

# Endocrine and reproductive outcome of men born with various degrees of hypospadias

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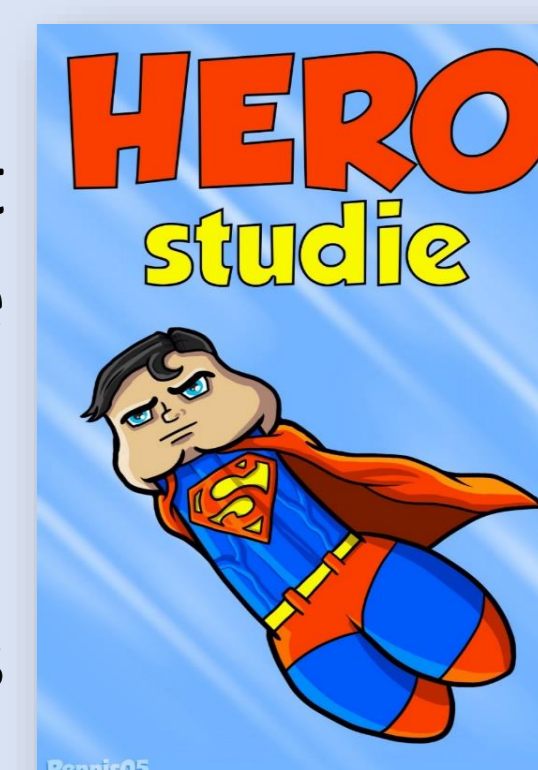
## Key messages

- 20% of hypospadias cases have reduced semen quality
- No difference in testosterone and LH levels was found between hypospadias cases and controls
- FSH and Inhibin B levels are not very reliable in predicting low sperm counts

## INTRODUCTION

**Background:** Limited, small-scale studies have revealed that men with proximal hypospadias or with other signs of undermasculinisation (i.e. complex hypospadias) are at risk of reduced fertility and/or impaired testicular hormone synthesis. However, the extent of this phenomenon and if milder forms of isolated hypospadias are also affected, remains unclear.

**Aims:** To explore reproductive hormones and semen quality of young men (16-21 years old) born with all forms of non-syndromic hypospadias in comparison to healthy controls.



## METHODOLOGY

**Design:** Cross-sectional assessment

**Centers:** Ghent University Hospital and Wien Medical University

**Tests:** Blood sampling between 8:00 and 9:00 AM for total and free testosterone, LH, FSH and Inhibin B. Spermogram on two independent semen samples, according to the WHO 2010 criteria.

**Statistics:** IBM SPSS® 25.0 using an unpaired Student t-test or Mann Whitney-U test as appropriate.

## PARTICIPANTS

Hypospadias	N= 192
Distal	132/192 (68,8%)
Midshaft	37/192 (19,3%)
Proximal	23/192 (12,0%)
Complex	20/192 (10,4%)
Controls	N=50

## RESULTS

### Spermograms

#### Samples:

- Two semen samples: 83,9% and 93,8% of cases and controls, respectively

#### Results:

- Azoospermia in 7 (4,1%) cases
- Oligozoospermia in 25 (14,7%) cases and 2 (4%) controls
- Normal spermogram: 86/170 (50,6%) cases; 30/50 (60%) controls
- In controls, mild astheno- and teratozoospermia were the most common causes of abnormalities

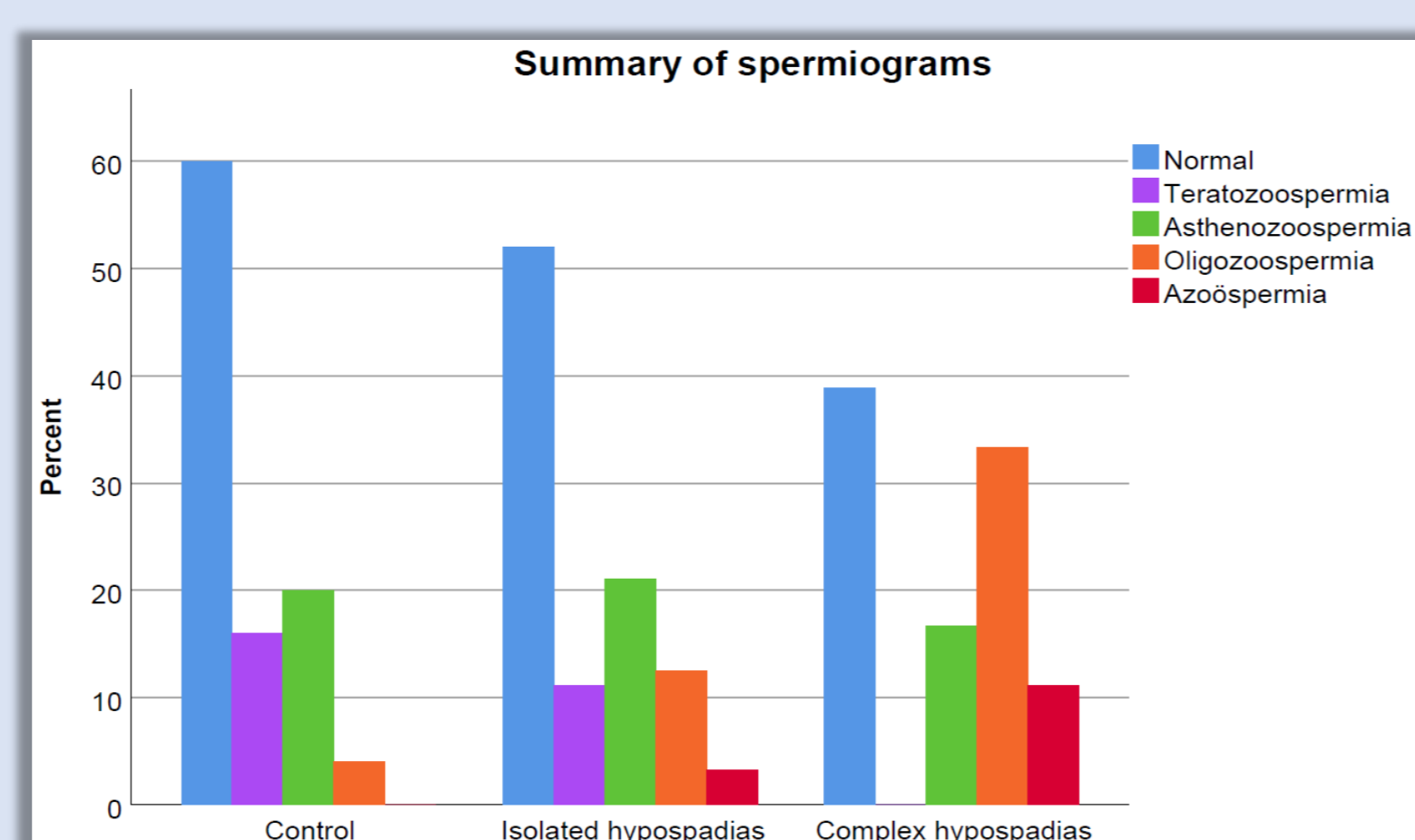


Figure: Summary of spermograms (according to WHO 2010 criteria). Complex, isolated hypospadias and controls.

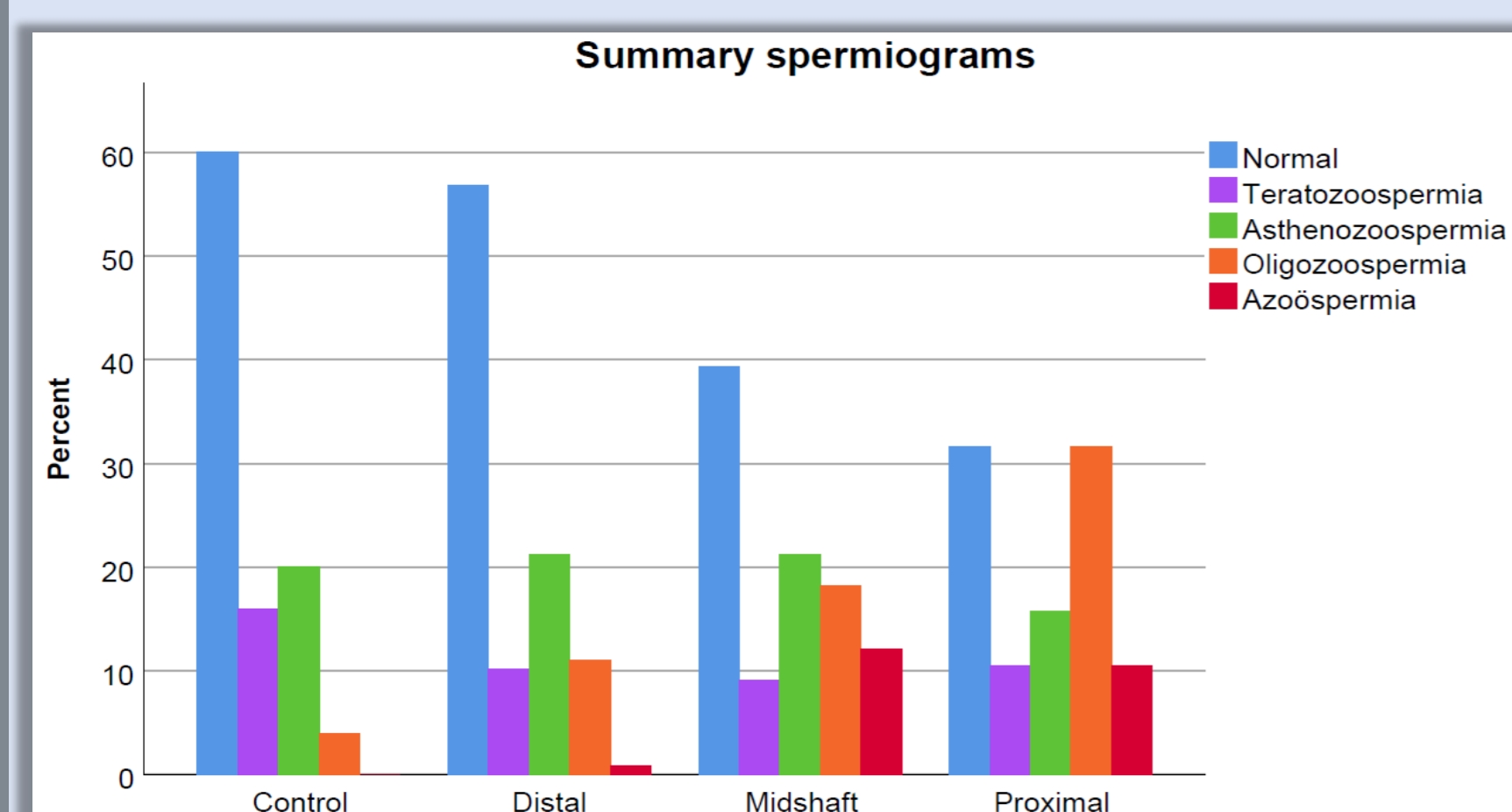


Figure: Summary of spermograms (according to WHO 2010 criteria). Proximal, midshaft, distal hypospadias and controls.

#### Spermatogenesis:

- Lower semen concentrations in proximal and complex hypospadias in comparison to controls (p=0,011 and <0,001, respectively)

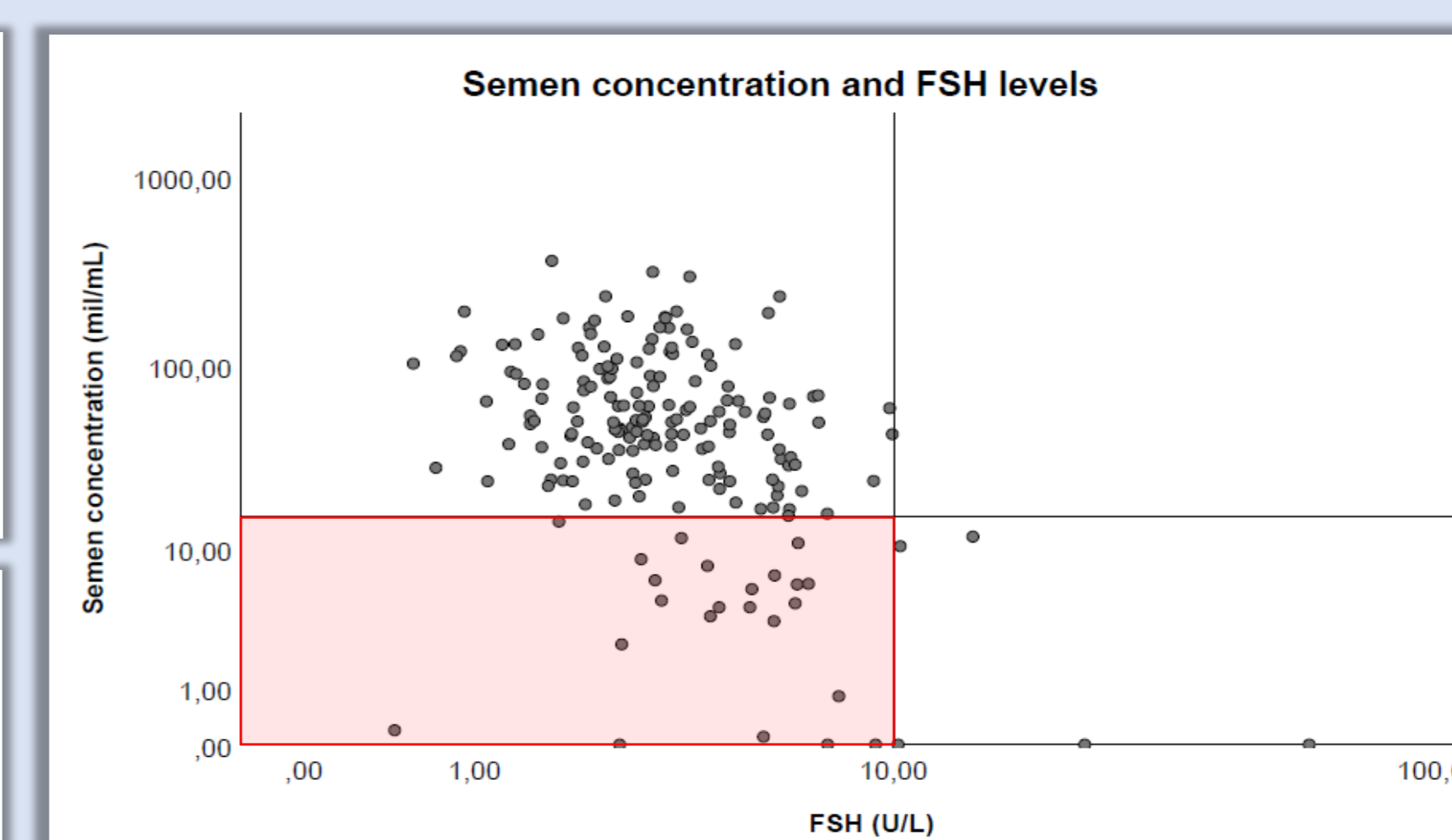
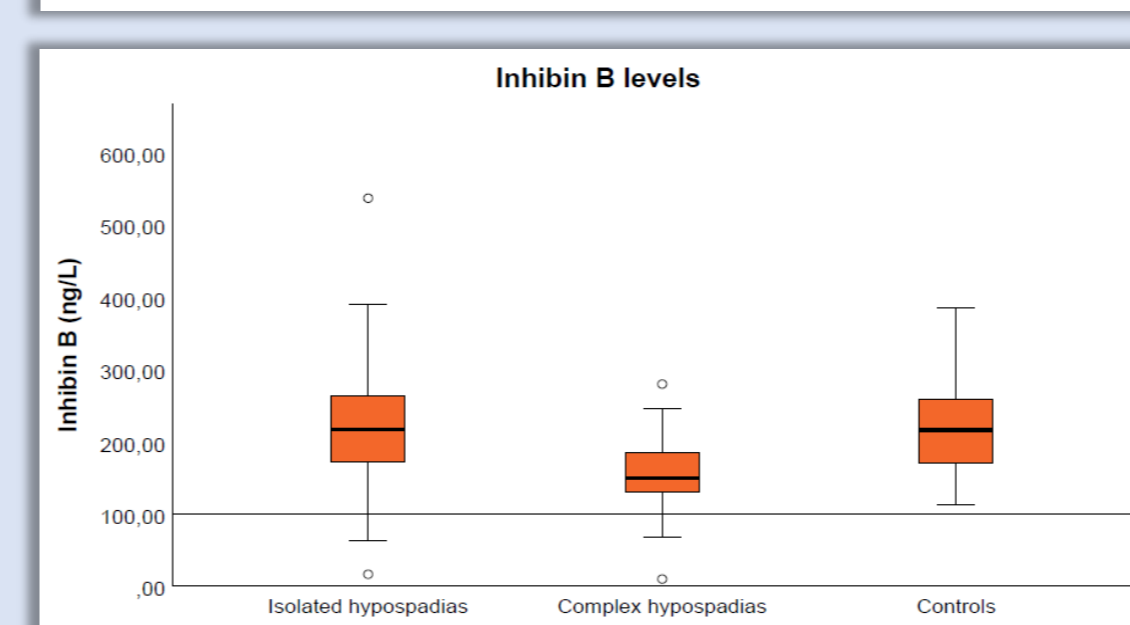
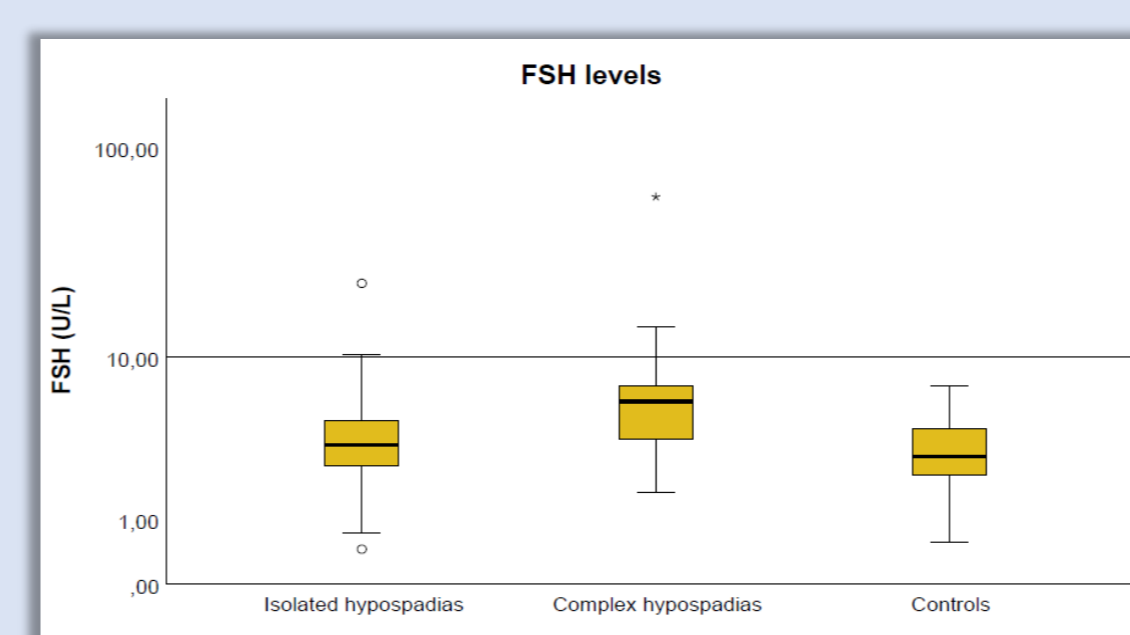
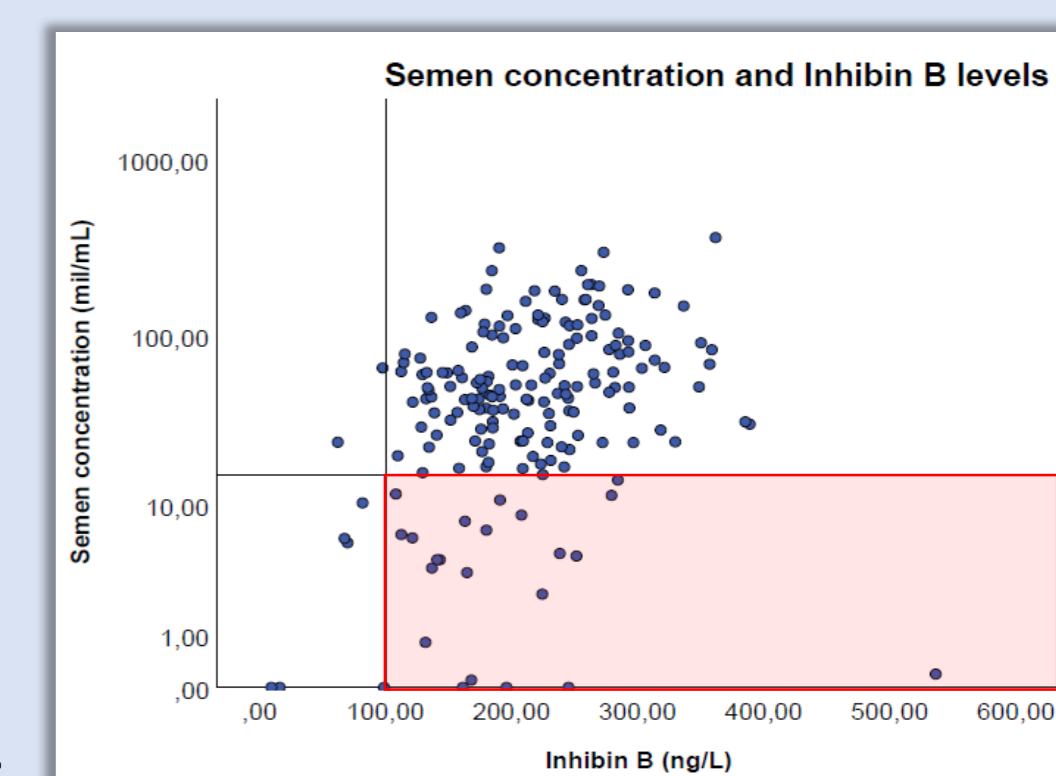
### Hormone levels

#### Androgens:

- Free / total testosterone & DHT levels
  - No differences:
    - Proximal / Distal / Controls
    - Complex / Isolated / Controls

#### Sertoli cell function:

- Higher FSH levels in complex hypospadias
  - Isolated hypospadias (p=0,011); Controls (p=0,005)
- Lower inhibin B levels in complex hypospadias
  - Isolated hypospadias (p=0,001); Controls (p=0,008)
- Both FSH and LH were poor predictors of oligo/azoospermia



Figures: FSH (top left) and Inhibin B (lower left) levels in complex, isolated hypospadias and controls. Inhibin B levels (top right) and FSH levels (lower right) and semen concentration. Red area marking oligo/azoospermia with normal hormone levels.

## CONCLUSION

In our cohort, over 20% of men born with hypospadias have reduced semen quality. Over 40% of complex and proximal hypospadias have oligo-/azoospermia as compared to 11,9% and 15,8% of distal and isolated hypospadias, respectively. There is no difference in testosterone or LH levels between hypospadias cases and controls. FSH and Inhibin B levels are not always predictive of a low sperm count.

## Funding

