

Surgical intervention in infertility management

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Despite improvement in the success of IVF, reproductive surgery will remain an important option and complement to assisted reproductive technologies (ART) for many couples. Reproductive surgery should be considered as the first-line treatment when the correction of infertility pathologies is simple and a good result is expected once corrected, when the pathology is causing symptoms such as pain or abnormal bleeding, or if uncorrected will compromise the results or increase the risks of ART. The success of surgical infertility treatment depends on the careful selection of cases using appropriate investigative techniques, with procedures performed in centres with sufficient expertise. For both specialized reproductive and general gynaecological surgery it is crucial to follow microsurgical principles to avoid adhesion formation and conserve normal tissues, especially tubal and ovarian. These aspects of reproductive surgery, and different surgical techniques used for various tubal, peritoneal, uterine and ovarian conditions to achieve the optimal reproductive outcome are discussed in this article.

Tubal and peritoneal factors account for 30–40% of cases of female infertility. Tubal factors include damage to or obstruction of the Fallopian tubes, usually related to previous pelvic inflammatory disease or pelvic or tubal surgery. Peritoneal factors include pelvic and peri-adnexal adhesions and endometriosis. Although some are correctable with surgery, the limitations of surgical repair in many cases have been the driving force behind the development of assisted reproduction technology (ART).

Although ART results are improving, operative endoscopy has also made advances, ensuring an ongoing place in the management of infertility. Improvement in instrumentation and the feasibility of laparoscopic suturing has allowed many reconstructive procedures that traditionally required a laparotomy to be accomplished safely with laparoscopy on a day-surgery basis with lower costs, shorter hospital stays and faster recovery times. Surgical infertility treatment offers other advantages over ART. Unlike ART, which bypasses pelvic pathologies, surgical approaches improve fertility by correcting them and potentially improving other related symptoms such as pain and abnormal menstruation. After surgery, couples can have unlimited attempts to conceive naturally without being subjected to the risks of multiple pregnancy and ovarian overstimulation, stress and cost associated with ART cycles. The fact that more than 75% of couples who commence ART but have not conceived by the end of the third cycle drop out for non-clinical reasons independent of cost illustrates the difficulty that ART procedures present (Land *et al.*, 1997; Meldrum *et al.*, 1998). ART cycles are expensive and the multiple cycles

needed to achieve a reasonable cumulative pregnancy rate are unaffordable for many couples and communities. The age of the patient has a major effect on the cost-effectiveness of IVF treatment. Women who are older than 38 years have cost per delivery by IVF of nearly three times that of younger women (Trad *et al.*, 1995; Van Voorhis *et al.*, 1997). Older patients (especially over 40 years of age) also have additional deleterious effects of *in vitro* culture compared with the tubal environment, such as zona pellucida hardening (Cohen *et al.*, 1992). In these women, some surgical treatments such as sterilization reversal can achieve pregnancy rates of 42–64%, with a median interval to conception of 5 months (Trimbos-Kemper, 1990; Kim JD *et al.*, 1997; Yoon *et al.*, 1999), whereas IVF success rates tend to be poor with a per cycle pregnancy rate of less than 10% (ASRM–SART Registry, 2002). In addition, ART outcomes can be adversely affected by some surgically correctable pathologies, such as uterine and endometrial lesions, hydrosalpinx and endometrioma. Fertility related surgeries are listed in Box 1.

Many gynaecological operations are performed for women of reproductive age for indications unrelated to fertility. Some of these procedures have consequences that may compromise the future fertility of the patient through tubal damage or adhesions or by affecting ovarian reserve. Every effort should be made to avoid these complications.

Surgical principles and adhesion prevention

Previous surgery is a significant contributor to pelvic adhesions, and 75% of women develop adhesions after pelvic surgery (DeCherney and Mezer, 1984). Adhesions can result in pelvic pain as well as infertility in women of reproductive age. The potentially serious risks of intestinal obstruction and injury to vessels and viscera in future procedures are increased as a result of adhesions.

In the normal healing process endogenous fibrinolytic enzymes, such as plasminogen activator, lyse and remove fibrin matrix formed at the site of injury within 4 days. Local tissue ischaemia, tissue or peritoneal surface desiccation or infection can result in the generation of inhibitors to plasminogen activity, reducing fibrinolysis. Generation of excessive fibrin clots as a result of poor haemostasis and the early apposition of damaged peritoneal surfaces further limit the fibrinolytic capability and enhance fibroblast invasion and angiogenesis. The key principle of adhesion prevention is therefore first to limit the inciting event of tissue or peritoneal damage through microsurgical principles and second to keep the damaged peritoneal surfaces apart for a minimum of 5–7 days to maximize the potential for fibrinolysis while healing occurs.

Microsurgery is a surgical philosophy, a delicate surgical approach designed to minimize peritoneal trauma and tissue

disruption while increasing the accuracy of the procedure and improving the outcome. These principles include:

- The use of techniques designed to minimize tissue injury including delicate handling of tissues and frequent intraoperative irrigation to prevent desiccation.
- Judicious dissection to minimize adjacent tissue damage and appropriate use of sutures, electrical or laser energy to achieve meticulous haemostasis.
- Avoiding the introduction of foreign bodies into the peritoneal cavity including the use of fine, inert or absorbable sutures, the use of powder-free gloves, peritoneal lavage and copious irrigation.
- Use of magnification that permits accurate identification of abnormal morphology for complete removal, recognition and avoidance of surgical injury, an application of the preceding principles with the use of fine microsurgical instruments and suture materials for precise alignment and approximation of tissue planes.

Endoscopic surgery has many inherent microsurgical advantages. Endoscopy provides some degree of magnification at all time and operating within a closed peritoneal cavity largely prevents desiccation of the peritoneal surfaces. It minimizes the need for tissue handling, eliminates the use of packs and prevents the introduction of foreign materials such as lint and talcum powder. Intraoperative irrigation can be performed easily to keep tissues moist and expose any bleeding vessels. Furthermore, the pressure effect of the pneumoperitoneum diminishes venous oozing and permits spontaneous coagulation of minor vessels. In addition it allows visualization of the operative field at angles that are not possible with open microsurgery. Both animal and human studies have shown that laparoscopic adhesiolysis is more effective in reducing adhesions, with less *de novo* adhesions formed after laparoscopic surgery as compared with laparotomy (Lundroff *et al.*, 1991).

Various substances have been used in an attempt to minimize postoperative adhesions. The administration of systemic anti-inflammatory adjuvants produces disappointing results. Adhesions tend to occur in devascularized sites where systemic agents penetrate poorly. Instillation of fluid at the time of surgery is also generally unsuccessful, as the peritoneal cavity rapidly absorbs fluid and electrolytes, with most fluid being absorbed within 24 h, well before the healing process of the peritoneal surface is completed. Heterologous barriers either of permanent (Gore-Tex) or delayed bioresorbable synthetic products based on hyaluronic acid compositions and regenerated cellulose (Septrafilm, Interceed) produce various degrees of success. After application they create a barrier effect during the re-epithelialization stages before undergoing absorption. Several barrier gels based on hyaluronic acid, which can be easily administered even during laparoscopy to cover large operative areas, have recently been introduced, with promising initial results. Most of these products have been demonstrated to be effective in laboratory models of adhesion formation and oxidized regenerated cellulose has been demonstrated to reduce adhesion formation in women in well-controlled randomized prospective studies (Nordic Adhesion Prevention Group, 1995; Franklin, 1995). However, none of these agents has yet been demonstrated to be effective in increasing the fecundability of women

Box 1. Commonly performed fertility related surgeries in males and females

Male	Female
Varicocele (varicocelectomy or embolization)	Tubal diseases
Reversal of male sterilization (vasovasostomy)	Pelvic adhesions
Ejaculatory duct obstruction (transurethral resection)	Endometriosis
Sperm extraction (PESA, MESA, TESE, TESA)	Uterine
	Fibroid
	Uterine septum and other congenital anomalies
	Endometrial polyp
	Intrauterine adhesions
	Ovarian
	Polycystic ovarian syndrome
	Ovarian cysts/accidents
	Cancer preservation surgery
	Ovarian transposition
	? Ovarian tissue harvest, freezing and grafting

PESA: percutaneous epididymal sperm aspiration; MESA: microsurgical epididymal sperm aspiration; TESA: testicular sperm aspiration; TESE: testicular sperm extraction.

undergoing infertility surgery. For now, we should strive to apply microsurgical principles to minimize the morbidity and costs caused by the postoperative formation of adhesions.

Tubal disease

Tubal function and testing

The tubal epithelium with its secretions provides a unique nurturing environment that enhances oocyte maturation and sperm function leading to improved fertilization, and is the site of early embryo development. Tubal patency testing alone cannot reflect the full functional roles of the Fallopian tubes. It was observed in both animal and human studies that fertility diminishes in a linear fashion with the shortening or mucosal damage of the ampulla.

Hysterosalpingogram (HSG) allows the visualization of the uterine cavity and tubes by X-ray using iodine-based contrast medium, which outlines the endometrium and tubal mucosa. A meta-analysis of 20 investigations comparing the accuracy of diagnosis of tubal patency by HSG with laparoscopy showed a low overall sensitivity of 65% (Swart *et al.*, 1995). Mucosal adhesions and flattening were seen at salpingoscopy (Marconi *et al.*, 1992) and falloposcopy (Surrey *et al.*, 1997) in about 40% of infertile patients with normal laparoscopy or HSG. Pregnancy outcome (after a 2-year follow-up) correlated with the status of the mucosa as determined by salpingoscopy (Surrey and Surrey, 1996) and falloposcopy (Kerin *et al.*, 1992) but not with laparoscopy or HSG findings.

Approximately 50% of patients with documented tubal damage have no identifiable risk factor for tubal disease (Rosenfeld *et al.*, 1983), and a considerable proportion of patients with endometriosis also present with no symptoms apart from reduced fertility. Laparoscopy with salpingoscopy should ideally be the preferred investigation over HSG, with additional advantages of permitting treatment at the same time. However, this is often not practical due to the need for general anaesthesia, the associated morbidity involved and the greater demand on resources. For the low-risk patients who want to avoid laparoscopy, HSG may be considered as the first line of investigation. The utilization of screening for *Chlamydia* antibodies may help to define this low-risk group (Mol *et al.*, 1997). The development of new techniques, such as hydro-laparoscopy, that can be performed under local anaesthetic and sedation in an outpatient setting shows great promise. Hydro-laparoscopy not only permits examination of peritubal and peritoneal conditions, but an examination of the ampullar mucosa can also be achieved (Brosens *et al.*, 1999).

Peritubal adhesiolysis and distal tubal disease

The effect of tubal and ovarian adhesions on fertility was investigated in a controlled study which evaluated the effect of salpingo-ovariolysis on subsequent fertility (Tulandi *et al.*, 1990). The cumulative pregnancy rate in the group that underwent salpingo-ovariolysis was three times higher than in the non-treated group (32% versus 11% at 12 months and 45% versus 16% at 24 months).

Laparoscopic salpingo-ovariolysis should be the treatment of choice for women with periadnexal adhesions as *de novo* adhesion is less likely to occur when compared with laparotomy (Lundroff *et al.*, 1991). High cumulative pregnancy rates of about 60% with low ectopic pregnancy (EP) rate (6%) have been reported, and almost half of the intrauterine pregnancies (IUP) occur within 6 months after surgery (Donnez, 1987; Gomel, 1989). Pregnancy rate fell sharply to below 20%, with an EP rate rising to a similar figure in the presence of dense or incomplete removal of adhesions, or when a good anatomical relationship between Fallopian tube and ovary failed to be restored. Adhesiolysis can often be performed at the time of the routine diagnostic laparoscopy during the infertility work-up. Lysis of dense adhesions is occasionally difficult by laparoscopy requiring microsurgical laparotomy, but this often carries a poorer prognosis and such patients should be referred for IVF.

The inflammatory process in tubal disease may be limited to the serosal surface of the Fallopian tube or may cause extensive destruction of mucosal folds and the microscopic cilia that line the ampullary portion. Results for distal tubal surgery in general are less satisfactory, and pregnancy rates are about 30% (Kasia *et al.*, 1997). However, the results vary widely, mainly because of bias in case selection in the absence of standardized assessment of the extent of tubal damage, especially the mucosal state. In a prospective study, mild disease was defined as absent or small hydrosalpinx (15 mm) with easily recognized fimbriae when tubal patency was restored, no significant peritubal or periovarian adhesions, and hysterosalpingogram demonstrating normal rugal folds. Up to 80% of these patients achieved IUP. Moderate disease consisted of a hydrosalpinx

15–30 mm in diameter, with a portion of fimbriae not easily identified, the presence of perifimbrial or periovarian adhesions, few cul-de-sac adhesions and absent rugal folds on preoperative hysterosalpingogram. The pregnancy rate in this group was only 21%. Severe disease was characterized by a large hydrosalpinx (≥ 30 mm), total absence of fimbriae, dense pelvic or adnexal adhesions with a fixed ovary and tube and obliteration of cul-de-sac. Pregnancy rates of only 3% could be achieved in this subgroup. EP occurred in 4–20% of these patients, dependent on severity of tubal damage (Bower-Meisel *et al.*, 1986; Schlaff *et al.*, 1990).

A thorough preoperative and intraoperative assessment is important to identify those patients with distal tubal or peritubal disease who would benefit most from surgery. If HSG films are obtainable, evidence of proximal disease, ampullary mucosal folds, strictures, intratubal adhesions and spillage, and dispersion of the contrast medium should be specifically investigated. When distal and proximal obstruction are both present (that is, bipolar disease), the surgical success rate is $\leq 5\%$. At laparoscopy direct inspection of the crucial distal one-third of the tubal lumen can be carried out by salpingoscopy using either a rigid hysteroscope or fine flexible endoscope capable of visualizing intratubal adhesions or loss of normal landmarks.

Proximal tubal disease and sterilization reversal

Up to 25% of female infertility is attributed to tubal occlusion, of which proximal tubal problems account for one-third. Mid-tubal obstruction is often a result of a previous sterilization procedure or management of an EP. Other tubal occlusions are attributed to intratubal adhesions and plugs, pelvic inflammatory disease, salpingitis isthmica nodosa or other intratubal causes such as polyps and endometriosis.

Non-structural occlusion can often be dislodged by simple measures. A meta-analysis of six randomized and six non-randomized controlled studies comprising a total of 2635 patients demonstrated that flushing the tubes with oil-soluble media at hysterosalpingography increases subsequent pregnancy rates in infertile patients (odds ratio (OR) 1.80, 95% confidence interval (CI) 1.29–2.50). Similarly, a review of 11 studies of transcervical Fallopian tube catheterization in 1990 described an 82% cannulation rate with IUP and EP rates of 24% and 6%, respectively (Thurmond and Rosch, 1990). However, the 60% discrepancy between the cannulation and pregnancy rates raises concerns regarding the effect of cannulation on tubal mucosal health or pre-existing tubo-peritoneal conditions in these women, despite tubal patency being achieved.

Microsurgery is considered to be an effective approach to non-sterilization related structural proximal tubal obstruction. Pregnancy rates as high as 70% can be achieved by microsurgical resection and re-anastomosis in a highly selected group of patients (Dubuisson *et al.*, 1997). However, the amount of affected portion that must be removed can be difficult to assess in patients with disorders such as salpingitis isthmica nodosa. Rates of subsequent IUP and EP can vary widely; adverse prognostic factors for future fertility include reduced residual length of tube, significant intramural damage and evidence of pelvic infection or inflammation.

Up to 14% of women who have had surgical tubal sterilization will request a reversal procedure (Schmidt *et al.*, 2000).

Tubal reanastomosis for reversal of sterilization is the most successful technique in microsurgery, as these patients are otherwise fertile and healthy tubal tissues are anastomosed after the localized damage is removed. Seven large series including a total of 2018 patients available for follow-up with many sterilized by cautery showed an average pregnancy rate of 68% and EP rate of 4% after microsurgical reversal either open or laparoscopically (Table 1). In two series about 80% of patients who conceived did so within 12 months with a median interval from surgery to pregnancy of about 5 months (Kim JD *et al.*, 1997; Yoon *et al.*, 1999). Even in women at or over 40 years of age reversal can achieve high pregnancy rates averaging 46% for 149 patients (Trimbos-Kemper, 1990; Glock *et al.*, 1996; Kim JD *et al.*, 1997; Kim SH *et al.*, 1997; Yoon *et al.*, 1999). The ability to perform tubal anastomosis by laparoscopy (Table 1) or mini-laparotomy (Daniell and McTavish, 1995) on a day-surgery basis with comparable results to laparotomy has made the procedure even more attractive to patients.

The success of tubal reanastomosis is dependent on the method of sterilization, site of anastomosis, length of tube remaining and the presence of other infertility factors. Reversal of Falope rings or clips sterilization produces higher conception rates than in those sterilized by the use of the Pomeroy technique, whereas the worst results are observed in those sterilized with tubal cautery (Rock *et al.*, 1987; Kim JD *et al.*, 1997). The prognosis is best when the anastomotic sites have no significant discrepancy in diameter (for example, isthmic-isthmic or cornual-isthmic anastomosis). Higher pregnancy rates and a lower median interval between surgery and pregnancy are expected with a longer tube remaining (Kim JD *et al.*, 1997; Kim SH *et al.*, 1997). Pregnancy rates decrease by 50% if the tubal length is 3–4 cm and pregnancy does not occur if there is less than 3 cm of tube (Silber and Cohen, 1980). The time between sterilization and reversal is not generally regarded as important but one study noted an increased risk of damaged mucosa with flattening of epithelium and polyp formation in the proximal portion of the tube after 5 years following sterilization (Vasquez *et al.*, 1980). When compared with the macro-surgical approach, the use of a microsurgical technique has significantly improved the outcome of tubal anastomosis with reduced EP rates (Lavy *et al.*, 1987).

The patient's ovarian function, tubal condition and male infertility factors need to be assessed and considered in counselling the patient of the outcomes and likelihood of future fertility after reversal. Occasionally, laparoscopy may be necessary to assess surgical prognosis in complicated cases.

Ectopic pregnancy

With improved awareness and earlier detection of EPs, the treatment focus for EPs has largely evolved from saving lives to preserving fertility. In carefully selected patients medical treatment for EP can be highly successful. Although one meta-analysis showed an inferior successful resolution rate of 71% after a single dose of intramuscular methotrexate (Parker *et al.*, 1998), a more recent meta-analysis showed a success rate of 87% (range 75–90%), which is similar to that of laparoscopic salpingostomy (91%, range 72–100%; Morlock *et al.*, 2000). The subsequent IUP and EP rates were 54% and 8%, respectively, after medical treatment (Morlock *et al.*, 2000). Medical treatment

Table 1. Results of microsurgical tubal sterilization reversal by laparotomy and laparoscopy

Approach	<i>n</i>	% Pregnant	% Ectopic	Reference
Open	922	55.0	5.0	Kim SH <i>et al.</i> , 1997
Open	364	90.0	2.0	Kim JD <i>et al.</i> , 1997
Open	206	70.0	2.0	Dubuisson <i>et al.</i> , 1995
Open	124	74.0	6.0	DeCherney <i>et al.</i> , 1983
Open	118	64.0	1.0	Gomel, 1980
Laparoscopy	186	80.6	3.2	Yoon <i>et al.</i> , 1999
Laparoscopy	98	70.0	7.0	Bissonnette <i>et al.</i> , 1999
Total	2018	67.6	3.9	

was found to have a more negative impact on patients' health-related quality of life (Nieuwkerk *et al.*, 1998), but was generally less expensive than surgery when appropriately selected (Nieuwkerk *et al.*, 1998; Morlock *et al.*, 2000), with initial human chorionic gonadotrophin concentrations < 3000 mIU ml⁻¹ (Mol *et al.*, 1999). As there is risk of tubal rupture and surgical intervention may be necessary in the post-therapy period, access to transportation, patient contact and patient assistance are all key factors to be considered.

Laparoscopic surgery has been shown to be superior to laparotomy in the treatment of EP in three prospective randomized trials, including a total of 231 patients (Vermesh *et al.*, 1989; Lundroff *et al.*, 1991; Murphy *et al.*, 1992). A meta-analysis with a total of 1514 patients published in 1997 showed that the subsequent IUP and recurrent EP rates after linear salpingostomy for treatment of EP were 61% and 15%, respectively (Yao and Tulandi, 1997). Lower IUP and EP rates of 38% and 9.8%, respectively, were observed in a total of 3584 patients from 18 studies who had salpingectomy (Yao and Tulandi, 1997). However, it is uncertain whether the lower IUP rates in those who had salpingectomy were the result of surgery or the severity of the tubal pathology. A history of previous infertility or tubal pathology detected at the time of surgery had the strongest impact on future fertility. Patients with neither of these risk factors had a 75% IUP rate after salpingectomy for EP, whereas patients with one or both risk factors had a 36.6% IUP rate (Dubuisson *et al.*, 1996). The rate of persistent EP after salpingostomy is 3–20% and is often cured with single-dose methotrexate (Yao and Tulandi, 1997). In the absence of a randomized study, salpingostomy if technically feasible should be the preferred treatment option for EP in women who wish to preserve their fertility and are haemodynamically stable.

Hydrosalpinx

Numerous retrospective studies in the 1990s indicated that the presence of extensive tubal disease, especially a hydrosalpinx, might have a deleterious effect on the outcome of IVF. These reports were summarized in two meta-analyses which showed that pregnancy rates are reduced by half and the rates of spontaneous abortion are more than doubled in the presence

of a hydrosalpinx (Zeyneloglu *et al.*, 1998; Camus *et al.*, 1999). Animal and *in vitro* studies have further demonstrated that hydrosalpingeal fluid can inhibit sperm motility and is embryotoxic, though the latter has not been confirmed in humans (Mukherjee *et al.*, 1996). Endometria in women with a hydrosalpinx expressed significantly lower concentrations of integrins than endometria of women without a hydrosalpinx, and normal expression was restored after removal of the hydrosalpinx (Meyer *et al.*, 1997).

A recent Cochrane review of three prospective randomized trials showed that laparoscopic salpingectomy for hydrosalpinges before IVF increased pregnancy (OR 1.75, 95% CI 1.07, 2.86) and live birth (OR 2.13, 95% CI 1.24, 3.65) rates (Johnson *et al.*, 2002). In the subset of women with hydrosalpinges visible on ultrasonography and those affected bilaterally, salpingectomy increased the delivery rates by more than twofold and 3.5-fold, respectively (Strandell *et al.*, 1999).

Despite these findings, there are concerns about the potentially negative impact of salpingectomy on ovarian function. Though numerous early retrospective studies reported no immediate adverse effects on ipsilateral or contralateral ovarian function, some indicated that salpingectomy when not properly performed may disrupt the normal blood flow to the ovary, resulting in fewer oocytes retrieved from the side of operation during IVF cycle in comparison with the side with intact adnexa (Lass *et al.*, 1998). Options other than salpingectomy have been considered. Studies on simple aspiration of the hydrosalpinx have shown conflicting results (Sowter *et al.*, 1997; Van Voorhis *et al.*, 1998). The roles of proximal tubal occlusion and distal salpingostomy have yet to be clarified.

Endometriosis and endometrioma

Endometriosis is a common finding in women of reproductive age. It often presents with pelvic pain and infertility. Although both medical and surgical treatments are generally effective in treatment of pain due to endometriosis, the surgical approach is associated with lower recurrence rates as the disease is removed rather than suppressed as in medical therapy. However, for patients who are infertile, medical treatment is nowadays accepted as ineffective in enhancing fertility, whereas its role as an adjuvant therapy for ART remains debatable. Earlier studies indicated that women with moderate or severe disease had improved fecundity with the removal of implants. Further studies, including meta-analyses of non-randomized studies (Adamson and Pasta, 1994) and a randomized study in Canada (Marcoux *et al.*, 1997), have shown small but significant increases in fecundity when implants in mild disease are removed.

ART is effective in the treatment of infertility in patients with endometriosis. Although ART pregnancy rates in patients with endometriosis in general are comparable with those of women with tubal disease, the results for women with tubal disease may be suboptimal due to the presence of a hydrosalpinx. There are reports of poorer ovarian responses to stimulation and poorer outcomes of ART cycles in women with advanced endometriosis, especially those with endometriomas (Dicker *et al.*, 1991; Loh *et al.*, 1999). Ovarian hyperstimulation during ART cycles can worsen the disease and patient's symptoms. There is also an increased risk of infection

during egg collection in the presence of endometrioma. Medical suppression of the endometriosis or the removal of the endometriotic cyst should thus be considered before IVF. Both approaches have been shown to improve the ovarian response to ovarian hyper-stimulation and improve pregnancy rates during the IVF cycle (Dicker *et al.*, 1991; Marcus and Edwards, 1994; Loh *et al.*, 1999).

Various strategies have been used to treat endometrioma including aspiration, cystectomy and coagulation of cyst capsules. Medical therapy or simple puncture and drainage of endometrioma offer a poor response rate with recurrence rates greater than 80% at 6-month follow-up (Vercellini *et al.*, 1992). Removal of the cyst lining, by either stripping or sharp excision of the capsule, has been shown to produce a good outcome with recurrence rates from 0 to 12.2% and a pregnancy rate of 43–45% (Bateman *et al.*, 1994; Catalano *et al.*, 1996). However, not all endometriotic cysts have a 'true' cyst wall or obvious cleavage plane, and in one study up to 81% of suspected endometriomas were in fact luteal cysts (Nezhat *et al.*, 1992). There are concerns about the damage that this technique may cause to the underlying ovary; changes in ovarian artery vascular dynamics as assessed by Doppler ultrasonography have been reported following laparoscopic stripping (La Torre *et al.*, 1998). Alternatively, the visible deposits on cyst lining can be ablated by bipolar or KTP laser energy after exposure and irrigation (Sutton *et al.*, 1997), or as a two-step procedure after initial drainage and a period of medical suppression (Brosens *et al.*, 1996). The latter is more invasive though possibly produces better long-term results. Retrospective studies have shown that women with endometriomas treated by fenestration and ablation had threefold higher recurrence and re-operation rates at 18 and 42 months compared with those treated by total excision of the cyst (Saleh and Tulandi, 1999). However, these findings were not confirmed in a small prospectively randomized study, although the power of that study was too low to distinguish all but a large difference in outcome (Beretta *et al.*, 1998). Pre-treatment with gonadotrophin-releasing hormone agonist (GnRH-a) has little to offer (Muzii *et al.*, 1996), and the benefit of postoperative use is still unclear. The one area in which GnRH-a treatment does appear to be beneficial is as an adjuvant to a two-stage ablative procedure (Donnez *et al.*, 1996).

Excision of endometriotic deposits or stripping of the capsule can result in a significant amount of adhesion formation, leading to decreased fecundity (Canis *et al.*, 1992; Gurgan *et al.*, 1996). On the other hand, patients may continue to suffer pain and compromised fertility as a result of residual infiltrating disease that is undiagnosed or under-treated. A carefully considered treatment plan according to the main surgical indication together with intraoperative assessment should be made to strike a balance. For endometrioma, stripping of its capsule would seem reasonable if the capsule is well-defined, but with resort to ablation in areas where the cyst wall is densely adherent. If there is no obvious tissue plane, it may be prudent to consider the use of an ablative procedure either in one or two stages in conjunction with GnRH-a therapy.

Uterine fibroids and endometrial lesions

Uterine fibroids are found in 25% of women older than 35 years with common symptoms ranging from excessive uterine

bleeding, pelvic pressure and pain, recurrent pregnancy loss and infertility. Pregnancy loss and infertility appear to be mainly related to submucous or intramural myomas that distort uterine cavity. A recent review of 11 observational studies showed that neither intramural nor subserosal fibroids with no intracavitary component had significant adverse effect upon implantation, pregnancy and delivery rates (Pritts, 2001). However, this remains an issue of debate as recent publications have indicated that intramural fibroids can impair results of ART despite the absence of intracavitary involvement (Bajekal and Li, 2000; Bernard *et al.*, 2000; Hart *et al.*, 2001). However, as significant morbidity and adhesion formation that may compromise future fertility can follow surgery, patients should carefully consider the risks and benefits of myomectomy before proceeding to surgery. For women who have uterine fibroids without intracavitary involvement, myomectomy should be considered only in those with otherwise unexplained infertility or recurrent miscarriage in which all other causes of infertility have been excluded. Laparoscopy may become the preferred approach as it is associated with reduced risks of haemorrhage and postoperative adhesions when compared with open myomectomy (Dubuisson *et al.*, 2000). These findings are supported by two randomized trials, involving a total of 171 patients, which demonstrated that the laparoscopic approach is associated with less febrile morbidity, haemoglobin drop, pain and shorter hospital stay, with compatible sizes of fibroids removed and similar subsequent pregnancy outcomes when compared with laparotomy (Mais *et al.*, 1996; Seracchioli *et al.*, 2000).

Conversely, women who had a submucosal or intramural fibroid resulting in abnormal endometrial cavities had significantly lower IVF pregnancy (relative risk (RR) 0.32, 95% CI 0.13–0.70) and implantation (RR 0.28, 95% CI 0.10–0.72) rates than infertile controls, with pregnancy and delivery rates increased above those of infertile controls after myomectomy (Bajekal and Li, 2000; Pritts, 2001). Similarly, other studies have demonstrated a significant reduction in ongoing pregnancy rates in patients with other intracavitary lesions such as intra-uterine adhesions (Pabuccu *et al.*, 1997), uterine septum (Simon *et al.*, 1991; Acien, 1997; Homer *et al.*, 2000) and endometrial polyps (Varasteh *et al.*, 1999). With the advent of hysteroscopic resection, such lesions can be removed easily via a relatively minor surgical procedure with reduced risks, shortened hospitalization, reduced costs and better results. Complete removal of submucosal fibroids can be achieved by hysteroscopic myomectomy in more than 90% of cases. In the remaining 10% with a significant intramural component, the lesion can be removed during a second operation optionally under laparoscopic guidance in up to 98% of cases (Mencaglia and Guidetti, 1995).

Septate uterus accounts for approximately 80–90% of congenital uterine pathologies and is associated with the poorest reproductive performance with delivery rates of only 6–28% and miscarriage rates of over 60% (Simon *et al.*, 1991). Hysteroscopic metroplasty can be highly successful with data showing a fall in overall miscarriage rates to 6% and a live birth rate of 80–90% achievable in the next pregnancy (Acien, 1997; Homer *et al.*, 2000). In patients with recurrent pregnancy loss and infertility resulting from intrauterine adhesions, hysteroscopic adhesiolysis restored normal menstrual flow in

81% and achieved a favourable obstetric outcome in up to 71% of the pregnancies (Pabuccu *et al.*, 1997).

Ovarian surgery for polycystic ovarian syndrome

Polycystic ovarian syndrome (PCOS) affects 5–10% of women of reproductive age and is the most common cause of anovulatory infertility (Lobo and Carmina, 2000). Medical treatment has been advocated as the first-line treatment for anovulation in women with PCOS. Although up to 80% of women ovulate in response to clomiphene, only 40–50% will conceive. Historically, those who do not conceive with clomiphene move on to intensive treatment with gonadotrophin either in the form of ovulation induction or superovulation for IVF. The ovarian response to gonadotrophin in this group of patients can be highly unpredictable and difficult to control. They may either not develop any large follicles or over-respond with a significant risk for hyperstimulation syndrome and multiple pregnancy. The need for prolonged stimulation and intensive monitoring, and the high cancellation and miscarriage rates can be extremely stressful to these patients.

In comparison, ovarian drilling is not only a much simpler form of treatment for patients with clomiphene-resistant PCOS but it may also correct other associated endocrinopathy, improving symptoms of hirsutism and lowering miscarriage rates. In a review of 27 studies with a total of 729 patients who underwent ovarian drilling, 84.2% ovulated and 55.7% conceived (Donesky and Adashi, 1995). Peri-ovarian adhesions are common even with an endoscopic approach but most are mild and do not have a significant adverse effect on fertility. One long-term follow-up study reported that most women with PCOS became ovulatory soon after surgery and 74% were still ovulating when studied 18–20 years later (Gjonnaess, 1998). Several pregnancies can be achieved after a single treatment with lower risk of early pregnancy loss compared with gonadotrophin therapy, and with no additional risk of multiple pregnancy or ovarian hyperstimulation. In addition, ovarian drilling also renders patients with PCOS more responsive to either clomiphene or gonadotrophin and may facilitate these treatments if required, with reduced risks of complications. Reports of successful ovulation and conceptions after the use of insulin-sensitizing agents such as metformin are encouraging; however, results are still inconsistent (Norman *et al.*, 2001). Further research is needed to identify the subgroup of PCOS patients who will benefit most from ovarian drilling and those who would be better treated with metformin.

Conclusion

Reproductive surgery will remain a significant part of the range of modern infertility treatments available to patients. Advanced surgical training should be part of the experience of all trainees in reproductive medicine and units should be prepared to audit and publish their success rates. Although it is undeniably difficult to conduct adequately powered randomized controlled trials in reproductive surgery, large centres should strive to undertake such work, thereby demonstrating the clinical- and cost-effectiveness of reproductive surgery in comparison with medical treatment and IVF in different groups of infertile patients.

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