (Supplemental Table 4). For boys undergoing surgery after 36 months, 4.1% (n = 206 of 4980) had a recorded diagnosis at birth, and 18% (n = 877) did not.

**DISCUSSION**

This is one of the largest population-based cohort studies examining the diagnosis and treatment of UDT with long-term follow-up of ≤12 years. We found that >1 in 50 boys had a recorded diagnosis of UDT, and
the prevalence of boys undergoing surgery was 1% in NSW. Both the prevalence and the age at surgery decreased over the past decade; however, nearly two-thirds of cases still underwent surgery later than the recommended age of 12 months. We also found that almost 1 in 10 boys required subsequent revision.

The proportion of UDTs diagnosed at birth (1.9%) and those subsequently treated (1%) in NSW is similar to that published by a recent Danish population-based study. We found a decrease in prevalence of treated UDT over the past decade, which appears consistent with worldwide reported trends in Germany, New Zealand, Norway, and Russia and the Australian states of Victoria and Western Australia.

To our knowledge, this is the first population-based study reporting rates for testicular atrophy (1.8%), torsion (1.2%), and subsequent outcomes of orchidopexy, with 9% of patients requiring revision surgery and a rate of testicular atrophy at second surgery of 3.6%. To date, the only other published multicenter study (10 sites, n = 281 patients) reported overall 7.2% and 6.1% rates of failure and atrophy, respectively, in patients undergoing laparoscopic orchidopexy.

We found that at least two-thirds of corrective surgeries for UDT were
performed later than recommended guidelines. This is of concern, as late orchidopexy may be associated with increased risk of testicular cancer and diminished fertility potential in boys.\textsuperscript{22–26} Two recent systematic reviews reported a 2- to 32-fold increased risk of testicular cancer in men with a history of UDT, with the risk lower for younger age at orchidopexy but higher for previous bilateral UDT.\textsuperscript{8,27} Such findings have prompted a significant reduction in the recommended age to perform orchidopexy in the past decades, to its current 12 months of age.\textsuperscript{28} Although we found an encouraging trend of decreasing age at surgery, consistent with recent reports from different settings,\textsuperscript{11,29} the current rates of late surgery are particularly concerning, especially for those with bilateral UDT. We found that the overall proportion of primary surgery performed after 12 months was 74%. But, when limited to complete follow-up of 36 months, it was 67%. Two studies that were limited to congenital cases have reported 70% to 82% of cases having surgery after 12 months.\textsuperscript{11,30} Other studies have also reported a high proportion of orchidopexy performed later than the recommended age, but most potentially included cases of acquired UDT, which are diagnosed and treated at later ages.\textsuperscript{11} To date, there are no current Australian guidelines that specify the importance of the timing of early surgery.

One of the motivations of our study was to try and differentiate congenital from late diagnosed and potentially acquired cases. Although we had information on recorded diagnosis at birth, we found identification of congenital cases to be difficult because of the underreporting of cases at birth (only 35%) and a majority reported at time of surgery, with no information on previous testis position. Our findings suggest that the characteristics of boys undergoing surgery before 36 months of age without diagnosis at birth are similar to those diagnosed at birth and may potentially be congenital cases. The 18% having surgery after 36 months (without a recorded UDT diagnosis at birth) may be more likely to be acquired cases. A previous longitudinal cohort study ($n = 1072$) with examination of testicular position at 0, 3, 18, 36, and 54 months of age reported an 18% rate of acquired UDT;\textsuperscript{31} however, the proportion of acquired UDT may have been overestimated because of a 45% loss of follow-up.

Strategies to reduce the age at orchidopexy require the identification of factors that may delay surgery. These may involve missed diagnosis at birth, delays in follow-up by parents, missed diagnosis or lack of timely referral by general practitioners, or limited availability of hospital resources to conduct surgery.\textsuperscript{32,33} Missed diagnosis at birth may have occurred because infants were not examined or UDT was missed owing to a lack of experience or training of maternity care providers. Previous studies reveal that when examination is performed by trained experts, the prevalence of UDT detected at birth is higher, as reported in cohorts from Denmark (9%)\textsuperscript{2} and Lithuania (5.4%),\textsuperscript{34} compared with the 2.1% reported here. After birth, delays in diagnosis may also occur and be due to lack of parental awareness or limited attendance to routine health visits in the child’s first year of life. Lack of assessment, follow-up, or timely referral by general practitioners may also be a contributing factor. Thus, adherence to monitoring during routine health checks is essential to ensure timely intervention. This is particularly important because testicular position may vary in early life.

Given that interventions providing advice to general practitioners and parents in the UK have been shown to be effective at reducing age at orchidopexy,\textsuperscript{35} our results suggest that these may need to be implemented for future improvements in practice in Australia. This necessity is further confirmed by results from a recent survey reporting that 75% of UK general practitioners still consider 2 years to be the optimal age for performing orchidopexy.\textsuperscript{36} Another potential influence on the age at surgery remains the limited capacity and availability of NSW hospitals to perform orchidopexy before 1 year of age, with NSW health policy considering this a major surgical procedure to be performed only by surgeons credentialed in pediatric surgery, and with specialist anesthetists involved.\textsuperscript{37} Given more urgent cases and limited resources, delays in surgery may occur in the public hospital system. Therefore, increasing availability and access to surgery in more NSW hospitals would reduce demand and redistribute the burden and access to surgery.\textsuperscript{38} One alternative is for surgery to be undertaken in private hospitals, which do not have the same demands on limited resources.

In Australia, patients without private health insurance are treated in the public health system and placed on a waiting list, which is determined by urgency, whereas those with private health insurance are managed more immediately according to the availability of the pediatric surgeon. This issue is supported by our findings that boys with private health insurance had earlier primary surgery, compared with those publicly insured and those from lower socioeconomic background or living in rural areas.

The main strength of this study was the use of a large record-linked population-based cohort of boys that allowed the assessment and follow-up from 65 hospitals across NSW for ≤12 years. The health datasets used are accurate and
The authors have indicated they have no potential conflicts of interest to disclose.

CONCLUSIONS

In conclusion, 1 in 48 boys is born with UDT; however, two thirds of cases had no report of corrective surgery. The prevalence of treated UDT was 1%, with almost 1 in 10 requiring subsequent revision. The median age at surgery is decreasing; however, two-thirds of cases were still performed after the recommended age of 12 months. Further interventions are needed to improve detection of UDT at birth and increase understanding of the diagnosis by parents and management of UDT by primary health care providers. Importantly, the significance of early diagnosis and surgical correction before 12 months of age should be implemented in Australian guidelines, and changes should be made in health policy to increase health care resources to perform surgery at earlier ages.

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The authors thank the NSW Ministry of Health for access to the population health data and the NSW Centre for Health Record Linkage for record linkage.

ABBREVIATIONS

ACHI: Australian Classification of Health Interventions
aOR: adjusted odds ratio
CI: confidence interval
ICD10-AM: International Classification of Diseases, 10th Revision, Australian Modification
NSW: New South Wales
UDT: undescended testis

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