This discussion incorporates early spay-neuter, defined crinologic disorders, obesity and urinary incontinence in increased risk of neoplasia of various organ systems, and on societal management of unowned dog and cat populations.

Introduction

Elective gonadectomy, castration in males and either ovariectomy (OV) or ovariohysterectomy (OHE) in females, is the most common small animal surgery performed in some countries and is unethical or illegal in other countries (Salmeri et al. 1991b; Gunzel-Apel 1998; Greenfield et al. 2004). There is no question that we alter animals when we perform gonadectomy. This is a review of the specific changes associated with gonadectomy in dogs and cats and the evidence we have for cause-and-effect. Thorough reviews of the literature have been published (Root Kustritz 2007, 2010; Reichler 2009).

Some recognized reasons for the promotion of gonadectomy and some stated concerns are societal in nature, some refer to populations of animals, and some are pertinent to health of individual animals. Veterinarians are encouraged to help clients and those who write public policy to understand which aspect is being addressed when specific recommendations for gonadectomy are being made. There also is great variability in number of studies supporting some of the findings described, and veterinarians are encouraged to be aware of the amount of evidence to support any given claim. Finally, breed predispositions exist for some of the concerns described and must be included in decisions about suitability of gonadectomy in a given animal.

Advantages of gonadectomy include decrease in reproductive tract disease, including pyometra and mammary neoplasia in bitches and queens, and testicular neoplasia and prostate disease in male dogs. Other advantages include: decrease in pregnancy and parturition-related disorders including metritis, mastitis and dystocia; decrease in hormone-associated disorders such as vaginal prolapse in bitches and mammary hypertrophy in queens; and decrease in undesirable sexual behaviours (Romagnoli 2008). Disadvantages of gonadectomy include surgical and anaesthetic complications, increased risk of neoplasia of various organ systems, increased incidence of some musculoskeletal and endocrinologic disorders, obesity and urinary incontinence in bitches. All of these will be described in more detail. This discussion incorporates early spay-neuter, defined as gonadectomy at 8–16 weeks of age (Root Kustritz 1999), and gonadectomy at any age after 16 weeks.

Surgical and Anaesthetic Risk

General recommendations for anaesthesia are to minimize anaesthetic dose and time under anaesthesia. A series of excellent studies evaluating morbidity and mortality of dogs and cats identified general risk factors and some specific risk factors that are easily countered (Brodbelt et al. 2008; Brodbelt 2009, 2010). Cats were at greater risk of morbidity or mortality than dogs because of difficulties with intubation and prevalence of inapparent cardiomyopathy. If 6 months to 5 years of age is set as a baseline, all other ages were at greater risk, with a slight risk for younger animals and much greater risk for geriatric animals. Hypothermia greatly increased risk of morbidity or mortality during an anaesthetic episode. Profound hypothermia may develop as early as during induction and surgery prep. Body temperature should be monitored regularly and external sources of heat applied as needed. All animals, especially very young animals, should be weighed to ensure overdosing of drugs does not occur. Intra-operative monitoring shown to be of great benefit included recording of body temperature, pain assessment and use of pulse oximetry.

Concerns about anaesthesia are most commonly voiced when considering gonadectomy of very young animals. Concerns specific to animals <4–5 months of age include altered ability to metabolize and excrete anaesthetic agents and pain medication because of immature renal and hepatic function, decreased protein binding of medications in the bloodstream, predisposition to hypoglycaemia because of decreased gluconeogenesis and minimal glycogen stores in skeletal muscle, and decreased ability to maintain body temperature (Root Kustritz 1999). Puppies and kittens <10 weeks of age should be fasted for no more than 3–4 h before surgery and older animals fasted for no more than 8 h (Faggella and Aronsohn 1993, 1994). All surgical preparation fluids should be warmed and clipping of hair kept to a minimum. In studies of 98 puppies and 98 kittens presented for elective gonadectomy, incidence of anaesthetic complications was 0% for both groups (Aronsohn and Faggella 1993; Faggella and Aronsohn 1994). When looking at anaesthetic complications for gonadectomy at all ages, cardiac dysrhythmias, gastric dilation and drug overdosage were identified as the most common anaesthetic complications (Howe 1997).

Intra-operative and post-surgical complications are reported usually to be mild and self-limiting. Reported incidence of surgical complications is 6.1–27% in bitches.
and dogs and 2.6–33% in queens and toms (Pollari et al. 1996; Romagnoli 2008). The wide range reported most likely is due to differing definitions of ‘complications’ in the papers cited. Most post-surgical complications are reported to be mild, with most requiring only nursing care and not veterinary intervention. Incidence of complications is lower in young animals than in older animals, with one study identifying fewer complications in animals gonadectomized prepuberally and another study identifying 2 years of age as a cut-off (Faggella and Aronsohn 1994; Pollari et al. 1996; Howe 1997; Romagnoli 2008).

Reported short-term complications of OV or OHE include haemorrhage of the ovarian or uterine pedicle, passage of bloody vulvar discharge, onset of pseudocyesis, dehiscence or infection of the incision and, very rarely, peritonitis or evisceration (Romagnoli 2008). Long-term complications of OV or OHE include ovarian remnant syndrome, ligation of a ureter with subsequent hydroureter and ipsilateral renal dysfunction, granuloma of the uterine or ovarian pedicle, and disease of the uterine stump (Romagnoli 2008). Reported incidence of complications is not correlated with age of the animal, ability of the surgeon or presence of reproductive disease (Miller 1995; Romagnoli 2008). Complications associated with haemorrhage may be more common when bitches and queens are spayed while in oestrus under the influence of oestrogen, and increased incidence of complications is associated with increased surgery time and increased body weight of the patient (Howe 1997; Howe et al. 2000, 2001; Burrow et al. 2005).

**Population Control**

Pet overpopulation may occur due to attitudes of the animal owning public permitting irresponsible breeding or ready surrender of animals or due to free-roaming or feral dogs and cats. Although it has been reported that 82% of cats and 64% of dogs in the US have undergone gonadectomy (Trevejo et al. 2011), a recent study reported that only 52% of people who had purchased a dog or cat within the last year had had that animal gonadectomized (PetSmart Charities 2002). In the United States, millions of dogs and cats with no owner or guardian are euthanized annually (Nassar et al. 1992; National Council on Pet Population Study and Policy 1994). Intact animals are more likely to be surrendered to humane organizations (Patronek et al. 1996a,b; New et al. 2000; Mondelli et al. 2004). Spay-neuter contracts are sent out with animals adopted while still intact but compliance with these contracts, even with low-cost surgical options readily available, is <60%, permitting intact animals to re-populate those shelters from which they were adopted (Alexander and Shane 1994; Patronek et al. 1996a; New et al. 2000; Mondelli et al. 2004). Lack of compliance with recommendations for surgery is multifactorial. Cultural and personal factors include religious affiliation, ethnic background, intended working life of the animal, urban or rural location of the household and literacy status (Manning and Rowan 1992; Mahlow 1999; Eze and Eze 2002). Cats are more likely to be spayed or castrated than are dogs, and females are more likely to be presented for gonadectomy than are males in the United States (Manning and Rowan 1992; Alexander and Shane 1994; Mahlow 1999; New et al. 2000).

Many pet owners have limited knowledge about normal reproduction in dogs and cats. In the PetSmart survey (PetSmart Charities A&U Barriers 2002), 13% of dogs and 19% of cats purchased within the previous year had borne litters, with over 50% of those unplanned. In one survey of dog- and cat-owning households, 56% of 154 canine litters and 68% of 317 feline litters were unplanned, with the majority of those owners reporting that they did not know the female had been in heat (New et al. 2004). Because of this, many humane organizations in the United States are working toward gonadectomy of all unowned animals prior to adoption. Other measures that will be required to stem pet overpopulation include education of pet owners and increased availability of affordable, safe, readily provided non-surgical contraceptive or sterilization options.

**Behaviour**

Many people request gonadectomy because they are unwilling or unable to tolerate normal breeding behaviours in their animals. Male cat normal breeding behaviour, which is aggressive and includes urine spraying, is incompatible with life as a house pet (Root Kustritz 1996). Male dogs may mount other animals, people or inanimate objects and may urine mark. Bitches and queens may show altered behaviours with oestrus, including increases in some forms of aggression, and bitches may show nesting and mothering behaviours and lactation with false pregnancy.

Gonadectomy, especially castration, is a commonly recommended treatment for behavioural problems in dogs including aggression. Not all behaviours are altered by gonadectomy. Those most commonly affected are sexually dimorphic behaviours, defined as behaviours that vary between genders, and are likely to be at least partially mediated by sex hormones.

In male cats, the undesirable behaviours decreased to the greatest extent by castration are sexual behaviours, roaming and urine spraying (Hart 1973, 1979; Hart and Barrett 1973). In male dogs, the undesirable behaviours decreased to the greatest extent by castration are mounting, urine marking and roaming (Hart 1979; Neilson et al. 1997). Likelihood of response to castration is not associated with age at castration or duration of the behaviour prior to castration in male dogs or cats (Hart 1973; Hart and Barrett 1973; Hopkins et al. 1976; Neilson et al. 1997). The hormonal basis of these behaviours is not always clear. Oestrogen stimulates arousal in females but also is associated with release of oxytocin and opioids, which may have an anti-anxiety effect (Mong and Pfaff 2003; Root Kustritz 2005). Concerns vary regarding trainability of dogs after gonadectomy, with some suggesting that dogs will be more attentive after gonadectomy as they are less likely to show behaviours related to copulation, such as roaming. It has been demonstrated in rats that attention is not affected by rising or falling concentrations of testosterone, but that acuity of spatial working memory.
is testosterone dependent (Johnson and Burk 2006; Kritzer et al. 2007).

Aggression may or may not be affected by gonadectomy (Hart and Eckstein 1997; Neilson et al. 1997; Duffy and Serpell 2006). A breed-specific increase in owner-directed aggression and in degree of reactivity was reported in both males and females after gonadectomy and in bitches after OHE; in one study, this increase in aggression was most common in bitches spayed at <1 year of age who were showing aggression prior to surgery (Wright and Nesselrote 1987; O’Farrell and Peachey 1990; Reisner et al. 2005; Duffy and Serpell 2006). The form of aggression best controlled by castration in both male dogs and cats is intermale aggression (Hart 1979; Borchelt and Voith 1987). Other forms of aggression in cats include predation, play, fear or defence, territorial and redirected aggression; these are much less likely to be decreased by castration (Borchelt 1983). Other forms of aggression in dogs include predation, fear or defence, dominance, protectiveness, punishment and pain-related aggression (Borchelt 1983).

In one study of older male dogs with mild cognitive impairment, those that were castrated were more likely to progress to severe cognitive impairment than were those left intact (Hart 2001). Androgen deprivation is associated with increased deposition of amyloid in brains of humans and rodents and with decreased number of synaptic connections in brains of rodents and primates (Janowsky 2006). However, studies directly evaluating the brain of aged dogs showed that among beagles aged approximately 10 years, intact dogs had more DNA damage in the brain than did castrated dogs (Waters et al. 2000).

Neoplasia

Mammary

Mammary neoplasia is very common in dogs and cats, with reported incidence of 3.4 and 2.5%, respectively, in the United States (Fidler and Brodey 1967; Dorn et al. 1968a; Moe 2001; Richards et al. 2001; Verstegen and Onclin 2003a). In dogs, approximately half of all mammary tumours are malignant; in cats, >90% of mammary tumours are malignant (Cotchin 1951; Dorn et al. 1968a; Moulton et al. 1970; Hampe and Misendorp 1974; Hayes et al. 1981; Brodey et al. 1983). Despite the fact that many mammary tumours in bitches are benign, it was reported in one U.S. study that 59.7% of bitches with mammary neoplasia were euthanized at the time of diagnosis (Misendorp and Hart 1979). It has been well demonstrated that incidence of mammary cancer when aged is greatly decreased in female dogs and cats by ovariecotmy, especially if performed before the first heat cycle. Intact dogs and cats are seven times more likely than are spayed dogs and cats to develop mammary neoplasia when aged (Dorn et al. 1968b). Risk for bitches has been defined as 0.5% before the first oestrus, 8% after one oestrus, and 26% after two or more oestrous cycles (Schneider et al. 1969). Some sparing effect may be present with OHE up to 9 years of age, although that effect may be greater for benign than for malignant tumours (Verstegen and Onclin 2003b; Reichler 2009). Siamese cats and Japanese domestic breed cats are reported to be at greater risk of developing mammary neoplasia, as are the boxer, Brittany spaniel, cocker spaniel, dachshund, English setter, English springer spaniel, German shepherd dog, Maltese, miniature poodle, pointer, toy poodle and Yorkshire terrier (Dorn et al. 1968b; Hayes et al. 1981; Sorenmo 2003; Verstegen and Onclin 2003a). Cause-and-effect relationship between gonadectomy and decreased incidence of mammary neoplasia is not clear. In human medicine, oestrogen is implicated in mammary neoplasia. Oestrogen receptors have been identified in neoplastic mammary tissue of dogs and cats, with fewer oestrogen receptors present as tissue becomes more anaplastic (Hamilton et al. 1977; Donnay et al. 1995; Verstegen and Onclin 2003b). One study suggested that mammary neoplasia may be more common in bitches that exhibited more than three episodes of overt false pregnancy in their lives, again supporting a role for hormonal stimulation of mammary tissue as a factor associated with neoplastic transformation of that tissue (Verstegen and Onclin 2003b).

Genitourinary

Prostate

Prostatic neoplasia is uncommon in dogs, with reported incidence of 0.2-0.6% (Bell et al. 1991; Teske et al. 2002). Prostatic neoplasia in dogs virtually always is malignant; both adenocarcinoma and transitional cell carcinoma are described. Several studies have documented increased incidence of prostatic cancer in castrated dogs compared with intact dogs, with reported increased risk ranging from 2.4 to 4.3 times (Obradovich et al. 1987; Bell et al. 1991; Teske et al. 2002; Sorenmo et al. 2003). Prostatic neoplasia is extremely uncommon in cats, but in one case series of eight cats with prostatic neoplasia, seven were castrated (Hawe 1983; Carpenter et al. 1987; Hubbard et al. 1990; Caney et al. 1998; LeRoy and Lech 2004). No breed predisposition is reported. Cause-and-effect relationship is not clear.

Urinary bladder

Incidence of transitional cell carcinoma is at most 1% of malignant tumours (Poirier et al. 2004). Incidence of transitional cell carcinoma of the urinary bladder and urethra of female dogs and urethra of male dogs is reported to be increased after gonadectomy, with increased risk of 2-4 times (Norris et al. 1992; Knapp et al. 2000). This has not been reported in cats. Dog breeds at increased risk are the Airedale terrier, beagle, collie, Scottish terrier, Shetland sheepdog, West Highland white terrier and wire-haired fox terrier (Ru et al. 1998). Cause-and-effect relationship is not clear.

Testicular

Testicular neoplasia is very common in aged male dogs, with reported incidence of 0.9% (Hahn et al. 1992). The three tumour types most commonly reported are Sertoli
cell tumours, seminomas and Leydig (interstitial) cell tumours. Testicular neoplasia is uncommon in male cats. Castration is curative.

Haematologic

Overall incidence of haemangiosarcoma in dogs is 0.2% and in cats is 0.03% (Reichler 2009). Incidence of both splenic- and heart-based haemangiosarcoma are reported to be increased in dogs after gonadectomy; in bitches, splenic haemangiosarcoma risk after OHE increases by a factor of 2.2 and cardiac haemangiosarcoma risk increases by a factor of 5, and in male dogs, overall risk of haemangiosarcoma increases by a factor 2.4 after castration (Prymak et al. 1988; Ware and Hopper 1999). This has not been reported in cats. Breeds at increased risk are the boxer, English setter, German shepherd dog, golden retriever, Great Dane, Labrador retriever, pointer, poodle and Siberian husky (Smith 2003). Cause-and-effect relationship is not clear.

Orthopaedic

Osteosarcoma is an uncommon tumour with overall incidence of 0.2% (Root Kustritz 2007). Incidence is increased with gonadectomy, by a factor of 1.3–2 (Priester and McKay 1980; Ru et al. 1998). Breeds at increased risk are the Doberman pinscher, Great Dane, Irish setter, Irish wolfhound, rottweiler and St. Bernard (Ru et al. 1998; Chunn and DeLorimer 2003). Other risk factors for osteosarcoma include increasing age and increasing body weight (Ru et al. 1998; Cooley et al. 2002). Cause-and-effect relationship is not clear.

Dermatologic

Increase incidence of cutaneous mast cells tumours has been reported in spayed female dogs compared with intact female dogs (White et al. 2011). Cause-and-effect relationship is not clear.

Musculoskeletal Disease

Growth plate closure

Growth plate closure in long bones is mediated by gonadal hormones. It has been well demonstrated that growth plate closure is delayed in dogs and cats spayed or castrated prepuberally (May et al. 1991; Salmeri et al. 1991a; Houlton and McGlennon 1992; Root et al. 1996c; Stubbs et al. 1996). Significance of this delay is not clear. Increased incidence of growth plate fractures in cats spayed prepuberally was demonstrated in one study; obesity may have contributed to the disorder in those cats (McNicholas et al. 2002).

Hip dysplasia

One large study evaluating incidence of various disorders in dogs spayed at various ages in a humane organization reported increased incidence of hip dysplasia in dogs spayed when young (Spain et al. 2004b). It was not clear in that study that all dogs were diagnosed with hip dysplasia by a veterinarian. In one study, it was demonstrated that neutered boxers were 1.5 times as likely as intact boxers to develop hip dysplasia; that study did not figure in possible contribution of excess body weight (VanHagen et al. 2005). Hip dysplasia has a strong genetic component. Breeds at risk include the Chesapeake Bay retriever, English setter, German shepherd dog, golden retriever, Labrador retriever, Samoyed and St. Bernard (Priester and Mulvihill 1972). A hypothesis for cause-and-effect is asymmetry in closure of growth plates causing deformity and laxity of the hip joint with subsequent arthritis.

Anterior Cruciate Ligament (ACL) Rupture

Rupture of the anterior or cranial cruciate ligament is more common after gonadectomy than in intact dogs (Whitehair et al. 1993; Duval et al. 1999; Slaumberk et al. 2004). Breeds at risk for rupture of the ACL include the Akita, American Staffordshire terrier, Chesapeake Bay retriever, German shepherd dog, golden retriever, Labrador retriever, mastiff, Neopolitan mastiff, Newfoundland, poodle, rottweiler and St. Bernard (Duval et al. 1999; Harasen 2003). Other risk factors include obesity and abnormal angulation of the stifle (Ragetly et al. 2011). One could argue that increased risk of ACL injury after gonadectomy is because of decreased athleticism and obesity in gonadectomized animals but the trend stands even in studies that statistically compensated for these effects in dogs. Joint laxity may differ under varying hormonal stimuli, suggesting one possible cause-and-effect mechanism. Another hypothesis is increasing stifle angulation with asymmetry of growth plate closure in the femur and tibia.

Genitourinary Disease

Urinary incontinence

Urethral sphincter mechanism incompetence, formerly called oestrogen-responsive urinary incontinence, is leakage of urine from female dogs, usually as they lie relaxed. A similar clinical manifestation is seen in castrated male dogs but is less common than in females. Urethral sphincter mechanism incompetence is much more common in spayed than in intact female dogs with reported incidence of 4.9–20.0% (Arnold 1997; Stocklin-Gautschi et al. 2001; Angioletti et al. 2004; Spain et al. 2004b). One study identified 3 months of age as the age before which OHE most greatly contributed to eventual clinical manifestation of urinary incontinence (Spain et al. 2004b). Other contributing factors include higher lean body weight (>20 kg); breed, with breeds at increased risk including the boxer, Doberman pinscher, giant schnauzer, Irish setter, old English sheepdog, rottweiler, springer spaniel and Weimaraner; and urethral length or resting position of the urinary bladder creating a pressure differential across the length of the urethra (Gregory et al. 1992, 1999; Holt and Thursfield 1993; Arnold 1997; Atalan et al. 1998; Holt 2004). Urethral sphincter tone is mediated by the sympathetic nervous system and function may be potentiated by
of 100 proliferative adrenal lesions in ferrets, almost 30% were from intact jills (Olson 1997). There have been no reports of increased adrenal disease in cat populations secondary to gonadectomy, and any impact on dog populations has not been reported.

Diabetes Mellitus in Cats
Cats have a 2- to 9-fold increased risk of developing diabetes mellitus after neutering, with Burmese cats reported to be especially susceptible (Panciera et al. 1990; Rand et al. 1997; McCann et al. 2007; Prahl et al. 2007). One hypothesis is that this is because of decreased insulin sensitivity (Hoenig and Ferguson 2002; Kanchuk et al. 2002). Obesity also is associated with decreased insulin sensitivity and may be a confounding factor.

Hypothyroidism
Risk of hypothyroidism in dogs was reported to be increased with neutering in some studies (Milne and Hayes 1981; Panciera 1994) and not to be associated with neutering in other studies (Dixon and Mooney 1999a; Dixon et al. 1999b) Gonadectomy is associated with persistent increase in blood concentrations of gonadotropins (Loftstedt and VanLeeuwen 2002), many of which share molecular configuration with TSH in protein subunit chains and in glycosylation. It can be hypothesized that persistent high concentrations of gonadotropins lead to altered formation or glycosylation of TSH, and subsequent change in thyroid function. However, one study showed no change in resting serum TSH or thyroxin concentrations, or alterations in stimulus testing with thyroid releasing hormone when comparing castrated to intact male beagles (Gunzel-Apel et al. 2009).

Metabolic Disorders
Obesity
In retrospective studies, up to 2.8% of the canine population has been demonstrated to be obese, with up to 50% of gonadectomized dogs and cats designated as obese (Mason 1970; David and Rajendran 1980). Increase in indiscriminate appetite was reported in spayed bitches in one study but in another study of spayed and castrated dogs, no change in food intake or depth of back fat was reported by 15 months of age (O’Farrell and Peachey 1990; Salmeri et al. 1991a). Risk factors other than gonadectomy include housing of the animal; increasing age; ownership by an overweight person or a person over 40 years of age; and breed, with the beagle, cairn terrier, cavalier King Charles spaniel, cocker spaniel, dachshund and Labrador retriever among those breeds at greatest risk (Mason 1970; Edney and Smith 1986; Crane 1991; Sloth 1992; Colliard et al. 2006). In cats, castration and OHE are associated with increased risk of obesity, with one paper describing a 3.4-fold risk factor (Fettman et al. 1997; Kanchuk et al. 2002; Nguyen et al. 2004). Gonadectomized cats have increased body weight, increased body mass index, increased depth of the

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falciform fat pad, decreased activity and decreased metabolic rate (Root 1995; Root et al. 1996b; Stubbs et al. 1996; Pettman et al. 1997). Castrated cats also have an increase in serum concentrations of insulin growth factor I, prolactin and leptin, all of which are associated with metabolic rate and fat metabolism (Martin et al. 2006).

**Lifespan**

Most studies evaluating lifespan suggest that neutered animals live longer than intact animals (Kraft 1998; Greer et al. 2007). Increased lifespan in castrated male dogs may be due to a decrease in risky sexually dimorphic behaviours such as roaming, or to increased owner investment in the animal’s care (Bronson 1982; Michell 1999; Moore et al. 2001). Castrated cats have reduced mortality compared with intact cats (Kalz 2001).

One study evaluating lifespan in a group of exceptionally long-lived rottweilers showed increased lifespan if left intact (Waters et al. 2009). This could be due to specific genetic factors in this population of dogs, to hormonal changes not yet identified, or to a lower incidence of obesity in this group; obesity itself has a negative effect on life expectancy (Greer et al. 2007).

It has been suggested that studies that separate dogs into intact or neutered categories at the time of disease onset or death create an artificial black-or-white evaluation of data regarding benefits and detriments of gonadectomy (Waters et al. 2011). Future studies that will guide this debate include those defining lifelong effects of exposure to gonadal hormones or better relating time of neuter to onset of disease.

**Conflicts of interest**

None of the authors have any conflicts of interest to declare.

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