

INFERTILITY/POLYCYSTIC OVARIAN SYNDROME

Introduction

Infertility is defined as the absence of pregnancy following 12 months of unprotected intercourse. Infertility may be caused by Ovulatory Dysfunction, Blocked Fallopian Tubes, Male Factor Infertility or Unexplained Causes. Ovulatory Dysfunction can be caused by hypothalamic causes, endocrinopathies (hyperprolactinemia, thyroid dysfunction) or ovarian causes (Polycystic Ovarian Syndrome (PCOS), ovarian failure). Only those causes of infertility which require a TUE will be addressed in this document.

Ovulatory Dysfunction: Polycystic Ovarian Syndrome (PCOS)

1. Diagnosis

A. Medical History

- Absent or irregular menstrual cycles;
- Clinical evidence of androgen excess (hirsutism, acne).

B. Diagnostic criteria

1. Hyperandrogenism: Hirsutism and/or hyperandrogenemia;
2. Ovarian dysfunction: Oligo-anovulation and/or polycystic ovaries and
3. Exclusion of other androgen excess or related disorders.

C. Relevant medical information

Some women with PCOS will have associated insulin resistance which may manifest as impaired glucose tolerance or overt diabetes.

2. Medical Best Practice Treatment

Prohibited substances:

- a. Clomiphene citrate and Letrozole
- b. Spironolactone

A. Name of prohibited substance

- a. First line therapy is clomiphene citrate, a weak anti estrogen, and letrozole, an aromatase inhibitor.

- b. Spironolactone may be used in some geographic regions of the world as a secondary treatment in the management of hirsutism caused by PCOS.

B. Route

Oral.

C. Dosage and Frequency

- a. 5 days per month
- b. Daily

D. Recommended duration of treatment

- c. 9-12 months
- d. Long-term use if necessary

3. Other Non-Prohibited Alternative Treatments

- a. Subcutaneous exogenous Follicle Stimulating Hormone (FSH) can be used as an alternative.

Metformin has not proven to be as effective as clomiphene or letrozole as a first line treatment. However, in women who are non-responsive to clomiphene or letrozole, or who demonstrate insulin resistance, an insulin sensitizer such as metformin may be added.

hCG and Progesterone may be used as adjuncts to first line therapy.

- b. Diane 35 (2 mg cyproterone acetate) and Yaz (3 mg drospirenone) are two oral contraceptives with anti-androgenic effects that are used as first line therapy for the treatment of hirsutism caused by PCOS. Any oral contraceptive or the NuvaRing (11.7 mg etonogestrel) will increase sex hormone binding globulin as a result of the increased estrogen. This will decrease free unbound, circulating androgens resulting in decreased hirsutism. For more severe or long standing cases, larger doses of cyproterone acetate (25-50mg) may be necessary. In some areas of the world, oral flutamide (non-steroidal anti-androgen) is used to treat hirsutism. Hormonal therapy can be combined with physical hair removal techniques such as laser or electrolysis.

A TUE may be granted for spironolactone should the athlete have proved:

The necessity [i.e. presence of hirsutism in the clinical picture of PCOS] and one or more of the following criteria:

- A contraindication to a non-prohibited method;

- An intolerance to a non-prohibited method;
- A failed response to a non-prohibited method;
- Inability to benefit from physical methods of hair removal due to prohibitive cost.

4. Consequences to Health If Treatment Is Withheld

- a. Significantly decreased quality of life if infertility is unresolved
- b. Significant decreased quality of life for women with hirsutism resulting from PCOS.

5. Treatment Monitoring

- a. Blood estrogen, and Luteinizing Hormone (LH) and ultrasound scan of the ovaries for follicular growth monitoring.
- b. Monitoring by gynaecologist, endocrinologist or dermatologist on a yearly basis is recommended.

6. TUE Validity and Recommended Review Process

- a. 2 years
- b. 10 years TUE with an annual review by a specialist can be granted for this substance as PCOS is a lifelong condition.

Unexplained Infertility

1. Diagnosis

A. Medical History

No pregnancy despite regular ovulatory cycles, open tubes, regular timed intercourse and normal semen analysis.

B. Relevant medical information

Nil.

2. Medical Best Practice Treatment

May be treated with clomiphene citrate (see PCOS), FSH/LH (TUE not required)

References

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